

最 終 報 告 書

ドデシルオキシラン：ラットを用いた経口投与による
反復投与毒性・生殖発生毒性併合試験

試験番号 R-1261

試験期間

2020 年 10 月 2 日-2021 年 3 月 23 日

試験施設

株式会社ボゾリサーチセンター 御殿場研究所
〒412-0039 静岡県御殿場市かまど 1284

試験委託者

厚生労働省 医薬・生活衛生局 医薬品審査管理課 化学物質安全対策室
〒100-8916 東京都千代田区霞が関 1-2-2

試験受託者

株式会社ボゾリサーチセンター
〒151-0065 東京都渋谷区大山町 36-7

R-1261

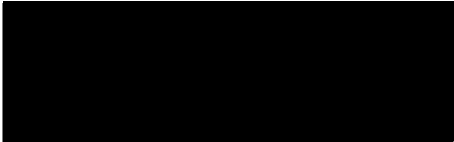
1. GLP 陳述書

試験番号 : R-1261

試験表題 : ドデシルオキシラン：ラットを用いた経口投与による
反復投与毒性・生殖発生毒性併合試験

本試験は以下の GLP 基準を遵守して実施した。

- 「新規化学物質等に係る試験を実施する試験施設に関する基準」
(平成 23 年 3 月 31 日：薬食発 0331 第 8 号、平成 23・03・29 製局第 6 号、
環保企発第 110331010 号)



2021 年 3 月 23 日

試験責任者

株式会社ボゾリサーチセンター 御殿場研究所

2. 目次

1.	GLP 陳述書	2
2.	目次	3
3.	試験実施概要	8
3.1	試験番号	8
3.2	試験表題	8
3.3	試験目的	8
3.4	試験委託者	8
3.5	試験受託者	8
3.6	試験施設及び試験場所	8
3.6.1	試験施設	8
3.6.2	試験場所（ホルモン測定）	8
3.7	被験物質	8
3.8	試験日程	9
3.9	試験責任者	9
3.10	試験担当者	9
3.11	試験成績の信頼性に影響を及ぼしたと思われる環境要因	9
3.12	予見することができなかった試験の信頼性に影響を及ぼす疑いの ある事態及び試験計画書に従わなかったこと	10
3.13	資料保存	10
3.14	試験責任者の署名	11
4.	要約	12
4.1	反復投与毒性	12
4.2	生殖発生毒性	12
5.	緒言	13
5.1	GLP	13
5.2	毒性試験ガイドライン	13
5.3	動物の福祉	13
6.	試験材料及び方法	14
6.1	被験物質及び媒体	14
6.1.1	被験物質	14
6.1.2	媒体	15
6.2	媒体及び被験液の調製	15
6.2.1	媒体の調製	15
6.2.1.1	調製方法	15
6.2.1.2	保存条件	15
6.2.2	被験液の調製	15

6.2.2.1	調製方法	15
6.2.2.2	調製頻度	16
6.2.2.3	保存条件	16
6.2.2.4	安定性	16
6.2.3	被験液の分析	16
6.3	試験系及び系統の選択理由	18
6.4	試験動物及び群分け	18
6.5	飼育条件	19
6.6	飼料、床敷及び飲料水中の混入物質	20
6.7	動物の識別	20
6.8	投与経路、投与期間、投与回数及び回復期間とそれらの選択理由	20
6.9	投与方法	21
6.10	投与量及び群構成	21
6.11	投与量の設定根拠	21
6.12	観察及び検査の方法	22
6.12.1	一般状態の観察	22
6.12.2	詳細な一般状態の観察、機能検査、握力及び自発運動量の測定	22
6.12.2.1	詳細な一般状態の観察	23
6.12.2.2	機能検査	23
6.12.2.3	握力測定	23
6.12.2.4	自発運動量測定	23
6.12.3	体重測定	24
6.12.4	摂餌量測定	24
6.12.5	腔垢検査	24
6.12.6	交配	24
6.13	分娩及び哺育観察	25
6.13.1	母動物の観察	25
6.13.2	出生児の観察	25
6.14	尿検査	26
6.15	血液学検査	27
6.16	血液化学検査	28
6.17	ホルモン測定	29
6.17.1	測定対象	29
6.17.2	試料の送付	29
6.17.3	ホルモン（T4）測定	30
6.18	病理学検査	30
6.18.1	剖検	30
6.18.2	器官重量	30

6.18.3	病理組織学検査	31
7.	データ採取に使用したコンピュータシステム	33
7.1	統計解析	33
7.1.1	パラメータの算出	33
7.1.2	検定	34
8.	試験結果	35
8.1	一般状態	35
8.2	詳細な一般状態の観察、機能検査、握力測定及び自発運動量の測定	35
8.2.1	詳細な一般状態の観察	35
8.2.1.1	ケージ内観察	35
8.2.1.2	手に持つての観察	35
8.2.1.3	オープンフィールド内観察	35
8.2.2	機能検査	36
8.2.3	握力	36
8.2.4	自発運動量	37
8.3	体重	37
8.4	摂餌量	37
8.5	尿検査（摂水量測定を含む）	38
8.6	血液学検査	38
8.7	血液化学検査	38
8.8	血清中ホルモン濃度測定	39
8.9	器官重量	39
8.10	剖検所見	40
8.11	病理組織学検査	40
8.12	性周期	41
8.13	交配成績	41
8.14	分娩成績並びに分娩及び哺育状態	41
8.15	出生児の Anogenital Distance (AGD)	41
8.16	出生児の生存性	41
8.17	出生児の性比	42
8.18	出生児の体重	42
8.19	死亡児の肉眼観察所見	42
8.20	雄新生児の乳頭/乳輪数	42
8.21	出生児の生後 13 日剖検所見	42
8.22	出生児の器官重量	42
9.	考察及び結論	42
9.1	反復投与毒性	43
9.2	生殖発生毒性	43

10. 参考文献	44
----------------	----

添付資料

Attachment 1	試験成績書（被験物質の特性）	45
Attachment 2	試験成績書（被験物質の安定性）	46
Attachment 3	試験成績書（被験液中ドデシルオキシランの安定性）	48
Attachment 4	被験液中のドデシルオキシラン濃度分析結果	50
Attachment 5	ホルモン濃度測定報告書	52

図

Fig. 1~5	自発運動量.....	68
Fig. 6~8	体重.....	73
Fig. 9~11	摂餌量	76

表

Table 1-1~1-12	一般状態.....	79
Table 2-1~2-24	ホームケージ内観察	91
Table 2-25~2-86	手に持つての観察.....	115
Table 2-87~2-158	オープンフィールド内観察	177
Table 2-159~2-163	機能検査.....	249
Table 2-164~2-168	握力.....	254
Table 2-169~2-173	自発運動量.....	259
Table 3-1~3-7	体重.....	264
Table 4-1~4-9	摂餌量	271
Table 5-1~5-21	尿検査	280
Table 6-1~6-5	血液学検査.....	301
Table 7-1~7-5	血液化学検査	306
Table 8-1~8-8	器官重量	311
Table 9-1~9-6	剖検所見	319
Table 10-1~10-18	病理組織学検査	325
Table 11	性周期	343
Table 12-1、12-2	交配成績	344
Table 13	分娩成績.....	346
Table 14	出生児の外表面観察.....	347
Table 15	出生児の Anogenital Distance (AGD)	348
Table 16	出生児の生存性.....	349
Table 17	出生児の性比	350
Table 18-1、18-2	出生児の体重	351

Table 19	死亡児の肉眼観察所見	353
Table 20	雄新生児の乳頭/乳輪数	354
Table 21	出生児の生後 13 日剖検所見	355
Table 22-1、22-2	出生児の器官重量	356

個体別表

Appendix 1-1~1-36	一般状態	358
Appendix 2-1~2-76	ホームケージ内観察	394
Appendix 2-77~2-152	手に持つての観察	470
Appendix 2-153~2-304	オープンフィールド内観察	546
Appendix 2-305~2-318	機能検査	698
Appendix 2-319~2-332	握力	712
Appendix 2-333~2-346	自発運動量	726
Appendix 3-1~3-22	体重	740
Appendix 4-1~4-28	摂餌量	762
Appendix 5-1~5-30	尿検査	790
Appendix 6-1~6-14	血液学検査	820
Appendix 7-1~7-14	血液化学検査	834
Appendix 8-1~8-30	器官重量	848
Appendix 9-1~9-116	剖検所見及び病理組織学検査	878
Appendix 10-1~10-8	性周期	994
Appendix 11-1~11-8	交配成績	1002
Appendix 12-1~12-4	分娩成績	1010
Appendix 13-1~13-4	出生児の外表観察	1014
Appendix 14-1~14-4	出生児の Anogenital Distance (AGD)	1018
Appendix 15-1~15-4	出生児の生存性	1022
Appendix 16-1~16-4	出生児の性比	1026
Appendix 17-1~17-8	出生児の体重	1030
Appendix 18-1~18-4	死亡児の肉眼観察所見	1038
Appendix 19-1~19-4	雄新生児の乳頭/乳輪数	1042
Appendix 20-1~20-4	出生児の生後 13 日剖検所見	1046
Appendix 21-1~21-8	出生児の器官重量	1050

信頼性保証書	1058
--------------	------

R-1261

3. 試験実施概要

3.1 試験番号

R-1261

3.2 試験表題

ドデシルオキシラン：ラットを用いた経口投与による反復投与毒性・生殖発生毒性併合試験

3.3 試験目的

本試験は、動物に被験物質を一定期間反復投与したときに現れる被験物質の一般毒性及び生殖発生毒性を明らかにすることを目的とした。

3.4 試験委託者

厚生労働省

医薬・生活衛生局 医薬品審査管理課 化学物質安全対策室

〒100-8916 東京都千代田区霞が関 1-2-2

3.5 試験受託者

株式会社ボゾリサーチセンター

〒151-0065 東京都渋谷区大山町 36-7

3.6 試験施設及び試験場所

3.6.1 試験施設

株式会社ボゾリサーチセンター 御殿場研究所

〒412-0039 静岡県御殿場市かまど 1284

3.6.2 試験場所（ホルモン測定）

株式会社ボゾリサーチセンター つくば研究所

〒300-2611 茨城県つくば市大久保 8 番

3.7 被験物質

製造者	:	████████████████████
名称	:	ドデシルオキシラン
英語名称	:	Dodecyloxirane
CAS 番号	:	3234-28-4
官報公示整理番号	:	(2)-220（化審法）
入手日	:	2020 年 9 月 29 日

3.8 試験日程

試験開始日 : 2020 年 10 月 2 日
被験物質配布日 : 2020 年 10 月 20 日
動物入荷日 : 2020 年 10 月 5 日
実験開始日（投与開始日）
: 2020 年 10 月 27 日
交配開始日 : 2020 年 11 月 10 日
分娩開始日 : 2020 年 12 月 2 日
投与終了剖検日 : 2020 年 12 月 8 日（雄及び非交配群）
回復開始日 : 2020 年 12 月 8 日
母動物剖検開始日 : 2020 年 12 月 16 日
回復終了剖検日 : 2020 年 12 月 22 日
動物試験終了日 : 2020 年 12 月 30 日
実験完了日（病理組織学検査終了日）
: 2021 年 2 月 24 日
試験終了日 : 2021 年 3 月 23 日

3.9 試験責任者

■■■■■■■■■■

株式会社ボゾリサーチセンター 御殿場研究所 研究部

3.10 試験担当者

被験物質管理責任者 : ■■■■■■■■■■
試験主担当者 : ■■■■■■■■■■
臨床検査責任者 : ■■■■■■■■■■
病理検査責任者 : ■■■■■■■■■■
化学分析（被験液分析）
: ■■■■■■■■■■
血清中ホルモン濃度測定責任者（試験主任者）*
: ■■■■■■■■■■

*：株式会社ボゾリサーチセンター つくば研究所

3.11 試験成績の信頼性に影響を及ぼしたと思われる環境要因

本試験に関し、試験成績の信頼性に影響を及ぼしたと思われる環境要因はなかった。

3.12 予見することができなかった試験の信頼性に影響を及ぼす疑いのある事態及び試験計画書に従わなかったこと

本試験に関し、試験の信頼性に影響を及ぼす疑いのある事態はなかった。

なお、試験計画書に従わなかったことを以下に記載した。

- 1) 病理組織学検査の標本作製において、1 例（動物番号 1012）で上皮小体（両側）がスライド標本から欠落した。しかし、媒体対照群の 1 例のみであり、他の動物では上皮小体への被験物質の影響は評価可能であったことから、本件による試験へ及ぼす影響はないと判断した。

3.13 資料保存

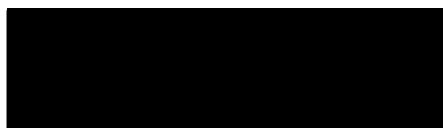
試験計画書（試験計画書変更書を含む）原本、記録文書、生データ、報告書類（最終報告書の原本を含む）及び標本（被験物質保存試料を含む）は株式会社ボゾリサーチセンター 御殿場研究所の資料保存施設に保存する。ただし、臨床検査後の残余生体試料（尿、血漿、血清）及び腔垢標本については試験終了後に廃棄する。また、固定した死産児及び死亡児については、出生率、生後 4 及び 13 日生存率に被験物質投与の影響が認められなかったため、廃棄した。資料の保存期間は試験終了後 5 年間とする。

ホルモン測定に関するホルモン測定計画書原本、生データ等の試験関係資料及びホルモン測定報告書原本は、試験場所（株式会社ボゾリサーチセンター つくば研究所）又は試験場所指定の GLP 保存施設に保存する。なお、その期間はホルモン測定報告書作成後 5 年間とする。

試験施設と試験場所における保存期間終了後の資料の取扱いについては厚生労働省 医薬・生活衛生局 医薬品審査管理課 化学物質安全対策室と株式会社ボゾリサーチセンター間で協議し、その処置を決定する。

R-1261

3.14 試験責任者の署名



2021 年 3 月 23 日

株式会社ボゾリサーチセンター 御殿場研究所

4. 要約

ドデシルオキシランをコーン油に溶解させ、100、300 及び 1000 mg/kg の投与量で Sprague-Dawley 系 SPF ラット（交配群として雌雄各 12 匹/群、雌には交配を行わない非交配群として媒体対照群及び 1000 mg/kg 群に各 10 匹を追加）の雄には交配前 14 日間に加え交配期間を通して剖検前日まで（42 日間）、交配群雌には交配前 14 日間に加え交配期間及び妊娠期間を通して授乳 13 日まで（50~64 日間）、非交配群雌には 42 日間、強制経口投与し、反復投与毒性及び生殖発生毒性を検討した。更に、媒体対照群及び 1000 mg/kg 群の一部の動物（雄 5 匹と非交配群雌 5 匹）については、42 日間投与した後 14 日間の回復期間を設け、毒性変化の可逆性を検討した。なお、媒体対照群には媒体であるコーン油を投与した。

4.1 反復投与毒性

いずれの投与群においても死亡動物はみられず、一般状態、詳細な一般状態観察、機能検査、握力、自発運動量、体重、摂餌量、血液学検査、血清中ホルモン(T4)及び剖検並びに病理組織学検査には、被験物質投与の影響は認められなかった。

1000 mg/kg 群では、雄で投与終了週に尿浸透圧の高値が、投与終了時に BUN、ALT 及び肝臓重量の高値が認められ、非交配群雌で ALT 及び肝臓重量の高値、交配群雌で肝臓重量の高値が認められた。

300 mg/kg 群では、交配群雌で投与終了時に肝臓重量の高値がみられた。

回復終了週に 1000 mg/kg 群の雄で尿浸透圧の高値が認められた。14 日間の回復期間により、その他の変化は消失し、回復性が認められた。

4.2 生殖発生毒性

親動物のいずれの検査においても、被験物質投与による影響は認められなかった。

児動物では、1000 mg/kg 群で雌雄の生後 13 日の体重並びに生後 4~13 日の間の体重増加量に低値が認められ、雄出生児の甲状腺重量の高値が認められた。

これらの結果から、本試験条件下では、本被験物質の反復投与毒性に対する無影響量及び無毒性量は、雄でいずれも 300 mg/kg、雌でいずれも 100 mg/kg と判断した。一方、生殖発生毒性に対する無影響量及び無毒性量は、いずれも親動物では 1000 mg/kg、児動物では 300 mg/kg と判断した。

5. 緒言

厚生労働省 医薬・生活衛生局 医薬品審査管理課 化学物質安全対策室の委託により、ドデシルオキシランのラットを用いた経口投与による反復投与毒性・生殖発生毒性併合試験を実施したので、その成績を報告する。

なお、本試験は以下の基準を遵守し、ガイドライン等に準拠して、株式会社ボゾリサーチセンターで実施した。

5.1 GLP

- 「新規化学物質等に係る試験を実施する試験施設に関する基準」
(平成 23 年 3 月 31 日：薬食発 0331 第 8 号、平成 23・03・29 製局第 6 号、環保企発第 110331010 号)

5.2 毒性試験ガイドライン

- 「新規化学物質等に係る試験の方法について：哺乳類を用いる反復投与毒性・生殖発生毒性併合試験」
(平成 23 年 3 月 31 日付け薬食発 0331 第 7 号厚生労働省医薬食品局長、平成 23・03・29 製局第 5 号経済産業省製造産業局長、環保企発第 110331009 号環境省総合環境政策局長連名通知) (最終改正：令和元年 7 月 1 日)

5.3 動物の福祉

- 「動物の愛護及び管理に関する法律」
(昭和 48 年 10 月 1 日法律第 105 号)
- 「実験動物の飼養及び保管並びに苦痛の軽減に関する基準」
(平成 18 年 4 月 28 日環境省告示第 88 号)
- 「動物実験の適正な実施に向けたガイドライン」
(日本学術会議 平成 18 年 6 月 1 日)
- 「厚生労働省の所管する実施機関における動物実験等の実施に関する基本指針について」
(平成 18 年 6 月 1 日科発第 0601001 号)

本試験は動物実験委員会 (IACUC) の承認を受けて実施した (承認番号: G200254)。

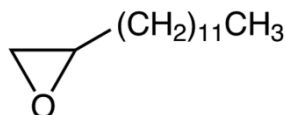
6. 試験材料及び方法

6.1 被験物質及び媒体

6.1.1 被験物質

被験物質は東京化成工業株式会社から、GMP 下で実施された分析結果に基づく以下の情報とともに購入した。また、投与開始前に赤外吸収スペクトルの測定により特性が確認されている（Attachment 1）。

製造者	: [REDACTED]
名称	: ドデシルオキシラン（略称：DDOR）
英語名称	: Dodecyloxirane
別名	: 1,2-エポキシテトラデカン
CAS 番号	: 3234-28-4
官報公示整理番号	: (2)-220（化審法）
構造式又は示性式	:



分子式	: C ₁₄ H ₂₈ O
分子量	: 212.38
外観	: 無色透明液体
沸点/沸騰範囲	: 96°C/0.05kPa
比重	: 0.84
ロット番号	: [REDACTED]
入手量	: 1,500 mL
純度（GC）	: 97.8%
保存条件	: 密栓、冷暗所、防湿、窒素置換
保存方法	: 気密容器に入れ窒素充填し、冷蔵庫内（許容範囲：1°C~10°C）に保存
取扱い上の注意	: 作業場の換気を十分に行い、マスク、保護眼鏡、保護手袋等の適切な保護具を着用し、直接の接触を防ぐ。取り扱い後は、手、顔等を良く洗い、うがいをする。
保存場所	: 被験物質保存室及び被験物質調製室
安定性確認	: 投与期間終了後に株式会社ボゾリサーチセンター 御殿場研究所で残余被験物質の赤外吸収スペクトルを確認した結果、投与期間開始前に確認したスペクトルとほぼ同様であり安定性に問題はなかった（Attachment 2）。
保存試料	: 被験物質約 1 g を保存試料として保存した。
残余品の処理	: 使用後の残余は全て廃棄した。

6.1.2 媒体

名称	: コーン油 ^注
規格	: 生化学用
メーカー	: 富士フイルム和光純薬株式会社
ロット番号（有効期限）	: CAR2149（2024年12月4日）及びSKP4098（2025年10月8日）
保存条件	: 室温
保存場所	: 被験物質調製室

注：媒体は、本試験に先立って実施した「ドデシルオキシラン：特性及び安定性試験並びにGC法を用いた被験液中濃度測定法バリデーション及び安定性試験（媒体：コーン油）」（株式会社ボゾリサーチセンター、試験番号：A-3221）¹⁾において、コーン油中での被験物質の安定性に良好な結果が得られていることから、コーン油を選択した。

6.2 媒体及び被験液の調製

6.2.1 媒体の調製

6.2.1.1 調製方法

コーン油をそのまま用いた。媒体対照群の投与に用いる媒体は、被験物質を取り扱う前に1日必要分ずつ褐色ガラス瓶に分注した。

6.2.1.2 保存条件

媒体対照群投与液として分注した媒体は、被験液と同じ条件で保存した。

6.2.2 被験液の調製

6.2.2.1 調製方法

先ず高用量群液を調製し、中及び低用量群液は高用量群液を媒体で段階的に希釈して調製した。調製手順を以下に示す。

＜高用量群液（200 mg/mL）＞

- 1) 被験物質を室温に戻した後、必要量をビーカーに秤取した。
- 2) これに適量（被験物質とほぼ同量）の媒体を加えて溶解した。
- 3) 溶解液をメスシリンダーに移した。
- 4) 容器を媒体で2、3回洗い、その液もメスシリンダーに入れた。
- 5) 媒体を加えて所定量とした後、ラップで蓋をして転倒混和し、高用量群液とした。
- 6) それをビーカーに移し、中用量群液の調製に必要な量をメスシリンダーに採取した後、残りを1日必要分ずつ褐色ガラス瓶に分注した。

＜中用量群液（60 mg/mL）＞

- 1) 採取した高用量群液を媒体で10/3倍希釈（3 mL→10 mLの割合）した後、ラップで蓋をして転倒混和し、中用量群液とした。
- 2) それをビーカーに移し、低用量群液の調製に必要な量をメスシリンダーに採取し

た後、残りを 1 日必要分ずつ褐色ガラス瓶に分注した。

＜低用量群液（20 mg/mL）＞

- 1) 採取した中用量群液を媒体で 3 倍希釈（1 mL→3 mL の割合）した後、ラップで蓋をして転倒混和し、低用量群液とした。
- 2) それをビーカーに移し、1 日必要分ずつ褐色ガラス瓶に分注した。

6.2.2.2 調製頻度

最大 7 日分を一括して調製し、調製後 7 日以内に使用した。

6.2.2.3 保存条件

冷蔵（許容範囲：1°C~10°C）保存した。

6.2.2.4 安定性

株式会社ボゾリサーチセンターにおいて、ドデシルオキシラン（ロット番号 OPEHE）の 1 及び 200 mg/mL 液（媒体：コーン油）は褐色ガラス瓶で冷蔵（1°C~10°C）8 日間保存後室温 24 時間の安定性が確認されている（試験番号 A-3221¹⁾、Attachment 3）。

6.2.3 被験液の分析

初回投与及び雄の最終投与に使用した各濃度の被験液（任意の場所から各約 10 mL 採取）について、株式会社ボゾリサーチセンター 御殿場研究所にてガスクロマトグラフ（GC）法により濃度測定を実施した。なお、初回投与に使用した被験液については、投与開始前に分析を行った。その結果、各濃度液における被験物質濃度の表示値に対する割合は 97.0%~103.0%であり、いずれも評価基準（濃度許容範囲：表示値に対する割合；100% ± 10%）を満たしたことから、被験液の調製に問題はなかったと判断した（Attachment 4）。なお、本測定法は株式会社ボゾリサーチセンター 御殿場研究所にて実施したバリデーション試験で確立された方法である（試験番号：A-3221）¹⁾。分析法の概略を以下に示した。

- 1) 測定対象物質

ドデシルオキシラン

- 2) 標準物質

被験物質から一部分取したものを標準物質として使用した。

名称 : ドデシルオキシラン

製造者 :

保存場所 : 被験物質保存室及び分析棟 分析室

- 3) 測定実測試料の調製

被験液を転倒混和した後、無作為に n=3 で採取し、以下の表に従い測定実測試料を用時に調製した。

被験液濃度 (mg/mL)	1 次希釈		2 次希釈		希釈率
	採取量 (mL)	定容量 (mL)	1 次希釈液 採取量 (mL)	定容量 (mL)	
20	1	20	1	20	400
60	1	100	2.5	30	1200
200	1	100	2.5	100	4000

溶媒：アセトン

4) GC システム

機器名及び型式		メーカー
GC	HP6890N	Agilent Technologies Inc.
インジェクタ	G2613A	
オートサンプラトレイ	G2614A	
データ処理ソフト	GC ChemStation G2070AJ	

5) GC 測定条件

カラム	: HP-1 (0.32 mm I.D.×30 m、膜厚 0.25 μm、 Agilent Technologies, Inc.)
オーブン温度	: 140°C (Hold 1 min) → 210°C (10°C/min、Hold 2 min)
モード (注入方式)	: スプリット
スプリット比	: 20 : 1
注入口温度	: 200°C
キャリアガス	: He
流量モード	: コンスタントフローモード
流量	: 2.00 mL/min
検出器	: Flame ionization detector (FID)
検出器温度	: 230°C
H ₂ 流量	: 45 mL/min
Air 流量	: 360 mL/min
メイクアップ流量 (N ₂)	: 30 mL/min
注入量	: 1 μL

6) 測定結果の保証

(1) 再注入及び再測定

再注入及び再測定を必要とする事象はなかった。

(2) 試験計画書からの逸脱及び測定結果の信頼性に影響を与える事象

測定開始前に実施したシステム適合性の結果は判定基準を満たした。

試験計画書からの逸脱及び測定結果の信頼性に影響を与える事象はなかった。

6.3 試験系及び系統の選択理由

毒性試験ガイドラインによりげっ歯類を用いた試験が必要とされている。ラットは医薬品などの安全性試験に広く用いられており、この試験に使用される系統のラットは特性がよく知られ、背景資料が豊富であることから選択した。

6.4 試験動物及び群分け

- | | |
|---------------|--|
| 動物種 | : ラット (SPF) |
| 系統 | : Sprague-Dawley [CrI:CD(SD)] |
| 供給源 | : 日本チャールス・リバー株式会社、厚木飼育センター |
| 入荷時週齢 | : 7 週齢 |
| 購入匹数 | : 雄 56 匹、雌 82 匹 |
| 検疫・馴化期間 | : 22 日間 |
| 投与開始時週齢 | : 10 週齢 |
| 供試動物数 | : 雄 48 匹、雌 68 匹 |
| 検疫・馴化期間中の観察 | : 全動物について、体重測定 (5 回: 馴化 1、3 日、以降は週 1 回以上及び群分け日)、生死及び体外表、栄養状態、姿勢、行動及び排泄物の異常などの一般状態観察 (1 回/日) を実施した。詳細な一般状態の観察 (詳細は 6.12.2 項参照) を検疫終了後、群分けまでの期間に実施した (1 回)。更に、雄は検疫期間終了後、群分け前までに陰囊内の精巣の触診 (1 回) 及び陰茎亀頭の観察 (1 回) を実施し、雌は検疫期間終了後、群分け前まで膣垢を採取して性周期を調べた (詳細は 6.12.5 項参照)。 |
| 供試動物の選択 | : 検疫・馴化期間中の一般状態、詳細な一般状態の観察、入荷日から群分け日までの体重推移、精巣の触診結果及び性周期 (馴化 6~19 日の 14 日間) に異常がなく、性成熟に達した陰茎亀頭であることを条件とし、健康と考えられる動物を選択した。 |
| 群分け | : 上記に示した手順で選択した動物のうち、体重増加 (検疫初回の体重~群分け時の体重) が順調な動物を群分け当日 (投与開始前日) の体重に基づいて、各群の平均体重ができるだけ均等となるように群分けを行った。動物の割付けはコンピュータを用いたブロック・ランダム化により行った (ブロック配置により必要な群を構成し、試験群及び群内の個体番号を無作為に割当てた)。 |
| 投与開始時の体重の変動範囲 | : 雄は 351 g~424 g (平均値: 383 g)、雌は 211 g~280 g |

(平均値：251 g)であり、雌雄それぞれ平均体重の±20%以内であった。

余剰動物の処置 : 供試動物の選択条件を満たして余剰となった雌 12 匹は交配予備動物（無処置動物）として継続飼育し、交配群動物の交配が終了した後、2020 年 12 月 2 日に試験から除外した。その他の残余動物の雄 8 匹、雌 2 匹（性周期異常）は、群分け後の 2020 年 10 月 28 日に試験から除外した。なお、試験から除外された動物は、動物管理部門に移管した。

6.5 飼育条件

飼育室番号 : 907 号室及び 908 号室。検疫・馴化期間は性ごと（雄：907 号室、雌：908 号室）に収容し、群分け後は各群を 2 分割して収容した（表 1 参照）。

表 1. 飼育室の動物振り分け表（群分け後）

飼育室番号	907		908	
性	雄	雌	雄	雌
動物番号	1001~1007	1101~1107	1008~1012	1108~1112
	2001~2007	2101~2107	2008~2012	2108~2112
	3001~3007	3101~3107	3008~3012	3108~3112
	4001~4007	4101~4107	4008~4012	4108~4112
		5101~5105		5106~5110
		6101~6105		6106~6110

温度	: 23°C ± 3°C
相対湿度	: 50% ± 20%
換気回数	: 1 時間 10~15 回
照明	: 1 日 12 時間 (07:00~19:00)
ケージ	: 床敷 (コンフィネスト: 株式会社ファルマ、ロット番号 2019-10、15T023061) を入れたプラスチックケージ (W440 × D275 × H180 mm: 株式会社羽生精密)
ケージ収容匹数	: 入荷から群分けまで; 同性 2 匹/ケージ 群分け以降; 雌雄とも 1 匹/ケージ 交配中; 雌雄各 1 匹の計 2 匹/ケージ 交尾成立雌; 授乳 13 日まで 1 腹単位/ケージ
飼料	: 固型飼料 CRF-1 (γ 線滅菌、オリエンタル酵母工業株式会社、ロット番号 200305、200520) をステンレス製給餌器に入れて自由に摂取させた。
飲料水	: 御殿場市営水道水を自動給水装置により自由に摂取させた。

環境エンリッチメント

: 動物実験委員会の指針に従って、適切な環境エンリッチメントを与えた。

6.6 飼料、床敷及び飲料水中の混入物質

下記の混入物質に関する分析報告書の結果が基準内であることを確認した。

- 飼料の分析報告書（Eurofins Food and Product Testing、供試全ロット）
- 床敷の分析報告書（一般社団法人日本食品分析センター、供試全ロット）
- 水道法に準拠した水質の分析報告書（芝浦セムテック株式会社、年4回）

6.7 動物の識別

- | | | |
|--------|---|--|
| 小動物用耳標 | : | 番号が刻印された小動物用耳標を入荷時に装着し、この番号を個体識別の基本とした。 |
| 動物番号 | : | 群分け後は、試験群、性及び群内の個体が区別できるよう、4桁の数字を割り当てた（表2及び表3の群構成参照）。出生児については生後13日の体重測定時に、母動物番号に続けて性別を示すアルファベット（雄はM、雌はF）、同腹内番号を示す2桁の出生児番号を体重測定順に割付け、油性インクによる記号で識別した。 |
| ケージラベル | : | 群分け後、各飼育ケージには投与量（群）ごとに色分けしたケージラベルを添付し、試験番号、投与経路、投与量、性別、動物番号、耳標番号、剖検予定日（雄及び非交配群の雌）、交尾成立日（交配群の雌雄）及び分娩日（交配群の雌）を明記した。ケージラベルの裏には試験番号、性別及び耳標番号のみを表示し、詳細な一般状態の観察、機能検査あるいは握力測定時には、ケージラベルを裏返して観察者への投与量の情報を制限した。 |

6.8 投与経路、投与期間、投与回数及び回復期間とそれらの選択理由

毒性試験ガイドラインに従い、投与経路は、経口投与を選択した。投与期間は、投与日数として、交配群雌雄ともに交配前14日間、更に、雄は交配開始後28日間、雌は交配期間及び妊娠期間を通して授乳13日までとし、非交配群の雌は42日間とした。投与回数は反復投与試験で一般的に行われている1日1回（7日/週）とした。

回復動物は、42日間の投与期間が終了した後、被験物質投与に起因した変化の可逆性、持続性並びに遅発性毒性等について観察するため、14日間の回復期間を設けた。

6.9 投与方法

投与方法は、げっ歯類の経口投与において一般的な強制経口投与とした。

投与容量は 5 mL/kg とし、フレキシブル胃ゾンデを用いて被験液を胃内に強制経口投与した（概ね午前中）。ただし、投与時に分娩途中であった場合、分娩の終了を待って投与を行った。媒体対照群には媒体（コーン油）を同様に投与した。動物ごとの投与液量（表示単位：0.1 mL）は直近の体重を基準に算出した。

6.10 投与量及び群構成

投与量は 100、300 及び 1000 mg/kg の 3 用量とし、試験群はこれらの用量群に媒体対照群を加えた 4 群構成とした。1 群当たりの動物数は交配を行う交配群として各群で雌雄各 12 匹、交配しない非交配群として媒体対照群及び高用量群で雌各 10 匹とした。なお、媒体対照群及び高用量群における雄の 5 匹、非交配群の雌の 5 匹は、最終投与終了日以降、回復動物とした。群構成を表 2 及び表 3 に示す。

表 2. 群構成（雄）

試験群	投与量 (mg/kg)	被験液 濃度 (mg/mL)	投与 容量 (mL/kg)	交配群			
				投与終了時剖検群		回復終了時剖検群	
				動物数	動物番号	動物数	動物番号
媒体対照	0	0	5	7	1001-1007	5	1008-1012
低用量	100	20	5	12	2001-2012		
中用量	300	60	5	12	3001-3012		
高用量	1000	200	5	7	4001-4007	5	4008-4012

表 3. 群構成（雌）

試験群	投与量 (mg/kg)	被験液 濃度 (mg/mL)	投与 容量 (mL/kg)	交配群		非交配群			
				投与終了時剖検群		投与終了時剖検群		回復終了時剖検群	
				動物数	動物番号	動物数	動物番号	動物数	動物番号
媒体対照	0	0	5	12	1101-1112	5	5101-5105	5	5106-5110
低用量	100	20	5	12	2101-2112				
中用量	300	60	5	12	3101-3112				
高用量	1000	200	5	12	4101-4112	5	6101-6105	5	6106-6110

6.11 投与量の設定根拠

本試験の投与量は、先に実施した「ドデシルオキシラン：ラットを用いた 14 日間反復経口投与毒性試験（投与量設定試験）（試験番号：N-R096、投与量：250、500 及び 1000 mg/kg、5 匹/群/性、株式会社ボゾリサーチセンター）」²⁾の結果を参考に設定した。投与量設定試験では、1000 mg/kg の雄で総コレステロールの増加、総タンパク質及びアルブミンの低下、肝臓重量の増加が認められ、250 mg/kg 以上の雌で赤血球数及びヘマトクリットの増加、500 mg/kg 以上の雌で脾臓重量の低下、1000 mg/kg の雌で赤血球容積分布幅の低下、尿素窒素及び肝臓重量の増加が認められた。一方、1000 mg/kg においても雌雄ともに死亡はみられず、一般状態、体重、摂餌量及び剖検

には被験物質投与による明らかな毒性変化は認められなかった。これらのことから、本試験の高用量群を、投与量設定試験と同様に毒性試験ガイドライン〔新規化学物質等に係る試験の方法について（平成 23 年 3 月 31 日）〕に示されている限度試験の最高投与量である 1000 mg/kg に設定し、中用量群以下を公比約 3 で除して 300 及び 100 mg/kg に設定した。

6.12 観察及び検査の方法

試験日の起算に関しては以下の通りとした。

投与開始日	: 投与 1 日 (Day of dosing 1、Day 1)
投与 1 日から 7 日	: 投与 1 週 (Week of dosing 1、Week 1)
回復開始日 (投与 42 日の翌日)	
	: 回復 1 日 (Day 1 of recovery)
回復 1 日から 7 日	: 回復 1 週 (Week 1 of recovery)
交尾成立日	: 妊娠 0 日 (Gestation Day 0)
分娩終了日 (出産日)	
	: 授乳 0 日 (Lactation Day 0)
出生日	: 生後 0 日 (Postnatal Day 0)

6.12.1 一般状態の観察

全動物について、投与期間中は毎日 3 回（投与前、投与直後及び投与 1~4 時間後、ただし、詳細な一般状態の観察実施日は投与前と投与直後の 2 回/日）、回復期間中は毎日 1 回（午前；07:00~12:30）、剖検日は 1 回（剖検のための動物搬出前）、それぞれ生死、身体の外観、排泄物（尿、糞）、栄養状態、姿勢及び行動などの一般状態における異常の有無をケージサイドから観察した。なお、動物を手にとっての詳細な観察は、投与開始日に 1 回、投与 2 日及び投与 2 週以降は詳細な一般状態の観察と兼ねて行った。

6.12.2 詳細な一般状態の観察、機能検査、握力及び自発運動量の測定

詳細な一般状態の観察は、全動物について行った（投与期間中は検査当日の投与後）。

雄及び非交配群雌は投与期間中及び回復期間中毎週 1 回、交配群雌は交配前投与期間中毎週 1 回、交配期間中の未交尾動物は交配開始後 6 及び 13 日、妊娠期間中及び授乳期間中は指定された日（交尾動物は妊娠 1、7、14 及び 20 日、分娩動物は授乳 6 及び 13 日）にそれぞれ行った。

機能検査、握力測定及び自発運動量測定は、各群の 5 匹（雄の群内番号 8~12 番、交配群雌は各群の分娩順で動物番号が若い順に 5 匹^{注)}、非交配群雌の群内番号 6~10 番）について行った。投与期間中は交配群雌では授乳 13 日（投与 50~53 日）に、その他は投与期間終了週（投与 37 日）に、回復期間中は媒体対照群及び高用量群の雌雄について回復期間終了週（回復 9 日）に行った。

詳細な一般状態の観察及び機能検査の観察・検査結果については実測値あるいはスコア化した評点法を用いて記録した。また、投与及び回復期間中の観察、検査及び測定は、観察者に対して投与量などの情報を制限（ブラインド化）し、動物をランダムに配置した状態で行った。ただし、自発運動量の測定はブラインド化しないで行った。

注)：交配群雌の検査対象動物を表 4 に示した。

表 4. 機能検査、握力測定及び自発運動量測定を実施した交配群雌動物

試験群	動物番号
媒体対照群	1101、1105、1107、1108、1111
低用量群	2104、2105、2107、2108、2112
中用量群	3101、3102、3104、3110、3111
高用量群	4102、4103、4107、4109、4111

6.12.2.1 詳細な一般状態の観察

1) ケージ内観察

姿勢、痙攣、異常行動

2) 手に持つての観察

ケージからの取り出し易さ、被毛・皮膚の状態、眼・鼻の分泌物、眼球突出、眼瞼閉鎖状態、可視粘膜、自律神経機能（流涙、流涎、立毛、瞳孔径、異常呼吸）、ハンドリングに対する反応、ハンドリング時の発声

3) オープンフィールド内観察

覚醒状態、痙攣、異常行動、常同行動、歩行、姿勢、身繕い、立ち上がり回数、排泄物（排糞数、排尿）

6.12.2.2 機能検査

聴覚反応、接近反応、接触反応、痛覚反応、瞳孔反射、空中正向反射

6.12.2.3 握力測定

機能検査に引き続き、CPU ゲージ MODEL-RX-5（アイコーエンジニアリング株式会社）を用いて前肢及び後肢の握力を測定した。測定は前肢及び後肢とも 2 回行い、それぞれ平均値を算出した。

6.12.2.4 自発運動量測定

握力測定に引き続き、実験動物用自発運動センサー NS-AS01（株式会社ニューロサイエンス）を用いて自発運動量を測定した。測定時間は 1 時間とし、10 分間隔及び 0~60 分の測定値を集計した。

6.12.3 体重測定

雄及び非交配群雌は投与 1、8、15、22、29、36 及び 42 日に、回復動物は更に回復 1、8 及び 14 日に、交配群雌は投与 1、8、15 及び 22 日、妊娠 0、7、14 及び 20 日、授乳 0、4、7 及び 13 日、並びに全動物は剖検日に体重を測定した。また、投与期間中の体重増加量（雄及び非交配群雌：投与 1 日から投与 42 日、交配群雌：投与 1 日から投与 15 日、妊娠 0 日から妊娠 20 日及び授乳 0 日から授乳 13 日）及び回復期間中の体重増加量（回復 1 日から回復 14 日）を算出した。分娩が午後となった母動物の授乳 0 日の体重は午後に、剖検日は前日から約 16 時間絶食させた後、相対器官重量算出用に搬出前に、その他は午前中（投与期間中は投与前）に測定した。交尾不成立雌動物（動物番号 2109、3108）、妊娠 25 日未分娩動物（動物番号 2106、3109、4101、4106）の剖検日の体重並びに交配期間中の雌の体重は評価の対象としないため、帳票には反映させなかった。

6.12.4 摂餌量測定

雄及び非交配群雌は投与 2、8、15、30、36 及び 42 日に、回復動物は更に回復 2、8 及び 14 日に、交配群雌は投与 2、8 及び 15 日、妊娠 1、7、14 及び 20 日並びに授乳 2、4、7 及び 13 日にそれぞれ残餌量を測定し、前日の給餌量との差から 1 匹当たりの 1 日摂餌量を算出した。また、各期間の総摂餌量（雄及び非交配群雌：投与 1 日から投与 15 日及び回復 1 日から回復 14 日、交配群雌：投与 1 日から投与 15 日、妊娠 0 日から妊娠 20 日及び授乳 0 日から授乳 13 日）を算出した。給餌量/残餌量の測定は測定日の午前中（投与期間中は投与前）に行った。

6.12.5 腔垢検査

交配群雌の全動物について、投与開始日から交尾成立まで毎日（午前）腔垢を採取した。交配前投与期間中の腔垢標本について、多数の角化上皮細胞から成る腔垢像を発情の指標とし、発情回数及び発情期から次の発情期までの日数（性周期）、観察雌動物数に対する性周期異常を示した動物数の割合を調べた。交配期間中は腔垢内の精子の有無を調べるのに加え、腔垢像を発情前期（P）、発情期（E）、発情後期（M）及び発情休止期（D）に分類した。

非交配群雌については、交配群の交配成績から交尾までに要した日数あるいは交尾率に被験物質投与の影響が認められなかったことから、腔垢検査は実施しなかった。

交配期間中の腔垢観察は交尾確認を目的とすることから、腔垢像の分類結果については評価の対象としないため、帳票には反映させなかった。

6.12.6 交配

交配前投与期間終了後、交配群の同投与群内の個体番号が同じ雌雄を 1:1 で終夜同居させ、翌朝、腔内の腔栓形成あるいは腔垢中に精子を確認したものを交尾成立とみなした。同居は雌を雄のケージへ移すことにより行った。交配開始日を 0 日と起算し

て交尾までに要した日数を求めた。同一雌雄による同居期間は最長 14 日間であった。交尾が成立しなかった雌は交配期間終了日を妊娠 0 日と仮定して、一般状態の観察、体重・摂餌量測定並びに投与を継続し、25 日後に剖検を行った。なお、交尾不成立雌動物については、仮定した妊娠日に基づくデータは評価から除外した。

6.13 分娩及び哺育観察

6.13.1 母動物の観察

交尾成立雌動物は、妊娠 21 日から妊娠 25 日の午前まで 1 日 2 回（午前・午後）、分娩の有無並びにその終了を確認するとともに分娩状態（胎盤の処理及び出産児の状態）を観察した。交尾成立から分娩終了までを妊娠期間とし、妊娠期間を 0.5 日単位で求めた。なお、分娩終了の確認は 17:00 までとした。分娩が終了した母動物は授乳 13 日まで出生児を哺育させ、児集め、営巣及び授乳を指標として哺育状態を観察し、授乳 14 日に病理学検査に供した。妊娠 25 日の午前までに分娩しなかった 4 例の雌動物（動物番号 2106、3109、4101、4106）は、イソフルラン吸入麻酔下で採血後、腹大動脈からの放血により安楽死させ、剖検を行った。その結果、3 例（動物番号 2106、3109 及び 4101）は、着床は認められなかったため不妊と判断した。これらの動物の妊娠期間中のデータは評価から除外した。残りの 1 例（動物番号 4106）は子宮内に着床の痕跡が認められたため、黄体数及び着床数を数え記録した。

6.13.2 出生児の観察

生後 0 日に出生児数及び死産児数を数え、合わせて出産児数とした。出産児は口腔内を含む外表異常の有無を肉眼的に観察し、性別を判定した。出生児は体重を個体別に測定し、更にスライドゲージを用いて、肛門縁から生殖突起の付け根までの距離（AGD ; anogenital distance、肛門生殖突起間距離）を測定（0.1 mm 単位）した後、母動物に哺育させた。死産児はブアン液で固定して保存した。

生後 13 日まで、母動物の一般状態観察時に死亡児及び異常行動の有無を確認した。死亡児は肉眼で異常の有無を観察した後、ブアン液で固定して保存した。生存児は生後 0、4、7 及び 13 日に体重を個体別に測定し、各腹単位で雌雄別に平均値を算出した。また、体重増加量（生後 0 日から生後 4 日、生後 4 日から生後 7 日及び生後 4 日から生後 13 日）を算出した。生後 4 日の性別の確認及び体重測定後、1 腹当たりの哺育児数を 8 匹（可能な限り雌雄 4 匹ずつ）となるように調整した。哺育児数が 8 匹に満たない腹については調整を行わなかった。除外された出生児のうち各群分娩順の 5 腹について、腹ごとに生後 4 日除外児を無作為に選び、イソフルラン吸入麻酔下に開腹し、腹大動脈から血液（目標：合計約 0.6 mL）を無処理のサンプルチューブに注射筒を用いて採取した^註。血液は遠心分離（3,000 rpm、約 1,600×g、10 分間）して血清を得た。血清は、腹ごとにプールした後（約 0.2 mL）、ポリプロピレン容器に移しホルモン測定に備え-80°C の冷凍庫（許容範囲：-70°C 以下）に保存し、試験場所に輸送したが、雄及び生後 13 日齢児のホルモン測定結果に影響が認められなかったため、

測定は実施せず試験終了までに廃棄した。採血した児は、残りの除外された出生児とともにイソフルラン吸入麻酔下で腹大動脈からの放血により安楽死させた後、廃棄した。

生後 13 日に全生存児の性別を確認し、雄児について乳頭/乳輪数を調べ、記録した。各群分娩順で両性の児が存在する 5 腹について、各腹から無作為に雌雄各 1 匹を選び、イソフルラン吸入麻酔下に開腹し、腹大動脈から血液（目標：合計約 2.0 mL）を無処理のサンプルチューブに注射筒を用いて採取した。血液は遠心分離（3,000 rpm、約 1,600×g、10 分間）して血清を得た。血清は腹ごとにプールした後、ポリプロピレン容器に分注（約 0.3 mL×3 本；それぞれ 1 本ずつ赤、青、緑に色分けした）し、試験場所への送付時まで-80°C の冷凍庫（許容範囲：-70°C 以下）に保存した。採血した児は、残りの生存児とともにイソフルラン吸入麻酔下で腹大動脈からの放血により安楽死させ、外部生殖器に注意を払い体外表、胸腔内及び腹腔内の主要器官/組織を肉眼的に観察した。全母体の各腹雌雄各 1 匹（出生児番号：M01 及び F01）の甲状腺は、摘出してリン酸緩衝 10%ホルマリン液で固定し、重量を測定後、同固定液に保存した。甲状腺重量（絶対重量）は左右別々に測定し、その合計値と剖検日の体重から体重 100 g 当たりの相対重量を算出した。

注： 媒体対照群の生後 4 日除外児（母動物番号 1101）の採血量が規定量に満たなかった（約 0.4 mL）が、血清は規定量採取することができた。

6.14 尿検査

以下の動物を検査対象とした。

投与 6 週（投与 38~39 日）

： 雄及び非交配群雌の各群 5 匹（雄の群内番号 8~12 番、
非交配群雌の群内番号 6~10 番）

回復 2 週（回復 10~11 日）

： 回復動物の全動物

採尿器をセットしたケージに動物を個別に収容し（投与 6 週検査では投与後に収容した）、絶食・自由摂水下で 4 時間尿を、次いで自由摂食・自由摂水下でその後の 20 時間尿を採取し、表 5 に記載した項目及び方法により検査した。なお、給水には給水瓶を用いた。また、尿検査時には、前日からの 1 日当たりの摂水量（単位：mL/24h）を、給水瓶を用いて重量法で測定した（1 g = 1 mL とみなした）。

表 5. 尿検査の項目、測定法及び使用機器など

1) 4 時間尿についての検査	
検査項目	測定方法
pH	マルティステックス試験紙 ^{a)}
タンパク質	マルティステックス試験紙 ^{a)}
ケトン体	マルティステックス試験紙 ^{a)}
グルコース	マルティステックス試験紙 ^{a)}
潜血	マルティステックス試験紙 ^{a)}
ビリルビン	マルティステックス試験紙 ^{a)}
ウロビリノーゲン	マルティステックス試験紙 ^{a)}
色調	肉眼観察
沈渣	鏡検法
尿量 (4 時間量) 注 1	容量測定 (単位: mL)
2) 20 時間尿についての検査	
検査項目	測定方法
尿量 (20 時間量) 注 1	容量測定 (単位: mL)
浸透圧	氷点降下法 ^{b)} (単位: mOsm/kg)
ナトリウム ^{注 2}	イオン選択電極法 ^{c)} (単位: mmol/24h)
カリウム ^{注 2}	イオン選択電極法 ^{c)} (単位: mmol/24h)
塩素 ^{注 2}	イオン選択電極法 ^{c)} (単位: mmol/24h)
使用測定機器	
a): 尿自動分析装置 クリニテック 500 (Siemens Healthcare Diagnostics Inc.)	
b): 自動浸透圧測定装置 オズモステーション OM-6060 (株式会社アークレイファクトリー)	
c): 臨床化学自動分析装置 TBA-120FR (キヤノンメディカルシステムズ株式会社)	
注 1: 4 時間の尿量と 20 時間の尿量を合計して 24 時間の尿量 (mL/24h) を算出した。	
注 2: 20 時間尿について濃度を測定し、測定濃度と 24 時間の尿量から 1 日当たりの排泄量を算出した。	

6.15 血液学検査

全動物について剖検前日 (最終投与日あるいは回復 14 日) から一夜 (16~21 時間) 絶食させた。雄及び交配群雌並びに非交配群雌の各群 5 匹 (動物番号が若い順、ただし、交配群雌は分娩順の 5 匹) について、イソフルラン吸入麻酔下に開腹し、腹大動脈から EDTA-2K 加採血瓶 (BD Vacutainer: Becton, Dickinson and Company) に血液 (約 1 mL) を採取した。得られた血液について表 6-1) に記載した項目及び方法により検査した。また、鏡検による確認に備え、全例について、May-Grünwald-Giemsa 染色法による血液塗抹標本を作製し、鏡検した^{注 1}。更に、凝固系の検査用として血液 (0.9 mL) を 3.8%クエン酸ナトリウム溶液加試験管 (血液 9 容に対し 1 容の割合) に採取し、遠心分離 (3,000 rpm、約 1,600×g、10 分間) により得られた血漿について表 6-2) に記載した項目及び方法により検査した。交尾不成立雌動物及び妊娠 25 日未分娩動物についても検査した^{注 2}。ただし、その成績は帳票には反映しなかった。

注 1: 回復 14 日の 1000 mg/kg 群の 1 例 (動物番号 6108) の血小板の測定において、ADVIA による血小板測定値が低値を示したが、血漿採取容器のパフィーコート層は薄く、血液塗抹標本においても血小板は非常に少ないことから、ADVIA 測定値の妥当性に問題はないと判断し、そのままの値を採用した。

注 2: 100 mg/kg 群の交尾不成立雌動物 1 例 (動物番号 2109) の ADVIA 全項目の測定において、初回測定時、機器により白血球百分率の測定不良を示す表示がなされたため再測定を実施し、表示のみられなかった再測定値を採用した。

表 6. 血液学検査の項目、測定法及び使用機器など

1) EDTA-2K 加血液についての検査		
検査項目	測定方法	単位
赤血球数(RBC)	フローサイトメトリー法 ^{a)}	10 ⁴ (E4)/ μ L
ヘモグロビン濃度(HGB)	シアンメトヘモグロビン変法 ^{a)}	g/dL
ヘマトクリット値(HCT)	RBC 及び MCV から算出 ^{a)}	%
平均赤血球容積(MCV)	フローサイトメトリー法 ^{a)}	fL
平均赤血球ヘモグロビン量(MCH)	RBC 及び HGB から算出 ^{a)}	pg
平均赤血球ヘモグロビン濃度(MCHC)	HGB 及び HCT から算出 ^{a)}	g/dL
赤血球容積分布幅(RDW)	赤血球容積ヒストグラムの標準偏差と MCV から算出 ^{a)}	%
網赤血球数(RETIC)	フローサイトメトリー法 ^{a)}	10 ⁹ (E9)/L
血小板数(PLT)	フローサイトメトリー法 ^{a)}	10 ⁴ (E4)/ μ L
白血球数(WBC)	フローサイトメトリー法 ^{a)}	10 ² (E2)/ μ L
白血球分類 ^注	フローサイトメトリー法 ^{a)}	10 ² (E2)/ μ L
2) クエン酸ナトリウム加血液から分離した血漿についての検査 (凝固系検査)		
検査項目	測定方法	単位
プロトロンビン時間(PT)	Coagulometric test ^{b)}	s
活性化部分トロンボプラスチン時間(APTT)	Coagulometric test ^{b)}	s
フィブリノーゲン量(FIB)	Coagulometric test ^{b)}	mg/dL
使用測定機器		
a) : 総合血液学検査装置 アドヴィア 2120i (Siemens Healthcare Diagnostics Inc.)		
b) : 血液凝固自動測定装置 ACL Elite Pro (Instrumentation Laboratory)		
注 : リンパ球(LYMP)、好中球(NEUT)、好酸球(EOS)、好塩基球(BASO)、単球(MONO)及び大型非染色球(LUC)		

6.16 血液化学検査

血液学検査のための採血と同時に、ヘパリン加試験管（ベノジェクト II ヘパリンナトリウム、5 mL 採血用：テルモ株式会社）に採取した血液（約 3 mL）を遠心分離（3,000 rpm、約 1,600×g、10 分間）し、上清を同じ条件で更に遠心して得られた血漿について、表 7 に記載の項目及び方法により検査した。交尾不成立雌動物及び妊娠 25 日未分娩動物についても検査した。ただし、その成績は帳票には反映しなかった。また、凝固促進剤入り試験管（ベノジェクト II オートセップ、6 mL 採血用：テルモ株式会社）に採取した血液（約 4 mL）を遠心分離（3,000 rpm、約 1,600×g、10 分間）し血清を得た。血清は T4（thyroxine）を測定するため、試験番号、採血時期、検査項目、試料番号、採取年月日を明記したポリプロピレン製容器に分注し（約 0.5 mL×3 本。それぞれ 1 本ずつ赤、青、緑に色分けした。）、試験場所への送付時まで-80℃の冷凍庫（許容範囲：-70℃以下）に保存した。

表 7. 血液化学検査の項目、測定法及び使用機器など

検査項目	測定方法	単位
アスパラギン酸アミノトランスフェラーゼ(AST)	UV-rate 法 ^{a)}	IU/L
アラニンアミノトランスフェラーゼ(ALT)	UV-rate 法 ^{a)}	IU/L
アルカリホスファターゼ(ALP)	Bessey-Lowry 法 ^{a)}	IU/L
γ -グルタミルトランスペプチダーゼ(γ -GTP)	L- γ -グルタミル-3-カルボキシ-4-ニトロアニリド法 ^{a)}	IU/L
総胆汁酸(TBA)	3 α -ヒドロキシステロイドデヒドロゲナーゼ(3 α -HSD)法 ^{a)}	μ mol/L
総コレステロール(T-CHO)	CEH-COD-POD 法 ^{a)}	mg/dL
トリグリセライド(TG)	LPL-GK-GPO-POD 法 ^{a)}	mg/dL
リン脂質(PL)	PLD-ChOD-POD 法 ^{a)}	mg/dL
総ビリルビン(T-BIL)	ビリルビンオキシダーゼ法 ^{a)}	mg/dL
グルコース(GLU)	グルコースデヒドロゲナーゼ法 ^{a)}	mg/dL
尿素窒素(BUN)	Urease-LEDH 法 ^{a)}	mg/dL
クレアチニン(CRNN)	Creatininase-creatinase-sarcosine oxidase-POD 法 ^{a)}	mg/dL
ナトリウム(Na)	イオン選択電極法 ^{a)}	mmol/L
カリウム(K)	イオン選択電極法 ^{a)}	mmol/L
塩素(Cl)	イオン選択電極法 ^{a)}	mmol/L
カルシウム(Ca)	OCPC 法 ^{a)}	mg/dL
無機リン(P)	PNP-XOD-POD 法 ^{a)}	mg/dL
総タンパク質(TP)	Biuret 法 ^{a)}	g/dL
アルブミン(ALB)	BCG 法 ^{a)}	g/dL
使用測定機器		
a) : 臨床化学自動分析装置 TBA-120FR (キヤノンメディカルシステムズ株式会社)		

6.17 ホルモン測定

6.17.1 測定対象

6.16項で得られた雄及び交配群雌並びに非交配群雌の血清及び6.13.2項で得られた児動物の血清についてホルモン (T4) を測定した。ただし、測定は雄及び生後 13 日の児動物の測定を優先して実施し、測定結果に被験物質投与の影響と推定される異常がみられなかったことから、非交配群雌、授乳 14 日の母動物及び生後 4 日の児動物並びに回復動物については測定を行わなかった。3 本に分注した血清はそのうち 1 本 (赤、青、緑の順に優先した) を測定し、残余の血清はホルモン測定用の予備として保存した。なお、交尾不成立雌動物及び妊娠 25 日未分娩動物の試料については測定を行わず廃棄した。

6.17.2 試料の送付

測定に供する血清は、凍結した状態でドライアイスと同梱させ、試験場所 (試験主任者宛) に送付した。

送付試料数 : 110 サンプル (交配群雄 30 サンプル×3 本、交配群雌 20 サンプル×3 本、非交配群雌 20 サンプル×3 本、生後 13 日の児動物 20 サンプル×3 本、生後 4 日の児動物 20 サ

サンプル)
送付日 : 2020 年 12 月 9 日 (2020 年 12 月 10 日着)
; 交配群雄 (投与終了時) のサンプル
2020 年 12 月 24 日 (2020 年 12 月 25 日着)
; 上記以外のサンプル

6.17.3 ホルモン (T4) 測定

- 1) ホルモン測定番号
OT-200011
- 2) 分析方法
RIA method (詳細は Attachment 5 参照、最終報告書に添付)
- 3) 測定項目
T4
- 4) 作業・操作
上記測定に関する操作手順は試験場所の SOP に従って行われた。
- 5) ホルモン測定報告書
試験場所においてホルモン測定報告書が作成された。試験責任者はホルモン測定報告書の PDF ファイルを入手し、当試験 (R-1261) の最終報告書に添付した。
また、その内容を当試験の最終報告書に反映させた。
- 6) 残余血清の処理
ホルモン測定終了後の残余血清は、ホルモン測定報告書作成後に試験責任者の指示に基づいて試験場所で廃棄した。

6.18 病理学検査

6.18.1 剖検

最終投与翌日及び回復期間終了日に、血液学検査、血液化学検査及びホルモン測定のために採血した動物は採血後、その他の動物はイソフルラン吸入麻酔下で、腹大動脈からの放血により安楽死させ、外表及び全ての器官/組織を詳細に観察した。交尾不成立雌動物及び妊娠 25 日未分娩動物については、採血して放血安楽死させた後、剖検を行った。なお、母動物については剖検時に着床痕数を数えた。

6.18.2 器官重量

全動物について、表 8 に示す器官の重量 (絶対重量) を測定するとともに、絶対重量と剖検日の体重から体重 100 g 当たりの相対重量を算出した。両側性の器官 (精嚢を除く) については左右別々に測定し、その合計値で評価した。なお、交尾不成立雌動物及び妊娠 25 日未分娩動物の測定結果については、帳票には反映しなかった。

6.18.3 病理組織学検査

検査対象器官/組織を表 8 に示した。全ての成熟動物について全検査対象器官/組織をリン酸緩衝 10%ホルマリン液で固定した。ただし、眼球及び視神経はリン酸緩衝 3% グルタルアルデヒド・2.5%ホルマリン液で固定、精巣及び精巣上体はブアン液で固定した後、リン酸緩衝 10%ホルマリン液で保存した。以降の検査は次の通りとした。

1) パラフィン包埋及びヘマトキシリン・エオジン（H・E）染色標本作製

以下に記載した動物の全検査対象器官/組織について標本作製した。

- － 投与終了時及び回復終了時の計画剖検時に採血を実施した雄及び交配群雌並びに非交配群雌の各群 5 匹（動物番号が若い順、ただし、交配群雌は分娩順の 5 匹）
- － 妊娠 25 日未分娩動物、その交配相手雄（ただし、未分娩動物が不妊の場合）
- － 交尾不成立雌雄動物

ただし、甲状腺に関しては、被験物質投与による変化が疑われなかったため（親動物に組織学変化及び甲状腺重量の変動はみられなかったため）、採血を実施しなかった雄及び交配群雌並びに雌雄の生後 13 日児動物（全母体の各腹雌雄各 1 例）については、標本作製を行わなかった。

2) 病理組織学検査

鏡検は以下の通り実施した。

＜投与終了時剖検群＞

媒体対照群及び高用量群

： 全検査対象器官/組織

中及び低用量群 ： 高用量群において、被験物質投与の影響と考えられる変化はみられなかったため、鏡検は実施しなかった。

＜回復終了時剖検群＞

全群 ： 全検査対象器官/組織

＜妊娠 25 日未分娩動物及びその交配相手雄(ただし、未分娩動物が不妊の場合のみ)、交尾不成立雌雄動物＞

全群 ： 全検査対象器官/組織

＜計画剖検時に採血を実施しなかった雄及び交配群雌並びに生後 13 日児動物＞
被験物質投与による変化が疑われなかったため、甲状腺の鏡検は実施しなかった。

表 8. 病理学検査対象器官/組織

器官/組織	病理組織学検査 (HE 染色)	器官重量測定
脳 (大脳、小脳、橋)	√	√ (脳として)
延髄	√	
坐骨神経	√*	
脊髄 (胸部)	√	
眼球	√*	
視神経	√*	
ハーダー腺	(保存のみ・両側)	
下垂体	√	
甲状腺	√	√ (甲状腺として)
上皮小体	√*	
副腎	√	√
胸腺	√	√
脾臓	√	√
顎下リンパ節	√*	
腸間膜リンパ節	√	
パイエル板 (回腸)	√	
心臓	√	√
胸大動脈	(保存のみ)	
気管	√	
肺 (気管支を含む)	√	
舌	(保存のみ)	
喉頭	(保存のみ)	
食道	(保存のみ)	
胃	√	
十二指腸	√	
空腸	√	
回腸	√	
盲腸	√	
結腸	√	
直腸	√	
顎下腺	√*	
舌下腺	(保存のみ・両側)	
肝臓	√	√
脾臓	√	
腎臓	√	√
膀胱	√	
精巣/卵巣	√/√	√/√
精巣上体/子宮	√/√	√/√
脛	√	
前立腺	√	√
精囊 (凝固腺含む)	√	√
皮膚 (鼠径部)	√*	
乳腺 (鼠径部)	√*	
胸骨	(保存のみ)	
大腿骨	√*	
胸骨骨髓	(保存のみ)	

器官/組織	病理組織学検査 (HE 染色)	器官重量測定
大腿骨骨髓	√*	
大腿四頭筋	√*	
肉眼的異常部位	√	
肛門挙筋・球海綿体筋 (LABC)	(重量測定のみ)	√
カウパー腺 (COW)	(重量測定のみ)	√
陰茎龟头 (GP)	(重量測定のみ)	√
個体識別部 (耳標を装着した耳介)	(保存のみ)	

各項目該当ある場合は√で示した。

*: 両側を摘出し、片側を鏡検した (他の両側性器官は両側を鏡検)。

7. データ採取に使用したコンピュータシステム

- MiTOX-BOZO システム (Version 9.3.1.1、三井 E&S システム技研株式会社)
- PATHOS5 システム (Pathology Operating Systems Ltd.)

7.1 統計解析

7.1.1 パラメータの算出

性周期異常率、交尾率、受胎率、出産率、異常児を有する母動物発現率、生後 0 及び 4 日の性比を群ごとに、妊娠期間、分娩率、出生率、外表異常率、生後 4 及び 13 日生存率を母動物 (腹) ごとに、以下の式により算出した。なお、出生児の体重及び AGD については母動物ごとに雌雄別に平均値を求めた。この際、AGD は生後 0 日の体重の三乗根で除して標準化 (小数点以下第 1 位まで表示) した。雄生存児の乳頭/乳輪数は母動物 (腹) ごとに平均値を求めた。空中正向反射は 3 回の動物ごとのトータルスコアを求めた。

性周期異常率(%) = (性周期異常を示した動物数/観察雌動物数) × 100

交尾率(%) = (交尾動物数/同居動物数) × 100

(雌又は雄の)受胎率(%) = [妊娠した雌(又は雌を妊娠させた雄)の数/交尾した雌(又は交尾した雄)の数] × 100

出産率(%) = (生児出産雌数/妊娠雌数) × 100

異常児を有する母動物発現率 (%) = (異常児を有する母動物数/母動物数) × 100

外表異常率(%) = (外表異常児数/出産児数) × 100

生後 0 日の性比(%) = (雄出生児数/出生児数) × 100

生後 4 日の性比(%) = (生後 4 日の雄生存児数/生後 4 日の生存児数) × 100

妊娠期間(日) = 妊娠 0 日から分娩した日までの日数

分娩率(%) = (出産児数/着床痕数) × 100

出生率(%) = (出生児数/出産児数) × 100

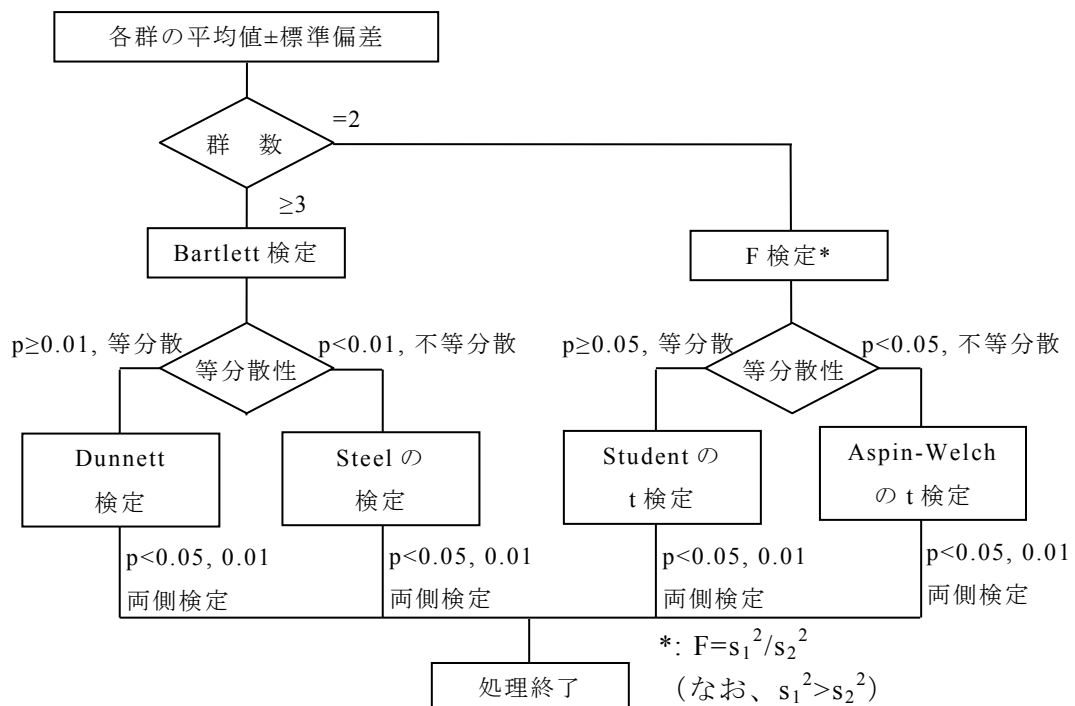
生後 4 日生存率(%) = (生後 4 日の生存児数/出生児数) × 100

生後 13 日生存率(%) = (生後 13 日の生存児数/生後 4 日調整後の児数) × 100

7.1.2 検定

媒体対照群と各被験物質投与群との間で検定を行った。解析には、SAS Release 9.1.3 (SAS Institute Inc.)を使用した。なお、交尾不成立雌は交尾率にのみ含め、交配開始後のデータは試験成績から除外した。

- 1) 体重、体重増加量（雄及び非交配群雌：投与 1 日から投与 42 日及び回復 1 日から回復 14 日、交配群雌：投与 1 日から投与 15 日、妊娠 0 日から妊娠 20 日及び授乳 0 日から授乳 13 日、出生児：生後 0 日から生後 4 日、生後 4 日から生後 7 日及び生後 4 日から生後 13 日）、摂餌量、総摂餌量（雄及び非交配群雌：投与 1 日から投与 15 日及び回復 1 日から回復 14 日、交配群雌：投与 1 日から投与 15 日、妊娠 0 日から妊娠 20 日及び授乳 0 日から授乳 13 日）、オープンフィールド内観察における排糞数と立ち上がり回数、握力及び自発運動量、尿検査の定量的項目及び摂水量、血液学検査、血液化学検査及び器官重量（含、剖検時体重）、発情期発現回数、性周期（発情周期）、交尾までに要した日数、妊娠期間、着床数、出産児数、出生児数、死産児数、生存児数、AGD、乳頭/乳輪数は、群ごとに平均値及び標準偏差を求めた。動物ごとに得られた値あるいは平均値を 1 標本単位とし、下記の模式図に従って検定した^{3),4),5)}。なお、いずれの雄新生児にも乳頭/乳輪はみられなかったため、乳頭/乳輪数について検定は行わなかった。



- 2) 分娩率、外表異常率、出生率、生後 4 及び 13 日生存率については、母動物ごとに得られた率を 1 標本単位とし、群ごとに平均値及び標準偏差を求め、Wilcoxon の順位和検定を行った（有意水準 0.05 及び 0.01、両側）⁶⁾。なお、いずれの群にも外表異常児はみられなかったため、外表異常率について検定は行わなかった。

- 3) 性周期異常率、交尾率、雌雄の受胎率、出産率及び児の性比（生後 0 及び 4 日）については、各群の性周期異常を示した動物数、交尾動物数、雌を妊娠させた雄動物数、妊娠雌動物数、出生児出産雌動物数、雌雄児数に基づき、Fisher の直接確率計算法により検定を行った（有意水準 0.05 及び 0.01、両側）⁷⁾。なお、いずれの群にも外表異常児はみられなかったため、異常児を有する母動物発現率について検定は行わなかった。

8. 試験結果

8.1 一般状態

結果を Table 1-1~1-12、Appendix 1-1~1-36 に示した。

- 1) 投与期間
雄及び非交配群雌では、投与期間を通じ、いずれの動物にも異常はみられなかった。
交配群雌では、交配前、妊娠及び授乳期の各投与期間を通じ、いずれの動物にも分娩/哺育状態を含め異常はみられなかった。
- 2) 回復期間
回復期間を通じ、いずれの動物にも異常は認められなかった。

8.2 詳細な一般状態の観察、機能検査、握力測定及び自発運動量の測定

8.2.1 詳細な一般状態の観察

8.2.1.1 ケージ内観察

結果を Table 2-1~2-24、Appendix 2-1~2-76 に示した。

- 1) 投与期間
いずれの動物にも妊娠中及び授乳中を含め、被験物質投与に関連すると考えられる異常はみられなかった。
- 2) 回復期間
いずれの動物にも被験物質投与に関連すると考えられる異常はみられなかった。

8.2.1.2 手に持っでの観察

結果を Table 2-25~2-86、Appendix 2-77~2-152 に示した。

- 1) 投与期間
いずれの動物にも妊娠中及び授乳中を含め、被験物質投与に関連すると考えられる異常はみられなかった。
- 2) 回復期間
いずれの動物にも被験物質投与に関連すると考えられる異常はみられなかった。

8.2.1.3 オープンフィールド内観察

結果を Table 2-87~2-158、Appendix 2-153~2-304 に示した。

1) 投与期間

いずれの動物にも妊娠中及び授乳中を含め、被験物質投与に関連すると考えられる異常はみられなかった。

交配群雌の 1000 mg/kg 群では、媒体対照群と比べ立ち上がり回数が投与 2 週に有意な高値、妊娠 14 日に有意な低値を示したが、非交配群雌の同観察時点あるいは妊娠期間の他の観察時点に異常はみられず、いずれも試験施設の背景データ (Min.-Max. ; 投与 2 週 : 6-8、妊娠 14 日 : 4-7、12 試験、2015-2020 年) の範囲内であることから、生理的変動範囲内の変化で偶発的有意差と判断した。その他に 100 mg/kg 群で授乳 13 日の立ち上がり回数に媒体対照群と比べ有意な高値が認められたが、用量に関連しない変化であり、偶発的な有意差と判断した。

非交配群雌の 1000 mg/kg 群では、投与 3 週の立ち上がり回数に媒体対照群と比べ有意な低値が認められたが、その他の観察時点に異常はみられず、試験施設の背景データ (Min.-Max. ; 6-9、12 試験、2015-2020 年) の範囲内であることから、生理的変動範囲内の変化で偶発的有意差と判断した。

その他には、雌雄とも立ち上がり回数及び排糞の個数に媒体対照群と各被験物質投与群との間に差は認められなかった。

2) 回復期間

いずれの動物にも被験物質投与に関連すると考えられる異常はみられなかった。雄の 1000 mg/kg 群で回復 2 週の立ち上がり回数に媒体対照群と比べ有意な低値が認められたが、回復 1 週には異常はみられず、試験施設の背景データ (Min.-Max. : 3-6、12 試験、2015-2020 年) の範囲内であることから、生理的変動範囲内の変化で偶発的有意差と判断した。

その他には、雌雄ともに立ち上がり回数及び排糞の個数に媒体対照群と 1000 mg/kg 群との間に差は認められなかった。

8.2.2 機能検査

結果を Table 2-159~2-163、Appendix 2-305~2-318 に示した。

1) 投与期間終了週検査

いずれの動物にも聴覚反応、接近反応、接触反応、痛覚反応、瞳孔反射及び空中正向反射に被験物質投与に関連すると考えられる異常はみられなかった。

2) 回復期間終了週検査

いずれの動物にも聴覚反応、接近反応、接触反応、痛覚反応、瞳孔反射及び空中正向反射に被験物質投与に関連すると考えられる異常はみられなかった。

8.2.3 握力

結果を Table 2-164~2-168、Appendix 2-319~2-332 に示した。

1) 投与期間終了週測定

雄、非交配群雌及び交配群雌では、媒体対照群と各被験物質投与群との間に差は

認められなかった。

2) 回復期間終了週測定

雌雄ともに媒体対照群と 1000 mg/kg 群との間に差は認められなかった。

8.2.4 自発運動量

結果を Fig. 1~5、Table 2-169~2-173、Appendix 2-333~2-346 に示した。

1) 投与期間終了週測定

雄及び非交配群雌では、媒体対照群と各被験物質投与群との間に差は認められなかった。なお、交配群雌では 100 mg/kg 群で測定開始後 20~30 分の運動量に媒体対照群と比べ有意な低値が認められたが、用量に関連しない変化であり、偶発的な有意差と判断した。

2) 回復期間終了週測定

雄では、1000 mg/kg 群と媒体対照群との間に差は認められなかった。なお、雌では、測定開始後 30~40 分の運動量に媒体対照群と比べ有意な高値が認められたが、その他の測定時点及び総運動量に異常はみられず、試験施設の背景データ (Min.-Max. ; 89-291、12 試験、2015-2020 年) の範囲内であることから、生理的変動範囲内の変化で偶発的な有意差と判断した。

8.3 体重

結果を Fig. 6~8、Table 3-1~3-7、Appendix 3-1~3-22 に示した。

1) 投与期間

雄では、媒体対照群と各被験物質投与群との間に差は認められなかった。

交配群雌では、交配前、妊娠及び授乳期の各期間を通じ、媒体対照群と各被験物質投与群との間に差は認められなかった。

非交配群雌では、媒体対照群と 1000 mg/kg 群との間に差は認められなかった。

2) 回復期間

雌雄ともに媒体対照群と 1000 mg/kg 群との間に差は認められなかった。

8.4 摂餌量

結果を Fig. 9~11、Table 4-1~4-9、Appendix 4-1~4-28 に示した。

1) 投与期間

雄では、媒体対照群と各被験物質投与群との間に差は認められなかった。

交配群雌では、交配前及び授乳期の各期間を通じ媒体対照群と各被験物質投与群との間に差は認められなかった。300 mg/kg 群で妊娠 14 日及び妊娠期間中の総摂餌量に媒体対照群と比べ有意な高値が認められたが、用量に関連しない変化であり、偶発的な有意差と判断した。

非交配群雌では、媒体対照群と 1000 mg/kg 群との間に差は認められなかった。

2) 回復期間

雌雄ともに媒体対照群と 1000 mg/kg 群との間に有意な差は認められなかった。

8.5 尿検査（摂水量測定を含む）

結果を Table 5-1~5-21、Appendix 5-1~5-30 に示した。

1) 投与期間終了週検査

定性項目については、いずれの動物にも異常はみられなかった。

定量項目については、雄の 1000 mg/kg 群で尿浸透圧に媒体対照群と比べ有意な高値が認められた。その他には、いずれの検査項目においても雌雄ともに媒体対照群と各被験物質投与群との間に差は認められなかった。

2) 回復期間終了週検査

定性項目については、いずれの動物にも異常はみられなかった。

定量項目については、雄の 1000 mg/kg 群で尿浸透圧に媒体対照群と比べ有意な高値が認められた。その他には、いずれの検査項目においても雌雄ともに媒体対照群と 1000 mg/kg 群との間に有意な差は認められなかった。

8.6 血液学検査

結果を Table 6-1~6-5、Appendix 6-1~6-14 に示した。

1) 投与期間終了時検査

雄及び交配群雌では、いずれの検査項目においても媒体対照群と各被験物質投与群との間に差は認められなかった。

非交配群雌では、1000 mg/kg 群で媒体対照群と比べ MCV 及び MCH に有意な高値が認められたが、関連するパラメータに異常はなく、いずれも試験施設の背景データ（Min.-Max. ; MCV : 51.7-55.7、MCH : 18.0-19.7、12 試験、2015-2020 年）の範囲内であることから、生理的変動範囲内の変化で偶発的有意差と判断した。

2) 回復期間終了時検査

雄では、1000 mg/kg 群で APTT に媒体対照群と比べ有意な高値が認められたが、関連するパラメータに異常はなく、試験施設の背景データ（Min.-Max. ; 15.3-20.8、12 試験、2015-2020 年）の範囲内であることから、生理的変動範囲内の変化で偶発的有意差と判断した。

雌では、いずれの検査項目においても媒体対照群と 1000 mg/kg 群との間に差は認められなかった。

8.7 血液化学検査

結果を Table 7-1~7-5、Appendix 7-1~7-14 に示した。

1) 投与期間終了時

雄では、1000 mg/kg 群で媒体対照群と比べ ALT 及び BUN に有意な高値が認め

られた。その他に 1000 mg/kg 群で媒体対照群と比べ Cl に有意な高値、Ca 及び TP に有意な低値が認められたが、いずれも媒体対照群と比べ 6%以下の軽微な変動であり、試験施設の背景データ (Min.-Max. ; Cl : 106-108、Ca : 9.6-10.4、TP : 5.6-6.3、12 試験、2015-2020 年) の範囲内であることから、生理的変動範囲内の変化で偶発的有意差と判断した。

交配群雌では、いずれの検査項目においても媒体対照群と各被験物質投与群との間に差は認められなかった。

非交配群雌では、1000 mg/kg 群で媒体対照群と比べ ALT に有意な高値が認められた。その他に 1000 mg/kg 群で媒体対照群と比べ TG 及び Na に有意な低値が認められたが、関連するパラメータに異常はなく、いずれも試験施設の背景データ (Min.-Max. ; TG : 13-52、Na : 140-143、12 試験、2015-2020 年) の範囲内であることから、生理的変動範囲内の変化で偶発的有意差と判断した。

2) 回復期間終了時

雄では、いずれの検査項目においても媒体対照群と 1000 mg/kg 群との間に差は認められなかった。

雌では、1000 mg/kg 群で媒体対照群と比べ T-BA に有意な低値、Cl に有意な高値が認められたが、関連するパラメータに異常はなく、いずれも試験施設の背景データ (Min.-Max. ; T-BA : 12.1-28.4、Cl : 107-110、12 試験、2015-2020 年) の範囲内であることから、生理的変動範囲内の変化で偶発的有意差と判断した。

8.8 血清中ホルモン濃度測定

結果をホルモン濃度測定報告書 (Attachment 5、最終報告書に添付) に示した。

1) 交配群雄投与期間終了時

T4 濃度には、媒体対照群と各被験物質投与群との間に差は認められなかった。

2) 生後 13 日齢児

T4 濃度には、被験物質の影響は認められなかった。なお、100 mg/kg 群で媒体対照群と比べ有意な低値が認められたが、用量に関連しない変化であり、偶発的な有意差と判断した。

8.9 器官重量

結果を Table 8-1~8-8、Appendix 8-1~8-30 に示した。

1) 投与期間終了時

雄では、1000 mg/kg 群で媒体対照群と比べ肝臓の相対重量に有意な高値が認められた。その他に、1000 mg/kg 群で陰茎亀頭の絶対重量に媒体対照群と比べ有意な低値が認められたが、相対重量に有意差はなく、他の生殖腺類の重量に異常はないこと、試験施設の背景データ (Min.-Max. : 108.2-140.4、6 試験、2015-2020 年) の範囲内であることから、生理的変動範囲内の変化で偶発的有意差と判断した。

交配群雌では、媒体対照群と比べ 1000 mg/kg 群で肝臓の絶対及び相対重量に有意な高値、300 mg/kg 群で肝臓の絶対重量に有意な高値が認められた。なお 300 mg/kg 群で脾臓の絶対重量に媒体対照群と比べ有意な高値が認められたが、用量に関連しない変化であり、偶発的な有意差と判断した。

非交配群雌では、1000 mg/kg 群で肝臓の相対重量に有意な高値が認められた。その他に 1000 mg/kg 群で心臓の相対重量に媒体対照群と比べ有意な低値が認められたが、媒体対照群と比べ 10%未満の軽微な変動であり、試験施設の背景データ (Min.-Max.: 0.30-0.34、10 試験、2015-2020 年) のほぼ範囲内であることから、生理的変動範囲内の変化で偶発的な有意差と判断した。

2) 回復期間終了時

雄では、いずれの器官重量においても媒体対照群と 1000 mg/kg 群との間に差は認められなかった。

雌では、1000 mg/kg 群で甲状腺の絶対及び相対重量に媒体対照群と比べ有意な高値が認められた。

8.10 剖検所見

結果を Table 9-1~9-6、Appendix 9-1~9-116 に示した。

1) 投与期間終了時及び回復期間終了時検査

交配群雌雄及び非交配群雌のいずれの動物にも被験物質投与の影響と考えられる肉眼所見はみられなかった。なお、いくつかの臓器に肉眼的変化がみられたが、その病理学的性状又は発現状況から被験物質投与に関連しない偶発的な変化と考えられた。

2) 交尾不成立雌動物及び妊娠 25 日未分娩動物

交尾が成立しなかった 100 及び 300 mg/kg 群の雌各 1 例 (動物番号 2109 及び 3108)、妊娠 25 日までに分娩しなかった 100 及び 300 mg/kg 群の各 1 例 (動物番号 2106 及び 3109) と 1000 mg/kg 群の 2 例 (動物番号 4101 及び 4106) では、いずれの臓器にも肉眼的異常はみられなかった。

8.11 病理組織学検査

結果を Table 10-1~10-18、Appendix 9-1~9-116 に示した。

1) 投与期間終了時及び回復期間終了時検査

交配群雌雄及び非交配群雌のいずれの動物にも被験物質投与の影響と考えられる変化はみられなかった。みられた変化はいずれも、その発生状況及び病理学的性状、動物種、週齢から自然発生病変もしくは偶発性変化と考えられた。

2) 交尾不成立雌雄動物、妊娠 25 日未分娩動物及びその交配相手雄

交尾不成立雌雄動物 (動物番号 2009 と 2109、3008 と 3108) では、特記すべき変化はみられなかった。

妊娠 25 日未分娩動物 (動物番号 2106、3109、4101、4106) 及び不妊であった

雌の交配相手雄（動物番号 2006、3009、4001）では、特記すべき変化はみられず、更に不妊の原因と考えられる病理組織学的変化もみられなかった。

8.12 性周期

結果を Table 11、Appendix 10-1~10-8 に示した。

性周期異常率、発情期像発現回数及び平均性周期（発情周期）には、媒体対照群と各被験物質投与群との間に差は認められなかった。

8.13 交配成績

結果を Table 12-1 及び 12-2、Appendix 11-1~11-8 に示した。

交尾不成立であった 100 及び 300 mg/kg 群の各 1 組を除き、交配 5 日までに各群 12 例中 10~11 組の交尾が成立し、交尾までに要した日数、交尾率及び受胎率には媒体対照群と各被験物質投与群との間に差は認められなかった。なお、妊娠 25 日に至っても未分娩であった 100 及び 300 mg/kg 群の各 1 例と 1000 mg/kg 群の 1 例は着床を確認できず不妊であった。

8.14 分娩成績並びに分娩及び哺育状態

結果を Table 13、14、Appendix 12-1~12-4、13-1~13-4 並びに Table 1-5~1-7、Appendix 1-17~1-28 に示した。

分娩成績では、妊娠期間、出産率、着床痕数、分娩率、出生率、出産児数、出生児数、性比（生後 0 日）及び死産児数に媒体対照群と各被験物質投与群との間で差は認められなかった。また、いずれの群にも外表異常児はみられなかった。

妊娠動物の分娩状態では、妊娠 25 日に至っても未分娩であった 100 及び 300 mg/kg 群の各 1 例と 1000 mg/kg 群の 2 例を除き、妊娠 21.5 から 23.0 日に全例が分娩し、胎盤及び羊膜の処理は正常に行われた。哺育状態では、いずれの母動物にも児集め、営巣及び授乳行動に異常はみられなかった。

8.15 出生児の Anogenital Distance (AGD)

結果を Table 15、Appendix 14-1~14-4 に示した。

生後 0 日の AGD には、雌雄とも媒体対照群と各被験物質投与群との間に差は認められなかった。

8.16 出生児の生存性

結果を Table 16、Appendix 15-1~15-4 に示した。

生後 4 及び 13 日の生存児数及び生存率には、媒体対照群と各被験物質投与群との間に差は認められなかった。

8.17 出生児の性比

結果を Table 17、Appendix 16-1~16-4 に示した。

生後 4 日の性比には、媒体対照群と各被験物質投与群との間に差は認められなかった。

8.18 出生児の体重

結果を Table 18-1 及び 18-2、Appendix 17-1~17-8 に示した。

1000 mg/kg 群では、雌雄の生後 13 日の体重並びに生後 4~13 日の間の体重増加量に媒体対照群と比べ有意な低値が認められた。

その他、出生児の雌雄体重並びに生後 0~4、4~7 及び 4~13 日の間の体重増加量には、媒体対照群と各被験物質投与群との間に差は認められなかった。

8.19 死亡児の肉眼観察所見

結果を Table 19、Appendix 18-1~18-4 に示した。

いずれの死亡児にも肉眼的異常はみられなかった。

8.20 雄新生児の乳頭/乳輪数

結果を Table 20、Appendix 19-1~19-4 に示した。

雄新生児の乳頭/乳輪数には、媒体対照群と各被験物質投与群との間に差は認められず、被験物質投与の影響と考えられる変化はみられなかった。

8.21 出生児の生後 13 日剖検所見

結果を Table 21、Appendix 20-1~20-4 に示した。

いずれの動物にも肉眼的異常はみられなかった。

8.22 出生児の器官重量

結果を Table 22-1 及び 22-2、Appendix 21-1~21-8 に示した。

1000 mg/kg 群では、媒体対照群と比べ雄の剖検時体重に有意な低値及び甲状腺相対重量に有意な高値が認められた。その他には、雌雄とも甲状腺重量には媒体対照群と各被験物質投与群との間に差は認められなかった。

9. 考察及び結論

ドデシルオキシランの 0（対照群：コーン油）、100、300 及び 1000 mg/kg を Sprague-Dawley 系 SPF ラットの雄には交配前 14 日間に加え交配期間を通して剖検前日まで（42 日間）、雌には交配前 14 日間に加え交配期間及び妊娠期間を通して授乳 13 日まで（50~64 日間）、非交配群雌には 42 日間、それぞれ強制経口投与し、反復投与毒性及び生殖発生毒性を検討した。更に、0 及び 1000 mg/kg 群の一部の動物については、雌雄ともに 42 日間投与した後 14 日間の回復期間を設け、毒性変化の可逆性

を検討した。

9.1 反復投与毒性

いずれの投与群においても死亡動物はみられず、一般状態、詳細な一般状態観察、機能検査、握力、自発運動量、体重、摂餌量、血液学検査、血清中ホルモン(T4)及び剖検並びに病理組織学検査には、被験物質投与の影響は認められなかった。

尿検査では、投与終了週に雄の 1000 mg/kg 群で尿浸透圧の高値が認められたが、摂水量、尿量及び尿中電解質に異常は認められず、腎臓の病理組織にも異常はみられないことから、毒性学的意義はないと判断した。なお、回復終了週にも雄の 1000 mg/kg 群で尿浸透圧の高値が認められたが、投与終了週と同様に他のパラメータに異常はないことから、毒性学的意義はないと判断した。

血液化学検査では、BUN の高値が雄の 1000 mg/kg 群で投与終了時に認められ、試験施設の背景データ (Min.-Max. : 13-19、12 試験、2015-2020 年) の範囲を上回り被験物質投与の影響と考えられたが、腎臓の病理組織に異常はみられないことから、毒性学的意義は低いと判断した。ALT の高値が雄及び非交配群雌の 1000 mg/kg 群で投与終了時に認められ、試験施設の背景データ (Min.-Max. : 雄 ; 32-47、雌 ; 28-93、12 試験、2015-2020 年) の範囲を雄は上回り被験物質投与の影響と考えられたが、雌雄とも肝臓の病理組織に異常はみられないことから、毒性学的意義は低いと判断した。これらの変化は回復期間終了時には消失し、回復性がみられた。

器官重量では、投与終了時に雄の 1000 mg/kg 群、交配群雌の 300 及び 1000 mg/kg 群並びに非交配群雌の 1000 mg/kg 群で肝臓重量の高値が認められ、試験施設の背景データ (Min.-Max.[絶対/相対] : 雄 ; 12.59-15.35/2.45-2.84、母動物 ; 9.73-12.28/3.12-3.49、雌 ; 6.70-8.88/2.40-2.78、10 試験、2015-2020 年) の範囲を上回り被験物質投与の影響と考えられたが、いずれも肝臓の病理組織に異常はみられないことから、毒性学的意義は低いと判断した。この変化は回復期間終了時には消失し、回復性がみられた。回復終了時に雌の 1000 mg/kg 群で甲状腺重量に高値が認められたが、投与終了時に異常は認められず、病理組織にも異常はみられないことから、被験物質投与に関連しない偶発的变化と判断した。

9.2 生殖発生毒性

1000 mg/kg 群においても雌雄動物の生殖機能に関するパラメータに被験物質投与の影響は認められないことから、雌雄動物の交尾能、授精能及び受胎能、母動物の妊娠維持、分娩及び哺育行動などへの影響はないと判断した。

1000 mg/kg 群で雌雄の生後 13 日の体重並びに生後 4~13 日の間の体重増加量に低値が認められ、児動物の発育への影響が考えられた。1000 mg/kg 群で雄出生児の甲状腺相対重量の高値が認められたが、前述の体重増加抑制に関連した変化の可能性が考えられるものの、甲状腺に肉眼的異常はみられず、血清中ホルモン(T4)にも異常は認められないことから、毒性学的意義のない変化と判断した。その他には 1000 mg/kg

群においても、出生児のいずれの検査項目にも被験物質投与による影響は認められないことから、児の子宮内発生及び分化並びに生存性などへの影響はないと判断した。

これらの結果から、本試験条件下でドデシルオキシランを反復経口投与することにより、被験物質に起因すると考えられる主な変化として、投与終了週に 1000 mg/kg 群の雄で尿浸透圧の高値、投与終了時に 1000 mg/kg 群の雄で BUN、ALT 及び肝臓重量の高値、非交配群雌で ALT 及び肝臓重量の高値、交配群雌の 300 及び 1000 mg/kg 群で肝臓重量の高値、回復終了週に 1000 mg/kg 群の雄で尿浸透圧の高値が認められた。したがって、本被験物質の反復投与毒性に対する無影響量及び無毒性量は、雄でいずれも 300 mg/kg、雌でいずれも 100 mg/kg と判断した。一方、生殖発生毒性に対する無影響量及び無毒性量は、雄親動物及び母動物のいずれも被験物質の影響が認められなかったが、1000 mg/kg 群で児動物に体重増加抑制が認められたことから、いずれも親動物では 1000 mg/kg、児動物では 300 mg/kg と判断した。

10. 参考文献

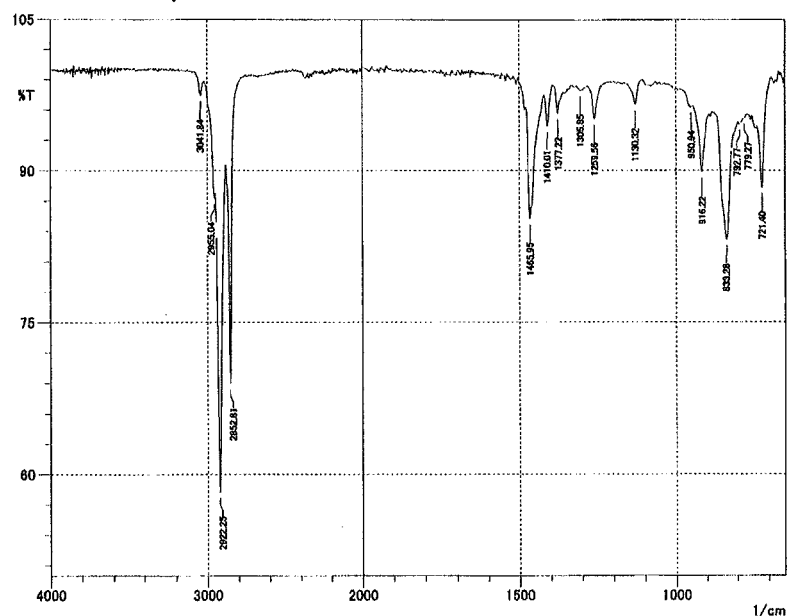
- 1) 鷹田信朗：ドデシルオキシラン：特性及び安定性試験並びに GC 法を用いた被験液中濃度測定法バリデーション及び安定性試験（媒体：コーン油）（株式会社ボゾリサーチセンター、試験番号：A-3221、2020 年）
- 2) 鷹野正生：ドデシルオキシラン：ラットを用いた 14 日間反復経口投与毒性試験（投与量設定試験）（株式会社ボゾリサーチセンター、試験番号：N-R096、2020 年）
- 3) Snedecor GW, Cochran WG. Statistical methods. 8th ed. Ames: Iowa State University Press; 1989.
- 4) Dunnett, CW. New tables for multiple comparisons with a control. Biometrics 20: 482-91. 1964.
- 5) Steel RGD. A multiple comparison rank sum test: Treatments versus control. Biometrics 15: 560-72. 1959.
- 6) Siegel S, Castellan NJ. Jr. Nonparametric statistics for the behavioral sciences. 2nd ed. New York: McGraw-Hill; 1988.
- 7) 佐久間昭（1981）：薬効評価－計画と分析－II, pp.23-27, 387-389, 東京大学出版会, 東京.

試験成績書
ドデシルオキシランの特性（投与期間開始前）

試験番号 : R-1261
被験物質 : ドデシルオキシラン
ロット番号 : XXXXXXXXXX
保存条件 : 密栓、冷暗所、防湿、窒素置換
試験施設 : 株式会社ボゾリサーチセンター 御殿場研究所
測定日 : 2020 年 10 月 16 日

測定項目 : 赤外吸収スペクトル測定法（ATR 法）

結果 :



遵守 GLP : 「新規化学物質等に係る試験を実施する試験施設に関する基準」（平成 23 年 3 月 31 日：薬食発 0331 第 8 号、平成 23・03・29 製局第 6 号、環保企発第 110331010 号）

XXXXXXXXXX
化学分析責任者
株式会社ボゾリサーチセンター 御殿場研究所

2020 年 10 月 23 日

試験成績書
ドデシルオキシランの安定性（投与期間終了後）

試験番号 : R-1261
被験物質 : ドデシルオキシラン
ロット番号 : 
保存条件 : 密栓、冷暗所、防湿、窒素置換
試験施設 : 株式会社ボゾリサーチセンター 御殿場研究所
測定日 : 2021 年 1 月 12 日

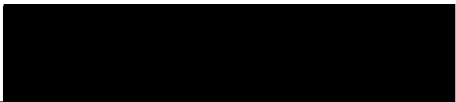
測定項目 : 赤外吸収スペクトル測定法（ATR 法）

判定基準 : 投与期間終了後のスペクトルを投与期間開始前のスペクトルと比較し、同等（主な吸収について、同一の波数域に同様の強度の吸収を認める）であること。

結果 : 投与期間開始前のスペクトルと同等であった。
赤外吸収スペクトルは次ページに示す。

判定 : 適

遵守 GLP : 「新規化学物質等に係る試験を実施する試験施設に関する基準」（平成 23 年 3 月 31 日：薬食発 0331 第 8 号、平成 23・03・29 製局第 6 号、環保企発第 110331010 号）

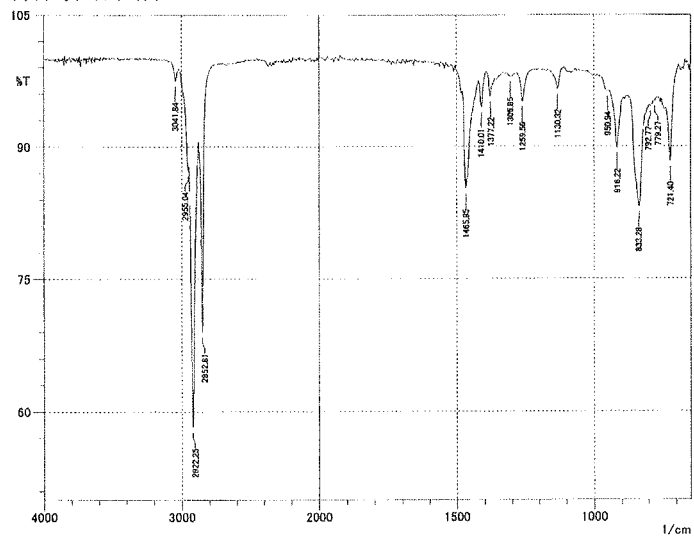

化学分析責任者

株式会社ボゾリサーチセンター 御殿場研究所

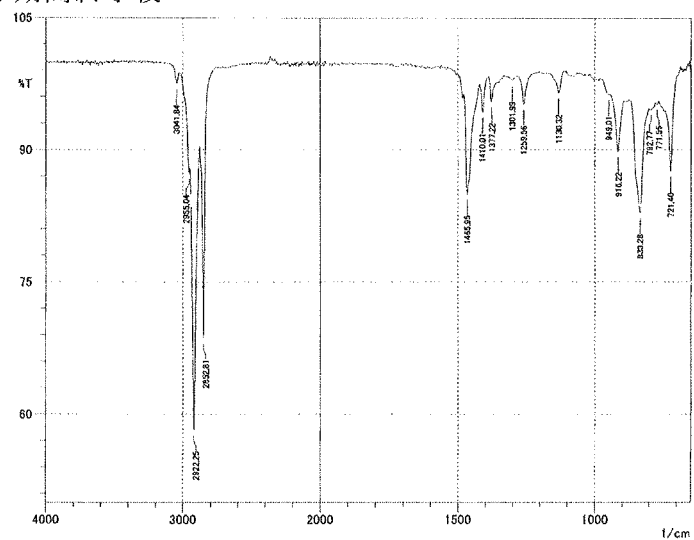
2021 年 1 月 15 日

試験成績書
ドデシルオキシランの安定性（投与期間終了後）

投与期間開始前



投与期間終了後



CERTIFICATE OF ANALYSIS
(Stability of Dodecyloxirane in Dose Formulations)

Study number: A-3221
Test article: Dodecyloxirane (Lot No. 
Vehicle: Corn oil
Form: Solution

Analyte: Dodecyloxirane

Storage conditions: Put into a brown glass bottle, stored under refrigeration (acceptable range: 1°C to 10°C) and then at room temperature (acceptable range: 1°C to 30°C)

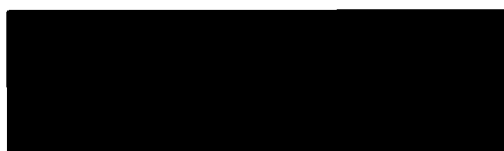
Storage period: Stored at room temperature for 24 hours after storage under refrigeration for 8 days

Acceptance criteria
Stability: The residual rate [the proportion of the mean concentration after storage to initial (100)] should be within 100.0% ± 10.0%.

Results: The results are described on the next page.
Judgment: Passed

This study was conducted in compliance with the following GLP regulations:

“The Good Laboratory Practice for New Chemicals”, (YakuShokuHatsu 0331 No. 8, Heisei 23-03-29 SeiKyoku No. 6, KanHoKiHatsu No. 110331010, the Ministry of Health, Labour and Welfare, the Ministry of Economy, Trade and Industry, and the Ministry of Environment, Japan, March 31, 2011)



Study director
Gotemba Laboratory, BoZo Research Center Inc.

October 19, 2020
Date

CERTIFICATE OF ANALYSIS
(Stability of Dodecyloxirane in Dose Formulations)

Study number: A-3221

Results:

Concentration (mg/mL)	No.	Measured concentration (mg/mL)	
		Initial	After storage
1	1	0.997	0.995
	2	0.978	0.997
	3	0.976	1.00
	Mean	0.984	0.997
	Residual rate (%)	(100)	101.3
200	1	210	203
	2	202	203
	3	203	203
	Mean	205	203
	Residual rate (%)	(100)	99.0

分析結果

試験番号 : R-1261
測定項目 : 濃度
ステージ : 初回投与

測定日 : 2020 年 10 月 22 日

測定試料

被験物質 : ドデシルオキシラン (ロット番号 XXXXXXXXXX)
媒体 : コーン油
形態 : 溶液
調製日 : 2020 年 10 月 22 日

測定対象物質 : ドデシルオキシラン

評価基準

濃度許容範囲 : 表示値に対する割合 ; $100\% \pm 10\%$

結果 :

表示値 (mg/mL)	測定濃度 (mg/mL)	平均値	表示値に対する割合 (%)
20	19.3	19.4	97.0
	19.6		
	19.4		
60	60.1	60.2	100.3
	60.2		
	60.4		
200	202	204	102.0
	205		
	204		

判定 : 適

分析結果

試験番号 : R-1261
測定項目 : 濃度
ステージ : 雄の最終投与

測定日 : 2020 年 12 月 1 日

測定試料

被験物質 : ドデシルオキシラン (ロット番号 XXXXXXXXXX)
媒体 : コーン油
形態 : 溶液
調製日 : 2020 年 12 月 1 日

測定対象物質 : ドデシルオキシラン

評価基準

濃度許容範囲 : 表示値に対する割合 ; 100% ± 10%

結果 :

表示値 (mg/mL)	測定濃度 (mg/mL)	平均値	表示値に対する割合 (%)
20	20.5	20.3	101.5
	20.2		
	20.3		
60	62.5	61.8	103.0
	61.7		
	61.2		
200	206	205	102.5
	204		
	206		

判定 : 適

R-1261
Attachment 5 (1/16)

OT-200011

ホルモン測定報告書

「ドデシルオキシラン：ラットを用いた経口投与による反復投与毒性・生殖発生毒性
併合試験（試験番号 R-1261）」における
ホルモン測定

OT-200011

株式会社ボゾリサーチセンター
〒151-0065 東京都渋谷区大山町 36-7

OT-200011

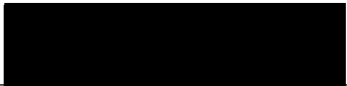
1. GLP 陳述書

ホルモン測定番号 : OT-200011

ホルモン測定表題 : 「ドデシルオキシラン：ラットを用いた経口投与による
反復投与毒性・生殖発生毒性併合試験(試験番号 R-1261)」
におけるホルモン測定

本作業は以下の GLP 基準を遵守して実施したものです。

- 「新規化学物質等に係る試験を実施する試験施設に関する基準」
(平成 23 年 3 月 31 日：薬食発 0331 第 8 号、平成 23・03・29 製局第 6 号、環
保企発第 110331010 号)

 2021 年 3 月 18 日

試験主任者
株式会社ボゾリサーチセンター つくば研究所

2. 目次

1.	GLP 陳述書	2
2.	目次	3
3.	ホルモン測定実施概要	5
3.1	ホルモン測定番号	5
3.2	ホルモン測定表題	5
3.3	目的	5
3.4	遵守した基準	5
3.4.1	GLP	5
3.5	試験施設	5
3.6	試験場所	5
3.7	試験責任者	5
3.8	試験主任者	5
3.9	担当者	6
3.10	日程	6
3.11	試験成績の信頼性に影響を及ぼしたと思われる環境要因	6
3.12	資料の保存	6
3.13	試験主任者の署名	7
4.	要約	8
5.	試験構成	9
5.1	概要	9
5.2	試料	10
6.	材料及び方法	10
6.1	アッセイ方法	11
6.1.1	T4 測定	11
6.1.1.1	Cal-1/2 及び Cal-1/4 の調製	11
6.1.1.2	測定操作	11
6.2	データ解析	12
6.3	測定成立条件の評価	12
6.4	再測定	12
7.	統計解析	13
8.	結果及び考察	14
8.1	EOA（交配群投与終了時雄動物）	14
8.2	PD13（生後 13 日児動物）	14
9.	結論	14

R-1261
Attachment 5 (4/16)

OT-200011

Tables

Table 1	Hormone concentrations in rat serum of EOA (Male)	15
Table 2	Hormone concentrations in rat serum of PD13 (Pups).....	15

信頼性保証書	16
--------------	----

OT-200011

3. ホルモン測定実施概要

3.1 ホルモン測定番号

OT-200011

3.2 ホルモン測定表題

「ドデシルオキシラン：ラットを用いた経口投与による反復投与毒性・生殖発生毒性併合試験（試験番号 R-1261）」におけるホルモン測定

3.3 目的

「ドデシルオキシラン：ラットを用いた経口投与による反復投与毒性・生殖発生毒性併合試験（試験番号 R-1261）」において採取された血清中のホルモン濃度を測定することを目的とした。

3.4 遵守した基準

3.4.1 GLP

- 「新規化学物質等に係る試験を実施する試験施設に関する基準」
（平成 23 年 3 月 31 日：薬食発 0331 第 8 号、平成 23・03・29 製局第 6 号、環
保企発第 110331010 号）

3.5 試験施設

株式会社ボゾリサーチセンター 御殿場研究所
〒412-0039 静岡県御殿場市かまど 1284

3.6 試験場所

株式会社ボゾリサーチセンター つくば研究所
〒300-2611 茨城県つくば市大久保 8 番

3.7 試験責任者

株式会社ボゾリサーチセンター 御殿場研究所 研究部
[REDACTED]

3.8 試験主任者

株式会社ボゾリサーチセンター つくば研究所 臨床検査部門
[REDACTED]

OT-200011

3.9 担当者

測定担当者 : 

3.10 日程

試料受領日	2020 年 12 月 10 日、2020 年 12 月 25 日
測定期間	2020 年 12 月 17 日 ~ 2020 年 12 月 28 日
ホルモン測定報告書作成日	2021 年 3 月 18 日

3.11 試験成績の信頼性に影響を及ぼしたと思われる環境要因

該当事象はなかった。

3.12 資料の保存

以下のホルモン測定関係資料をファイルに綴じ込み、株式会社ボゾリサーチセンターつくば研究所の資料保存施設に保存する。その期間はホルモン測定報告書作成後 5 年間とする。期間終了後の取扱いについては、試験施設と株式会社ボゾリサーチセンターつくば研究所の間で協議し、その処置を決定する。

- 1) ホルモン測定計画書の原本
- 2) 試験計画書の写し
- 3) 試験計画書変更書の写し
- 4) 生データ及びその他の記録文書
- 5) ホルモン測定報告書の原本
- 6) 通信記録

R-1261
Attachment 5 (7/16)

OT-200011

3.13 試験主任者の署名

 2021 年 3 月 18 日

株式会社ボゾリサーチセンター つくば研究所

OT-200011

4. 要約

「ドデシルオキシラン：ラットを用いた経口投与による反復投与毒性・生殖発生毒性併合試験（試験番号 R-1261）」において採取された EOA（交配群投与終了時雄動物）及び PD13（生後 13 日児動物）の血清中のホルモン（T4）濃度を測定した。

その結果、EOA では T4 濃度は媒体対照群と全ての投与群との間に有意差は認められなかった。一方、PD13 では低用量群で有意な低下が認められた。

OT-200011

5. 試験構成

5.1 概要

試験番号 R-1261 (ドデシルオキシラン: ラットを用いた経口投与による反復投与毒性・生殖発生毒性併合試験) の群構成を下表に示す。

群構成表 (雄)

試験群	投与量 (mg/kg)	交配群			
		投与終了時剖検群		回復終了時剖検群	
		動物数	動物番号	動物数	動物番号
媒体対照	0	7	1001-1007	5	1008-1012
低用量	100	12	2001-2012		
中用量	300	12	3001-3012		
高用量	1000	7	4001-4007	5	4008-4012

群構成表 (雌)

試験群	投与量 (mg/kg)	交配群		非交配群			
		投与終了時剖検群		投与終了時剖検群		回復終了時剖検群	
		動物数	動物番号	動物数	動物番号	動物数	動物番号
媒体対照	0	12	1101-1112	5	5101-5105	5	5106-5110
低用量	100	12	2101-2112				
中用量	300	12	3101-3112				
高用量	1000	12	4101-4112	5	6101-6105	5	6106-6110

OT-200011

5.2 試料

Text-Table 1

ホルモン 測定試料 (血清)	試験群	動物数	血清試料数			
			交配群		非交配群	
			投与終了 時剖検群	回復終了 時剖検群	投与終了 時剖検群	回復終了 時剖検群
測定対象	媒体対照	雄 10	5	5 ^{a)}	—	—
		雌 15	5 ^{a)}	—	5 ^{a)}	5 ^{a)}
	低用量	雄 5	5	—	—	—
		雌 5	5 ^{a)}	—	—	—
	中用量	雄 5	5	—	—	—
		雌 5	5 ^{a)}	—	—	—
	高用量	雄 10	5	5 ^{a)}	—	—
		雌 15	5 ^{a)}	—	5 ^{a)}	5 ^{a)}
	上記に加え、生後 4 日及び 13 日の児動物（交配群の各用量群の母動物から 5 腹ずつ、計 40 腹分）。					
	a)：本動物及び生後 4 日児動物は、試験責任者の判断により測定しなかった。					
受領チューブ数	雌雄：3 本、児（生後 4 日）：1 本、児（生後 13 日）：3 本、合計 290 本					
試験施設での 保存条件	-80°C の冷凍庫（許容範囲：-70°C 以下）、実測値は許容範囲内であった。					
輸送条件	ドライアイス同梱下で輸送された。					
識別	ラベルに明記された情報（試験番号、採血時期、検査項目、試料番号、試料名、色分け及び採取年月日）をもとに、試験場所で血清試料名を割り付けた。					
試験場所での 保存条件	-80°C の冷凍庫（許容範囲：-70°C 以下）、実測値は許容範囲内であった。					
血清試料安定性 (-70°C 以下)	保存安定性：16 週間、繰返し凍結融解回数：3 回まで。 測定は上記の条件内で実施した。					
残余の取扱い	測定後の残余の血清は、ホルモン測定報告書作成後に試験責任者の指示に基づいて試験場所で廃棄する。					

6. 材料及び方法

受領した試料（Text-Table 1）を用いて下記 6.1 に従い、ホルモン（T4）濃度を測定した。

測定方法及び測定キットに関する情報を下表（Text-Table 2、Text-Table 3）に示す。

OT-200011

Text-Table 2

ホルモン測定				
検査項目	測定方法	単位	表示桁数	測定範囲
T4	RIA 法	ng/mL	小数点第 1 位	5.7 ~ 340.0 ng/mL
使用測定機器 ガンマカウンター (AccuFLEX γ 7010、日立)				

Text-Table 3

T4	
キット	: RIA-gnost®-T4 (Cisbio Bioassays)
保存条件	: 2~8°C (実測値は許容範囲内であった。)
Lot No. (使用期限)	: 273U (1/26/2021)
Calibrator (表示濃度)	: Cal-0、1 (22.6 ng/mL)、2 (49.0 ng/mL)、3 (100.0 ng/mL)、 4 (196.0 ng/mL)、5 (340.0 ng/mL)

6.1 アッセイ方法

6.1.1 T4 測定

6.1.1.1 Cal-1/2 及び Cal-1/4 の調製

Cal-1 と Cal-0 を 1 : 1 (60 μ L : 60 μ L) で混合し、Cal-1/2 (表示濃度 : 11.3 ng/mL) とした。さらに、Cal-1/2 と Cal-0 を 1 : 1 (30 μ L : 30 μ L) で混合し、Cal-1/4 (表示濃度 : 5.7 ng/mL) とした。調製当日を使用期限とした。

6.1.1.2 測定操作

- 1) 各試料 20 μ L を Ab チューブ (Antibody-coated tube) に加えた (TC を除く)。
- 2) 125 I-T4 tracer (1 mL) をチューブに添加し、ふたをして攪拌した。
- 3) TC については、tracer をプラスチックチューブ (丸底テストチューブ、WATSON) に加えた。
- 4) 23°C 設定のインキュベータ内で 45~55 分間 (実時間 : 50 分間) 振とうした。
- 5) チューブ内の溶液をアスピレータで除去した (TC を除く)。
- 6) ガンマカウンターでチューブの 125 I (60 秒間) の放射線計数率 (cpm) を測定した。

OT-200011

6.2 データ解析

Magellan Tracker ver. 7.0 ソフトウェア (Tecan) を用いて、放射線計数率 (cpm) から各試料の T4 濃度を算出した。

X 軸を濃度 (ng/mL、対数表示)、Y 軸を放射線計数率 (cpm) の測定値とし、表示濃度のプロットから検量線を作成した。

解析法、表示桁数を下表に示した。

検査項目	解析法	表示桁数	
		放射線計数率 (cpm)	測定濃度
T4	4 パラメーター Marquardt (重み付け: 1/Y ²)	小数点第 1 位	小数点第 1 位

6.3 測定成立条件の評価

本測定は、以下の測定成立条件を全て満たしていたことから測定成立とした。計算は Microsoft office Excel 2010 (Ver.14.0) で実施 (表示桁数で計算) した。

1) TC 及び Cal-0

- TC 及び Cal-0 の平均放射線計数率が Cal-1/4 の平均放射線計数率を超えている。
- 平均放射線計数率は以下の計算式から算出した (小数点第 1 位まで表示)。

$$\text{平均放射線計数率 (cpm)} = \frac{\text{個別放射線計数率 (cpm) の和}}{2}$$

2) 標準試料

- duplicate での個別放射線計数率の差が 2 倍未満である。
- 測定濃度 (個別濃度の平均) の表示濃度に対する真度が Cal-1/4、Cal-5 で 100±25.0%以内、Cal-1/2~4 のうち 4 試料以上で 100±20.0%以内である。
- 真度 (%) は以下の計算式から算出した (小数点第 1 位まで表示)。

$$\text{真度 (\%)} = \frac{\text{測定濃度}}{\text{表示濃度}} \times 100$$

3) QC

duplicate での個別放射線計数率の差が 2 倍未満で、かつ測定濃度 (個別濃度の平均) が表示濃度範囲内である。

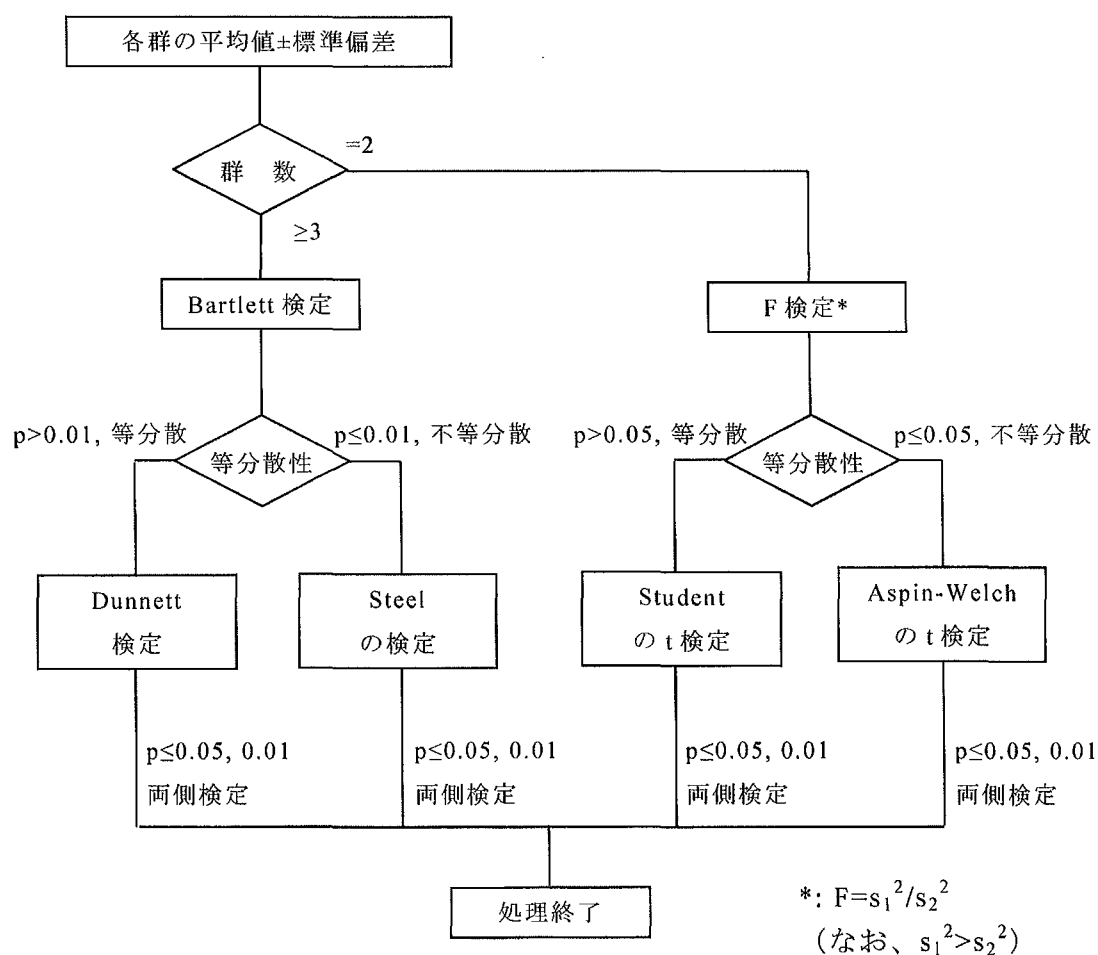
6.4 再測定

再測定は実施しなかった。

OT-200011

7. 統計解析

検査項目について下記の模式図に示す方法に従って検定した。各群の平均値 (Mean) 及び標準偏差 (SD) の算出には、それぞれ Microsoft Office Excel 2010 (Ver.14) SP2 における AVERAGE 関数及び STDEV 関数を用いた。解析には SAS Release 9.1.3 を使用した。



OT-200011

8. 結果及び考察

8.1 EOA（交配群投与終了時雄動物）

T4 濃度測定の結果を Table 1 に示した。

T4 濃度は媒体対照群と全ての投与群との間に有意差は認められなかった。

8.2 PD13（生後 13 日児動物）

T4 濃度測定の結果を Table 2 に示した。

T4 濃度は低用量群において対照群との間に有意な低下が認められたが、中用量及び高用量群では有意差は認められなかった。

9. 結論

「ドデシルオキシラン：ラットを用いた経口投与による反復投与毒性・生殖発生毒性併合試験（試験番号 R-1261）」において採取された EOA（交配群投与終了時雄動物）及び PD13（生後 13 日児動物）の血清中のホルモン（T4）濃度を測定した。

その結果、EOA では T4 濃度は媒体対照群と全ての投与群との間に有意差は認められなかった。一方、PD13 では低用量群で有意な低下が認められた。

OT-200011

Table 1 Hormone concentrations in rat serum of EOA (Male)

Animal No.	T4 (ng/mL)	Animal No.	T4 (ng/mL)	Animal No.	T4 (ng/mL)	Animal No.	T4 (ng/mL)
1001	31.6	2001	24.3	3001	30.9	4001	27.1
1002	26.1	2002	31.0	3002	27.4	4002	25.8
1003	27.2	2003	23.0	3003	26.0	4003	38.4
1004	30.4	2004	34.5	3004	37.0	4004	21.5
1005	28.4	2005	20.9	3005	33.3	4005	20.5
Mean	28.7	Mean	26.7	Mean	30.9	Mean	26.7
SD	2.3	SD	5.8	SD	4.5	SD	7.1
n	5	n	5	n	5	n	5

EOA: End of administration

No significant difference in any treated groups from vehicle control group.

Table 2 Hormone concentrations in rat serum of PD13 (Pups)

Animal No.	T4 (ng/mL)	Animal No.	T4 (ng/mL)	Animal No.	T4 (ng/mL)	Animal No.	T4 (ng/mL)
1101	48.1	2104	38.9	3101	45.1	4102	39.6
1105	40.1	2105	36.9	3102	40.3	4103	41.0
1107	38.7	2107	34.4	3104	38.5	4107	33.5
1108	40.2	2108	31.8	3110	36.3	4109	40.2
1111	42.7	2112	35.6	3111	32.5	4111	36.9
Mean	42.0	Mean	35.5 *	Mean	38.5	Mean	38.2
SD	3.7	SD	2.7	SD	4.7	SD	3.1
n	5	n	5	n	5	n	5

PD13: Postnatal day 13

Animal No.: Dam No. of pups

*: $p \leq 0.05$, Dunnett's test

OT-200011

信頼性保証書

ホルモン測定番号 : OT-200011

ホルモン測定表題 : 「ドデシルオキシラン：ラットを用いた経口投与による反復投与毒性・生殖発生毒性併合試験（試験番号 R-1261）」におけるホルモン測定

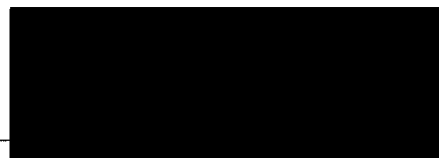
本測定は以下に示す基準に従って実施されたことを保証致します。

- 「新規化学物質等に係る試験を実施する試験施設に関する基準」
（平成 23 年 3 月 31 日：薬食発 0331 第 8 号、平成 23・03・29 製局第 6 号、環
保企発第 110331010 号）

なお、調査は下記の通り実施し、報告致しました。

調査項目	調査担当者	調査日	試験主任者 及び試験場所 管理責任者 への報告日	試験責任者、 運営管理者及び 主信頼性保証 部門責任者 への報告日
ホルモン測定 計画書		2020 年 10 月 20 日 2020 年 10 月 22 日	2020 年 10 月 22 日	2020 年 10 月 23 日
試料受領（雄動物）		2020 年 12 月 10 日	2020 年 12 月 10 日	2020 年 12 月 10 日
ホルモン測定（T4）		2020 年 12 月 17 日	2020 年 12 月 17 日	2020 年 12 月 17 日
試料受領（雌雄 及び児動物）		2020 年 12 月 25 日	2020 年 12 月 28 日	2020 年 12 月 28 日
ホルモン測定報告 書草案、生データ		2021 年 2 月 18 日 2021 年 2 月 19 日	2021 年 2 月 25 日	2021 年 2 月 25 日
改善確認		2021 年 3 月 15 日	2021 年 3 月 15 日	2021 年 3 月 15 日
ホルモン測定 報告書		2021 年 3 月 17 日	2021 年 3 月 18 日	2021 年 3 月 18 日

2021 年 3 月 18 日
株式会社ボゾリサーチセンター
つくば研究所
試験場所信頼性保証部門責任者



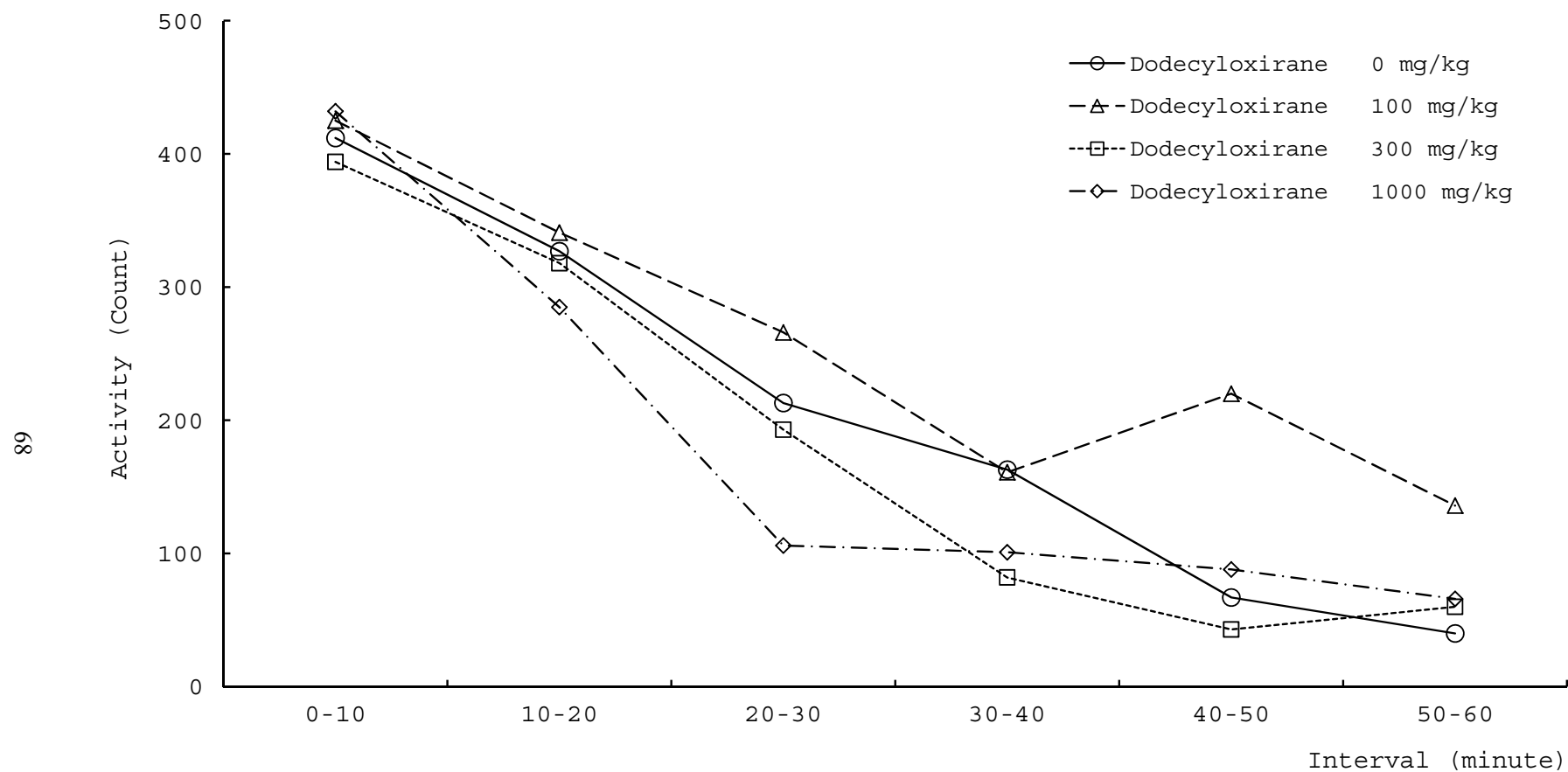


Fig.1 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Motor activity of male rats (Week 6 of administration)

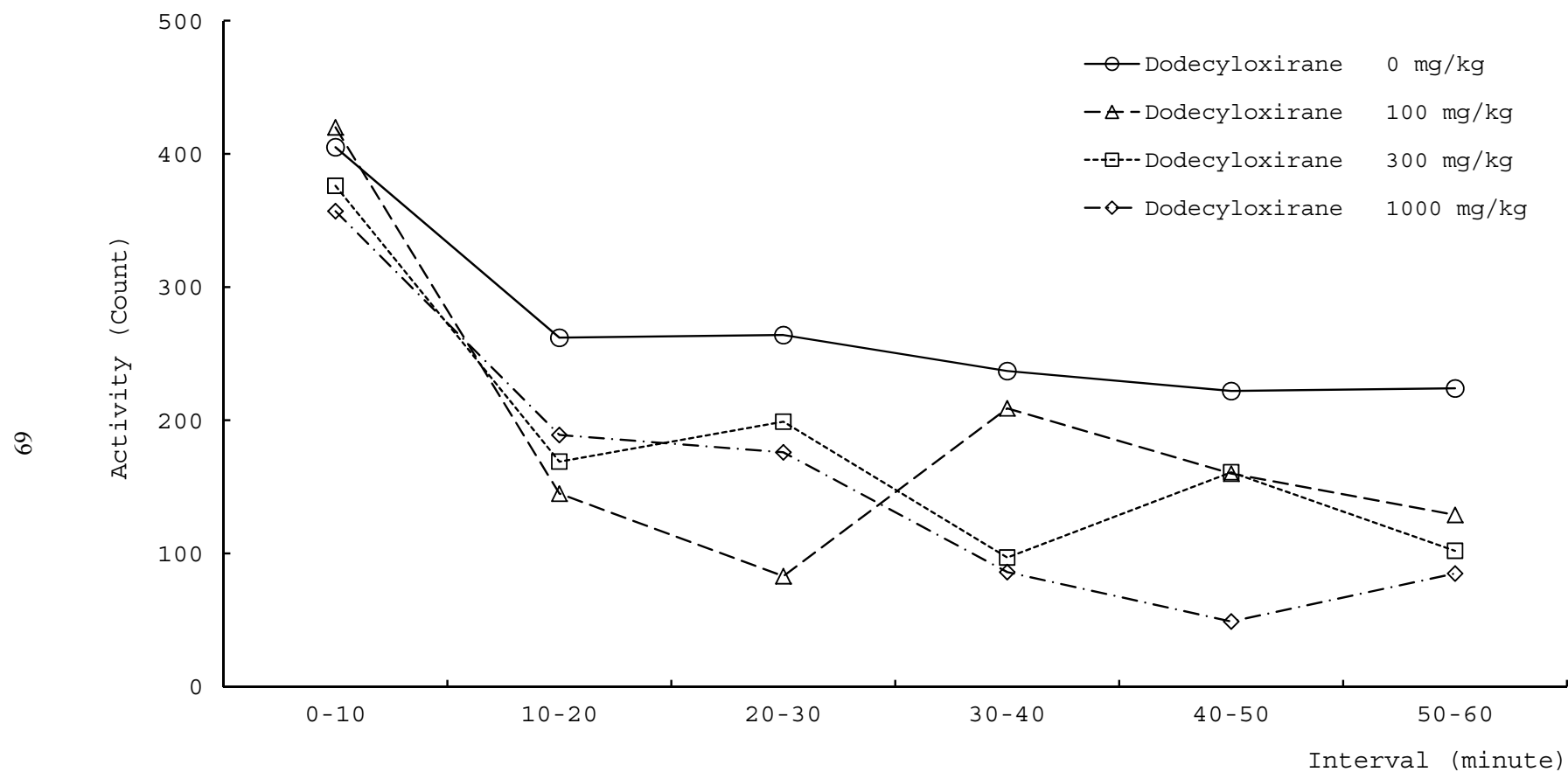


Fig.2 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Motor activity of female rats (Main group, Lactation Day13)

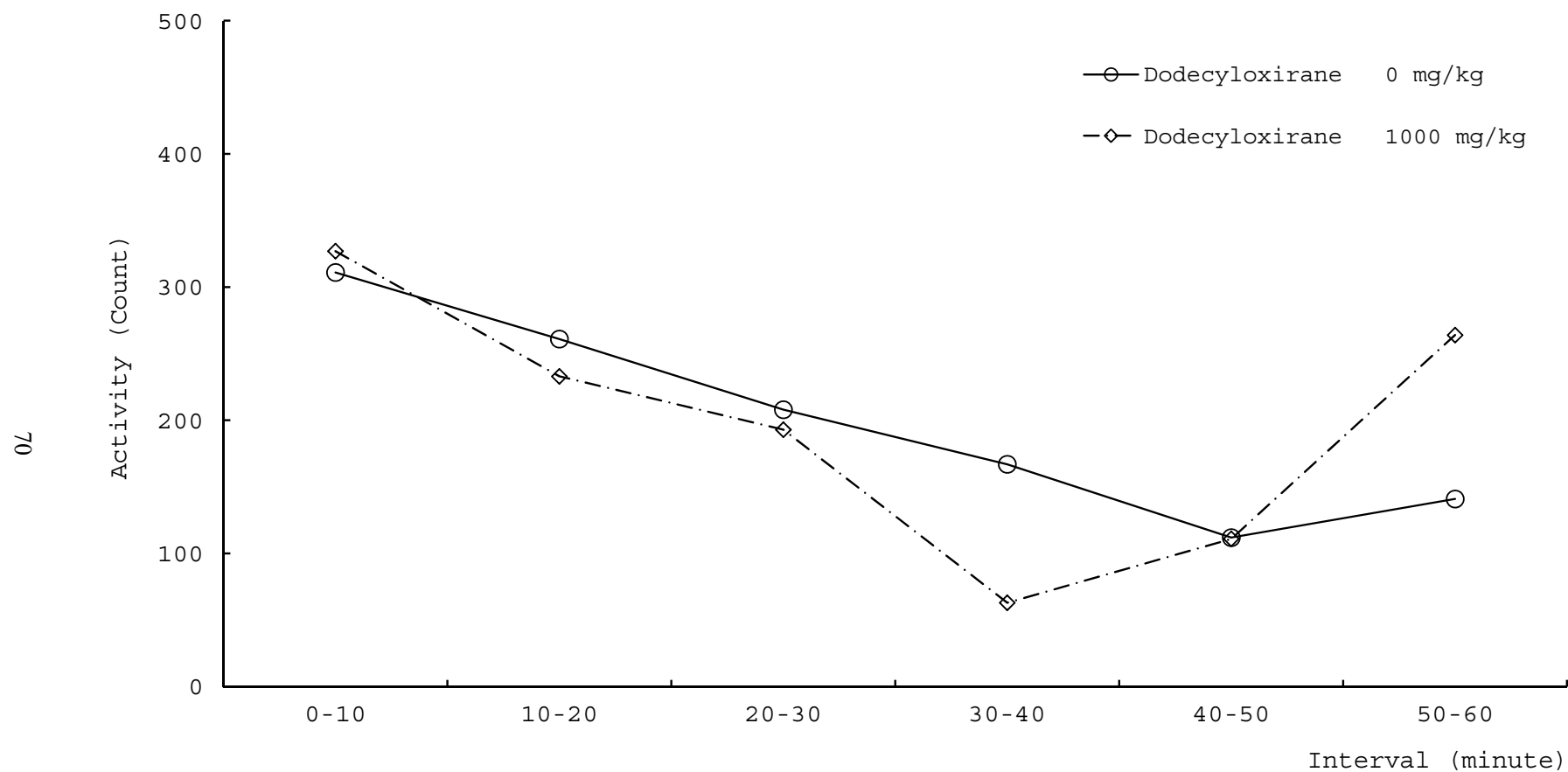


Fig.3 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Motor activity of female rats (Non-mating group, Week 6 of administration)

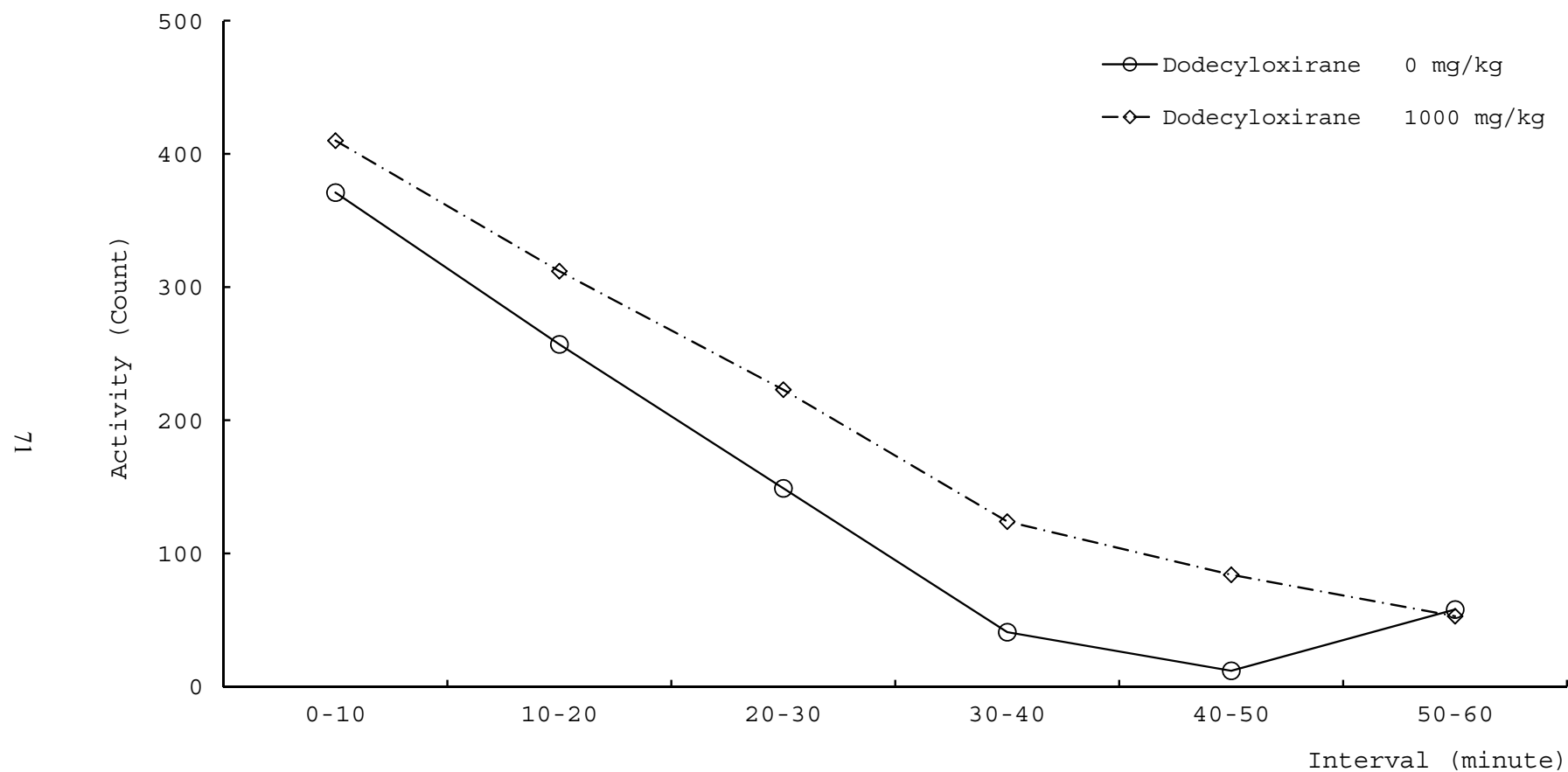


Fig.4 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Motor activity of male rats (Week 2 of recovery)

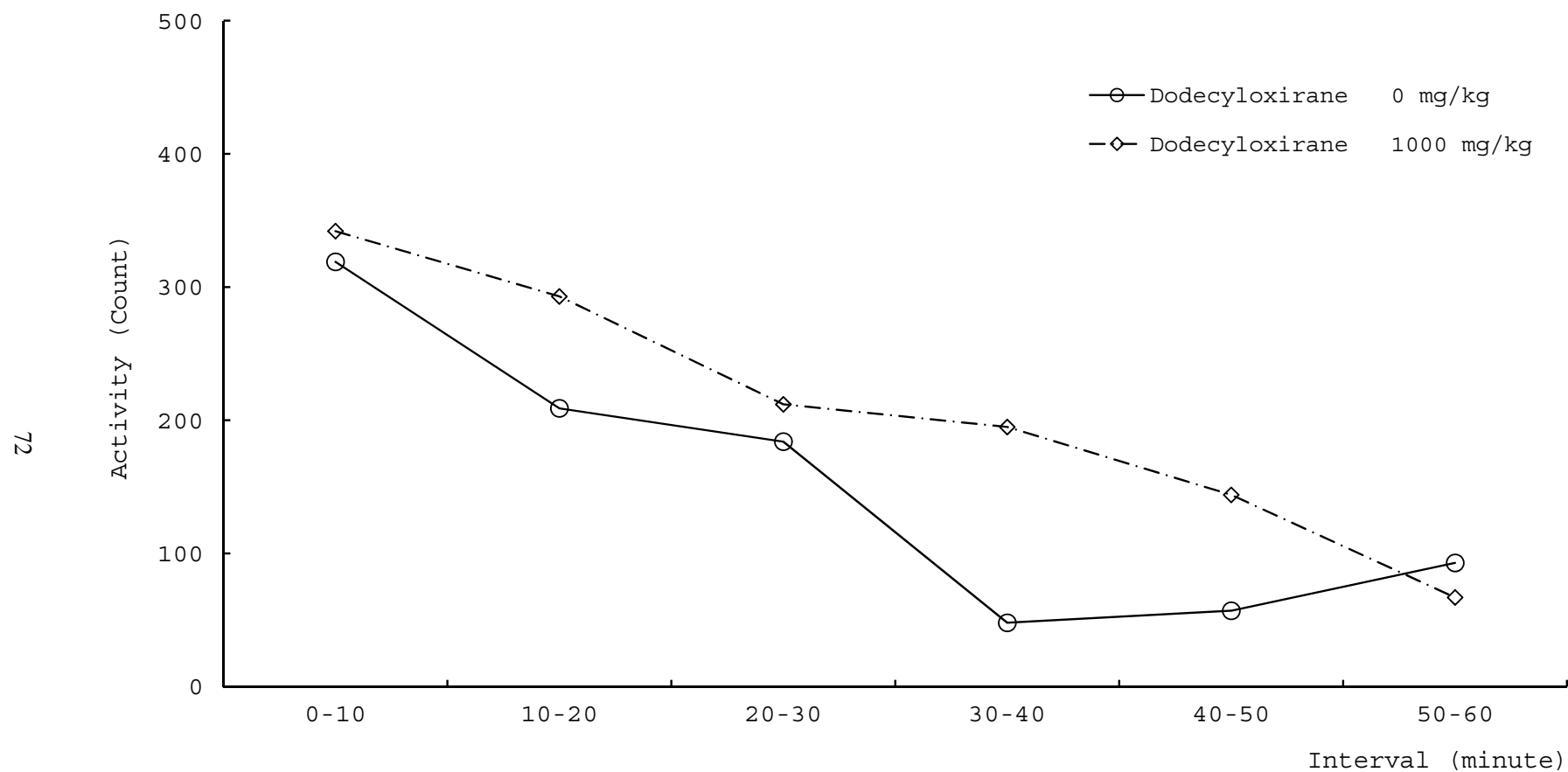


Fig.5 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Motor activity of female rats (Non-mating group, Week 2 of recovery)

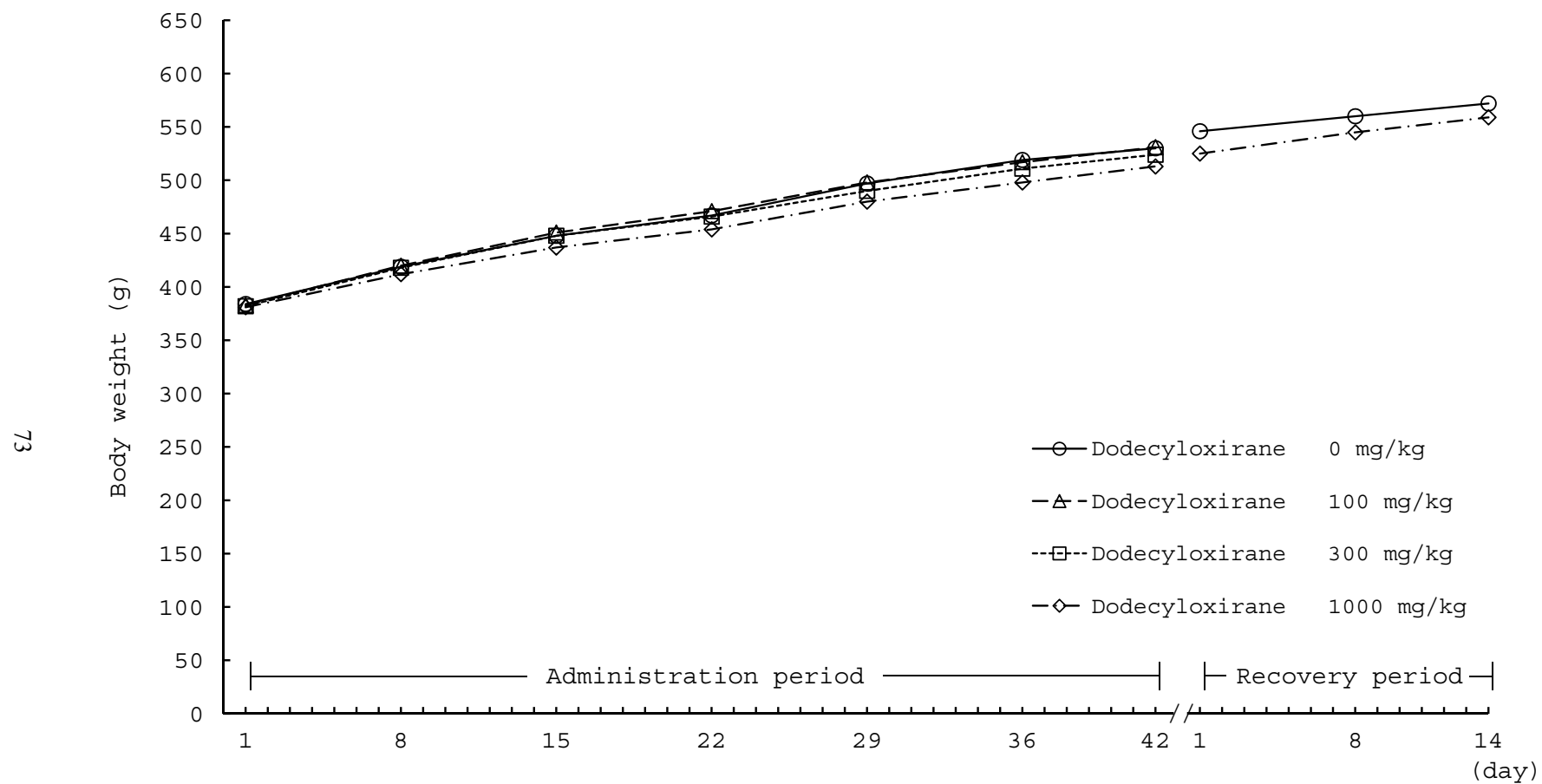


Fig.6 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Body weight of male rats

74

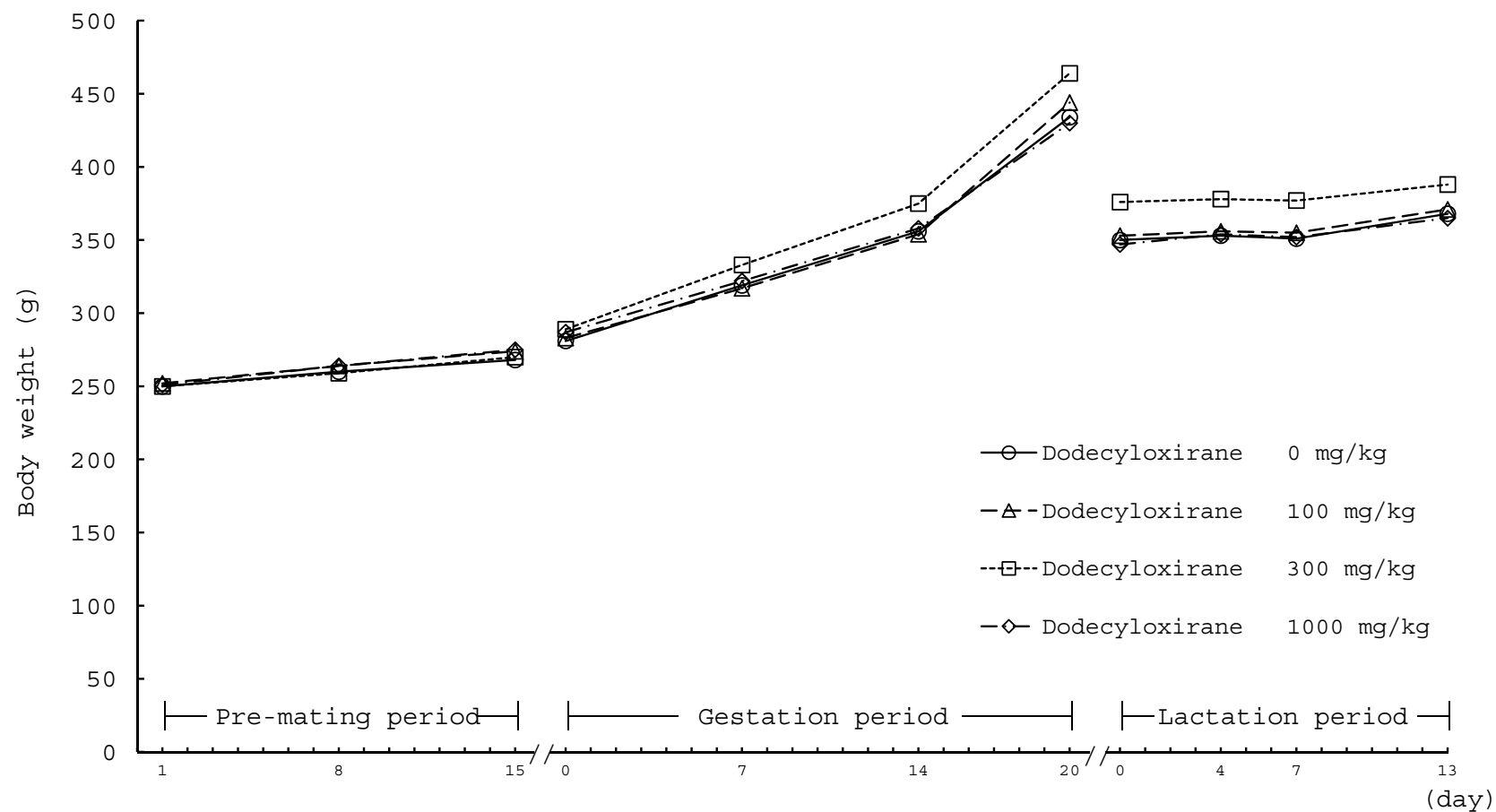


Fig.7 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Body weight of female rats (Mating group)

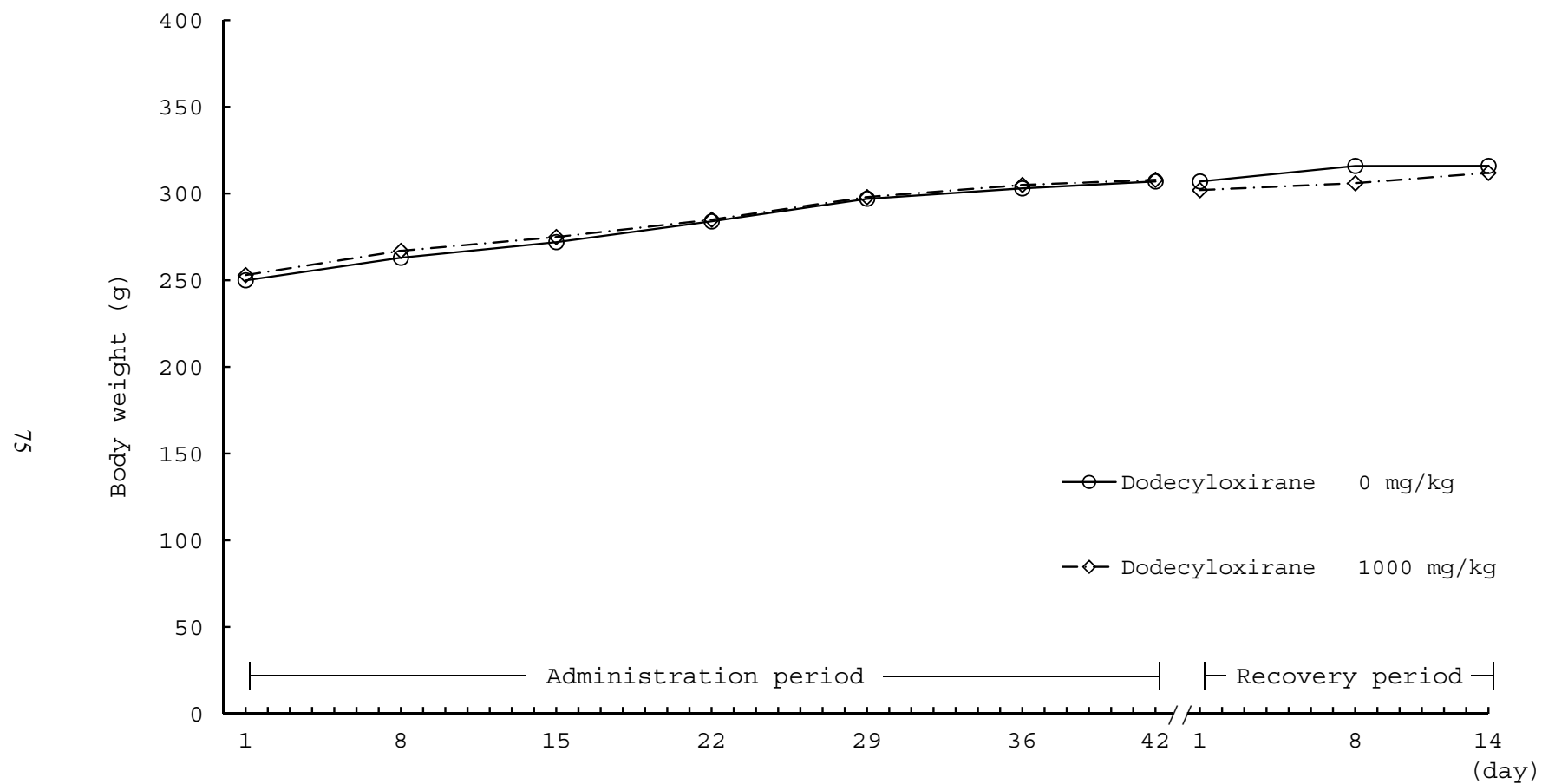


Fig.8 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Body weight of female rats (Non-mating group)

96

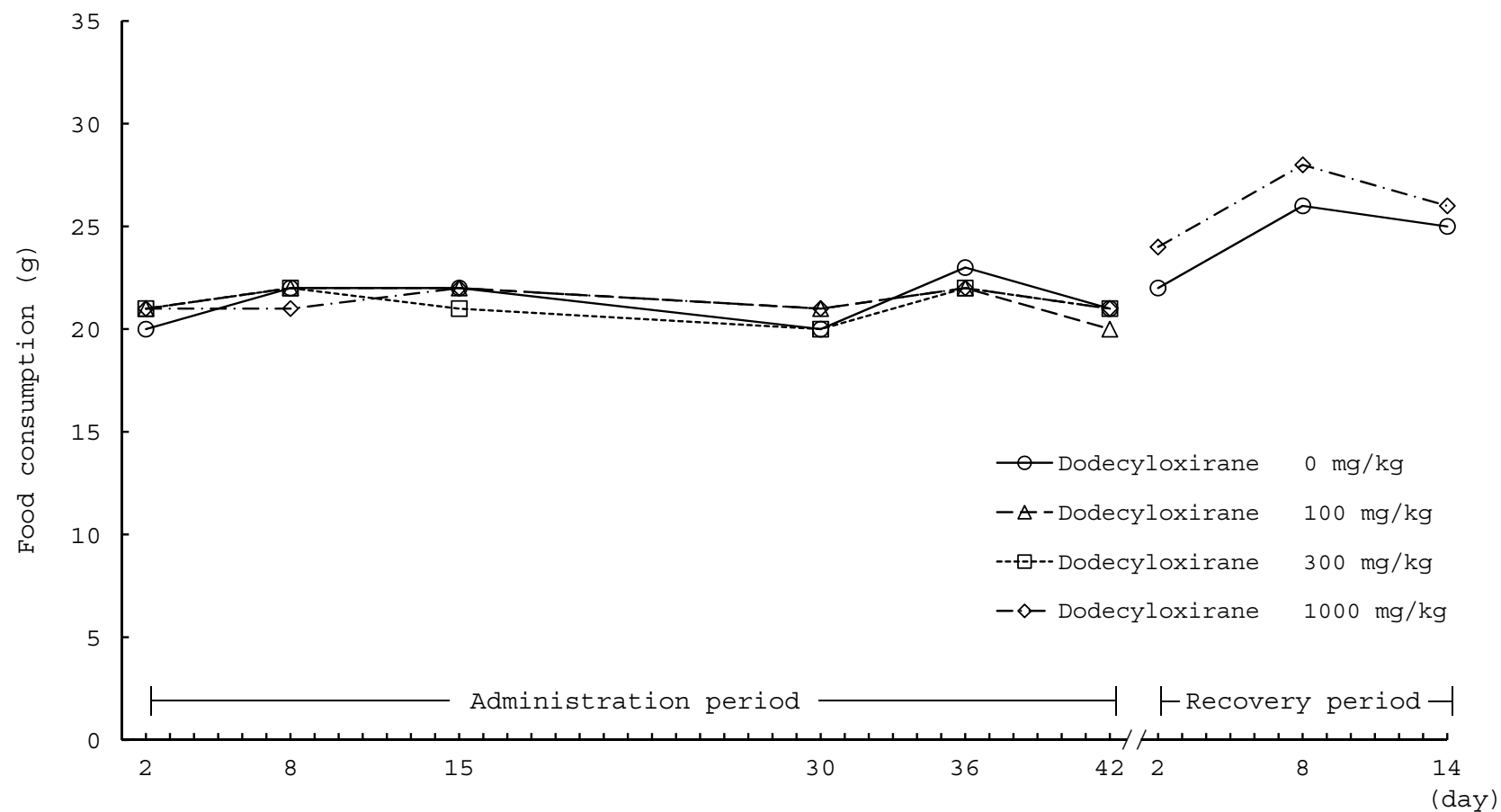


Fig.9 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Food consumption of male rats

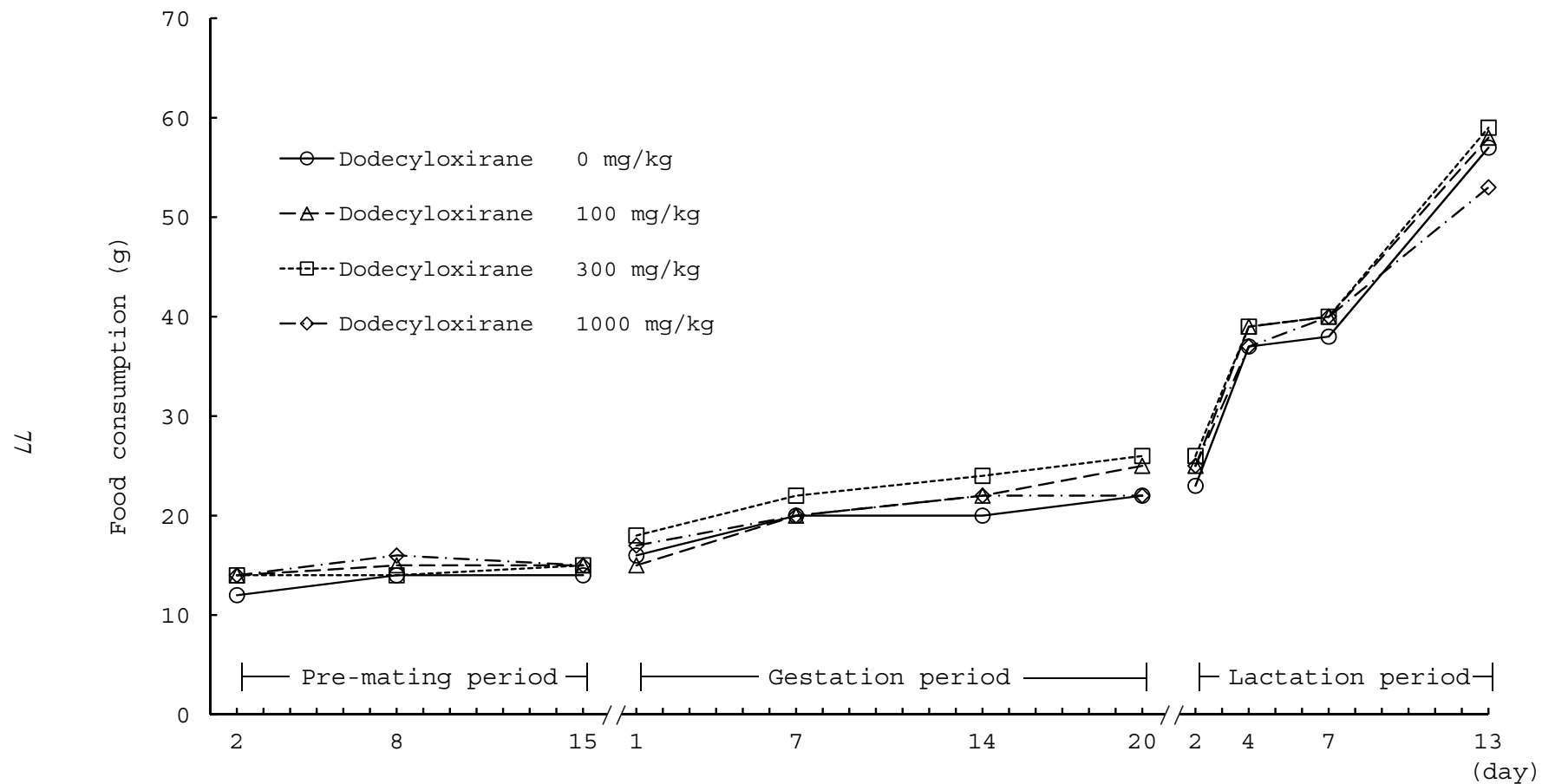


Fig.10 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Food consumption of female rats (Mating group)

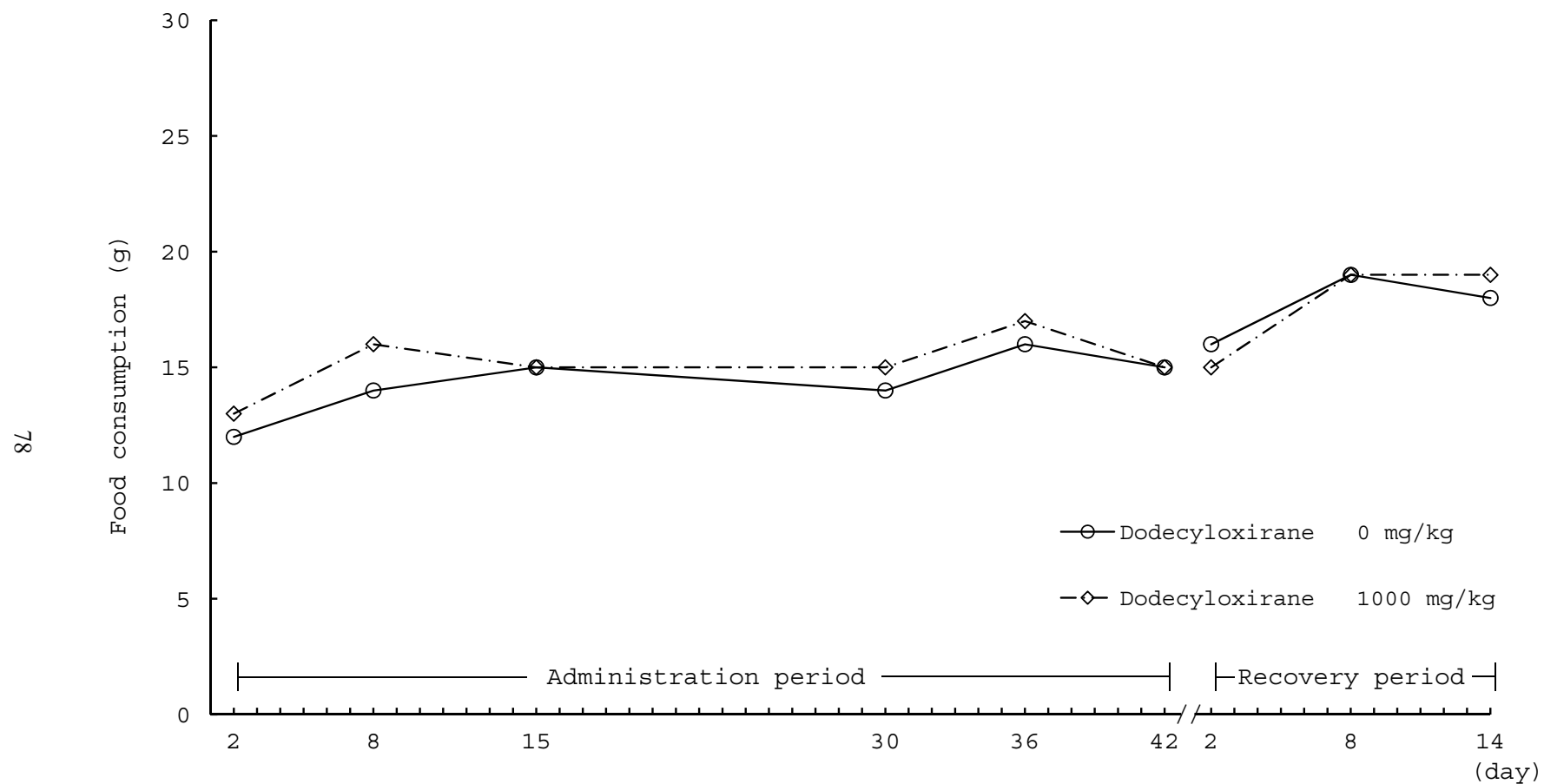


Fig.11 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Food consumption of female rats (Non-mating group)

Table 1 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Male

Study No. : R-1261

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Species : Rat

Test article Dose	Signs	/Day																				
		1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21
Dodecyloxirane 0 mg/kg	Number of animals	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 100 mg/kg	Number of animals	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 300 mg/kg	Number of animals	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	Number of animals	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

Table 1 - 2

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Male

Study No. : R-1261

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Species : Rat

Test article Dose	Signs	/Day																				
		22	23	24	25	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	42
Dodecyloxirane 0 mg/kg	Number of animals	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 100 mg/kg	Number of animals	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 300 mg/kg	Number of animals	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	Number of animals	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

Table 1 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Male

Study No. : R-1261

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Species : Rat

Test article Dose	Signs	/Day 43 a)
Dodecyloxirane 0 mg/kg	Number of animals Number of animals with abnormal findings	7 0
Dodecyloxirane 100 mg/kg	Number of animals Number of animals with abnormal findings	12 0
Dodecyloxirane 300 mg/kg	Number of animals Number of animals with abnormal findings	12 0
Dodecyloxirane 1000 mg/kg	Number of animals Number of animals with abnormal findings	7 0

a): Day of necropsy

Table 1 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Female

Study No. : R-1261

Period : F0 before mating Day 1-15

Species : Rat

Test article Dose	Signs	/Day														
		1	2	3	4	5	6	7	8	9	10	11	12	13	14	15
Dodecyloxirane 0 mg/kg	Number of dams	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 100 mg/kg	Number of dams	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 300 mg/kg	Number of dams	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	Number of dams	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

Table 1 - 5

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs in dams
Sex : Female

Study No. : R-1261

Period : F0 gestation Day 0-25

Species : Rat

Sex : Female		Species : Rat																				
Test article		/Day																				
Dose	Signs	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20
Dodecyloxirane 0 mg/kg	Number of dams	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 100 mg/kg	Number of dams	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 300 mg/kg	Number of dams	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	Number of dams	11	11	11	11	11	11	11	11	11	11	11	11	11	11	11	11	11	11	11	11	11
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

Table 1 - 6

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs in dams
Sex : Female

Study No. : R-1261

Period : F0 gestation Day 0-25

Species : Rat

Test article Dose	Signs	/Day				
		21	22	23	24	25
Dodecyloxirane 0 mg/kg	Number of dams	12	4	0	0	0
	Number of dams with abnormal findings	0	0	0	0	0
Dodecyloxirane 100 mg/kg	Number of dams	10	1	0	0	0
	Number of dams with abnormal findings	0	0	0	0	0
Dodecyloxirane 300 mg/kg	Number of dams	10	0	0	0	0
	Number of dams with abnormal findings	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	Number of dams	11	2	1	1	1
	Number of dams with abnormal findings	0	0	0	0	0

Table 1 - 7

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs in dams
Sex : Female

Study No. : R-1261

Period : F0 lactation Day 0-14

Species : Rat

Test article Dose	Signs	/Day														
		0	1	2	3	4	5	6	7	8	9	10	11	12	13	14 a)
Dodecyloxirane 0 mg/kg	Number of dams	7	12	12	12	12	12	12	12	12	12	12	12	12	12	12
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 100 mg/kg	Number of dams	8	10	10	10	10	10	10	10	10	10	10	10	10	10	10
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 300 mg/kg	Number of dams	7	10	10	10	10	10	10	10	10	10	10	10	10	10	10
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	Number of dams	9	10	10	10	10	10	10	10	10	10	10	10	10	10	10
	Number of dams with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

a): Day of necropsy

Table 1 - 8

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Female

Study No. : R-1261

Period : Administration day 1-43(non-mating group)

Species : Rat

Test article Dose	Signs	/Day																				
		1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21
Dodecyloxirane 0 mg/kg	Number of animals	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	Number of animals	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

Table 1 - 9

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Female

Study No. : R-1261

Period : Administration day 1-43(non-mating group)

Species : Rat

		Sex : Female																				Species : Rat	
Test article Dose	Signs	/Day																					
		22	23	24	25	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	42	
Dodecyloxirane 0 mg/kg	Number of animals	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
Dodecyloxirane 1000 mg/kg	Number of animals	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10	
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	

Table 1 - 10

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Female

Study No. : R-1261

Period : Administration day 1-43(non-mating group)

Species : Rat

Test article		/Day
Dose	Signs	43 a)
Dodecyloxirane	Number of animals	5
0 mg/kg	Number of animals with abnormal findings	0
Dodecyloxirane	Number of animals	5
1000 mg/kg	Number of animals with abnormal findings	0

a): Day of necropsy

Table 1 - 11

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Male

Study No. : R-1261

Period : Recovery Day 1-15

Species : Rat

Test article Dose	Signs	/Day														
		1	2	3	4	5	6	7	8	9	10	11	12	13	14	15 a)
Dodecyloxirane 0 mg/kg	Number of animals	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	Number of animals	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

a): Day of necropsy

Table 1 - 12

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Female

Study No. : R-1261

Period : Recovery Day 1-15 (non-mating group)

Species : Rat

Test article Dose	Signs	/Day														
		1	2	3	4	5	6	7	8	9	10	11	12	13	14	15 a)
Dodecyloxirane 0 mg/kg	Number of animals	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	Number of animals	5	5	5	5	5	5	5	5	5	5	5	5	5	5	5
	Number of animals with abnormal findings	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

a): Day of necropsy

Species : Rat

Generation 1		Posture NC			Convulsion NC				Spontaneous NC	
Test article	Dose									
		N	F	H	0	1	2	3		
Dodecyloxirane	0 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane	100 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane	300 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane	1000 mg/kg	n	12	0	0	12	0	0	0	

		Abnormal NC behavior			
Test article		0	1	2	3
Dose	n				
Dodecyloxirane 0 mg/kg	n	12	0	0	0
Dodecyloxirane 100 mg/kg	n	12	0	0	0
Dodecyloxirane 300 mg/kg	n	12	0	0	0
Dodecyloxirane 1000 mg/kg	n	12	0	0	0

Posture) N: Normal, F: Flattened, H: Hunched
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe
Not calculated : NC

Species : Rat

Test article	Dose	Posture NC			Convulsion NC				Species NC	
		N	F	H	0	1	2	3		
Dodecyloxirane	0 mg/kg	n	12	0	0	0	12	0	0	0
Dodecyloxirane	100 mg/kg	n	12	0	0	0	12	0	0	0
Dodecyloxirane	300 mg/kg	n	12	0	0	0	12	0	0	0
Dodecyloxirane	1000 mg/kg	n	12	0	0	0	12	0	0	0

Test article	Dose	Abnormal NC behavior			
		0	1	2	3
Dodecyloxirane	n	12	0	0	0
0 mg/kg					
Dodecyloxirane	n	12	0	0	0
100 mg/kg					
Dodecyloxirane	n	12	0	0	0
300 mg/kg					
Dodecyloxirane	n	12	0	0	0
1000 mg/kg					

Posture) N: Normal, F: Flattened, H: Hunched
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe
Not calculated : NC

Table 2 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation Period : F0 mating
Generation : F0 Sex : Male Week : 3

Study No. : R-1261

Home cage observation		Sex : Male			Week : 3		Period : P 0 mating			Species : Rat	
		Posture NC				Convulsion NC					
Test article											
Dose		N	F	H		0	1	2		3	
Dodecyloxirane 0 mg/kg	n	12	0	0		12	0	0		0	
Dodecyloxirane 100 mg/kg	n	12	0	0		12	0	0		0	
Dodecyloxirane 300 mg/kg	n	12	0	0		12	0	0		0	
Dodecyloxirane 1000 mg/kg	n	12	0	0		12	0	0		0	
Abnormal NC behavior											
Test article											
Dose		0	1	2		3					
Dodecyloxirane 0 mg/kg	n	12	0	0		0					
Dodecyloxirane 100 mg/kg	n	12	0	0		0					
Dodecyloxirane 300 mg/kg	n	12	0	0		0					
Dodecyloxirane 1000 mg/kg	n	12	0	0		0					
Posture) N: Normal, F: Flattened, H: Hunched											
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe											
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe											
Not calculated : NC											

Table 2 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation Period : F0 mating
Generation : F0 Sex : Male Week : 4

Study No. : R-1261

Home cage observation		Sex : Male			Week : 4		Period : P 0 mating			Species : Rat	
		Posture NC				Convulsion NC					
Test article											
Dose		N	F	H		0	1	2		3	
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	0	0	
Dodecyloxirane 100 mg/kg	n	12	0	0	0	12	0	0	0	0	
Dodecyloxirane 300 mg/kg	n	12	0	0	0	12	0	0	0	0	
Dodecyloxirane 1000 mg/kg	n	12	0	0	0	12	0	0	0	0	
Abnormal NC behavior											
Test article											
Dose		0	1	2		3					
Dodecyloxirane 0 mg/kg	n	12	0	0	0	0					
Dodecyloxirane 100 mg/kg	n	12	0	0	0	0					
Dodecyloxirane 300 mg/kg	n	12	0	0	0	0					
Dodecyloxirane 1000 mg/kg	n	12	0	0	0	0					
Posture) N: Normal, F: Flattened, H: Hunched											
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe											
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe											
Not calculated : NC											

Table 2 - 5

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation Period : F0 mating
Generation : F0 Sex : Male Week : 5

Study No. : R-1261

Generation : F0		Sex : Male		Week : 5		Posture NC		Convulsion NC		Species : Rat					
Test article															
Dose		N		F		H		0		1		2		3	
Dodecyloxirane 0 mg/kg		n	12	0	0	0	12	0	0	0	0				
Dodecyloxirane 100 mg/kg		n	12	0	0	0	12	0	0	0	0				
Dodecyloxirane 300 mg/kg		n	12	0	0	0	12	0	0	0	0				
Dodecyloxirane 1000 mg/kg		n	12	0	0	0	12	0	0	0	0				
Abnormal NC behavior															
Test article															
Dose		0		1		2		3							
Dodecyloxirane 0 mg/kg		n	12	0	0	0	0								
Dodecyloxirane 100 mg/kg		n	12	0	0	0	0								
Dodecyloxirane 300 mg/kg		n	12	0	0	0	0								
Dodecyloxirane 1000 mg/kg		n	12	0	0	0	0								
Posture) N: Normal, F: Flattened, H: Hunched															
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe															
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe															
Not calculated : NC															

Species : Rat

Generation : F0		Sex : male			Week : 6			Species : Rat			
Test article Dose	n	Posture NC			Convulsion NC						
		N	F	H	0	1	2	3			
Dodecyloxirane 0 mg/kg	n	12	0	0	12	0	0	0			
Dodecyloxirane 100 mg/kg	n	12	0	0	12	0	0	0			
Dodecyloxirane 300 mg/kg	n	12	0	0	12	0	0	0			
Dodecyloxirane 1000 mg/kg	n	12	0	0	12	0	0	0			

Test article	Dose	n	Abnormal NC behavior				
			0	1	2	3	4
Dodecyloxirane	0 mg/kg	n	12	0	0	0	0
Dodecyloxirane	100 mg/kg	n	12	0	0	0	0
Dodecyloxirane	300 mg/kg	n	12	0	0	0	0
Dodecyloxirane	1000 mg/kg	n	12	0	0	0	0

Posture) N: Normal, F: Flattened, H: Hunched
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe
Not calculated : NC

Table 2 - 7

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation Period : F0 before mating
Generation : F0 Sex : Female Week : 1

Study No. : R-1261

Home cage observation		Sex : Female		Week : 1		Period : 10 before mating				Species : Rat	
		Posture NC				Convulsion NC					
Test article											
Dose		N	F	H		0	1	2		3	
Dodecyloxirane 0 mg/kg	n	12	0	0		12	0	0		0	
Dodecyloxirane 100 mg/kg	n	12	0	0		12	0	0		0	
Dodecyloxirane 300 mg/kg	n	12	0	0		12	0	0		0	
Dodecyloxirane 1000 mg/kg	n	12	0	0		12	0	0		0	
Abnormal NC behavior											
Test article											
Dose		0	1	2		3					
Dodecyloxirane 0 mg/kg	n	12	0	0		0					
Dodecyloxirane 100 mg/kg	n	12	0	0		0					
Dodecyloxirane 300 mg/kg	n	12	0	0		0					
Dodecyloxirane 1000 mg/kg	n	12	0	0		0					
Posture) N: Normal, F: Flattened, H: Hunched											
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe											
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe											
Not calculated : NC											

Species : Rat

Test article	Dose	Posture NC			Convulsion NC				Species : Rat
		N	F	H	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane 300 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane 1000 mg/kg	n	12	0	0	12	0	0	0	

Abnormal NC
behavior

Test article	Dose				
Dose	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	0
Dodecyloxirane 100 mg/kg	n	12	0	0	0
Dodecyloxirane 300 mg/kg	n	12	0	0	0
Dodecyloxirane 1000 mg/kg	n	12	0	0	0

Posture) N: Normal, F: Flattened, H: Hunched
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe
Not calculated : NC

Species : Rat

Test article Dose	Generation : F0	Sex : Female	Day : 1	Posture NC				Convulsion NC				Species : Rat
				N	F	H	0	1	2	3		
Dodecyloxirane 0 mg/kg	n			12	0	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n			10	0	0	0	10	0	0	0	
Dodecyloxirane 300 mg/kg	n			10	0	0	0	10	0	0	0	
Dodecyloxirane 1000 mg/kg	n			11	0	0	0	11	0	0	0	

Test article	Dose	Abnormal NC behavior				
		0	1	2	3	
Dodecyloxirane	n	12	0	0	0	
0 mg/kg						
Dodecyloxirane	n	10	0	0	0	
100 mg/kg						
Dodecyloxirane	n	10	0	0	0	
300 mg/kg						
Dodecyloxirane	n	11	0	0	0	
1000 mg/kg						

Posture) N: Normal, F: Flattened, H: Hunched
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe
Not calculated : NC

Table 2 - 10

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Home cage observation
Generation : F0 Sex : Female Day : 7 Period : F0 gestation

Study No. : R-1261

Test article	Dose	Posture NC			Convulsion NC			
		N	F	H	0	1	2	3
Dodecyloxirane	n	12	0	0	12	0	0	0
0 mg/kg								
Dodecyloxirane	n	10	0	0	10	0	0	0
100 mg/kg								
Dodecyloxirane	n	10	0	0	10	0	0	0
300 mg/kg								
Dodecyloxirane	n	11	0	0	11	0	0	0
1000 mg/kg								
Test article	Dose	Abnormal NC behavior						
		0	1	2	3			
Dodecyloxirane	n	12	0	0	0			
0 mg/kg								
Dodecyloxirane	n	10	0	0	0			
100 mg/kg								
Dodecyloxirane	n	10	0	0	0			
300 mg/kg								
Dodecyloxirane	n	11	0	0	0			
1000 mg/kg								
Posture) N: Normal, F: Flattened, H: Hunched								
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe								
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe								
Not calculated : NC								

Table 2 - 11

Dodecylloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation
Generation : F0 Sex : Female Day : 14 Period : F0 gestation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 14		Period : Postnatal		Species : Rat	
		Posture NC				Convulsion NC			
Test article									
Dose		N	F	H		0	1	2	3
Dodecyloxirane	n	12	0	0		12	0	0	0
0 mg/kg									
Dodecyloxirane	n	10	0	0		10	0	0	0
100 mg/kg									
Dodecyloxirane	n	10	0	0		10	0	0	0
300 mg/kg									
Dodecyloxirane	n	11	0	0		11	0	0	0
1000 mg/kg									
Abnormal NC behavior									
Test article									
Dose		0	1	2	3				
Dodecyloxirane	n	12	0	0	0				
0 mg/kg									
Dodecyloxirane	n	10	0	0	0				
100 mg/kg									
Dodecyloxirane	n	10	0	0	0				
300 mg/kg									
Dodecyloxirane	n	11	0	0	0				
1000 mg/kg									
Posture) N: Normal, F: Flattened, H: Hunched									
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe									
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe									
Not calculated : NC									

Table 2 - 12

Dodecylloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation
Generation : F0 Sex : Female Day : 20 Period : F0 gestation

Study No. : R-1261

Test article Dose	Generation : F0	Posture NC			Convulsion NC				Species : Rat	
		N	F	H	0	1	2	3		
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	10	0	0	0	10	0	0	0	
Dodecyloxirane 300 mg/kg	n	10	0	0	0	10	0	0	0	
Dodecyloxirane 1000 mg/kg	n	11	0	0	0	11	0	0	0	
Test article Dose	Generation : F0	Abnormal NC behavior				Abnormal NC behavior				Species : Rat
		0	1	2	3	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	0	0	0	0	0	
Dodecyloxirane 100 mg/kg	n	10	0	0	0	0	0	0	0	
Dodecyloxirane 300 mg/kg	n	10	0	0	0	0	0	0	0	
Dodecyloxirane 1000 mg/kg	n	11	0	0	0	0	0	0	0	
Posture) N: Normal, F: Flattened, H: Hunched										
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe										
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe										
Not calculated : NC										

Table 2 - 13

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Home cage observation
Generation : F0 Sex : Female Day : 6 Period : F0 lactation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 6		Purified F0 isolation		Species : Rat	
		Posture NC			Convulsion NC				
Test article									
Dose		N	F	H	0	1	2	3	
Dodecyloxirane	n	12	0	0	12	0	0	0	
0 mg/kg									
Dodecyloxirane	n	10	0	0	10	0	0	0	
100 mg/kg									
Dodecyloxirane	n	10	0	0	10	0	0	0	
300 mg/kg									
Dodecyloxirane	n	10	0	0	10	0	0	0	
1000 mg/kg									
Abnormal NC behavior									
Test article		0	1	2	3				
Dose									
Dodecyloxirane	n	12	0	0	0				
0 mg/kg									
Dodecyloxirane	n	10	0	0	0				
100 mg/kg									
Dodecyloxirane	n	10	0	0	0				
300 mg/kg									
Dodecyloxirane	n	10	0	0	0				
1000 mg/kg									
Posture) N: Normal, F: Flattened, H: Hunched									
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe									
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe									
Not calculated : NC									

Table 2 - 14

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Home cage observation
Generation : F0 Sex : Female Day : 13 Period : F0 lactation

Study No. : R-1261

Test article	Dose	Posture NC				Convulsion NC			
		N	F	H	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	10	0	0	10	0	0	0	
Dodecyloxirane 300 mg/kg	n	10	0	0	10	0	0	0	
Dodecyloxirane 1000 mg/kg	n	10	0	0	10	0	0	0	
Abnormal NC behavior									
Test article	Dose	0	1	2	3				
Dodecyloxirane 0 mg/kg	n	12	0	0	0				
Dodecyloxirane 100 mg/kg	n	10	0	0	0				
Dodecyloxirane 300 mg/kg	n	10	0	0	0				
Dodecyloxirane 1000 mg/kg	n	10	0	0	0				
Posture) N: Normal, F: Flattened, H: Hunched Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe Not calculated : NC									

Table 2 - 15

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female Week : 1

Species : Rat

		Posture NC				Convulsion NC			
Test article									
Dose		N	F	H		0	1	2	3
Dodecyloxirane	n	10	0	0		10	0	0	0
0 mg/kg									
Dodecyloxirane	n	10	0	0		10	0	0	0
1000 mg/kg									
		Abnormal NC behavior							
Test article									
Dose		0	1	2	3				
Dodecyloxirane	n	10	0	0	0				
0 mg/kg									
Dodecyloxirane	n	10	0	0	0				
1000 mg/kg									

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Not calculated : NC

Table 2 - 16

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female Week : 2

Species : Rat

		Posture NC				Convulsion NC			
Test article									
Dose		N	F	H		0	1	2	3
Dodecyloxirane	n	10	0	0		10	0	0	0
0 mg/kg									
Dodecyloxirane	n	10	0	0		10	0	0	0
1000 mg/kg									
		Abnormal NC behavior							
Test article									
Dose		0	1	2	3				
Dodecyloxirane	n	10	0	0	0				
0 mg/kg									
Dodecyloxirane	n	10	0	0	0				
1000 mg/kg									

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Not calculated : NC

Table 2 - 17

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation
Generation : F0 Sex : Female Week : 3 Period : Administration (non-mating group)

Study No. : R-1261

		Generation : F0		Sex : Female		Week : 3		Dose administration (non-mating group)				Species : Rat			
		Posture NC						Convulsion NC							
Test article															
Dose		N		F		H		0		1		2		3	
Dodecyloxirane		n	10	0	0	0		10	0	0	0	0			
0 mg/kg															
Dodecyloxirane		n	10	0	0	0		10	0	0	0	0			
1000 mg/kg															
		Abnormal NC behavior													
Test article															
Dose		0		1		2		3							
Dodecyloxirane		n	10	0	0	0	0								
0 mg/kg															
Dodecyloxirane		n	10	0	0	0	0								
1000 mg/kg															
Posture) N: Normal, F: Flattened, H: Hunched															
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe															
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe															
Not calculated : NC															

Table 2 - 18

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation
Generation : F0 Sex : Female Week : 4 Period : Administration (non-mating group)

Study No. : R-1261

		Generation : F0		Sex : Female		Week : 4		Dose administration (non-mating group)				Species : Rat			
		Posture NC						Convulsion NC							
Test article															
Dose		N		F		H		0		1		2		3	
Dodecyloxirane 0 mg/kg		n	10	0	0	0		10	0	0	0	0			
Dodecyloxirane 1000 mg/kg		n	10	0	0	0		10	0	0	0	0			
		Abnormal NC behavior													
Test article															
Dose		0		1		2		3							
Dodecyloxirane 0 mg/kg		n	10	0	0	0	0	0							
Dodecyloxirane 1000 mg/kg		n	10	0	0	0	0	0							
Posture) N: Normal, F: Flattened, H: Hunched															
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe															
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe															
Not calculated : NC															

Table 2 - 19

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female Week : 5

Species : Rat

		Posture NC				Convulsion NC			
Test article									
Dose		N	F	H		0	1	2	3
Dodecyloxirane	n	10	0	0		10	0	0	0
0 mg/kg									
Dodecyloxirane	n	10	0	0		10	0	0	0
1000 mg/kg									
		Abnormal NC behavior							
Test article									
Dose		0	1	2	3				
Dodecyloxirane	n	10	0	0	0				
0 mg/kg									
Dodecyloxirane	n	10	0	0	0				
1000 mg/kg									

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Not calculated : NC

Table 2 - 20

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation
Generation : F0 Sex : Female Week : 6 Period : Administration (non-mating group)

Study No. : R-1261

		Home cage observation		Sex : Female		Week : 6		Period : Administration (non mating group)					Species : Rat		
		Posture NC				Convulsion NC									
Test article															
Dose		N		F		H		0		1		2		3	
Dodecyloxirane		n	10	0	0	0	10	0	0	0	0	0	0		
0 mg/kg															
Dodecyloxirane		n	10	0	0	0	10	0	0	0	0	0	0		
1000 mg/kg															

Table 2 - 21

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation Period : Recovery
Generation : F0 Sex : Male Week : 1

Study No. : R-1261

Home cage observation		Sex : Male	Week : 1	Period : Recovery					Species : Rat
		Posture NC			Convulsion NC				
Test article									
Dose		N	F	H	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	5	0	0	5	0	0	0	
Dodecyloxirane 1000 mg/kg	n	5	0	0	5	0	0	0	
Abnormal NC behavior									
Test article									
Dose		0	1	2	3				
Dodecyloxirane 0 mg/kg	n	5	0	0	0				
Dodecyloxirane 1000 mg/kg	n	5	0	0	0				
Posture) N: Normal, F: Flattened, H: Hunched Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe Not calculated : NC									

Table 2 - 22

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation Period : Recovery
Generation : F0 Sex : Male Week : 2

Study No. : R-1261

		Home cage observation		Sex : Male	Week : 2	Period : Recovery		Species : Rat		
		Posture NC				Convulsion NC				
Test article										
Dose		N		F	H	0		1	2	3
Dodecyloxirane 0 mg/kg	n	5	0	0		5	0	0	0	
Dodecyloxirane 1000 mg/kg	n	5	0	0		5	0	0	0	
		Abnormal NC behavior								
Test article										
Dose		0	1	2	3					
Dodecyloxirane 0 mg/kg	n	5	0	0	0					
Dodecyloxirane 1000 mg/kg	n	5	0	0	0					
Posture) N: Normal, F: Flattened, H: Hunched										
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe										
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe										
Not calculated : NC										

Table 2 - 23

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation
Generation : F0 Sex : Female Week : 1 Period : Recovery (non-mating group)

Study No. : R-1261

		Home cage observation		Sex : Female		Week : 1		Period : Recovery (non mating group)				Species : Rat			
		Posture NC				Convulsion NC									
Test article															
Dose		N		F		H		0		1		2		3	
Dodecyloxirane		n	5	0	0	5	0	0	0	0	0	0			
0 mg/kg															
Dodecyloxirane		n	5	0	0	5	0	0	0	0					
1000 mg/kg															
		Abnormal NC behavior													
Test article															
Dose		0		1		2		3							
Dodecyloxirane		n	5	0	0	0	0	0							
0 mg/kg															
Dodecyloxirane		n	5	0	0	0									
1000 mg/kg															
Posture) N: Normal, F: Flattened, H: Hunched															
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe															
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe															
Not calculated : NC															

Table 2 - 24

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Home cage observation
Generation : F0 Sex : Female Week : 2 Period : Recovery (non-mating group)

Study No. : R-1261

		Generation : F0		Sex : Female		Week : 2		Vehicle : Vehicle (new mating group)		Species : Rat					
		Posture NC				Convulsion NC									
Test article															
Dose		N		F		H		0		1		2		3	
Dodecyloxirane	n	5		0		0		5		0		0		0	
0 mg/kg															
Dodecyloxirane	n	5		0		0		5		0		0		0	
1000 mg/kg															
		Abnormal NC behavior													
Test article															
Dose		0		1		2		3							
Dodecyloxirane	n	5		0		0		0							
0 mg/kg															
Dodecyloxirane	n	5		0		0		0							
1000 mg/kg															
Posture) N: Normal, F: Flattened, H: Hunched															
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe															
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe															
Not calculated : NC															

Table 2 - 25

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 before mating

Study No. : R-1261

		Generation : F0		Sex : Male		Week : 1								Species : Rat			
		Ease of removal from cage						Fur condition NC									
Test article																	
Dose																	
Dodecyloxirane	n	-1	0	12	1	0	2	0	3	0	12	1	0	2	0	3	0
0 mg/kg																	
Dodecyloxirane	n		0	12		0		0		0	12		0		0		0
100 mg/kg																	
Dodecyloxirane	n		0	12		0		0		0	12		0		0		0
300 mg/kg																	
Dodecyloxirane	n		0	12		0		0		0	12		0		0		0
1000 mg/kg																	
		Skin NC						Secretions- NC Eye, Nose		Exophthalmos NC							
Test article																	
Dose																	
Dodecyloxirane	n	0	1	2	3			0	1	0	12	1					
0 mg/kg																	
Dodecyloxirane	n	12	0	0	0			12	0		12		0				
100 mg/kg																	
Dodecyloxirane	n	12	0	0	0			12	0		12		0				
300 mg/kg																	
Dodecyloxirane	n	12	0	0	0			12	0		12		0				
1000 mg/kg																	

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 26

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Male Week : 1 Period : F0 before mating

Study No. : R-1261

		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
0 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
100 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
300 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
1000 mg/kg													
		Piloerection NC			Pupil size NC				Salivation NC				
Test article													
Dose		0	1		-1	0	1	2	0	1	2	3	
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
0 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
100 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
300 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
1000 mg/kg													

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 27

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats		
In-the-hand observation		
Generation : F0	Sex : Male	Week : 1
Period : F0 before mating		

Study No. : R-1261

Control (NC)		Abnormal NC respiration				Vocalization NC				Species (Rat)
Test article										
Dose		0	1	2	3	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	12	0	0	0	12	0	0	0	
Dodecyloxirane 300 mg/kg	n	12	0	0	0	12	0	0	0	
Dodecyloxirane 1000 mg/kg	n	12	0	0	0	12	0	0	0	
		Reactivity to NC handling								
Test article										
Dose		-1	0	1	2	3				
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0				
Dodecyloxirane 100 mg/kg	n	0	12	0	0	0				
Dodecyloxirane 300 mg/kg	n	0	12	0	0	0				
Dodecyloxirane 1000 mg/kg	n	0	12	0	0	0				

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe
Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive
Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward
Not calculated : NC

Table 2 - 28

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 before mating

Study No. : R-1261

		Generation : F0		Sex : Male		Week : 2								Species : Rat				
		Ease of removal from cage						Fur condition NC										
Test article																		
Dose																		
Dodecyloxirane	n	-1	0	12	1	0	2	0	3	0	0	12	1	0	2	0	3	0
0 mg/kg																		
Dodecyloxirane	n		0	12		0		0		0		12		0		0		0
100 mg/kg																		
Dodecyloxirane	n		0	12		0		0		0		12		0		0		0
300 mg/kg																		
Dodecyloxirane	n		0	12		0		0		0		12		0		0		0
1000 mg/kg																		
		Skin NC						Secretions- NC		Exophthalmos NC								
								Eye, Nose										
Test article																		
Dose																		
Dodecyloxirane	n	0	1	2	3		0	1	0	0	1							
0 mg/kg		12	0	0	0	0	12	0	12	0								
Dodecyloxirane	n	12	0	0	0	0	12	0	12	0								
100 mg/kg																		
Dodecyloxirane	n	12	0	0	0	0	12	0	12	0								
300 mg/kg																		
Dodecyloxirane	n	12	0	0	0	0	12	0	12	0								
1000 mg/kg																		

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 29

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Male Week : 2 Period : F0 before mating

Study No. : R-1261

		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
0 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
100 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
300 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
1000 mg/kg													
		Piloerection NC			Pupil size NC				Salivation NC				
Test article													
Dose		0	1		-1	0	1	2	0	1	2	3	
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
0 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
100 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
300 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
1000 mg/kg													

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 30

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

In-the-hand observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 2

Study No. : R-1261

[illegible]

Table 2 - 31

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Male Week : 3 Period : F0 mating

Study No. : R-1261

		Ease of removal NC al from cage								Fur condition NC				Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 100 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 300 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
		Skin NC					Secretions- NC Eye, Nose			Exophthalmos NC				
Test article		0	1	2	3		0	1		0	1			
Dose														
Dodecyloxirane 0 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		
Dodecyloxirane 100 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		
Dodecyloxirane 300 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		
Dodecyloxirane 1000 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 32

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Male Week : 3 Period : F0 mating

Study No. : R-1261

		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
0 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
100 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
300 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
1000 mg/kg													
		Piloerection NC			Pupil size NC				Salivation NC				
Test article													
Dose		0	1		-1	0	1	2	0	1	2	3	
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
0 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
100 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
300 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
1000 mg/kg													

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 33

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats			
In-hand observation		Period : F0 mating	
Generation : F0	Sex : Male	Week : 3	

Study No. : R-1261

[illegible]

Table 2 - 34

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Male Week : 4 Period : F0 mating

Study No. : R-1261

		Ease of removal NC al from cage								Fur condition NC				Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 100 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 300 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
		Skin NC				Secretions- NC Eye, Nose			Exophthalmos NC					
Test article														
Dose		0	1	2	3	0	1		0	1				
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	12	0	0			
Dodecyloxirane 100 mg/kg	n	12	0	0	0	12	0	0	12	0	0			
Dodecyloxirane 300 mg/kg	n	12	0	0	0	12	0	0	12	0	0			
Dodecyloxirane 1000 mg/kg	n	12	0	0	0	12	0	0	12	0	0			

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 35

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 mating

Study No. : R-1261

		Generation : F0		Sex : Male		Week : 4						Species : Rat			
		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC			
Test article	Dose	0			1		2		3			0		1	
Dodecyloxirane	0 mg/kg	n	12	0	0	0	0	0	12	0	0	0	12	0	
Dodecyloxirane	100 mg/kg	n	12	0	0	0	0	12	0	0	0	12	0		
Dodecyloxirane	300 mg/kg	n	12	0	0	0	12	0	0	0	12	0			
Dodecyloxirane	1000 mg/kg	n	12	0	0	0	12	0	0	0	12	0			
		Piloerection NC				Pupil size NC				Salivation NC					
Test article	Dose	0		1		-1		0		1		2		3	
Dodecyloxirane	0 mg/kg	n	12	0	0	12	0	0	12	0	12	0	0	0	
Dodecyloxirane	100 mg/kg	n	12	0	0	12	0	0	12	0	12	0	0	0	
Dodecyloxirane	300 mg/kg	n	12	0	0	12	0	0	12	0	12	0	0	0	
Dodecyloxirane	1000 mg/kg	n	12	0	0	12	0	0	12	0	12	0	0	0	

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 36

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

In-hand observation	Period : F0 mating
Generation : F0	Sex : Male
	Week : 4

Study No. : R-1261

Condition : F0		Sex : Male				Species : Rat			
		Abnormal NC respiration				Vocalization NC			
Test article									
Dose		0	1	2	3	0	1	2	3
Dodecylloxirane 0 mg/kg	n	12	0	0	0	12	0	0	0
Dodecylloxirane 100 mg/kg	n	12	0	0	0	12	0	0	0
Dodecylloxirane 300 mg/kg	n	12	0	0	0	12	0	0	0
Dodecylloxirane 1000 mg/kg	n	12	0	0	0	12	0	0	0
		Reactivity to NC handling							
Test article									
Dose		-1	0	1	2	3			
Dodecylloxirane 0 mg/kg	n	0	12	0	0	0			
Dodecylloxirane 100 mg/kg	n	0	12	0	0	0			
Dodecylloxirane 300 mg/kg	n	0	12	0	0	0			
Dodecylloxirane 1000 mg/kg	n	0	12	0	0	0			
Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe									
Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive									
Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward									
Not calculated : NC									

Table 2 - 37

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Male Week : 5 Period : F0 mating

Study No. : R-1261

		Ease of removal- NC al from cage								Fur condition NC				Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 100 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 300 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
		Skin NC					Secretions- NC Eye, Nose			Exophthalmos NC				
Test article		0	1	2	3		0	1		0	1			
Dose														
Dodecyloxirane 0 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		
Dodecyloxirane 100 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		
Dodecyloxirane 300 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		
Dodecyloxirane 1000 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 38

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Male Week : 5 Period : F0 mating

Study No. : R-1261

Generation : F0		Sex : Male		Week : 5		Species : Rat							
		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane 0 mg/kg	n	12	0	0	0		12	0	0	0		12	0
Dodecyloxirane 100 mg/kg	n	12	0	0	0		12	0	0	0		12	0
Dodecyloxirane 300 mg/kg	n	12	0	0	0		12	0	0	0		12	0
Dodecyloxirane 1000 mg/kg	n	12	0	0	0		12	0	0	0		12	0
		Piloerection NC				Pupil size NC				Salivation NC			
Test article		0	1	-1	0	1	2		0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	12	0	0		12	0	0	0	
Dodecyloxirane 100 mg/kg	n	12	0	0	12	0	0		12	0	0	0	
Dodecyloxirane 300 mg/kg	n	12	0	0	12	0	0		12	0	0	0	
Dodecyloxirane 1000 mg/kg	n	12	0	0	12	0	0		12	0	0	0	

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 39

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 mating
Generation : F0 Sex : Male Week : 5

Study No. : R-1261

		Abnormal NC respiration					Vocalization NC					Species : Rat
Test article												
Dose		0	1	2	3		0	1	2	3		
Dodecyloxirane 0 mg/kg	n	12	0	0	0		12	0	0	0		
Dodecyloxirane 100 mg/kg	n	12	0	0	0		12	0	0	0		
Dodecyloxirane 300 mg/kg	n	12	0	0	0		12	0	0	0		
Dodecyloxirane 1000 mg/kg	n	12	0	0	0		12	0	0	0		
		Reactivity to NC handling										
Test article		-1	0	1	2	3						
Dose												
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0						
Dodecyloxirane 100 mg/kg	n	0	12	0	0	0						
Dodecyloxirane 300 mg/kg	n	0	12	0	0	0						
Dodecyloxirane 1000 mg/kg	n	0	12	0	0	0						

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 40

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Male Week : 6 Period : F0 mating

Study No. : R-1261

		Ease of removal NC al from cage								Fur condition NC				Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0		12	0	0	0			
Dodecyloxirane 100 mg/kg	n	0	12	0	0	0		12	0	0	0			
Dodecyloxirane 300 mg/kg	n	0	12	0	0	0		12	0	0	0			
Dodecyloxirane 1000 mg/kg	n	0	12	0	0	0		12	0	0	0			
		Skin NC				Secretions- NC Eye, Nose			Exophthalmos NC					
Test article														
Dose		0	1	2	3	0	1		0	1				
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0		12	0				
Dodecyloxirane 100 mg/kg	n	12	0	0	0	12	0		12	0				
Dodecyloxirane 300 mg/kg	n	12	0	0	0	12	0		12	0				
Dodecyloxirane 1000 mg/kg	n	12	0	0	0	12	0		12	0				

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 41

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Male Week : 6 Period : F0 mating

Study No. : R-1261

		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
0 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
100 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
300 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
1000 mg/kg													
		Piloerection NC				Pupil size NC				Salivation NC			
Test article													
Dose		0	1	-1	0	1	2	0	1	2	3		
Dodecyloxirane	n	12	0	0	12	0	0	12	0	0	0		
0 mg/kg													
Dodecyloxirane	n	12	0	0	12	0	0	12	0	0	0		
100 mg/kg													
Dodecyloxirane	n	12	0	0	12	0	0	12	0	0	0		
300 mg/kg													
Dodecyloxirane	n	12	0	0	12	0	0	12	0	0	0		
1000 mg/kg													

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 42

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation

Generation : F0	Sex : Male	Week : 6	Period : F0 mating
-----------------	------------	----------	--------------------

Study No. : R-1261

Generation : F0		Sex : Male		Week : 0		Abnormal NC respiration		Vocalization NC				Species : Rat	
Test article	Dose												
		0	1	2	3	0	1	2	3				
Dodecyloxirane	n	12	0	0	0	12	0	0	0				
0 mg/kg													
Dodecyloxirane	n	12	0	0	0	12	0	0	0				
100 mg/kg													
Dodecyloxirane	n	12	0	0	0	12	0	0	0				
300 mg/kg													
Dodecyloxirane	n	12	0	0	0	12	0	0	0				
1000 mg/kg													
		Reactivity to NC handling											
Test article	Dose	-1	0	1	2	3							
Dodecyloxirane	n	0	12	0	0	0							
0 mg/kg													
Dodecyloxirane	n	0	12	0	0	0							
100 mg/kg													
Dodecyloxirane	n	0	12	0	0	0							
300 mg/kg													
Dodecyloxirane	n	0	12	0	0	0							
1000 mg/kg													
Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe													
Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive													
Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward													
Not calculated : NC													

Table 2 - 43

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 before mating

Study No. : R-1261

		Generation : F0		Sex : Female		Week : 1		Period : F0 before mating				Species : Rat	
		Ease of removal from cage						Fur condition NC					
Test article	Dose												
		-1	0	1	2	3	0	1	2	3			
Dodecyloxirane	0 mg/kg	n	0	12	0	0	0	12	0	0	0		
Dodecyloxirane	100 mg/kg	n	0	12	0	0	0	12	0	0	0		
Dodecyloxirane	300 mg/kg	n	0	12	0	0	0	12	0	0	0		
Dodecyloxirane	1000 mg/kg	n	0	12	0	0	0	12	0	0	0		
		Skin NC				Secretions- NC Eye, Nose		Exophthalmos NC					
Test article	Dose	0	1	2	3	0	1	0	1				
Dodecyloxirane	0 mg/kg	n	12	0	0	0	12	0	12	0			
Dodecyloxirane	100 mg/kg	n	12	0	0	0	12	0	12	0			
Dodecyloxirane	300 mg/kg	n	12	0	0	0	12	0	12	0			
Dodecyloxirane	1000 mg/kg	n	12	0	0	0	12	0	12	0			

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 44

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Week : 1 Period : F0 before mating

Study No. : R-1261

		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
0 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
100 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
300 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
1000 mg/kg													
		Piloerection NC				Pupil size NC				Salivation NC			
Test article													
Dose		0	1	-1	0	1	2	0	1	2	3		
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
0 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
100 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
300 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
1000 mg/kg													

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 45

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Week : 1 Period : F0 before mating

Study No. : R-1261

Generation : F0		Sex : Female	Abnormal NC respiration				Vocalization NC				Species : Rat	
Test article	Dose	n	0	1	2	3	0	1	2	3		
Dodecyloxirane	0 mg/kg	n	12	0	0	0	12	0	0	0		
Dodecyloxirane	100 mg/kg	n	12	0	0	0	12	0	0	0		
Dodecyloxirane	300 mg/kg	n	12	0	0	0	12	0	0	0		
Dodecyloxirane	1000 mg/kg	n	12	0	0	0	12	0	0	0		
Test article			Reactivity to NC handling									
Dose	n	-1	0	1	2	3						
Dodecyloxirane	0 mg/kg	n	0	12	0	0	0					
Dodecyloxirane	100 mg/kg	n	0	12	0	0	0					
Dodecyloxirane	300 mg/kg	n	0	12	0	0	0					
Dodecyloxirane	1000 mg/kg	n	0	12	0	0	0					

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe
Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive
Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward
Not calculated : NC

Table 2 - 46

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Week : 2 Period : F0 before mating

Study No. : R-1261

		Ease of removal NC al from cage								Fur condition NC				Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 100 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 300 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
		Skin NC					Secretions- NC Eye, Nose			Exophthalmos NC				
Test article		0	1	2	3		0	1		0	1			
Dose														
Dodecyloxirane 0 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		
Dodecyloxirane 100 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		
Dodecyloxirane 300 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		
Dodecyloxirane 1000 mg/kg	n	12	0	0	0	0	12	0	0	12	0	0		

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 47

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Week : 2 Period : F0 before mating

Study No. : R-1261

		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
0 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
100 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
300 mg/kg													
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
1000 mg/kg													
		Piloerection NC			Pupil size NC				Salivation NC				
Test article													
Dose		0	1		-1	0	1	2	0	1	2	3	
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
0 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
100 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
300 mg/kg													
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	
1000 mg/kg													

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 48

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Week : 2 Period : F0 before mating

Study No. : R-1261

	Generation : F0	Sex : Female	Week : 2				Vocalization NC				Species : Rat
			Abnormal NC respiration								
Test article Dose			0	1	2	3	0	1	2	3	
Dodecyloxirane 0 mg/kg	n		12	0	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n		12	0	0	0	12	0	0	0	
Dodecyloxirane 300 mg/kg	n		12	0	0	0	12	0	0	0	
Dodecyloxirane 1000 mg/kg	n		12	0	0	0	12	0	0	0	
			Reactivity to NC handling								
Test article Dose			-1	0	1	2	3				
Dodecyloxirane 0 mg/kg	n		0	12	0	0	0				
Dodecyloxirane 100 mg/kg	n		0	12	0	0	0				
Dodecyloxirane 300 mg/kg	n		0	12	0	0	0				
Dodecyloxirane 1000 mg/kg	n		0	12	0	0	0				

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 49

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 gestation
Generation : F0 Sex : Female Day : 1

Study No. : R-1261

		Ease of removal- NC al from cage								Fur condition NC				Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0	0	10	0	0	0	0		
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0	0	10	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	11	0	0	0	0	11	0	0	0	0		
		Skin NC				Secretions- NC Eye, Nose			Exophthalmos NC					
Test article														
Dose		0	1	2	3	0	1		0	1				
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	12	0	0			
Dodecyloxirane 100 mg/kg	n	10	0	0	0	10	0	0	10	0	0			
Dodecyloxirane 300 mg/kg	n	10	0	0	0	10	0	0	10	0	0			
Dodecyloxirane 1000 mg/kg	n	11	0	0	0	11	0	0	11	0	0			

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 50

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Day : 1 Period : F0 gestation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 1						Species : Rat			
		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane 0 mg/kg	n	12	0	0	0		12	0	0	0		12	0
Dodecyloxirane 100 mg/kg	n	10	0	0	0		10	0	0	0		10	0
Dodecyloxirane 300 mg/kg	n	10	0	0	0		10	0	0	0		10	0
Dodecyloxirane 1000 mg/kg	n	11	0	0	0		11	0	0	0		11	0
		Piloerection NC				Pupil size NC				Salivation NC			
Test article													
Dose		0	1	-1	0	1	2	0	1	2	3		
Dodecyloxirane 0 mg/kg	n	12	0	0	12	0	0	12	0	0	0		
Dodecyloxirane 100 mg/kg	n	10	0	0	10	0	0	10	0	0	0		
Dodecyloxirane 300 mg/kg	n	10	0	0	10	0	0	10	0	0	0		
Dodecyloxirane 1000 mg/kg	n	11	0	0	11	0	0	11	0	0	0		

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 51

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Day : 1 Period : F0 gestation

Study No. : R-1261

		Abnormal NC respiration					Vocalization NC					Species : Rat
Test article												
Dose		0	1	2	3		0	1	2	3		
Dodecyloxirane	n	12	0	0	0		12	0	0	0		
0 mg/kg												
Dodecyloxirane	n	10	0	0	0		10	0	0	0		
100 mg/kg												
Dodecyloxirane	n	10	0	0	0		10	0	0	0		
300 mg/kg												
Dodecyloxirane	n	11	0	0	0		11	0	0	0		
1000 mg/kg												
		Reactivity to NC handling										
Test article												
Dose		-1	0	1	2	3						
Dodecyloxirane	n	0	12	0	0	0						
0 mg/kg												
Dodecyloxirane	n	0	10	0	0	0						
100 mg/kg												
Dodecyloxirane	n	0	10	0	0	0						
300 mg/kg												
Dodecyloxirane	n	0	11	0	0	0						
1000 mg/kg												

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 52

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 gestation
Generation : F0 Sex : Female Day : 7

Study No. : R-1261

		Ease of removal from cage								Fur condition NC				Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane	n	0	12	0	0	0		12	0	0	0			
0 mg/kg														
Dodecyloxirane	n	0	10	0	0	0		10	0	0	0			
100 mg/kg														
Dodecyloxirane	n	0	10	0	0	0		10	0	0	0			
300 mg/kg														
Dodecyloxirane	n	0	11	0	0	0		11	0	0	0			
1000 mg/kg														
		Skin NC				Secretions- NC Eye, Nose		Exophthalmos NC						
Test article														
Dose		0	1	2	3	0	1	0	1					
Dodecyloxirane	n	12	0	0	0	12	0	12	0					
0 mg/kg														
Dodecyloxirane	n	10	0	0	0	10	0	10	0					
100 mg/kg														
Dodecyloxirane	n	10	0	0	0	10	0	10	0					
300 mg/kg														
Dodecyloxirane	n	11	0	0	0	11	0	11	0					
1000 mg/kg														

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 53

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Day : 7 Period : F0 gestation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 7						Species : Rat			
		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane 0 mg/kg	n	12	0	0	0		12	0	0	0		12	0
Dodecyloxirane 100 mg/kg	n	10	0	0	0		10	0	0	0		10	0
Dodecyloxirane 300 mg/kg	n	10	0	0	0		10	0	0	0		10	0
Dodecyloxirane 1000 mg/kg	n	11	0	0	0		11	0	0	0		11	0
		Piloerection NC				Pupil size NC				Salivation NC			
Test article													
Dose		0	1	-1	0	1	2	0	1	2	3		
Dodecyloxirane 0 mg/kg	n	12	0	0	12	0	0	12	0	0	0		
Dodecyloxirane 100 mg/kg	n	10	0	0	10	0	0	10	0	0	0		
Dodecyloxirane 300 mg/kg	n	10	0	0	10	0	0	10	0	0	0		
Dodecyloxirane 1000 mg/kg	n	11	0	0	11	0	0	11	0	0	0		

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 54

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Day : 7 Period : F0 gestation

Study No. : R-1261

Generation : F1		Sex : Female Day : 1					Abnormal NC respiration					Vocalization NC					Species : Rat
Test article	Dose	n	0	1	2	3	n	0	1	2	3						
Dodecylloxirane	0 mg/kg	n	12	0	0	0	12	0	0	0	0						
Dodecylloxirane	100 mg/kg	n	10	0	0	0	10	0	0	0	0						
Dodecylloxirane	300 mg/kg	n	10	0	0	0	10	0	0	0	0						
Dodecylloxirane	1000 mg/kg	n	11	0	0	0	11	0	0	0	0						
		Reactivity to NC handling															
Test article	Dose	n	-1	0	1	2	3										
Dodecylloxirane	0 mg/kg	n	0	12	0	0	0										
Dodecylloxirane	100 mg/kg	n	0	10	0	0	0										
Dodecylloxirane	300 mg/kg	n	0	10	0	0	0										
Dodecylloxirane	1000 mg/kg	n	0	11	0	0	0										
Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe																	
Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive																	
Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward																	
Not calculated : NC																	

Table 2 - 55

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 gestation
Generation : F0 Sex : Female Day : 14

Study No. : R-1261

		Generation : F0		Sex : Female		Day : 14		Ease of removal from cage		Fur condition NC		Species : Rat							
Test article																			
Dose																			
Dodecyloxirane	n	-1	0	12	1	0	2	0	3	0	0	12	1	0	2	0	3	0	
0 mg/kg																			
Dodecyloxirane	n		0	10		0		0		0		10		0		0		0	
100 mg/kg																			
Dodecyloxirane	n		0	10		0		0		0		10		0		0		0	
300 mg/kg																			
Dodecyloxirane	n		0	11		0		0		0		11		0		0		0	
1000 mg/kg																			
		Skin NC					Secretions- NC Eye, Nose			Exophthalmos NC									
Test article																			
Dose																			
Dodecyloxirane	n	0	12	1	0	2	0	3	0	0	12	1	0	0	12	1	0		
0 mg/kg																			
Dodecyloxirane	n		10		0		0		0		10		0		10		0		
100 mg/kg																			
Dodecyloxirane	n		10		0		0		0		10		0		10		0		
300 mg/kg																			
Dodecyloxirane	n		11		0		0		0		11		0		11		0		
1000 mg/kg																			

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 56

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Day : 14 Period : F0 gestation

Study No. : R-1261

		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
0 mg/kg													
Dodecyloxirane	n	10	0	0	0		10	0	0	0		10	0
100 mg/kg													
Dodecyloxirane	n	10	0	0	0		10	0	0	0		10	0
300 mg/kg													
Dodecyloxirane	n	11	0	0	0		11	0	0	0		11	0
1000 mg/kg													
		Piloerection NC				Pupil size NC				Salivation NC			
Test article													
Dose		0	1	-1	0	1	2	0	1	2	3		
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	0
0 mg/kg													
Dodecyloxirane	n	10	0		0	10	0	0	10	0	0	0	0
100 mg/kg													
Dodecyloxirane	n	10	0		0	10	0	0	10	0	0	0	0
300 mg/kg													
Dodecyloxirane	n	11	0		0	11	0	0	11	0	0	0	0
1000 mg/kg													

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 57

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

In-the-hand observation

Generation : F0 Sex : Female Day : 14 Period : F0 gestation

Study No. : R-1261

Condition 1: F		Abnormal NC respiration				Vocalization NC				Species: Rat
Test article										
Dose		0	1	2	3	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	10	0	0	0	10	0	0	0	
Dodecyloxirane 300 mg/kg	n	10	0	0	0	10	0	0	0	
Dodecyloxirane 1000 mg/kg	n	11	0	0	0	11	0	0	0	
Reactivity to NC handling										
Test article										
Dose		-1	0	1	2	3				
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0				
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0				
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0				
Dodecyloxirane 1000 mg/kg	n	0	11	0	0	0				
Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward Not calculated : NC										

Table 2 - 58

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Day : 20 Period : F0 gestation

Study No. : R-1261

		Ease of removal- NC al from cage								Fur condition NC				Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0	0	10	0	0	0	0		
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0	0	10	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	11	0	0	0	0	11	0	0	0	0		
		Skin NC				Secretions- NC Eye, Nose			Exophthalmos NC					
Test article														
Dose		0	1	2	3	0	1		0	1				
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	12	0	0			
Dodecyloxirane 100 mg/kg	n	10	0	0	0	10	0	0	10	0	0			
Dodecyloxirane 300 mg/kg	n	10	0	0	0	10	0	0	10	0	0			
Dodecyloxirane 1000 mg/kg	n	11	0	0	0	11	0	0	11	0	0			

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 59

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Day : 20 Period : F0 gestation

Study No. : R-1261

		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
0 mg/kg													
Dodecyloxirane	n	10	0	0	0		10	0	0	0		10	0
100 mg/kg													
Dodecyloxirane	n	10	0	0	0		10	0	0	0		10	0
300 mg/kg													
Dodecyloxirane	n	11	0	0	0		11	0	0	0		11	0
1000 mg/kg													
		Piloerection NC			Pupil size NC				Salivation NC				
Test article													
Dose		0	1		-1	0	1	2	0	1	2	3	
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	0
0 mg/kg													
Dodecyloxirane	n	10	0		0	10	0	0	10	0	0	0	0
100 mg/kg													
Dodecyloxirane	n	10	0		0	10	0	0	10	0	0	0	0
300 mg/kg													
Dodecyloxirane	n	11	0		0	11	0	0	11	0	0	0	0
1000 mg/kg													

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 60

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

In-the-hand observation

Generation : F0 Sex : Female Day : 20 Period : F0 gestation

Study No. : R-1261

Condition 1: F		Abnormal NC respiration				Vocalization NC				Species: Rat
Test article										
Dose		0	1	2	3	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	10	0	0	0	10	0	0	0	
Dodecyloxirane 300 mg/kg	n	10	0	0	0	10	0	0	0	
Dodecyloxirane 1000 mg/kg	n	11	0	0	0	11	0	0	0	
		Reactivity to NC handling								
Test article										
Dose		-1	0	1	2	3				
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0				
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0				
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0				
Dodecyloxirane 1000 mg/kg	n	0	11	0	0	0				
Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe										
Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive										
Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward										
Not calculated : NC										

Table 2 - 61

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Day : 6 Period : F0 lactation

Study No. : R-1261

		Ease of removal- NC al from cage								Fur condition NC				Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0		12	0	0	0			
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0		10	0	0	0			
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0		10	0	0	0			
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0		10	0	0	0			
		Skin NC				Secretions- NC Eye, Nose			Exophthalmos NC					
Test article														
Dose		0	1	2	3	0	1		0	1				
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0		12	0				
Dodecyloxirane 100 mg/kg	n	10	0	0	0	10	0		10	0				
Dodecyloxirane 300 mg/kg	n	10	0	0	0	10	0		10	0				
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10	0		10	0				

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 62

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Day : 6 Period : F0 lactation

Study No. : R-1261

		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC		
Test article														
Dose		0	1	2	3		0	1	2	3		0	1	
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0	
0 mg/kg														
Dodecyloxirane	n	10	0	0	0		10	0	0	0		10	0	
100 mg/kg														
Dodecyloxirane	n	10	0	0	0		10	0	0	0		10	0	
300 mg/kg														
Dodecyloxirane	n	10	0	0	0		10	0	0	0		10	0	
1000 mg/kg														
		Piloerection NC			Pupil size NC				Salivation NC					
Test article														
Dose		0	1		-1	0	1	2	0	1	2	3		
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0		
0 mg/kg														
Dodecyloxirane	n	10	0		0	10	0	0	10	0	0	0		
100 mg/kg														
Dodecyloxirane	n	10	0		0	10	0	0	10	0	0	0		
300 mg/kg														
Dodecyloxirane	n	10	0		0	10	0	0	10	0	0	0		
1000 mg/kg														

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 63

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 lactation

Study No. : R-1261

		In-the-hand observation		Sex : Female		Day : 6		Period : P0 lactation				Species : Rat	
		Abnormal NC respiration					Vocalization NC						
Test article													
Dose													
Dodecyloxirane	n	0	1	2	3	0	0	1	2	3	0		
0 mg/kg		12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane	n	10	0	0	0	0	10	0	0	0	0		
100 mg/kg													
Dodecyloxirane	n	10	0	0	0	0	10	0	0	0	0		
300 mg/kg													
Dodecyloxirane	n	10	0	0	0	0	10	0	0	0	0		
1000 mg/kg													
		Reactivity to NC handling											
Test article													
Dose													
Dodecyloxirane	n	-1	0	1	2	3	0						
0 mg/kg		0	12	0	0	0	0						
Dodecyloxirane	n	0	10	0	0	0	0						
100 mg/kg													
Dodecyloxirane	n	0	10	0	0	0	0						
300 mg/kg													
Dodecyloxirane	n	0	10	0	0	0	0						
1000 mg/kg													

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 64

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 lactation
Generation : F0 Sex : Female Day : 13

Study No. : R-1261

		Ease of removal- NC al from cage								Fur condition NC				Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	0	12	0	0	0	0		
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0	0	10	0	0	0	0		
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0	0	10	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0	0	10	0	0	0	0		
		Skin NC				Secretions- NC Eye, Nose			Exophthalmos NC					
Test article														
Dose		0	1	2	3	0	1		0	1				
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	12	0	0			
Dodecyloxirane 100 mg/kg	n	10	0	0	0	10	0	10	10	0				
Dodecyloxirane 300 mg/kg	n	10	0	0	0	10	0	10	10	0				
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10	0	10	10	0				

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 65

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Day : 13 Period : F0 lactation

Study No. : R-1261

		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC	
Test article													
Dose		0	1	2	3		0	1	2	3		0	1
Dodecyloxirane	n	12	0	0	0		12	0	0	0		12	0
0 mg/kg													
Dodecyloxirane	n	10	0	0	0		10	0	0	0		10	0
100 mg/kg													
Dodecyloxirane	n	10	0	0	0		10	0	0	0		10	0
300 mg/kg													
Dodecyloxirane	n	10	0	0	0		10	0	0	0		10	0
1000 mg/kg													
		Piloerection NC			Pupil size NC				Salivation NC				
Test article													
Dose		0	1		-1	0	1	2	0	1	2	3	
Dodecyloxirane	n	12	0		0	12	0	0	12	0	0	0	0
0 mg/kg													
Dodecyloxirane	n	10	0		0	10	0	0	10	0	0	0	0
100 mg/kg													
Dodecyloxirane	n	10	0		0	10	0	0	10	0	0	0	0
300 mg/kg													
Dodecyloxirane	n	10	0		0	10	0	0	10	0	0	0	0
1000 mg/kg													

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Not calculated : NC

Table 2 - 66

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : F0 lactation
Generation : F0 Sex : Female Day : 13

Study No. : R-1261

		Abnormal NC respiration					Vocalization NC					Species : Rat
Test article												
Dose		0	1	2	3		0	1	2	3		
Dodecyloxirane 0 mg/kg	n	12	0	0	0		12	0	0	0		
Dodecyloxirane 100 mg/kg	n	10	0	0	0		10	0	0	0		
Dodecyloxirane 300 mg/kg	n	10	0	0	0		10	0	0	0		
Dodecyloxirane 1000 mg/kg	n	10	0	0	0		10	0	0	0		
		Reactivity to NC handling										
Test article		-1	0	1	2	3						
Dose												
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0						
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0						
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0						
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0						

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 67

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Administration (non-mating group)

Study No. : R-1261

		Generation : F0	Sex : Female	Week : 1					Species : Rat				
		Ease of removal from cage NC					Fur condition NC						
Test article													
Dose													
Dodecyloxirane 0 mg/kg	n	-1	0	10	0	0	0	0	10	0	0	0	
Dodecyloxirane 1000 mg/kg	n		0	10	0	0	0	0	10	0	0	0	
		Skin NC					Secretions- NC Eye, Nose		Exophthalmos NC				
Test article													
Dose													
Dodecyloxirane 0 mg/kg	n	0	1	2	3	0	0	10	0	0	10	0	
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	0	10	0	10	0			
		Palpebral NC closure					Mucosal NC membranes				Lacrimation NC		
Test article													
Dose													
Dodecyloxirane 0 mg/kg	n	0	1	2	3	0	0	10	0	0	0	10	0
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	0	10	0	0	0	0	10	0

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 68

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Week : 1 Period : Administration (non-mating group)

Study No. : R-1261

		Piloerection NC				Pupil size NC				Salivation NC			
Test article													
Dose		0	1	-1	0	1	2	0	1	2	3		
Dodecyloxirane 0 mg/kg	n	10	0	0	10	0	0	10	0	0	0		
Dodecyloxirane 1000 mg/kg	n	10	0	0	10	0	0	10	0	0	0		
		Abnormal NC respiration				Vocalization NC							
Test article													
Dose		0	1	2	3	0	1	2	3				
Dodecyloxirane 0 mg/kg	n	10	0	0	0	10	0	0	0				
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10	0	0	0				
		Reactivity to NC handling											
Test article													
Dose		-1	0	1	2	3							
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0							
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0							

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 69

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Administration (non-mating group)
Generation : F0 Sex : Female Week : 2

Study No. : R-1261

		Ease of removal from cage NC						Fur condition NC				Species : Rat
Test article												
Dose		-1	0	1	2	3		0	1	2	3	
Dodecyloxirane	n	0	10	0	0	0		10	0	0	0	
0 mg/kg												
Dodecyloxirane	n	0	10	0	0	0		10	0	0	0	
1000 mg/kg												
		Skin NC				Secretions- NC Eye, Nose		Exophthalmos NC				
Test article												
Dose		0	1	2	3	0	1	0	1			
Dodecyloxirane	n	10	0	0	0	10	0	10	0			
0 mg/kg												
Dodecyloxirane	n	10	0	0	0	10	0	10	0			
1000 mg/kg												
		Palpebral NC closure				Mucosal NC membranes				Lacrimation NC		
Test article												
Dose		0	1	2	3	0	1	2	3	0	1	
Dodecyloxirane	n	10	0	0	0	10	0	0	0	10	0	
0 mg/kg												
Dodecyloxirane	n	10	0	0	0	10	0	0	0	10	0	
1000 mg/kg												

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 70

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Week : 2 Period : Administration (non-mating group)

Study No. : R-1261

In-the-hand observation		Sex : Female		Week : 2		Period : Administration (non-mating group)						Species : Rat	
		Piloerection NC			Pupil size NC			Salivation NC					
Test article													
Dose		01			-1012			0123					
Dodecyloxirane 0 mg/kg	n	10	0	0	10	0	0	10	0	0	0	0	
Dodecyloxirane 1000 mg/kg	n	10	0	0	10	0	0	10	0	0	0	0	
Abnormal NC respiration													
Vocalization NC													
Test article													
Dose		0123			0123								
Dodecyloxirane 0 mg/kg	n	10	0	0	0	10	0	0	0	0	0	0	
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10	0	0	0	0	0	0	
Reactivity to NC handling													
Test article													
Dose		-10123											
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0	0						
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0	0						

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 71

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Administration (non-mating group)
Generation : F0 Sex : Female Week : 3

Study No. : R-1261

		Ease of removal from cage NC						Fur condition NC						Species : Rat
Test article														
Dose		-1	0	1	2	3		0	1	2	3			
Dodecyloxirane 0 mg/kg		n	0	10	0	0	0	10	0	0	0			
Dodecyloxirane 1000 mg/kg		n	0	10	0	0	0	10	0	0	0			
		Skin NC						Secretions- NC Eye, Nose		Exophthalmos NC				
Test article														
Dose		0	1	2	3			0	1	0	1			
Dodecyloxirane 0 mg/kg		n	10	0	0	0		10	0	10	0			
Dodecyloxirane 1000 mg/kg		n	10	0	0	0		10	0	10	0			
		Palpebral NC closure						Mucosal NC membranes				Lacrimation NC		
Test article														
Dose		0	1	2	3			0	1	2	3	0	1	
Dodecyloxirane 0 mg/kg		n	10	0	0	0		10	0	0	0	10	0	
Dodecyloxirane 1000 mg/kg		n	10	0	0	0		10	0	0	0	10	0	

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 72

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Week : 3 Period : Administration (non-mating group)

Study No. : R-1261

In-the-hand observation		Sex : Female		Week : 3		Period : Administration (non-mating group)		Species : Rat			
		Piloerection NC		Pupil size NC		Salivation NC					
Test article											
Dose		0	1	-1	0	1	2	0	1	2	3
Dodecyloxirane 0 mg/kg	n	10	0	0	10	0	0	10	0	0	0
Dodecyloxirane 1000 mg/kg	n	10	0	0	10	0	0	10	0	0	0
Abnormal NC respiration											
Vocalization NC											
Test article											
Dose		0	1	2	3	0	1	2	3		
Dodecyloxirane 0 mg/kg	n	10	0	0	0	10	0	0	0		
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10	0	0	0		
Reactivity to NC handling											
Test article											
Dose		-1	0	1	2	3					
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0					
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0					

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 73

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Administration (non-mating group)
Generation : F0 Sex : Female Week : 4

Study No. : R-1261

		Ease of removal NC al from cage						Fur condition NC				Species : Rat
Test article												
Dose		-1	0	1	2	3		0	1	2	3	
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0		10	0	0	0	
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0		10	0	0	0	
		Skin NC				Secretions- NC Eye, Nose			Exophthalmos NC			
Test article												
Dose		0	1	2	3	0	1		0	1		
Dodecyloxirane 0 mg/kg	n	10	0	0	0	10	0		10	0		
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10	0		10	0		
		Palpebral NC closure				Mucosal NC membranes				Lacrimation NC		
Test article												
Dose		0	1	2	3	0	1	2	3	0	1	
Dodecyloxirane 0 mg/kg	n	10	0	0	0	10	0	0	0	10	0	
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10	0	0	0	10	0	

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 74

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Week : 4 Period : Administration (non-mating group)

Study No. : R-1261

In-the-hand observation		Sex : Female		Week : 4		Period : Administration (non-mating group)		Species : Rat			
		Piloerection NC		Pupil size NC		Salivation NC					
Test article											
Dose		0	1	-1	0	1	2	0	1	2	3
Dodecyloxirane 0 mg/kg	n	10	0	0	10	0	0	10	0	0	0
Dodecyloxirane 1000 mg/kg	n	10	0	0	10	0	0	10	0	0	0

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 75

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Administration (non-mating group)

Study No. : R-1261

		Generation : F0	Sex : Female	Week : 5							Species : Rat				
		Ease of removal from cage NC							Fur condition NC						
Test article															
Dose															
Dodecyloxirane 0 mg/kg	n	-1	0	10	0	0	0	0	10	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n		0	10	0	0	0	0	10	0	0	0	0		
		Skin NC					Secretions- NC Eye, Nose			Exophthalmos NC					
Test article															
Dose															
Dodecyloxirane 0 mg/kg	n	0	1	2	3	0	0	1	0	0	1	0			
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10	0	10	0	10	0				
		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC			
Test article															
Dose															
Dodecyloxirane 0 mg/kg	n	0	1	2	3	0	10	0	0	0	0	10	0		
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10	0	0	0	0	10	0			

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 76

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation
Generation : F0 Sex : Female Week : 5 Period : Administration (non-mating group)

Study No. : R-1261

In-the-hand observation		Sex : Female		Week : 5		Period : Administration (non-mating group)						Species : Rat	
		Piloerection NC			Pupil size NC			Salivation NC					
Test article													
Dose		0	1		-1	0	1	2	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	10	0		0	10	0	0	10	0	0	0	
Dodecyloxirane 1000 mg/kg	n	10	0		0	10	0	0	10	0	0	0	

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 77

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Administration (non-mating group)

Study No. : R-1261

		Generation : F0	Sex : Female	Week : 6										Species : Rat	
		Ease of removal from cage NC										Fur condition NC			
Test article															
Dose															
Dodecyloxirane	0 mg/kg	n	0	10	0	0	0	0	10	0	0	0	0		
Dodecyloxirane	1000 mg/kg	n	0	10	0	0	0	0	10	0	0	0	0		
		Skin NC					Secretions- NC Eye, Nose			Exophthalmos NC					
Test article															
Dose															
Dodecyloxirane	0 mg/kg	n	10	0	0	0	10	0	10	0					
Dodecyloxirane	1000 mg/kg	n	10	0	0	0	10	0	10	0					
		Palpebral NC closure					Mucosal NC membranes					Lacrimation NC			
Test article															
Dose															
Dodecyloxirane	0 mg/kg	n	10	0	0	0	10	0	0	0	10	0			
Dodecyloxirane	1000 mg/kg	n	10	0	0	0	10	0	0	0	10	0			

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 78

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Administration (non-mating group)
Generation : F0 Sex : Female Week : 6

Study No. : R-1261

Generation : F0		Sex : Female		Week : 6		Species : Rat					
		Piloerection NC		Pupil size NC		Salivation NC					
Test article											
Dose		0	1	-1	0	1	2	3			
Dodecyloxirane 0 mg/kg	n	10	0	0	10	0	0	10	0	0	0
Dodecyloxirane 1000 mg/kg	n	10	0	0	10	0	0	10	0	0	0
		Abnormal NC respiration				Vocalization NC					
Test article											
Dose		0	1	2	3	0	1	2	3		
Dodecyloxirane 0 mg/kg	n	10	0	0	0	10	0	0	0		
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10	0	0	0		
		Reactivity to NC handling									
Test article											
Dose		-1	0	1	2	3					
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0					
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0					

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 79

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Recovery

Study No. : R-1261

		Generation : F0	Sex : Male	Week : 1						Species : Rat						
		Ease of removal from cage NC						Fur condition NC								
Test article																
Dose																
Dodecyloxirane 0 mg/kg	n	-1	0	5	0	0	0	0	5	0	0	0	0			
Dodecyloxirane 1000 mg/kg	n		0	5	0	0	0	5	0	0	0		0			
		Skin NC						Secretions- NC Eye, Nose			Exophthalmos NC					
Test article																
Dose																
Dodecyloxirane 0 mg/kg	n	0	1	0	2	0	3	0	0	5	1	0	0			
Dodecyloxirane 1000 mg/kg	n		5	0	0	0	5	0	5		0					
		Palpebral NC closure						Mucosal NC membranes				Lacrimation NC				
Test article																
Dose																
Dodecyloxirane 0 mg/kg	n	0	1	0	2	0	3	0	5	0	0	0	0	5	1	0
Dodecyloxirane 1000 mg/kg	n		5	0	0	0	5	0	0	0	0	5		0		

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 80

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Recovery

Study No. : R-1261

		In-the-hand observation		Sex : Male		Week : 1		Period : Recovery		Species : Rat			
		Piloerection NC			Pupil size NC				Salivation NC				
Test article													
Dose		0 1			-1 0 1 2				0 1 2 3				
Dodecyloxirane 0 mg/kg	n	5 0			0 5 0 0				5 0 0 0				
Dodecyloxirane 1000 mg/kg	n	5 0			0 5 0 0				5 0 0 0				
		Abnormal NC respiration					Vocalization NC						
Test article													
Dose		0 1 2 3					0 1 2 3						
Dodecyloxirane 0 mg/kg	n	5 0 0 0					5 0 0 0						
Dodecyloxirane 1000 mg/kg	n	5 0 0 0					5 0 0 0						
		Reactivity to NC handling											
Test article													
Dose		-1 0 1 2 3											
Dodecyloxirane 0 mg/kg	n	0 5 0 0 0											
Dodecyloxirane 1000 mg/kg	n	0 5 0 0 0											

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 81

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Recovery

Study No. : R-1261

		Generation : F0	Sex : Male	Week : 2					Species : Rat				
		Ease of removal from cage NC					Fur condition NC						
Test article													
Dose		-1	0	1	2	3	0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	5	0	0	0	5	0	0	0			
Dodecyloxirane 1000 mg/kg	n	0	5	0	0	0	5	0	0	0			
		Skin NC				Secretions- NC Eye, Nose		Exophthalmos NC					
Test article													
Dose		0	1	2	3	0	1	0	1				
Dodecyloxirane 0 mg/kg	n	5	0	0	0	5	0	5	0				
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	5	0	5	0				
		Palpebral NC closure				Mucosal NC membranes			Lacrimation NC				
Test article													
Dose		0	1	2	3	0	1	2	3	0	1		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	5	0	0	0	5	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	5	0	0	0	5	0		

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 82

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Recovery

Study No. : R-1261

		Generation : F0		Sex : Male		Week : 2		Pended : Recovery		Species : Rat			
		Piloerection NC				Pupil size NC				Salivation NC			
Test article													
Dose		0 1				-1 0 1 2				0 1 2 3			
Dodecyloxirane 0 mg/kg	n	5 0				0 5 0 0				5 0 0 0			
Dodecyloxirane 1000 mg/kg	n	5 0				0 5 0 0				5 0 0 0			
		Abnormal NC respiration				Vocalization NC							
Test article													
Dose		0 1 2 3				0 1 2 3							
Dodecyloxirane 0 mg/kg	n	5 0 0 0				5 0 0 0							
Dodecyloxirane 1000 mg/kg	n	5 0 0 0				5 0 0 0							
		Reactivity to NC handling											
Test article													
Dose		-1 0 1 2 3											
Dodecyloxirane 0 mg/kg	n	0 5 0 0 0											
Dodecyloxirane 1000 mg/kg	n	0 5 0 0 0											

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 83

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Recovery (non-mating group)

Study No. : R-1261

		Generation : F0					Sex : Female					Week : 1					Period : Recovery (non-mating group)					Species : Rat														
		Ease of removal from cage NC															Fur condition NC																			
Test article																																				
Dose		-1					0					1					2					3														
Dodecyloxirane 0 mg/kg		n	0	5	0	0	0	0	0	0	5	0	0	0	0	0																				
Dodecyloxirane 1000 mg/kg		n	0	5	0	0	0	0	5	0	0	0	0	0	0																					
		Skin NC										Secretions- NC Eye, Nose					Exophthalmos NC																			
Test article																																				
Dose		0					1					2					3					0					1					0				
Dodecyloxirane 0 mg/kg		n	5	0	0	0	0	0	5	1	0	5	0	5	1	0																				
Dodecyloxirane 1000 mg/kg		n	5	0	0	0	5	0	5	0	5	0																								
		Palpebral NC closure										Mucosal NC membranes					Lacrimation NC																			
Test article																																				
Dose		0					1					2					3					0					1					0				
Dodecyloxirane 0 mg/kg		n	5	0	0	0	0	5	1	0	0	0	0	5	1	0																				
Dodecyloxirane 1000 mg/kg		n	5	0	0	0	5	0	0	0	5	0	0	0	5	0																				

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 84

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Recovery (non-mating group)

Study No. : R-1261

		Generation : F0		Sex : Female		Week : 1		Phase : Recovery (non mating group)				Species : Rat	
		Piloerection NC				Pupil size NC				Salivation NC			
Test article													
Dose		0 1				-1 0 1 2				0 1 2 3			
Dodecyloxirane	n	5 0				0 5 0 0				5 0 0 0			
0 mg/kg													
Dodecyloxirane	n	5 0				0 5 0 0				5 0 0 0			
1000 mg/kg													
		Abnormal NC respiration				Vocalization NC							
Test article													
Dose		0 1 2 3				0 1 2 3							
Dodecyloxirane	n	5 0 0 0				5 0 0 0				0			
0 mg/kg													
Dodecyloxirane	n	5 0 0 0				5 0 0 0				0			
1000 mg/kg													
		Reactivity to NC handling											
Test article													
Dose		-1 0 1 2 3											
Dodecyloxirane	n	0 5 0 0 0											
0 mg/kg													
Dodecyloxirane	n	0 5 0 0 0											
1000 mg/kg													

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 85

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Recovery (non-mating group)

Study No. : R-1261

		Generation : F0					Sex : Female					Week : 2					Period : Recovery (non-mating group)					Species : Rat														
		Ease of removal from cage NC															Fur condition NC																			
Test article																																				
Dose		-1					0					1					2					3														
Dodecyloxirane 0 mg/kg		n		0		5		0		0		0		0		5		0		0		0		0												
Dodecyloxirane 1000 mg/kg		n		0		5		0		0		0		0		5		0		0		0		0												
		Skin NC										Secretions- NC Eye, Nose					Exophthalmos NC																			
Test article																																				
Dose		0					1					2					3					0					1					0				
Dodecyloxirane 0 mg/kg		n		5		0		0		0		0		0		5		0		5		0		5		0										
Dodecyloxirane 1000 mg/kg		n		5		0		0		0		0		5		0		5		0		5		0		0										
		Palpebral NC closure										Mucosal NC membranes										Lacrimation NC														
Test article																																				
Dose		0					1					2					3					0					1					0				
Dodecyloxirane 0 mg/kg		n		5		0		0		0		0		5		0		0		0		0		0		5		0								
Dodecyloxirane 1000 mg/kg		n		5		0		0		0		0		5		0		0		0		0		0		5		0								

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Not calculated : NC

Table 2 - 86

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation Period : Recovery (non-mating group)

Study No. : R-1261

Generation : F0		Sex : Female		Week : 2		Phase : Recovery (non mating group)		Species : Rat			
		Piloerection NC		Pupil size NC		Salivation NC					
Test article											
Dose		0	1	-1	0	1	2	0	1	2	3
Dodecyloxirane	n	5	0	0	5	0	0	5	0	0	0
0 mg/kg											
Dodecyloxirane	n	5	0	0	5	0	0	5	0	0	0
1000 mg/kg											
Abnormal NC respiration											
Vocalization NC											
Test article											
Dose		0	1	2	3	0	1	2	3		
Dodecyloxirane	n	5	0	0	0	5	0	0	0		
0 mg/kg											
Dodecyloxirane	n	5	0	0	0	5	0	0	0		
1000 mg/kg											
Reactivity to NC handling											
Test article											
Dose		-1	0	1	2	3					
Dodecyloxirane	n	0	5	0	0	0					
0 mg/kg											
Dodecyloxirane	n	0	5	0	0	0					
1000 mg/kg											

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Not calculated : NC

Table 2 - 87

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Open field observation

Generation : F0 Sex : Male Week : 1 Period : F0 before mating

Study No. : R-1261

Generation : F0		Sex : Male		Week : 1		Arousal NC		Convulsion NC				Species : Rat	
Test article	Dose	Arousal NC						Convulsion NC					
		-2	-1	0	1	2	0	1	2	3			
Dodecyloxirane	n	0	0	12	0	0	12	0	0	0			
0 mg/kg													
Dodecyloxirane	n	0	0	12	0	0	12	0	0	0			
100 mg/kg													
Dodecyloxirane	n	0	0	12	0	0	12	0	0	0			
300 mg/kg													
Dodecyloxirane	n	0	0	12	0	0	12	0	0	0			
1000 mg/kg													
		Abnormal NC behavior					Stereotypy NC						
Test article	Dose	0	1	2	3	0	1	2	3				
Dodecyloxirane	n	12	0	0	0	12	0	0	0				
0 mg/kg													
Dodecyloxirane	n	12	0	0	0	12	0	0	0				
100 mg/kg													
Dodecyloxirane	n	12	0	0	0	12	0	0	0				
300 mg/kg													
Dodecyloxirane	n	12	0	0	0	12	0	0	0				
1000 mg/kg													
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe Not calculated : NC													

Table 2 - 88

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Open field observation	Period : F0 before mating
------------------------	---------------------------

Study No. : R-1261

Generation : F0		Sex : Male		Week : 1		Gait NC		Posture NC		Species : Rat															
Test article	Dose																								
		U			0			1			2			3			N			F			H		
Dodecyloxirane 0 mg/kg	n	0			12			0			0			0			12			0			0		
Dodecyloxirane 100 mg/kg	n	0			12			0			0			0			12			0			0		
Dodecyloxirane 300 mg/kg	n	0			12			0			0			0			12			0			0		
Dodecyloxirane 1000 mg/kg	n	0			12			0			0			0			12			0			0		
Grooming NC													Urination NC												
Test article	Dose																								
		0		1		2		0		1		2		3		0		1		2		3			
Dodecyloxirane 0 mg/kg	n	12		0		0		10		2		0		0		12		0		0		0			
Dodecyloxirane 100 mg/kg	n	12		0		0		11		1		0		0		12		0		0		0			
Dodecyloxirane 300 mg/kg	n	12		0		0		12		0		0		0		12		0		0		0			
Dodecyloxirane 1000 mg/kg	n	12		0		0		10		2		0		0		12		0		0		0			
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly																									
Posture) N: Normal, F: Flattened, H: Hunched																									
Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)																									
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount																									
Not calculated : NC																									

Table 2 - 89

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Male Week : 1 Period : F0 before mating

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	2	0	
Dodecyloxirane 100 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	2	0	
Dodecyloxirane 300 mg/kg	n	12	12	
	Mean	5	0	
	S.D.	3	0	
Dodecyloxirane 1000 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	3	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Species : Rat

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe
Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe
Not calculated : NC

Table 2 - 91

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation Period : F0 before mating

Study No. : R-1261

Open field locomotion		Sex : Male		Week : 2		Pended 1 h before mating		Species : Rat	
		Gait NC				Posture NC			
Test article									
Dose		U	0	1	2	3	N	F	H
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	12	0	0
Dodecyloxirane 100 mg/kg	n	0	12	0	0	0	12	0	0
Dodecyloxirane 300 mg/kg	n	0	12	0	0	0	12	0	0
Dodecyloxirane 1000 mg/kg	n	0	12	0	0	0	12	0	0
Grooming NC					Urination NC				
Test article									
Dose		0	1	2	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	12	0	0	11	1	0	0	
Dodecyloxirane 300 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane 1000 mg/kg	n	12	0	0	11	1	0	0	
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly Posture) N: Normal, F: Flattened, H: Hunched Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four) Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount Not calculated : NC									

Table 2 - 92

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Male Week : 2 Period : F0 before mating

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	2	0	
Dodecyloxirane 100 mg/kg	n	12	12	
	Mean	4	0	
	S.D.	2	0	
Dodecyloxirane 300 mg/kg	n	12	12	
	Mean	5	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	12	12	
	Mean	4	0	
	S.D.	2	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 93

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Male Week : 3 Period : F0 mating

Study No. : R-1261

Generation : F0		Sex : male		Week : 5		Arousal NC		Convulsion NC				Species : Rat	
Test article													
Dose		-2	-1	0	1	2		0	1	2	3		
Dodecylloxirane 0 mg/kg	n	0	0	12	0	0		12	0	0	0		
Dodecylloxirane 100 mg/kg	n	0	0	12	0	0		12	0	0	0		
Dodecylloxirane 300 mg/kg	n	0	0	12	0	0		12	0	0	0		
Dodecylloxirane 1000 mg/kg	n	0	0	12	0	0		12	0	0	0		
		Abnormal NC behavior					Stereotypy NC						
Test article													
Dose		0	1	2	3		0	1	2	3			
Dodecylloxirane 0 mg/kg	n	12	0	0	0		12	0	0	0			
Dodecylloxirane 100 mg/kg	n	12	0	0	0		12	0	0	0			
Dodecylloxirane 300 mg/kg	n	12	0	0	0		12	0	0	0			
Dodecylloxirane 1000 mg/kg	n	12	0	0	0		12	0	0	0			
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert													
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe													
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe													
Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe													
Not calculated : NC													

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Male Week : 3 Period : F0 mating

Open field observation		Sex : Male		Week : 3		Period : 10 min		Species : Rat	
		Gait NC						Posture NC	
Test article									
Dose		U	0	1	2	3	N	F	H
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	12	0	0
Dodecyloxirane 100 mg/kg	n	0	12	0	0	0	12	0	0
Dodecyloxirane 300 mg/kg	n	0	12	0	0	0	12	0	0
Dodecyloxirane 1000 mg/kg	n	0	12	0	0	0	12	0	0
Grooming NC					Urination NC				
Test article									
Dose		0	1	2	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	12	0	0	11	1	0	0	
Dodecyloxirane 300 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane 1000 mg/kg	n	12	0	0	11	1	0	0	
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly									
Posture) N: Normal, F: Flattened, H: Hunched									
Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)									
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount									
Not calculated : NC									

Table 2 - 95

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Male Week : 3 Period : F0 mating

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	4	0	
	S.D.	2	0	
Dodecyloxirane 100 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	2	0	
Dodecyloxirane 300 mg/kg	n	12	12	
	Mean	5	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	2	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 96

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Open field observation

Generation : F0 Sex : Male Week : 4 Period : F0 mating

Study No. : R-1261

	Generation : F0	Sex : male	Week : 4	Arousal NC	Convulsion NC	Species : Rat
Test article Dose				-2 -1 0 1 2	0 1 2 3	
Dodecylloxirane 0 mg/kg	n			0 0 12 0 0	12 0 0 0	
Dodecylloxirane 100 mg/kg	n			0 0 12 0 0	12 0 0 0	
Dodecylloxirane 300 mg/kg	n			0 0 12 0 0	12 0 0 0	
Dodecylloxirane 1000 mg/kg	n			0 0 12 0 0	12 0 0 0	
				Abnormal NC behavior	Stereotypy NC	
Test article Dose				0 1 2 3	0 1 2 3	
Dodecylloxirane 0 mg/kg	n			12 0 0 0	12 0 0 0	
Dodecylloxirane 100 mg/kg	n			12 0 0 0	12 0 0 0	
Dodecylloxirane 300 mg/kg	n			12 0 0 0	12 0 0 0	
Dodecylloxirane 1000 mg/kg	n			12 0 0 0	12 0 0 0	
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe Not calculated : NC						

Table 2 - 97

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Open field observation

Generation : F0 Sex : Male Week : 4 Period : F0 mating

Study No. : R-1261

Generation : F0		Sex : Male		Week : 4		Gait NC		Posture NC		Species : Rat	
Test article											
Dose											
		U	0	1	2	3	N	F	H		
Dodecyloxirane	n	0	12	0	0	0	12	0	0		
0 mg/kg											
Dodecyloxirane	n	0	12	0	0	0	12	0	0		
100 mg/kg											
Dodecyloxirane	n	0	12	0	0	0	12	0	0		
300 mg/kg											
Dodecyloxirane	n	0	12	0	0	0	12	0	0		
1000 mg/kg											
Grooming NC						Urination NC					
Test article											
Dose		0	1	2		0	1	2	3		
Dodecyloxirane	n	12	0	0	10	2	0	0			
0 mg/kg											
Dodecyloxirane	n	12	0	0	10	2	0	0			
100 mg/kg											
Dodecyloxirane	n	12	0	0	12	0	0	0			
300 mg/kg											
Dodecyloxirane	n	12	0	0	10	2	0	0			
1000 mg/kg											
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly											
Posture) N: Normal, F: Flattened, H: Hunched											
Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)											
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount											
Not calculated : NC											

Table 2 - 98

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Male Week : 4 Period : F0 mating

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	1	0	
Dodecyloxirane 100 mg/kg	n	12	12	
	Mean	4	0	
	S.D.	2	0	
Dodecyloxirane 300 mg/kg	n	12	12	
	Mean	4	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	2	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 99

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Male Week : 5 Period : F0 mating

Study No. : R-1261

Generation : F0		Sex : Male		week : 5				Arousal NC				Convulsion NC				Species : Rat																	
Test article	Dose	-2				-1				0				1				2				3											
Dodecyloxirane	0 mg/kg	n	0	0	0	12	0	0	0	0	12	0	0	0	0	0	0																
Dodecyloxirane	100 mg/kg	n	0	0	12	0	0	12	0	0	0	0	0	0	0	0	0																
Dodecyloxirane	300 mg/kg	n	0	0	12	0	0	12	0	0	0	0	0	0	0	0	0																
Dodecyloxirane	1000 mg/kg	n	0	0	12	0	0	12	0	0	0	0	0	0	0	0	0																
		Abnormal NC behavior								Stereotypy NC																							
Test article	Dose	0				1				2				3				0				1				2				3			
Dodecyloxirane	0 mg/kg	n	12	0	0	0	0	12	0	0	0	0	12	0	0	0	0																
Dodecyloxirane	100 mg/kg	n	12	0	0	0	0	12	0	0	0	0	12	0	0	0	0																
Dodecyloxirane	300 mg/kg	n	12	0	0	0	0	12	0	0	0	0	12	0	0	0	0																
Dodecyloxirane	1000 mg/kg	n	12	0	0	0	0	12	0	0	0	0	12	0	0	0	0																
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert																																	
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe																																	
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe																																	
Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe																																	
Not calculated : NC																																	

Generation : F0

Sex : Male

Week : 5

Period : F0 mating

Species : Rat

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly
Posture) N: Normal, F: Flattened, H: Hunched
Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount
Not calculated : NC

Table 2 - 101

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Male Week : 5 Period : F0 mating

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	2	0	
Dodecyloxirane 100 mg/kg	n	12	12	
	Mean	4	0	
	S.D.	2	0	
Dodecyloxirane 300 mg/kg	n	12	12	
	Mean	4	0	
	S.D.	1	0	
Dodecyloxirane 1000 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	2	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 102

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Male Week : 6 Period : F0 mating

Study No. : R-1261

Generation : F0		Sex : male		week : 0				week : 1				week : 2				week : 3				Species : Rat													
		Arousal NC								Convulsion NC																							
Test article		-2				-1				0				1				2				3											
Dose	n																																
Dodecylloxirane 0 mg/kg	n																																
Dodecylloxirane 100 mg/kg	n																																
Dodecylloxirane 300 mg/kg	n																																
Dodecylloxirane 1000 mg/kg	n																																
		Abnormal NC behavior								Stereotypy NC																							
Test article		0				1				2				3				0				1				2				3			
Dose	n																																
Dodecylloxirane 0 mg/kg	n																																
Dodecylloxirane 100 mg/kg	n																																
Dodecylloxirane 300 mg/kg	n																																
Dodecylloxirane 1000 mg/kg	n																																
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe Not calculated : NC																																	

Table 2 - 103

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Open field observation			Period : F0 mating
Generation : F0	Sex : Male	Week : 6	

Study No. : R-1261

Generation : F0		Sex : Male		Week : 8		Gait NC		Posture NC			Species : Rat	
Test article Dose		Gait NC						Posture NC				
		U	0	1	2	3	N	F	H			
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	12	0	0			
Dodecyloxirane 100 mg/kg	n	0	12	0	0	0	12	0	0			
Dodecyloxirane 300 mg/kg	n	0	12	0	0	0	12	0	0			
Dodecyloxirane 1000 mg/kg	n	0	12	0	0	0	12	0	0			
		Grooming NC				Urination NC						
Test article Dose		Grooming NC			Urination NC							
		0	1	2	0	1	2	3				
Dodecyloxirane 0 mg/kg	n	12	0	0	11	1	0	0				
Dodecyloxirane 100 mg/kg	n	12	0	0	9	3	0	0				
Dodecyloxirane 300 mg/kg	n	12	0	0	12	0	0	0				
Dodecyloxirane 1000 mg/kg	n	12	0	0	11	1	0	0				
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly Posture) N: Normal, F: Flattened, H: Hunched Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four) Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount Not calculated : NC												

Table 2 - 104

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Male Week : 6 Period : F0 mating

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	4	0	
	S.D.	2	0	
Dodecyloxirane 100 mg/kg	n	12	12	
	Mean	5	0	
	S.D.	2	0	
Dodecyloxirane 300 mg/kg	n	12	12	
	Mean	5	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	12	12	
	Mean	4	0	
	S.D.	2	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 105

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Week : 1 Period : F0 before mating

Study No. : R-1261

		Arousal NC						Convulsion NC				Species : Rat	
Test article													
Dose		-2	-1	0	1	2		0	1	2	3		
Dodecyloxirane 0 mg/kg	n	0	0	12	0	0		12	0	0	0		
Dodecyloxirane 100 mg/kg	n	0	0	12	0	0		12	0	0	0		
Dodecyloxirane 300 mg/kg	n	0	0	12	0	0		12	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	0	12	0	0		12	0	0	0		
		Abnormal NC behavior						Stereotypy NC					
Test article													
Dose		0	1	2	3			0	1	2	3		
Dodecyloxirane 0 mg/kg	n	12	0	0	0			12	0	0	0		
Dodecyloxirane 100 mg/kg	n	12	0	0	0			12	0	0	0		
Dodecyloxirane 300 mg/kg	n	12	0	0	0			12	0	0	0		
Dodecyloxirane 1000 mg/kg	n	12	0	0	0			12	0	0	0		

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe
Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe
Not calculated : NC

Table 2 - 106

Dodecylloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Week : 1 Period : F0 before mating

Study No. : R-1261

Generation : F0		Sex : Female		Week : 1		Gait NC		Posture NC		Species : Rat		
Test article	Dose											
		U	0	1	2	3	N	F	H			
Dodecyloxirane	0 mg/kg	n	0	12	0	0	0	12	0	0		
Dodecyloxirane	100 mg/kg	n	0	12	0	0	0	12	0	0		
Dodecyloxirane	300 mg/kg	n	0	12	0	0	0	12	0	0		
Dodecyloxirane	1000 mg/kg	n	0	12	0	0	0	12	0	0		
Grooming NC						Urination NC						
Test article	Dose											
		0	1	2		0	1	2	3			
Dodecyloxirane	0 mg/kg	n	12	0	0	11	1	0	0			
Dodecyloxirane	100 mg/kg	n	12	0	0	12	0	0	0			
Dodecyloxirane	300 mg/kg	n	12	0	0	12	0	0	0			
Dodecyloxirane	1000 mg/kg	n	12	0	0	12	0	0	0			
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly												
Posture) N: Normal, F: Flattened, H: Hunched												
Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)												
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount												
Not calculated : NC												

Table 2 - 107

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Female Week : 1 Period : F0 before mating

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	5	0	
	S.D.	3	0	
Dodecyloxirane 100 mg/kg	n	12	12	
	Mean	6	0	
	S.D.	2	0	
Dodecyloxirane 300 mg/kg	n	12	12	
	Mean	7	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	12	12	
	Mean	5	0	
	S.D.	2	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 108

Dodecylloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Week : 2 Period : F0 before mating

Study No. : R-1261

Generation : F0		Sex : Female		Week : 2		Arousal NC		Convulsion NC				Species : Rat	
Test article													
Dose		-2	-1	0	1	2		0	1	2	3		
Dodecylloxirane 0 mg/kg	n	0	0	12	0	0		12	0	0	0		
Dodecylloxirane 100 mg/kg	n	0	0	12	0	0		12	0	0	0		
Dodecylloxirane 300 mg/kg	n	0	0	12	0	0		12	0	0	0		
Dodecylloxirane 1000 mg/kg	n	0	0	12	0	0		12	0	0	0		
		Abnormal NC behavior					Stereotypy NC						
Test article													
Dose		0	1	2	3		0	1	2	3			
Dodecylloxirane 0 mg/kg	n	12	0	0	0		12	0	0	0			
Dodecylloxirane 100 mg/kg	n	12	0	0	0		12	0	0	0			
Dodecylloxirane 300 mg/kg	n	12	0	0	0		12	0	0	0			
Dodecylloxirane 1000 mg/kg	n	12	0	0	0		12	0	0	0			
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert													
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe													
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe													
Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe													
Not calculated : NC													

Species : Rat

199

Table 2 - 110

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Female Week : 2 Period : F0 before mating

Study No. : R-1261

		Species : Rat	
		Rearing	Defecation count
Test article			
Dose			
Dodecyloxirane 0 mg/kg	n	12	12
	Mean	6	0
	S.D.	3	0
Dodecyloxirane 100 mg/kg	n	12	12
	Mean	7	0
	S.D.	1	0
Dodecyloxirane 300 mg/kg	n	12	12
	Mean	7	0
	S.D.	1	0
Dodecyloxirane 1000 mg/kg	n	12	12
	Mean	8	0
	S.D.	1	0

DT *

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05
 DT : Dunnett test (two-side)

Table 2 - 111

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 1 Period : F0 gestation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 1						Convulsion NC						Species : Rat	
		Arousal NC															
Test article																	
Dose		-2	-1	0	1	2	0	1	2	3							
Dodecyloxirane 0 mg/kg	n	0	0	12	0	0	12	0	0	0							
Dodecyloxirane 100 mg/kg	n	0	0	10	0	0	10	0	0	0							
Dodecyloxirane 300 mg/kg	n	0	0	10	0	0	10	0	0	0							
Dodecyloxirane 1000 mg/kg	n	0	0	11	0	0	11	0	0	0							
Abnormal NC behavior						Stereotypy NC											
Test article																	
Dose		0	1	2	3	0	1	2	3								
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	0								
Dodecyloxirane 100 mg/kg	n	10	0	0	0	10	0	0	0								
Dodecyloxirane 300 mg/kg	n	10	0	0	0	10	0	0	0								
Dodecyloxirane 1000 mg/kg	n	11	0	0	0	11	0	0	0								
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe Not calculated : NC																	

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 1 Period : F0 gestation

Open field observation		Sex : Female		Day : 1		Period : 10 gestation		Species : Rat	
		Gait NC				Posture NC			
Test article									
Dose		U	0	1	2	3	N	F	H
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	12	0	0
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0	10	0	0
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0	10	0	0
Dodecyloxirane 1000 mg/kg	n	0	11	0	0	0	11	0	0
Grooming NC					Urination NC				
Test article									
Dose		0	1	2	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	10	0	0	10	0	0	0	
Dodecyloxirane 300 mg/kg	n	10	0	0	10	0	0	0	
Dodecyloxirane 1000 mg/kg	n	11	0	0	11	0	0	0	
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly Posture) N: Normal, F: Flattened, H: Hunched Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four) Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount Not calculated : NC									

Table 2 - 113

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 1 Period : F0 gestation

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	7	0	
	S.D.	2	0	
Dodecyloxirane 100 mg/kg	n	10	10	
	Mean	7	0	
	S.D.	1	0	
Dodecyloxirane 300 mg/kg	n	10	10	
	Mean	7	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	11	11	
	Mean	6	0	
	S.D.	1	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 114

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 7 Period : F0 gestation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 1		Arousal NC					Convulsion NC					Species : Rat	
Test article	Dose	n	-2	-1	0	1	2	0	1	2	3						
Dodecyloxirane	0 mg/kg	n	0	0	12	0	0	0	12	0	0	0	0				
Dodecyloxirane	100 mg/kg	n	0	0	10	0	0	10	0	0	0		0				
Dodecyloxirane	300 mg/kg	n	0	0	10	0	0	10	0	0	0		0				
Dodecyloxirane	1000 mg/kg	n	0	0	11	0	0	11	0	0	0		0				

		Abnormal NC behavior					Stereotypy NC				
Test article	Dose	n	0	1	2	3	0	1	2	3	
Dodecyloxirane	0 mg/kg	n	12	0	0	0	12	0	0	0	
Dodecyloxirane	100 mg/kg	n	10	0	0	0	10	0	0	0	
Dodecyloxirane	300 mg/kg	n	10	0	0	0	10	0	0	0	
Dodecyloxirane	1000 mg/kg	n	11	0	0	0	11	0	0	0	

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Not calculated : NC

Table 2 - 115

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 7 Period : F0 gestation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 1		Gait NC		Posture NC			Species : Rat		
Test article Dose													
		U	0	1	2	3	NC	N	F	H	NC		
Dodecyloxirane 0 mg/kg	n	0	0	12	0	0	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0	0	0	10	0	0	0	
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0	0	0	10	0	0	0	
Dodecyloxirane 1000 mg/kg	n	0	11	0	0	0	0	0	11	0	0	0	
		Grooming NC				Urination NC							
Test article Dose													
		0	1	2	NC	0	1	2	3	NC			
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	0	0	0	0	
Dodecyloxirane 100 mg/kg	n	10	0	0	0	10	0	0	0	0	0	0	
Dodecyloxirane 300 mg/kg	n	10	0	0	0	10	0	0	0	0	0	0	
Dodecyloxirane 1000 mg/kg	n	11	0	0	0	11	0	0	0	0	0	0	
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly Posture) N: Normal, F: Flattened, H: Hunched Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four) Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount Not calculated : NC													

Table 2 - 116

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 7 Period : F0 gestation

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	7	0	
	S.D.	3	0	
Dodecyloxirane 100 mg/kg	n	10	10	
	Mean	7	0	
	S.D.	2	0	
Dodecyloxirane 300 mg/kg	n	10	10	
	Mean	6	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	11	11	
	Mean	6	0	
	S.D.	1	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 117

Dodecylloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 14 Period : F0 gestation

Study No. : R-1261

Generation : F0		Sex : Female		Day : F1		Arousal NC		Convulsion NC		Species : Rat	
Test article	Dose										
		-2	-1	0	1	2	0	1	2	3	
Dodecylloxirane	n	0	0	12	0	0	12	0	0	0	0
0 mg/kg											
Dodecylloxirane	n	0	0	10	0	0	10	0	0	0	
100 mg/kg											
Dodecylloxirane	n	0	0	10	0	0	10	0	0	0	
300 mg/kg											
Dodecylloxirane	n	0	0	11	0	0	11	0	0	0	
1000 mg/kg											
		Abnormal NC behavior					Stereotypy NC				
Test article	Dose										
		0	1	2	3	0	1	2	3		
Dodecylloxirane	n	12	0	0	0	12	0	0	0	0	
0 mg/kg											
Dodecylloxirane	n	10	0	0	0	10	0	0	0		
100 mg/kg											
Dodecylloxirane	n	10	0	0	0	10	0	0	0		
300 mg/kg											
Dodecylloxirane	n	11	0	0	0	11	0	0	0		
1000 mg/kg											
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert											
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe											
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe											
Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe											
Not calculated : NC											

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 14 Period : F0 gestation

Generation : F0		Sex : Female		Day : 14		Prenatal gestation		Species : Rat	
		Gait NC				Posture NC			
Test article									
Dose		U	0	1	2	3	N	F	H
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	12	0	0
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0	10	0	0
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0	10	0	0
Dodecyloxirane 1000 mg/kg	n	0	11	0	0	0	11	0	0
Grooming NC					Urination NC				
Test article									
Dose		0	1	2	0	1	2	3	
Dodecyloxirane 0 mg/kg	n	12	0	0	12	0	0	0	
Dodecyloxirane 100 mg/kg	n	10	0	0	10	0	0	0	
Dodecyloxirane 300 mg/kg	n	10	0	0	10	0	0	0	
Dodecyloxirane 1000 mg/kg	n	11	0	0	10	1	0	0	
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly Posture) N: Normal, F: Flattened, H: Hunched Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four) Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount Not calculated : NC									

Table 2 - 119

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Female Day : 14 Period : F0 gestation

Study No. : R-1261

		Species : Rat	
		Rearing	Defecation count
Test article			
Dose			
Dodecyloxirane	n	12	12
0 mg/kg	Mean	6	0
	S.D.	2	0
Dodecyloxirane	n	10	10
100 mg/kg	Mean	5	0
	S.D.	1	0
Dodecyloxirane	n	10	10
300 mg/kg	Mean	5	0
	S.D.	1	0
Dodecyloxirane	n	11	11
1000 mg/kg	Mean	4	0
	S.D.	2	0
		DT *	

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05
 DT : Dunnett test (two-side)

Table 2 - 120

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 20 Period : F0 gestation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 20						Convulsion NC						Species : Rat	
		Arousal NC															
Test article																	
Dose		-2	-1	0	1	2	0	1	2	3							
Dodecyloxirane 0 mg/kg	n	0	0	12	0	0	12	0	0	0							
Dodecyloxirane 100 mg/kg	n	0	0	10	0	0	10	0	0	0							
Dodecyloxirane 300 mg/kg	n	0	0	10	0	0	10	0	0	0							
Dodecyloxirane 1000 mg/kg	n	0	0	11	0	0	11	0	0	0							
Abnormal NC behavior																	
Stereotypy NC																	
Test article																	
Dose		0	1	2	3	0	1	2	3								
Dodecyloxirane 0 mg/kg	n	12	0	0	0	12	0	0	0								
Dodecyloxirane 100 mg/kg	n	10	0	0	0	10	0	0	0								
Dodecyloxirane 300 mg/kg	n	10	0	0	0	10	0	0	0								
Dodecyloxirane 1000 mg/kg	n	11	0	0	0	11	0	0	0								
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe Not calculated : NC																	

Table 2 - 121

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 20 Period : F0 gestation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 20		Gait NC		Posture NC		Species : Rat	
Test article											
Dose		U	0	1	2	3	N	F	H		
Dodecyloxirane 0 mg/kg	n	0	12	0	0	0	12	0	0		
Dodecyloxirane 100 mg/kg	n	0	10	0	0	0	10	0	0		
Dodecyloxirane 300 mg/kg	n	0	10	0	0	0	10	0	0		
Dodecyloxirane 1000 mg/kg	n	0	11	0	0	0	11	0	0		
Grooming NC						Urination NC					
Test article											
Dose		0	1	2		0	1	2	3		
Dodecyloxirane 0 mg/kg	n	12	0	0		12	0	0	0		
Dodecyloxirane 100 mg/kg	n	10	0	0		10	0	0	0		
Dodecyloxirane 300 mg/kg	n	10	0	0		9	1	0	0		
Dodecyloxirane 1000 mg/kg	n	11	0	0		11	0	0	0		
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly											
Posture) N: Normal, F: Flattened, H: Hunched											
Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)											
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount											
Not calculated : NC											

Table 2 - 122

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Female Day : 20 Period : F0 gestation

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	3	0	
	S.D.	2	0	
Dodecyloxirane 100 mg/kg	n	10	10	
	Mean	4	0	
	S.D.	2	0	
Dodecyloxirane 300 mg/kg	n	10	10	
	Mean	4	0	
	S.D.	1	0	
Dodecyloxirane 1000 mg/kg	n	11	11	
	Mean	4	0	
	S.D.	1	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 123

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 6 Period : F0 lactation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 6		Dose : 0 mg/kg		Species : Rat					
		Arousal NC						Convulsion NC					
Test article	Dose												
Dodecyloxirane	n	-2	-1	0	1	2	0	0	1	2	3	0	
0 mg/kg		0	0	12	0	0	12	0	0	0	0	0	
Dodecyloxirane	n	0	0	10	0	0	10	0	0	0	0	0	
100 mg/kg													
Dodecyloxirane	n	0	0	10	0	0	10	0	0	0	0	0	
300 mg/kg													
Dodecyloxirane	n	0	0	10	0	0	10	0	0	0	0	0	
1000 mg/kg													
		Abnormal NC behavior						Stereotypy NC					
Test article	Dose												
Dodecyloxirane	n	0	1	2	3	0	0	1	2	3	0	0	
0 mg/kg		12	0	0	0	10	0	0	0	0	0	0	
Dodecyloxirane	n	10	0	0	0	10	0	0	0	0	0	0	
100 mg/kg													
Dodecyloxirane	n	10	0	0	0	10	0	0	0	0	0	0	
300 mg/kg													
Dodecyloxirane	n	10	0	0	0	10	0	0	0	0	0	0	
1000 mg/kg													
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe Not calculated : NC													

Table 2 - 124

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 6 Period : F0 lactation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 0		Gait NC			Posture NC			Species : Rat	
Test article Dose													
		U	0	1	2	3	N	F	H				
Dodecylloxirane 0 mg/kg	n	0	0	12	0	0	0	0	12	0	0		
Dodecylloxirane 100 mg/kg	n	0	10	0	0	0	0	0	10	0	0		
Dodecylloxirane 300 mg/kg	n	0	10	0	0	0	0	0	10	0	0		
Dodecylloxirane 1000 mg/kg	n	0	10	0	0	0	0	0	10	0	0		
		Grooming NC				Urination NC							
Test article Dose													
		0	1	2	0	1	2	3					
Dodecylloxirane 0 mg/kg	n	12	0	0	12	0	0	0	0	0			
Dodecylloxirane 100 mg/kg	n	10	0	0	10	0	0	0	0	0			
Dodecylloxirane 300 mg/kg	n	10	0	0	10	0	0	0	0	0			
Dodecylloxirane 1000 mg/kg	n	10	0	0	9	1	0	0	0	0			
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly													
Posture) N: Normal, F: Flattened, H: Hunched													
Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)													
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount													
Not calculated : NC													

Table 2 - 125

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 6 Period : F0 lactation

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	5	0	
	S.D.	2	0	
Dodecyloxirane 100 mg/kg	n	10	10	
	Mean	6	0	
	S.D.	2	0	
Dodecyloxirane 300 mg/kg	n	10	10	
	Mean	5	0	
	S.D.	4	0	
Dodecyloxirane 1000 mg/kg	n	10	10	
	Mean	4	0	
	S.D.	2	0	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 126

Dodecylloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 13 Period : F0 lactation

Study No. : R-1261

		Generation : F0					Sex : Female					Day : 10					Convulsion NC					Species : Rat																								
		Arousal NC																																												
Test article		-2					-1					0					1					2					3																			
Dose	n																																													
Dodecylloxirane 0 mg/kg	n	0					0					12					0					0					12					0					0					0				
Dodecylloxirane 100 mg/kg	n	0					0					10					0					0					10					0					0					0				
Dodecylloxirane 300 mg/kg	n	0					0					10					0					0					10					0					0					0				
Dodecylloxirane 1000 mg/kg	n	0					0					10					0					0					10					0					0					0				
Abnormal NC behavior															Stereotypy NC																															
Test article		0					1					2					3					0					1					2					3									
Dose	n																																													
Dodecylloxirane 0 mg/kg	n	12					0					0					0					12					0					0					0					0				
Dodecylloxirane 100 mg/kg	n	10					0					0					0					10					0					0					0					0				
Dodecylloxirane 300 mg/kg	n	10					0					0					0					10					0					0					0					0				
Dodecylloxirane 1000 mg/kg	n	10					0					0					0					10					0					0					0					0				
Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert																																														
Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe																																														
Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe																																														
Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe																																														
Not calculated : NC																																														

Table 2 - 127

Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 13 Period : F0 lactation

Study No. : R-1261

	Generation : F0	Sex : Female	Day : D5	Gait NC						Posture NC			Species : Rat
Test article Dose													
				U	0	1	2	3		N	F	H	
Dodecyloxirane 0 mg/kg	n			0	12	0	0	0		12	0	0	
Dodecyloxirane 100 mg/kg	n			0	10	0	0	0		10	0	0	
Dodecyloxirane 300 mg/kg	n			0	10	0	0	0		10	0	0	
Dodecyloxirane 1000 mg/kg	n			0	10	0	0	0		10	0	0	
				Grooming NC				Urination NC					
Test article Dose													
				0	1	2		0	1	2	3		
Dodecyloxirane 0 mg/kg	n			12	0	0		12	0	0	0		
Dodecyloxirane 100 mg/kg	n			10	0	0		10	0	0	0		
Dodecyloxirane 300 mg/kg	n			10	0	0		10	0	0	0		
Dodecyloxirane 1000 mg/kg	n			10	0	0		10	0	0	0		
Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly Posture) N: Normal, F: Flattened, H: Hunched Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four) Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount Not calculated : NC													

Table 2 - 128

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Day : 13 Period : F0 lactation

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	12	12	
	Mean	6	0	
	S.D.	2	0	
Dodecyloxirane 100 mg/kg	n	10	10	
	Mean	7	0	
	S.D.	1	0	
		DT *		
Dodecyloxirane 300 mg/kg	n	10	10	
	Mean	6	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	10	10	
	Mean	6	0	
	S.D.	2	0	

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05
DT : Dunnett test (two-side)

Table 2 - 129

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Week : 1 Period : Administration (non-mating group)

Study No. : R-1261

		Arousal NC						Convulsion NC						Species : Rat
Test article														
Dose		-2	-1	0	1	2		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	0	10	0	0		10	0	0	0			
Dodecyloxirane 1000 mg/kg	n	0	0	10	0	0		10	0	0	0			
		Abnormal NC behavior						Stereotypy NC						
Test article														
Dose		0	1	2	3			0	1	2	3			
Dodecyloxirane 0 mg/kg	n	10	0	0	0			10	0	0	0			
Dodecyloxirane 1000 mg/kg	n	10	0	0	0			10	0	0	0			
		Gait NC						Posture NC						
Test article														
Dose		U	0	1	2	3		N	F	H				
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0		10	0	0				
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0		10	0	0				

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Not calculated : NC

Table 2 - 130

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female Week : 1

Species : Rat

		Grooming NC				Urination NC			
Test article		0	1	2		0	1	2	3
Dose	n								
Dodecyloxirane 0 mg/kg	n	10	0	0		10	0	0	0
Dodecyloxirane 1000 mg/kg	n	10	0	0		10	0	0	0

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Not calculated : NC

Table 2 - 131

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Female Week : 1
 Period : Administration (non-mating group)

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	10	10	
	Mean	6	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	10	10	
	Mean	6	0	
	S.D.	2	0	

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 132

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Week : 2 Period : Administration (non-mating group)

Study No. : R-1261

		Arousal NC						Convulsion NC						Species : Rat
Test article														
Dose		-2	-1	0	1	2		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	0	10	0	0		10	0	0	0			
Dodecyloxirane 1000 mg/kg	n	0	0	10	0	0		10	0	0	0			
		Abnormal NC behavior						Stereotypy NC						
Test article														
Dose		0	1	2	3			0	1	2	3			
Dodecyloxirane 0 mg/kg	n	10	0	0	0			10	0	0	0			
Dodecyloxirane 1000 mg/kg	n	10	0	0	0			10	0	0	0			
		Gait NC						Posture NC						
Test article														
Dose		U	0	1	2	3		N	F	H				
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0		10	0	0				
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0		10	0	0				

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Not calculated : NC

Table 2 - 133

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female Week : 2

Species : Rat

		Grooming NC				Urination NC			
Test article		0	1	2		0	1	2	3
Dose	n								
Dodecyloxirane 0 mg/kg	n	10	0	0		10	0	0	0
Dodecyloxirane 1000 mg/kg	n	10	0	0		10	0	0	0

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Not calculated : NC

Table 2 - 134

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Female Week : 2 Period : Administration (non-mating group)

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	10	10	
	Mean	7	0	
	S.D.	3	0	
Dodecyloxirane 1000 mg/kg	n	10	10	
	Mean	7	0	
	S.D.	1	0	

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 135

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Week : 3 Period : Administration (non-mating group)

Study No. : R-1261

		Arousal NC						Convulsion NC						Species : Rat
Test article														
Dose		-2	-1	0	1	2		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	0	10	0	0		10	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	0	10	0	0		10	0	0	0			
		Abnormal NC behavior						Stereotypy NC						
Test article														
Dose		0	1	2	3			0	1	2	3			
Dodecyloxirane 0 mg/kg	n	10	0	0	0	0		10	0	0	0			
Dodecyloxirane 1000 mg/kg	n	10	0	0	0	10		0	0	0				
		Gait NC						Posture NC						
Test article														
Dose		U	0	1	2	3		N	F	H				
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0		10	0	0				
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0		10	0	0				

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Not calculated : NC

Table 2 - 136

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female Week : 3

Species : Rat

		Grooming NC				Urination NC			
Test article		0	1	2		0	1	2	3
Dose	n								
Dodecyloxirane 0 mg/kg	n	10	0	0		10	0	0	0
Dodecyloxirane 1000 mg/kg	n	10	0	0		10	0	0	0

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Not calculated : NC

Table 2 - 137

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Female Week : 3 Period : Administration (non-mating group)

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	10	10	
	Mean	9	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	10	10	
	Mean	7	0	
	S.D.	2	0	
		T2 *		

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

T2 : Student t-test (two-side)

Table 2 - 138

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation Period : Administration (non-mating group)
Generation : F0 Sex : Female Week : 4

Study No. : R-1261

		Arousal NC						Convulsion NC						Species : Rat
Test article														
Dose		-2	-1	0	1	2		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	0	10	0	0		10	0	0	0			
Dodecyloxirane 1000 mg/kg	n	0	0	10	0	0		10	0	0	0			
		Abnormal NC behavior						Stereotypy NC						
Test article														
Dose		0	1	2	3			0	1	2	3			
Dodecyloxirane 0 mg/kg	n	10	0	0	0			10	0	0	0			
Dodecyloxirane 1000 mg/kg	n	10	0	0	0			10	0	0	0			
		Gait NC						Posture NC						
Test article														
Dose		U	0	1	2	3		N	F	H				
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0		10	0	0				
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0		10	0	0				

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Not calculated : NC

Table 2 - 139

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female Week : 4

Species : Rat

		Grooming NC				Urination NC			
Test article		0	1	2		0	1	2	3
Dose	n								
Dodecyloxirane 0 mg/kg	n	10	0	0		10	0	0	0
Dodecyloxirane 1000 mg/kg	n	10	0	0		10	0	0	0

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Not calculated : NC

Table 2 - 140

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Female Week : 4 Period : Administration (non-mating group)

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	10	10	
	Mean	7	0	
	S.D.	3	0	
Dodecyloxirane 1000 mg/kg	n	10	10	
	Mean	6	0	
	S.D.	2	0	

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 141

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation Period : Administration (non-mating group)
Generation : F0 Sex : Female Week : 5

Study No. : R-1261

		Arousal NC						Convulsion NC						Species : Rat
Test article														
Dose		-2	-1	0	1	2		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	0	10	0	0		10	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	0	10	0	0		10	0	0	0			
		Abnormal NC behavior						Stereotypy NC						
Test article														
Dose		0	1	2	3			0	1	2	3			
Dodecyloxirane 0 mg/kg	n	10	0	0	0			10	0	0	0			
Dodecyloxirane 1000 mg/kg	n	10	0	0	0			10	0	0	0			
		Gait NC						Posture NC						
Test article														
Dose		U	0	1	2	3		N	F	H				
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0		10	0	0				
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0		10	0	0				

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Not calculated : NC

Table 2 - 142

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female Week : 5

Species : Rat

		Grooming NC				Urination NC			
Test article		0	1	2		0	1	2	3
Dose	n								
Dodecyloxirane 0 mg/kg	n	10	0	0		10	0	0	0
Dodecyloxirane 1000 mg/kg	n	10	0	0		10	0	0	0

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Not calculated : NC

Table 2 - 143

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Female Week : 5 Period : Administration (non-mating group)

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	10	10	
	Mean	7	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	10	10	
	Mean	8	0	
	S.D.	2	0	

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 144

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation Period : Administration (non-mating group)
Generation : F0 Sex : Female Week : 6

Study No. : R-1261

		Arousal NC						Convulsion NC						Species : Rat
Test article														
Dose		-2	-1	0	1	2		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	0	10	0	0		10	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	0	10	0	0		10	0	0	0			
		Abnormal NC behavior						Stereotypy NC						
Test article														
Dose		0	1	2	3			0	1	2	3			
Dodecyloxirane 0 mg/kg	n	10	0	0	0			10	0	0	0			
Dodecyloxirane 1000 mg/kg	n	10	0	0	0			10	0	0	0			
		Gait NC						Posture NC						
Test article														
Dose		U	0	1	2	3		N	F	H				
Dodecyloxirane 0 mg/kg	n	0	10	0	0	0		10	0	0				
Dodecyloxirane 1000 mg/kg	n	0	10	0	0	0		10	0	0				

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Not calculated : NC

Table 2 - 145

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female Week : 6

Species : Rat

		Grooming NC				Urination NC			
Test article		0	1	2		0	1	2	3
Dose	n								
Dodecyloxirane 0 mg/kg	n	10	0	0		10	0	0	0
Dodecyloxirane 1000 mg/kg	n	10	0	0		10	0	0	0

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Not calculated : NC

Table 2 - 146

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Week : 6 Period : Administration (non-mating group)

Study No. : R-1261

		Species : Rat	
		Rearing	Defecation count
Test article			
Dose			
Dodecyloxirane	n	10	10
0 mg/kg	Mean	8	0
	S.D.	1	0
Dodecyloxirane	n	10	10
1000 mg/kg	Mean	8	0
	S.D.	2	0

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 147

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation Period : Recovery
Generation : F0 Sex : Male Week : 1

Study No. : R-1261

		Arousal NC							Convulsion NC							Species : Rat
Test article																
Dose		-2	-1	0	1	2			0	1	2	3				
Dodecyloxirane 0 mg/kg	n	0	0	5	0	0			5	0	0	0				
Dodecyloxirane 1000 mg/kg	n	0	0	5	0	0			5	0	0	0				
		Abnormal NC behavior							Stereotypy NC							
Test article																
Dose		0	1	2	3				0	1	2	3				
Dodecyloxirane 0 mg/kg	n	5	0	0	0				5	0	0	0				
Dodecyloxirane 1000 mg/kg	n	5	0	0	0				5	0	0	0				
		Gait NC							Posture NC							
Test article																
Dose		U	0	1	2	3			N	F	H					
Dodecyloxirane 0 mg/kg	n	0	5	0	0	0			5	0	0					
Dodecyloxirane 1000 mg/kg	n	0	5	0	0	0			5	0	0					

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Not calculated : NC

Table 2 - 148

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Male Week : 1 Period : Recovery

Study No. : R-1261

		Grooming NC				Urination NC				Species : Rat
Test article		0	1	2		0	1	2	3	
Dose	n									
Dodecyloxirane	n	5	0	0		3	2	0	0	
0 mg/kg										
Dodecyloxirane	n	5	0	0		4	1	0	0	
1000 mg/kg										

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount
Not calculated : NC

Table 2 - 149

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Male Week : 1 Period : Recovery

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article				
Dose				
Dodecyloxirane 0 mg/kg	n	5	5	
	Mean	4	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	5	5	
	Mean	3	0	
	S.D.	2	0	

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 150

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation Period : Recovery
Generation : F0 Sex : Male Week : 2

Study No. : R-1261

		Arousal NC							Convulsion NC							Species : Rat
Test article																
Dose		-2	-1	0	1	2			0	1	2	3				
Dodecyloxirane 0 mg/kg	n	0	0	5	0	0			5	0	0	0				
Dodecyloxirane 1000 mg/kg	n	0	0	5	0	0			5	0	0	0				
		Abnormal NC behavior							Stereotypy NC							
Test article																
Dose		0	1	2	3				0	1	2	3				
Dodecyloxirane 0 mg/kg	n	5	0	0	0				5	0	0	0				
Dodecyloxirane 1000 mg/kg	n	5	0	0	0				5	0	0	0				
		Gait NC							Posture NC							
Test article																
Dose		U	0	1	2	3			N	F	H					
Dodecyloxirane 0 mg/kg	n	0	5	0	0	0			5	0	0					
Dodecyloxirane 1000 mg/kg	n	0	5	0	0	0			5	0	0					

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Not calculated : NC

Table 2 - 151

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Male Week : 2 Period : Recovery

Study No. : R-1261

		Grooming NC				Urination NC				Species : Rat
Test article		0	1	2		0	1	2	3	
Dose	n									
Dodecyloxirane 0 mg/kg	n	5	0	0		4	1	0	0	
Dodecyloxirane 1000 mg/kg	n	5	0	0		4	1	0	0	

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount
Not calculated : NC

Table 2 - 152

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Male Week : 2 Period : Recovery

Study No. : R-1261

		Species : Rat	
		Rearing	Defecation count
Test article	Dose		
Dodecyloxirane 0 mg/kg	n	5	5
	Mean	6	0
	S.D.	1	0
Dodecyloxirane 1000 mg/kg	n	5	5
	Mean	4	0
	S.D.	2	0
		T2 **	

Significantly different from Dodecyloxirane 0 mg/kg : ** P<0.01

T2 : Student t-test (two-side)

Table 2 - 153

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation Period : Recovery (non-mating group)
Generation : F0 Sex : Female Week : 1

Study No. : R-1261

		Arousal NC						Convulsion NC						Species : Rat
Test article														
Dose		-2	-1	0	1	2		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	0	5	0	0		5	0	0	0			
Dodecyloxirane 1000 mg/kg	n	0	0	5	0	0		5	0	0	0			
		Abnormal NC behavior						Stereotypy NC						
Test article														
Dose		0	1	2	3			0	1	2	3			
Dodecyloxirane 0 mg/kg	n	5	0	0	0			5	0	0	0			
Dodecyloxirane 1000 mg/kg	n	5	0	0	0			5	0	0	0			
		Gait NC						Posture NC						
Test article														
Dose		U	0	1	2	3		N	F	H				
Dodecyloxirane 0 mg/kg	n	0	5	0	0	0		5	0	0				
Dodecyloxirane 1000 mg/kg	n	0	5	0	0	0		5	0	0				

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Not calculated : NC

Table 2 - 154

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Week : 1 Period : Recovery (non-mating group)

Study No. : R-1261

		Grooming NC				Urination NC				Species : Rat
Test article		0	1	2		0	1	2	3	
Dose	n									
Dodecyloxirane 0 mg/kg	n	5	0	0		5	0	0	0	
Dodecyloxirane 1000 mg/kg	n	5	0	0		5	0	0	0	

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount
Not calculated : NC

Table 2 - 155

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Week : 1 Period : Recovery (non-mating group)

Study No. : R-1261

		Species : Rat	
		Rearing	Defecation count
Test article			
Dose			
Dodecyloxirane 0 mg/kg	n	5	5
	Mean	6	0
	S.D.	2	0
Dodecyloxirane 1000 mg/kg	n	5	5
	Mean	4	0
	S.D.	1	0

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 156

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation Period : Recovery (non-mating group)
Generation : F0 Sex : Female Week : 2

Study No. : R-1261

		Arousal NC						Convulsion NC						Species : Rat
Test article														
Dose		-2	-1	0	1	2		0	1	2	3			
Dodecyloxirane 0 mg/kg	n	0	0	5	0	0		5	0	0	0			
Dodecyloxirane 1000 mg/kg	n	0	0	5	0	0		5	0	0	0			
		Abnormal NC behavior						Stereotypy NC						
Test article														
Dose		0	1	2	3			0	1	2	3			
Dodecyloxirane 0 mg/kg	n	5	0	0	0			5	0	0	0			
Dodecyloxirane 1000 mg/kg	n	5	0	0	0			5	0	0	0			
		Gait NC						Posture NC						
Test article														
Dose		U	0	1	2	3		N	F	H				
Dodecyloxirane 0 mg/kg	n	0	5	0	0	0		5	0	0				
Dodecyloxirane 1000 mg/kg	n	0	5	0	0	0		5	0	0				

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Not calculated : NC

Table 2 - 157

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Open field observation
Generation : F0 Sex : Female Week : 2 Period : Recovery (non-mating group)

Study No. : R-1261

		Grooming NC				Urination NC				Species : Rat
Test article		0	1	2		0	1	2	3	
Dose	n									
Dodecyloxirane 0 mg/kg	n	5	0	0		5	0	0	0	
Dodecyloxirane 1000 mg/kg	n	5	0	0		5	0	0	0	

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)
Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount
Not calculated : NC

Table 2 - 158

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Open field observation
 Generation : F0 Sex : Female Week : 2 Period : Recovery (non-mating group)

Study No. : R-1261

		Rearing	Defecation count	Species : Rat
Test article	Dose			
Dodecyloxirane 0 mg/kg	n	5	5	
	Mean	7	0	
	S.D.	2	0	
Dodecyloxirane 1000 mg/kg	n	5	5	
	Mean	7	0	
	S.D.	1	0	

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 159

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Manipulative test
Generation : F0 Sex : Male Week : 6 Period : F0 mating

Study No. : R-1261

		Generation : F0		Sex : Male		Week : 6		Species : Rat							
		Auditory response						Approach response						Touch response	
Test article	Dose	-2	-1	0	1	-1	0	0	5	1	-1	0	0	5	1
Dodecyloxirane	0 mg/kg	n	0	0	5	0	0	5	0	0	0	5	0	5	0
Dodecyloxirane	100 mg/kg	n	0	0	5	0	0	5	0	0	0	5	0	5	0
Dodecyloxirane	300 mg/kg	n	0	0	5	0	0	5	0	0	0	5	0	5	0
Dodecyloxirane	1000 mg/kg	n	0	0	5	0	0	5	0	0	0	5	0	5	0
		Tail pinch response						Pupillary reflex							
Test article	Dose	-2	-1	0	1	P	F	L	R						
Dodecyloxirane	0 mg/kg	n	0	0	5	0	5	0	0	0					
Dodecyloxirane	100 mg/kg	n	0	0	5	0	5	0	0	0					
Dodecyloxirane	300 mg/kg	n	0	0	5	0	5	0	0	0					
Dodecyloxirane	1000 mg/kg	n	0	0	5	0	5	0	0	0					
		Aerial right-ing reflex-total													
Test article	Dose	0	1	2	3	4	5	6							
Dodecyloxirane	0 mg/kg	n	5	0	0	0	0	0							
Dodecyloxirane	100 mg/kg	n	5	0	0	0	0	0							
Dodecyloxirane	300 mg/kg	n	5	0	0	0	0	0							
Dodecyloxirane	1000 mg/kg	n	5	0	0	0	0	0							

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-total) Total score of 3 times

Table 2 - 160

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Manipulative test
Generation : F0 Sex : Female Day : 13
Period : F0 lactation

Study No. : R-1261

Generation : F0		Sex : Female		Day : 13		Species : Rat							
Test article Dose		Auditory response				Approach response				Touch response			
		-2	-1	0	1	-1	0	1	-1	0	1		
Dodecyloxirane 0 mg/kg	n	0	0	5	0	0	5	0	0	5	0		
Dodecyloxirane 100 mg/kg	n	0	0	5	0	0	5	0	0	5	0		
Dodecyloxirane 300 mg/kg	n	0	0	5	0	0	5	0	0	5	0		
Dodecyloxirane 1000 mg/kg	n	0	0	5	0	0	5	0	0	5	0		
Test article Dose		Tail pinch response				Pupillary reflex							
		-2	-1	0	1	P	F	L	R				
Dodecyloxirane 0 mg/kg	n	0	0	5	0	5	0	0	0				
Dodecyloxirane 100 mg/kg	n	0	0	5	0	5	0	0	0				
Dodecyloxirane 300 mg/kg	n	0	0	5	0	5	0	0	0				
Dodecyloxirane 1000 mg/kg	n	0	0	5	0	5	0	0	0				
Test article Dose		Aerial right-ing reflex-total											
		0	1	2	3	4	5	6					
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	0	0					
Dodecyloxirane 100 mg/kg	n	5	0	0	0	0	0	0					
Dodecyloxirane 300 mg/kg	n	5	0	0	0	0	0	0					
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	0	0					

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate
Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction
Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction
Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate
Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds
Aerial righting reflex-total) Total score of 3 times

Table 2 - 161

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Manipulative test Period : Administration (non-mating group)
Generation : F0 Sex : Female Week : 6

Study No. : R-1261

		Auditory response					Approach response					Touch response				
Test article		-2	-1	0	1		-1	0	1			-1	0	1		
Dodecyloxirane	n	0	0	5	0		0	5	0			0	5	0		
0 mg/kg																
Dodecyloxirane	n	0	0	5	0		0	5	0			0	5	0		
1000 mg/kg																
		Tail pinch response					Pupillary reflex									
Test article		-2	-1	0	1		P	F	L	R						
Dodecyloxirane	n	0	0	5	0		5	0	0	0						
0 mg/kg																
Dodecyloxirane	n	0	0	5	0		5	0	0	0						
1000 mg/kg																
		Aerial right-ing reflex-total														
Test article		0	1	2	3	4	5	6								
Dodecyloxirane	n	5	0	0	0	0	0	0								
0 mg/kg																
Dodecyloxirane	n	5	0	0	0	0	0	0								
1000 mg/kg																

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-total) Total score of 3 times

Table 2 - 162

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Manipulative test
Generation : F0 Sex : Male Week : 2
Period : Recovery

Study No. : R-1261

		Auditory response					Approach response					Touch response				
Test article		-2	-1	0	1		-1	0	1			-1	0	1		
Dodecyloxirane	n	0	0	5	0		0	5	0			0	5	0		
0 mg/kg																
Dodecyloxirane	n	0	0	5	0		0	5	0			0	5	0		
1000 mg/kg																
		Tail pinch response					Pupillary reflex									
Test article		-2	-1	0	1		P	F	L	R						
Dodecyloxirane	n	0	0	5	0		5	0	0	0						
0 mg/kg																
Dodecyloxirane	n	0	0	5	0		5	0	0	0						
1000 mg/kg																
		Aerial right-ing reflex-total														
Test article		0	1	2	3	4	5	6								
Dodecyloxirane	n	5	0	0	0	0	0	0								
0 mg/kg																
Dodecyloxirane	n	5	0	0	0	0	0	0								
1000 mg/kg																

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-total) Total score of 3 times

Table 2 - 163

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Manipulative test
Generation : F0 Sex : Female Week : 2 Period : Recovery (non-mating group)

Study No. : R-1261

Manipulative test		Sex : Female		Week : 2		Period : Recovery (non mating group)								Species : Rat	
		Auditory response						Approach response				Touch response			
Test article															
Dose		-2	-1	0	1	0	-1	0	1	0	-1	0	1	0	
Dodecyloxirane 0 mg/kg	n	0	0	5	0	0	0	5	0	0	0	5	0	0	
Dodecyloxirane 1000 mg/kg	n	0	0	5	0	0	0	5	0	0	0	5	0	0	
		Tail pinch response						Pupillary reflex							
Test article		-2	-1	0	1	0	P	F	L	R					
Dose		-2	-1	0	1	0	P	F	L	R					
Dodecyloxirane 0 mg/kg	n	0	0	5	0	0	5	0	0	0					
Dodecyloxirane 1000 mg/kg	n	0	0	5	0	0	5	0	0	0					
		Aerial right-ing reflex-total													
Test article		0	1	2	3	4	5	6							
Dose		0	1	2	3	4	5	6							
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	0	0							
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	0	0							

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-total) Total score of 3 times

Table 2 - 164

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Grip strength
Generation : F0 Sex : Male Week : 6 Period : F0 mating

Study No. : R-1261

		Species : Rat	
Test article Dose		Grip strength of fore limb g	Grip strength of hind limb g
Dodecyloxirane 0 mg/kg	n	5	5
	Mean	1419	688
	S.D.	203	190
Dodecyloxirane 100 mg/kg	n	5	5
	Mean	1361	702
	S.D.	199	67
Dodecyloxirane 300 mg/kg	n	5	5
	Mean	1310	619
	S.D.	124	90
Dodecyloxirane 1000 mg/kg	n	5	5
	Mean	1442	698
	S.D.	149	104
Not significantly different from Dodecyloxirane 0 mg/kg			

Table 2 - 165

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Grip strength
Generation : F0 Sex : Female Day : 13
Period : F0 lactation

Study No. : R-1261

Test article Dose		Grip strength of fore limb g	Grip strength of hind limb g	Species : Rat
Dodecyloxirane 0 mg/kg	n	5	5	
	Mean	1092	495	
	S.D.	260	130	
Dodecyloxirane 100 mg/kg	n	5	5	
	Mean	884	394	
	S.D.	107	63	
Dodecyloxirane 300 mg/kg	n	5	5	
	Mean	1159	606	
	S.D.	272	181	
Dodecyloxirane 1000 mg/kg	n	5	5	
	Mean	1048	505	
	S.D.	235	138	
Not significantly different from Dodecyloxirane 0 mg/kg				

Table 2 - 166

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Grip strength

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Species : Rat

Test article Dose		Grip strength of fore limb g	Grip strength of hind limb g
Dodecyloxirane 0 mg/kg	n	5	5
	Mean	1181	628
	S.D.	300	149
Dodecyloxirane 1000 mg/kg	n	5	5
	Mean	1243	693
	S.D.	177	54

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 167

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Grip strength
Generation : F0 Sex : Male Week : 2 Period : Recovery

Study No. : R-1261

		Species : Rat	
		Grip strength of fore limb g	Grip strength of hind limb g
Test article			
Dose			
Dodecyloxirane	n	5	5
0 mg/kg	Mean	1802	938
	S.D.	169	185
Dodecyloxirane	n	5	5
1000 mg/kg	Mean	1633	784
	S.D.	194	205

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 168

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Grip strength
 Generation : F0 Sex : Female Week : 2 Period : Recovery (non-mating group)

Study No. : R-1261

		Species : Rat	
		Grip strength of fore limb g	Grip strength of hind limb g
Test article			
Dose			
Dodecyloxirane 0 mg/kg	n	5	5
	Mean	1295	726
	S.D.	176	116
Dodecyloxirane 1000 mg/kg	n	5	5
	Mean	1431	620
	S.D.	182	77

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 169

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Motor activity
Generation : F0 Sex : Male Week : 6 Period : F0 mating

Study No. : R-1261

		0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')	Species : Rat
Test article		Count	Count	Count	Count	Count	Count	Count	
Dose									
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	
	Mean	412	327	213	163	67	40	1221	
	S.D.	44	107	104	81	74	36	347	
Dodecyloxirane 100 mg/kg	n	5	5	5	5	5	5	5	
	Mean	425	341	266	161	220	136	1548	
	S.D.	27	35	115	117	131	132	365	
Dodecyloxirane 300 mg/kg	n	5	5	5	5	5	5	5	
	Mean	394	318	193	82	43	60	1089	
	S.D.	99	91	115	107	87	53	439	
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	
	Mean	432	285	106	101	88	66	1077	
	S.D.	74	135	84	72	145	65	191	
Not significantly different from Dodecyloxirane 0 mg/kg									

Table 2 - 170

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Motor activity
Generation : F0 Sex : Female Day : 13
Period : F0 lactation

Study No. : R-1261

		Species : Rat						
		0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Test article		Count	Count	Count	Count	Count	Count	Count
Dose								
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5
	Mean	405	262	264	237	222	224	1614
	S.D.	36	78	85	110	92	98	322
Dodecyloxirane 100 mg/kg	n	5	5	5	5	5	5	5
	Mean	420	145	83	209	160	129	1146
	S.D.	50	95	99	56	150	107	331
DT *								
Dodecyloxirane 300 mg/kg	n	5	5	5	5	5	5	5
	Mean	376	169	199	97	161	102	1105
	S.D.	48	75	113	127	163	125	586
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5
	Mean	357	189	176	86	49	85	942
	S.D.	71	165	86	127	96	157	583

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

DT : Dunnett test (two-side)

Table 2 - 171

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

		Period : Administration (non-mating group)						Species : Rat
Motor activity		Sex : Female	Week : 6					
Generation : F0		0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Test article		Count	Count	Count	Count	Count	Count	Count
Dose								
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5
	Mean	311	261	208	167	112	141	1200
	S.D.	110	101	102	176	175	171	752
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5
	Mean	327	233	193	63	111	264	1191
	S.D.	28	91	38	97	28	53	234

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 172

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Motor activity
Generation : F0 Sex : Male Week : 2 Period : Recovery

Study No. : R-1261

		0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')	Species : Rat
Test article		Count	Count	Count	Count	Count	Count	Count	
Dose									
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	
	Mean	371	257	149	41	12	58	888	
	S.D.	67	73	125	70	15	107	377	
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	
	Mean	410	312	223	124	84	53	1206	
	S.D.	57	47	100	89	81	33	302	

Not significantly different from Dodecyloxirane 0 mg/kg

Table 2 - 173

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Motor activity
Generation : F0 Sex : Female Week : 2 Period : Recovery (non-mating group)

Study No. : R-1261

	Motor activity Generation : F0	Sex : Female	Week : 2	Period : Recovery (non-mating group)					Species : Rat
	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')		
Test article Dose	Count	Count	Count	Count	Count	Count	Count		
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	
	Mean	319	209	184	48	57	93	910	
	S.D.	46	127	96	61	119	127	451	
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	
	Mean	342	293	212	195	144	67	1253	
	S.D.	79	178	201	98	130	109	573	
T2 *									

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

T2 : Student t-test (two-side)

Table 3 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Body weight

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Sex : Male

Unit : g

Species : Rat

Test article		/Day							Body weight
Dose		1	8	15	22	29	36	42	gain
Dodecyloxirane 0 mg/kg	n	12	12	12	12	12	12	12	12
	Mean	384	419	448	467	497	519	530	146
	S.D.	23	22	26	24	27	32	33	26
Dodecyloxirane 100 mg/kg	n	12	12	12	12	12	12	12	12
	Mean	383	420	451	471	498	517	531	148
	S.D.	18	25	32	37	40	44	44	30
Dodecyloxirane 300 mg/kg	n	12	12	12	12	12	12	12	12
	Mean	382	418	448	466	490	511	524	142
	S.D.	19	24	29	29	33	41	44	33
Dodecyloxirane 1000 mg/kg	n	12	12	12	12	12	12	12	12
	Mean	381	412	437	454	480	498	513	131
	S.D.	21	23	26	26	29	30	35	23
Not significantly different from Dodecyloxirane 0 mg/kg									

Table 3 - 2

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Body weight
Sex : Female
Period : F0 before mating Day 1-15

Study No. : R-1261

Test article		/Day			Body weight	Unit : g	Species : Rat
Dose		1	8	15	gain		
Dodecyloxirane 0 mg/kg	n	12	12	12	12		
	Mean	250	260	268	18		
	S.D.	17	19	19	8		
Dodecyloxirane 100 mg/kg	n	12	12	12	12		
	Mean	252	264	274	22		
	S.D.	14	17	18	11		
Dodecyloxirane 300 mg/kg	n	12	12	12	12		
	Mean	250	259	270	20		
	S.D.	14	14	17	12		
Dodecyloxirane 1000 mg/kg	n	12	12	12	12		
	Mean	251	264	275	24		
	S.D.	18	20	20	9		
Not significantly different from Dodecyloxirane 0 mg/kg							

Table 3 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Body weight
Sex : Female
Period : F0 gestation Day 0-25

Study No. : R-1261

		/Day				Unit : g	Species : Rat
Test article	Dose	0	7	14	20	Body weight gain	
Dodecyloxirane 0 mg/kg	n	12	12	12	12	12	
	Mean	281	319	356	434	153	
	S.D.	21	24	28	40	34	
Dodecyloxirane 100 mg/kg	n	10	10	10	10	10	
	Mean	283	317	354	444	161	
	S.D.	18	25	25	30	18	
Dodecyloxirane 300 mg/kg	n	10	10	10	10	10	
	Mean	289	333	375	464	175	
	S.D.	19	19	26	35	25	
Dodecyloxirane 1000 mg/kg	n	11	11	11	11	11	
	Mean	287	322	358	430	143	
	S.D.	29	27	25	42	38	
Not significantly different from Dodecyloxirane 0 mg/kg							

Table 3 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Body weight
Sex : Female
Period : F0 lactation Day 0-14

Study No. : R-1261

		/Day				Unit : g	Species : Rat
Test article	Dose	0	4	7	13	Body weight gain	
Dodecyloxirane 0 mg/kg	n	12	12	12	12	12	
	Mean	350	353	351	368	18	
	S.D.	29	25	26	26	11	
Dodecyloxirane 100 mg/kg	n	10	10	10	10	10	
	Mean	353	356	355	371	18	
	S.D.	25	23	21	21	16	
Dodecyloxirane 300 mg/kg	n	10	10	10	10	10	
	Mean	376	378	377	388	13	
	S.D.	27	32	32	29	18	
Dodecyloxirane 1000 mg/kg	n	10	10	10	10	10	
	Mean	347	354	352	365	17	
	S.D.	30	22	25	23	13	
Not significantly different from Dodecyloxirane 0 mg/kg							

Table 3 - 5

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Body weight
 Sex : Female
 Period : Administration (non-mating group)

Study No. : R-1261

		Unit : g								Species : Rat
Test article	/Day	Body weight								
Dose		1	8	15	22	29	36	42	gain	
Dodecyloxirane 0 mg/kg	n	10	10	10	10	10	10	10	10	
	Mean	250	263	272	284	297	303	307	58	
	S.D.	14	13	12	19	21	21	20	13	
Dodecyloxirane 1000 mg/kg	n	10	10	10	10	10	10	10	10	
	Mean	253	267	275	285	298	305	308	55	
	S.D.	15	17	19	19	19	25	22	11	

Not significantly different from Dodecyloxirane 0 mg/kg

Table 3 - 6

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Body weight
 Sex : Male
 Period : Recovery Day 1-15

Study No. : R-1261

		Unit : g			Species : Rat	
Test article Dose		/Day			Body weight	
		1	8	14	gain	
Dodecyloxirane 0 mg/kg	n	5	5	5	5	
	Mean	546	560	572	27	
	S.D.	25	30	26	8	
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	
	Mean	525	545	559	34	
	S.D.	44	46	47	8	

Not significantly different from Dodecyloxirane 0 mg/kg

Table 3 - 7

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Body weight
 Sex : Female
 Period : Recovery Day 1-15 (non-mating group)

Study No. : R-1261

		/Day			Body weight		Unit : g	Species : Rat
Test article	Dose	1	8	14	gain			
Dodecyloxirane 0 mg/kg	n	5	5	5	5			
	Mean	307	316	316	10			
	S.D.	18	19	19	6			
Dodecyloxirane 1000 mg/kg	n	5	5	5	5			
	Mean	302	306	312	11			
	S.D.	19	16	15	7			

Not significantly different from Dodecyloxirane 0 mg/kg

Table 4 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption
Sex : Male
Period : F0 before mating Day 1-15

Study No. : R-1261

		/Day			Total food consumption	Unit : g	Species : Rat
Test article Dose		2	8	15			
Dodecyloxirane 0 mg/kg	n	12	12	12	12		
	Mean	20	22	22	64		
	S.D.	2	3	4	6		
Dodecyloxirane 100 mg/kg	n	12	12	12	12		
	Mean	21	22	22	64		
	S.D.	2	3	4	9		
Dodecyloxirane 300 mg/kg	n	12	12	12	12		
	Mean	21	22	21	64		
	S.D.	2	3	4	8		
Dodecyloxirane 1000 mg/kg	n	12	12	12	12		
	Mean	21	21	22	64		
	S.D.	2	2	3	5		
Not significantly different from Dodecyloxirane 0 mg/kg							

Table 4 - 2

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption Period : F0 mating Day 15-43
Sex : Male

Study No. : R-1261

		Food consumption			Period : 14 mating Day 10-15		Unit : g	Species : Rat
Test article		/Day						
Dose		30	36	42				
Dodecyloxirane 0 mg/kg	n	12	12	12				
	Mean	20	23	21				
	S.D.	2	2	2				
Dodecyloxirane 100 mg/kg	n	12	12	12				
	Mean	21	22	20				
	S.D.	3	3	3				
Dodecyloxirane 300 mg/kg	n	12	12	12				
	Mean	20	22	21				
	S.D.	3	4	3				
Dodecyloxirane 1000 mg/kg	n	12	12	12				
	Mean	21	22	21				
	S.D.	2	2	2				
Not significantly different from Dodecyloxirane 0 mg/kg								

Table 4 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption
Sex : Female
Period : F0 before mating Day 1-15

Study No. : R-1261

		/Day			Total food consumption	Unit : g	Species : Rat
Test article Dose		2	8	15			
Dodecyloxirane 0 mg/kg	n	12	12	12	12		
	Mean	12	14	14	40		
	S.D.	3	2	3	5		
Dodecyloxirane 100 mg/kg	n	12	12	12	12		
	Mean	14	15	15	43		
	S.D.	3	3	2	5		
Dodecyloxirane 300 mg/kg	n	12	12	12	12		
	Mean	14	14	15	43		
	S.D.	3	2	3	6		
Dodecyloxirane 1000 mg/kg	n	12	12	12	12		
	Mean	14	16	15	45		
	S.D.	3	3	3	6		
Not significantly different from Dodecyloxirane 0 mg/kg							

Table 4 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption Period : F0 gestation Day 0-25
Sex : Female

Study No. : R-1261

		/Day				Unit : g	Species : Rat
Test article	Dose	1	7	14	20	Total food consumption	
Dodecyloxirane 0 mg/kg	n	12	12	12	12	12	
	Mean	16	20	20	22	78	
	S.D.	3	4	3	4	12	
Dodecyloxirane 100 mg/kg	n	10	10	10	10	10	
	Mean	15	20	22	25	82	
	S.D.	7	3	3	3	11	
Dodecyloxirane 300 mg/kg	n	10	10	10	10	10	
	Mean	18	22	24	26	89	
	S.D.	3	3	4	5	11	
				DT *		DT *	
Dodecyloxirane 1000 mg/kg	n	11	11	11	11	11	
	Mean	17	20	22	22	81	
	S.D.	2	3	2	3	6	

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

DT : Dunnett test (two-side)

Table 4 - 5

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption
Sex : Female
Period : F0 lactation Day 0-14

Study No. : R-1261

		Unit : g					Species : Rat
Test article	/Day	Total food consumption					
Dose		2	4	7	13		
Dodecyloxirane	n	12	12	12	12	12	
0 mg/kg	Mean	23	37	38	57	156	
	S.D.	5	6	5	5	17	
Dodecyloxirane	n	10	10	10	10	10	
100 mg/kg	Mean	25	39	40	58	162	
	S.D.	5	6	4	7	18	
Dodecyloxirane	n	10	10	10	10	10	
300 mg/kg	Mean	26	39	40	59	164	
	S.D.	4	5	5	5	15	
Dodecyloxirane	n	10	10	10	10	10	
1000 mg/kg	Mean	25	37	40	53	156	
	S.D.	3	6	7	6	18	
Not significantly different from Dodecyloxirane 0 mg/kg							

Table 4 - 6

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Food consumption
 Sex : Female
 Period : Administration Day 1-15(non-mating group)

Study No. : R-1261

		/Day			Total food consumption	Unit : g	Species : Rat
Test article	Dose	2	8	15			
Dodecyloxirane 0 mg/kg	n	10	10	10	10		
	Mean	12	14	15	42		
	S.D.	2	3	2	3		
Dodecyloxirane 1000 mg/kg	n	10	10	10	10		
	Mean	13	16	15	44		
	S.D.	2	2	2	4		

Not significantly different from Dodecyloxirane 0 mg/kg

Table 4 - 7

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption
Sex : Female
Period : Administration Day 15-43(non-mating group)

Study No. : R-1261

		Unit : g			Species : Rat	
Test article		/Day				
Dose		30	36	42		
Dodecyloxirane 0 mg/kg	n	10	10	10		
	Mean	14	16	15		
	S.D.	2	2	2		
Dodecyloxirane 1000 mg/kg	n	10	10	10		
	Mean	15	17	15		
	S.D.	2	2	2		

Not significantly different from Dodecyloxirane 0 mg/kg

Table 4 - 8

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Food consumption
 Sex : Male
 Period : Recovery Day 1-15

Study No. : R-1261

		Unit : g			Species : Rat
Test article	/Day				Total food
Dose		2	8	14	consumption
Dodecyloxirane	n	5	5	5	5
0 mg/kg	Mean	22	26	25	73
	S.D.	1	3	2	6
Dodecyloxirane	n	5	5	5	5
1000 mg/kg	Mean	24	28	26	78
	S.D.	3	2	2	6

Not significantly different from Dodecyloxirane 0 mg/kg

Table 4 - 9

Dodecyloxirane: An oral combined repeated dose toxicity study
 with the reproduction/ developmental toxicity screening test in rats
 Food consumption
 Sex : Female
 Period : Recovery Day 1-15 (non-mating group)

Study No. : R-1261

		/Day			Total food consumption	Unit : g	Species : Rat
Test article	Dose	2	8	14			
Dodecyloxirane 0 mg/kg	n	5	5	5	5		
	Mean	16	19	18	53		
	S.D.	4	1	2	6		
Dodecyloxirane 1000 mg/kg	n	5	5	5	5		
	Mean	15	19	19	53		
	S.D.	2	3	2	3		

Not significantly different from Dodecyloxirane 0 mg/kg

Table 5 - 1

Dodecylloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Sex : Male

Study No. : R-1261

	Sex : Male	Stage : Week 8 of administration						Species : Rat			
		pH									
Test article											
Dose		5.0	5.5	6.0	6.5	7.0	7.5	8.0	8.5	>=9.0	
Dodecyloxirane 0 mg/kg	n	0	0	0	0	1	0	0	3	1	
Dodecyloxirane 100 mg/kg	n	0	0	0	0	0	1	2	2	0	
Dodecyloxirane 300 mg/kg	n	0	0	0	0	1	2	1	1	0	
Dodecyloxirane 1000 mg/kg	n	0	0	0	1	0	0	1	2	1	
		Protein					Ketones				
Test article		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+
Dodecyloxirane 0 mg/kg	n	1	0	3	1	0	0	0	4	1	0
Dodecyloxirane 100 mg/kg	n	2	1	1	1	0	0	2	1	2	0
Dodecyloxirane 300 mg/kg	n	0	2	3	0	0	0	0	4	1	0
Dodecyloxirane 1000 mg/kg	n	0	0	1	4	0	0	0	0	4	1
Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL											

Table 5 - 2

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Stage : Week 6 of administration
Sex : Male

Study No. : R-1261

Sex : Male		Glucose						Occult Blood					Species : Rat		
Test article															
Dose		-	1+	2+	3+	4+		-	+/-	1+	2+	3+			
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0		4	1	0	0	0			
Dodecyloxirane 100 mg/kg	n	5	0	0	0	0		2	3	0	0	0			
Dodecyloxirane 300 mg/kg	n	5	0	0	0	0		4	1	0	0	0			
Dodecyloxirane 1000 mg/kg	n	3	2	0	0	0		4	1	0	0	0			
Bilirubin							Urobilinogen								
Test article															
Dose		-	1+	2+	3+		+/-	1+	2+	3+					
Dodecyloxirane 0 mg/kg	n	4	1	0	0		5	0	0	0					
Dodecyloxirane 100 mg/kg	n	5	0	0	0		5	0	0	0					
Dodecyloxirane 300 mg/kg	n	3	2	0	0		5	0	0	0					
Dodecyloxirane 1000 mg/kg	n	2	3	0	0		5	0	0	0					
Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL															
Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL															
Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL															
Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL															

Table 5 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Stage : Week 6 of administration
Sex : Male

Study No. : R-1261

		Color											Species : Rat
Test article		LY	Y	DY	BR	RE	RB	WH	BL	CL	OT		
Dodecyloxirane Dose	n	0	5	0	0	0	0	0	0	0	0	0	
Dodecyloxirane 0 mg/kg	n	0	5	0	0	0	0	0	0	0	0	0	
Dodecyloxirane 100 mg/kg	n	0	5	0	0	0	0	0	0	0	0	0	
Dodecyloxirane 300 mg/kg	n	0	5	0	0	0	0	0	0	0	0	0	
Dodecyloxirane 1000 mg/kg	n	0	5	0	0	0	0	0	0	0	0	0	
Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other													

Table 5 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Stage : Week 6 of administration (non-mating group)
Sex : Female

Study No. : R-1261

Sex : Female		pH										Species : Rat	
Test article													
Dose		5.0	5.5	6.0	6.5	7.0	7.5	8.0	8.5	>=9.0			
Dodecyloxirane 0 mg/kg	n	0	0	1	2	1	0	0	1	0			
Dodecyloxirane 1000 mg/kg	n	0	0	1	4	0	0	0	0	0			
Protein													
Test article							Ketones						
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	1	3	0	1	0	1	1	3	0	0		
Dodecyloxirane 1000 mg/kg	n	1	1	3	0	0	0	0	0	3	2		
Glucose													
Test article							Occult Blood						
Dose		-	1+	2+	3+	4+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	5	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	5	0	0	0	0		

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Table 5 - 5

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Sex : Female

Stage : Week 6 of administration (non-mating group)

Study No. : R-1261

		Bilirubin					Urobilinogen					Species : Rat
Test article		-	1+	2+	3+		+/-	1+	2+	3+		
Dose												
Dodecyloxirane 0 mg/kg	n	5	0	0	0		5	0	0	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0		5	0	0	0		
		Color										
Test article		LY	Y	DY	BR		RE	RB	WH	BL	CL	OT
Dose												
Dodecyloxirane 0 mg/kg	n	0	5	0	0		0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	n	0	5	0	0		0	0	0	0	0	0

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Table 5 - 6

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Sex : Male

Study No. : R-1261

Sex : Male		RBC					WBC					Species : Rat	
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	5	0	0	0	0	0	
Dodecyloxirane 100 mg/kg	n	5	0	0	0	0	5	0	0	0	0	0	
Dodecyloxirane 300 mg/kg	n	5	0	0	0	0	5	0	0	0	0	0	
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	5	0	0	0	0	0	
Squamous epithelial cells						Small round epithelial cells							
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	0	5	0	0	0	5	0	0	0	0	0	
Dodecyloxirane 100 mg/kg	n	0	5	0	0	0	5	0	0	0	0	0	
Dodecyloxirane 300 mg/kg	n	0	5	0	0	0	5	0	0	0	0	0	
Dodecyloxirane 1000 mg/kg	n	0	5	0	0	0	5	0	0	0	0	0	
RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe													

Table 5 - 7

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Sex : Male

Study No. : R-1261

Stage 1: Week 1 of administration											
Sex : Male		Cast					Species : Rat				
Test article											
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	3	2	0	0	0
Dodecyloxirane 100 mg/kg	n	5	0	0	0	0	4	1	0	0	0
Dodecyloxirane 300 mg/kg	n	5	0	0	0	0	5	0	0	0	0
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	3	2	0	0	0
Crystal calcium oxalate											
Test article											
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0					
Dodecyloxirane 100 mg/kg	n	5	0	0	0	0					
Dodecyloxirane 300 mg/kg	n	5	0	0	0	0					
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0					
Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe											
Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe											
Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe											

Table 5 - 8

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Stage : Week 6 of administration (non-mating group)
Sex : Female

Study No. : R-1261

		RBC					WBC					Species : Rat	
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	5	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	5	0	0	0	0		
		Squamous epithelial cells					Small round epithelial cells						
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	0	5	0	0	0	5	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	5	0	0	0	5	0	0	0	0		
		Cast					Crystal phosphate salts						
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	4	1	0	0	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	4	1	0	0	0		

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Table 5 - 9

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Stage : Week 6 of administration (non-mating group)
Sex : Female

Study No. : R-1261

		Crystal calcium oxalate						Species : Rat
Test article								
Dose		-	+/-	1+	2+	3+		
Dodecyloxirane	n	5	0	0	0	0		
0 mg/kg								
Dodecyloxirane	n	5	0	0	0	0		
1000 mg/kg								

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Table 5 - 10

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Stage : Week 6 of administration
Sex : Male

Study No. : R-1261

		Species : Rat					
Test article Dose		Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5
	Mean	42	13.4	1424	1.4	2.9	2.0
	S.D.	10	5.0	416	0.5	0.7	0.6
Dodecyloxirane 100 mg/kg	n	5	5	5	5	5	5
	Mean	39	16.6	1548	1.8	3.6	2.6
	S.D.	14	10.4	533	0.6	0.9	0.6
Dodecyloxirane 300 mg/kg	n	5	5	5	5	5	5
	Mean	37	11.1	1743	1.3	3.3	2.0
	S.D.	5	2.2	256	0.3	1.1	0.6
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5
	Mean	36	7.9	2180	1.1	2.6	1.7
	S.D.	4	2.0	211	0.3	0.6	0.4

DT *

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

DT : Dunnett test (two-side)

Table 5 - 11

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Urinalysis

Stage : Week 6 of administration (non-mating group)

Sex : Female

Species : Rat

		Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
Test article	Dose						
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5
	Mean	33	8.9	1694	0.9	1.9	1.3
	S.D.	10	4.4	806	0.3	0.5	0.4
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5
	Mean	31	5.7	1943	0.9	1.6	1.1
	S.D.	5	2.1	32	0.4	0.6	0.5

Not significantly different from Dodecyloxirane 0 mg/kg

Table 5 - 12

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Sex : Male
Stage : Week 2 of recovery

Study No. : R-1261

		pH										Species : Rat
Test article												
Dose		5.0	5.5	6.0	6.5	7.0	7.5	8.0	8.5	>=9.0		
Dodecyloxirane 0 mg/kg	n	0	0	0	0	0	0	0	0	3	2	
Dodecyloxirane 1000 mg/kg	n	0	0	0	0	1	0	0	0		4	
		Protein					Ketones					
Test article												
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+	
Dodecyloxirane 0 mg/kg	n	2	1	2	0	0	1	3	1	0	0	
Dodecyloxirane 1000 mg/kg	n	0	0	4	1	0	0	0	3	2	0	
		Glucose					Occult Blood					
Test article												
Dose		-	1+	2+	3+	4+	-	+/-	1+	2+	3+	
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	1	3	1	0	0	
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	5	0	0	0	0	

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Table 5 - 13

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Sex : Male
Stage : Week 2 of recovery

Study No. : R-1261

		Bilirubin					Urobilinogen					Species : Rat
Test article												
Dose		-	1+	2+	3+		+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	4	1	0	0		5	0	0	0		
Dodecyloxirane 1000 mg/kg	n	3	2	0	0		5	0	0	0		
		Color										
Test article												
Dose		LY	Y	DY	BR		RE	RB	WH	BL	CL	OT
Dodecyloxirane 0 mg/kg	n	0	5	0	0		0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	n	0	5	0	0		0	0	0	0	0	0

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Table 5 - 14

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Sex : Female
Stage : Week 2 of recovery (non-mating group)

Study No. : R-1261

Sex : Female		pH										Species : Rat	
Test article													
Dose		5.0	5.5	6.0	6.5	7.0	7.5	8.0	8.5	>=9.0			
Dodecyloxirane 0 mg/kg	n	0	0	0	1	1	2	0	0	1			
Dodecyloxirane 1000 mg/kg	n	0	0	0	0	0	0	0	3	2			
Protein							Ketones						
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	1	2	1	1	0	2	2	1	0	0		
Dodecyloxirane 1000 mg/kg	n	2	1	2	0	0	2	3	0	0	0		
Glucose						Occult Blood							
Test article													
Dose		-	1+	2+	3+	4+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	4	1	0	0	0	5	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	4	1	0	0	0	5	0	0	0	0		

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Table 5 - 15

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Sex : Female
Stage : Week 2 of recovery (non-mating group)

Study No. : R-1261

		Bilirubin					Urobilinogen					Species : Rat
Test article												
Dose		-	1+	2+	3+		+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	4	1	0	0		5	0	0	0		
Dodecyloxirane 1000 mg/kg	n	4	1	0	0		5	0	0	0		
		Color										
Test article												
Dose		LY	Y	DY	BR		RE	RB	WH	BL	CL	OT
Dodecyloxirane 0 mg/kg	n	0	5	0	0		0	0	0	0	0	0
Dodecyloxirane 1000 mg/kg	n	0	5	0	0		0	0	0	0	0	0

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Table 5 - 16

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Sex : Male
Stage : Week 2 of recovery

Study No. : R-1261

		RBC					WBC					Species : Rat	
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	5	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	5	0	0	0	0		
		Squamous epithelial cells					Small round epithelial cells						
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	0	5	0	0	0	5	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	5	0	0	0	5	0	0	0	0		
		Cast					Crystal phosphate salt						
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	4	1	0	0	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	3	2	0	0	0		

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Table 5 - 17

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Stage : Week 2 of recovery
Sex : Male

Study No. : R-1261

		Crystal calcium oxalate						Species : Rat
Test article								
Dose		-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0		
Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe								

Table 5 - 18

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Sex : Female
Stage : Week 2 of recovery (non-mating group)

Study No. : R-1261

		RBC					WBC					Species : Rat	
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	5	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	5	0	0	0	0		
		Squamous epithelial cells					Small round epithelial cells						
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	0	5	0	0	0	5	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	0	5	0	0	0	5	0	0	0	0		
		Cast					Crystal phosphate salt						
Test article													
Dose		-	+/-	1+	2+	3+	-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0	4	1	0	0	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0	4	1	0	0	0		

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Table 5 - 19

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Stage : Week 2 of recovery (non-mating group)
Sex : Female

Study No. : R-1261

		Crystal calcium oxalate						Species : Rat
Test article								
Dose		-	+/-	1+	2+	3+		
Dodecyloxirane 0 mg/kg	n	5	0	0	0	0		
Dodecyloxirane 1000 mg/kg	n	5	0	0	0	0		

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Table 5 - 20

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Stage : Week 2 of recovery
Sex : Male

Study No. : R-1261

		Species : Rat					
		Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
Test article	Dose						
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5
	Mean	57	19.1	1586	2.0	4.2	2.7
	S.D.	19	12.7	452	0.6	1.0	0.7
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5
	Mean	41	11.5	2296	1.9	4.2	2.7
	S.D.	4	2.0	182	0.3	0.6	0.4

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

T2 : Student t-test (two-side)

Table 5 - 21

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Urinalysis
Stage : Week 2 of recovery (non-mating group)
Sex : Female

Study No. : R-1261

		Species : Rat					
		Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
Test article	Dose						
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5
	Mean	37	8.0	1852	1.0	2.1	1.4
	S.D.	4	2.3	481	0.2	0.2	0.2
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5
	Mean	33	7.5	1998	1.1	2.2	1.6
	S.D.	2	2.0	254	0.2	0.5	0.3

Not significantly different from Dodecyloxirane 0 mg/kg

Table 6 - 1

Dodecylloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Hematology
Sex : Male
Stage : End of administration

Study No. : R-1261

Hematology Sex : Male		Stage : End of administration							Species : Rat		
		RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC
Test article		10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL
Dose											
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	864	15.3	44.7	51.7	17.7	34.2	12.5	193.2	103.4	74.7
	S.D.	16	0.1	0.9	1.5	0.3	0.5	0.4	28.5	10.1	20.9
Dodecyloxirane 100 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	825	14.8	43.0	52.1	18.0	34.5	12.6	174.3	109.8	72.4
	S.D.	44	0.5	1.6	1.9	0.6	0.3	0.3	20.7	4.5	14.2
Dodecyloxirane 300 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	825	15.2	44.2	53.6	18.4	34.4	12.2	179.1	105.5	93.4
	S.D.	11	0.3	1.6	2.5	0.6	0.5	0.2	37.2	7.8	19.4
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	838	15.1	44.0	52.5	18.1	34.4	12.7	158.3	109.0	71.9
	S.D.	20	0.3	0.9	1.2	0.5	0.5	0.7	26.2	18.8	8.1
		#LYMP	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB	
Test article		10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL	
Dose											
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	57.6	14.1	1.0	0.1	1.6	0.4	13.4	16.0	320	
	S.D.	14.8	6.4	0.6	0.1	0.8	0.2	0.7	1.7	23	
Dodecyloxirane 100 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	57.6	12.0	1.0	0.1	1.4	0.3	13.8	15.0	337	
	S.D.	15.2	2.7	0.4	0.0	0.3	0.2	0.9	2.5	33	
Dodecyloxirane 300 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	74.6	14.7	1.2	0.1	2.1	0.6	13.0	15.0	291	
	S.D.	16.0	4.4	0.4	0.1	1.0	0.2	0.7	1.7	21	
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	54.7	14.1	0.9	0.1	1.9	0.3	13.7	15.0	297	
	S.D.	9.7	3.5	0.1	0.0	0.9	0.1	0.7	0.6	27	
Not significantly different from Dodecyloxirane 0 mg/kg											

Table 6 - 2

Dodecylloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Hematology
Sex : Female
Stage : End of lactation (F0 dams)

Study No. : R-1261

Hematology Sex : Female		Stage : End of incubation (16 days)							Species : Rat		
		RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC
Test article		10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL
Dose											
Dodecylxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	738	15.2	43.2	58.5	20.6	35.2	13.6	263.8	124.7	99.1
	S.D.	16	0.9	2.1	2.5	1.1	1.0	0.2	39.5	10.6	32.3
Dodecylxirane 100 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	756	15.1	43.4	57.4	20.0	34.9	13.3	266.6	129.6	90.3
	S.D.	16	0.3	1.2	1.2	0.3	0.8	0.4	38.7	14.3	21.2
Dodecylxirane 300 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	745	15.7	44.0	59.1	21.1	35.7	13.3	266.4	122.2	89.6
	S.D.	37	0.5	1.4	1.5	0.5	0.9	0.7	28.5	16.1	35.2
Dodecylxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	736	15.8	43.6	59.3	21.4	36.2	14.4	224.9	116.4	93.3
	S.D.	34	0.6	1.1	1.4	0.2	0.6	0.7	30.8	9.6	24.3
		#LYMP	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB	
Test article		10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL	
Dose											
Dodecylxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	45.0	48.8	0.9	0.1	3.3	1.0	12.6	14.3	515	
	S.D.	18.3	17.7	0.5	0.1	2.2	0.7	1.1	1.7	120	
Dodecylxirane 100 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	43.7	41.7	0.7	0.1	3.3	0.9	12.7	14.0	451	
	S.D.	12.5	8.5	0.1	0.1	0.9	0.5	1.6	2.8	64	
Dodecylxirane 300 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	48.6	36.1	0.8	0.1	3.3	0.7	12.4	13.9	445	
	S.D.	27.0	8.4	0.4	0.1	1.9	0.8	0.8	1.1	96	
Dodecylxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	46.4	40.5	0.9	0.1	4.5	0.8	12.0	12.9	386	
	S.D.	15.5	10.5	0.3	0.1	2.6	0.8	1.6	1.5	82	
Not significantly different from Dodecylxirane 0 mg/kg											

Table 6 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Hematology
Sex : Female

Stage : End of administration (non-mating group)

Sex : Female		Stage : End of administration (non mating group)							Species : Rat		
		RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC
Test article		10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL
Dose											
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	793	14.5	41.6	52.5	18.3	34.7	11.9	175.9	110.4	70.8
	S.D.	22	0.4	1.1	1.3	0.4	0.7	0.5	32.7	11.0	10.1
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	764	14.5	42.2	55.2	19.0	34.4	11.8	159.7	112.3	60.5
	S.D.	30	0.4	0.8	1.3	0.2	0.4	0.4	40.8	9.0	19.3
					T2 *	T2 **					
		#LYMP	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB	
Test article		10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL	
Dose											
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	59.4	8.9	0.8	0.1	1.2	0.5	13.7	13.3	216	
	S.D.	8.8	2.7	0.3	0.0	0.4	0.2	0.8	1.4	21	
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	49.1	8.4	0.6	0.0	2.0	0.4	13.5	12.4	207	
	S.D.	17.5	2.0	0.2	0.1	0.9	0.2	0.6	1.0	12	

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05, ** P<0.01

T2 : Student t-test (two-side)

Table 6 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Hematology
Sex : Male
Stage : End of recovery

Study No. : R-1261

		Stage : End of recovery								Species : Rat	
Hematology		RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC
Sex : Male											
Test article		10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL
Dose											
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	876	15.8	44.7	50.9	18.0	35.4	12.5	153.2	103.1	69.4
	S.D.	39	1.0	2.8	1.0	0.4	0.6	0.4	20.5	11.4	15.2
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	844	15.5	43.9	52.1	18.4	35.3	12.9	177.1	107.9	77.2
	S.D.	42	0.7	1.6	1.3	0.6	0.4	0.6	25.7	8.3	6.8
		#LYMP	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB	
Test article		10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL	
Dose											
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	53.1	12.7	1.2	0.1	1.9	0.5	14.5	16.1	319	
	S.D.	13.8	2.7	0.3	0.1	0.6	0.1	0.5	2.4	19	
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	58.9	14.4	1.4	0.1	2.1	0.4	14.0	19.0	312	
	S.D.	7.6	1.9	0.1	0.0	0.5	0.2	0.4	0.8	27	
		A2 *									

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

A2 : Welch test (two-side)

A2 *

Table 6 - 5

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Hematology
Sex : Female
Stage : End of recovery (non-mating group)

Study No. : R-1261

Hematology		Stage : End of recovery (non mating group)							Species : Rat		
Sex : Female		RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC
Test article		10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL
Dose											
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	761	15.4	41.6	54.6	20.2	37.0	11.5	149.8	103.9	59.3
	S.D.	30	0.7	1.7	1.4	0.6	0.4	0.9	27.5	13.3	24.1
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	747	15.3	41.2	55.3	20.6	37.2	12.1	164.1	87.6	36.7
	S.D.	57	0.5	1.7	2.5	1.1	0.4	0.8	24.1	48.1	10.8
		#LYMP	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB	
Test article		10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL	
Dose											
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	46.7	9.5	1.0	0.1	1.6	0.3	14.5	14.0	218	
	S.D.	19.1	5.4	0.4	0.1	0.8	0.2	1.3	1.7	20	
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	29.5	5.1	0.7	0.0	1.1	0.2	14.8	14.0	232	
	S.D.	8.5	2.5	0.3	0.0	0.8	0.1	0.9	2.2	26	
Not significantly different from Dodecyloxirane 0 mg/kg											

Not significantly different from Dodecyloxirane 0 mg/kg

Table 7 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Blood chemistry
Sex : Male
Stage : End of administration

Study No. : R-1261

		Species : Rat								
		AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL
		IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL
Test article	Dose									GLU
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	60	42	626	0	44.2	65	32	106	0.1
	S.D.	8	8	238	0	14.2	14	11	15	0.0
Dodecyloxirane 100 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	64	41	583	0	27.0	64	43	105	0.1
	S.D.	13	9	139	0	15.7	10	22	13	0.0
Dodecyloxirane 300 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	58	39	523	0	37.2	78	56	121	0.1
	S.D.	9	7	152	0	19.7	11	22	9	0.0
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	69	62	585	0	27.4	82	32	119	0.1
	S.D.	5	8	174	0	13.9	3	11	6	0.0
DT **										
		BUN	CRNN	Na	K	Cl	Ca	P	TP	ALB
		mg/dL	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
Test article	Dose									
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	15	0.28	144	3.8	105	10.5	5.8	6.4	3.3
	S.D.	3	0.02	2	0.1	1	0.3	0.4	0.3	0.1
Dodecyloxirane 100 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	17	0.30	143	3.8	107	10.3	5.4	6.2	3.2
	S.D.	2	0.03	1	0.1	1	0.2	0.3	0.2	0.1
Dodecyloxirane 300 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	15	0.29	143	3.7	105	10.5	5.8	6.4	3.3
	S.D.	2	0.06	1	0.1	1	0.1	0.6	0.1	0.0
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	20	0.31	143	3.7	107	10.0	5.6	6.0	3.2
	S.D.	2	0.03	1	0.2	1	0.1	0.5	0.2	0.1
		DT *				DT *		DT **		DT *

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05, ** P<0.01

DT : Dunnett test (two-side)

Table 7 - 2

Dodecylloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Blood chemistry
Sex : Female
Stage : End of lactation (F0 dams)

Study No. : R-1261

Sex : Female		Stage 1 End Point Evaluation (n = 22/mg)								Species : Rat	
		AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU
Test article		IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
Dose											
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	126	75	382	1	51.1	102	96	194	0.1	129
	S.D.	37	12	91	0	14.9	13	16	19	0.0	5
Dodecyloxirane 100 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	97	66	420	1	54.4	93	94	177	0.1	129
	S.D.	31	21	40	0	22.8	15	26	20	0.0	14
Dodecyloxirane 300 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	107	74	400	1	50.0	108	125	191	0.1	142
	S.D.	38	23	107	0	32.9	13	97	23	0.0	15
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	5
	Mean	98	71	430	1	66.4	107	112	192	0.1	137
	S.D.	39	34	94	0	53.9	20	42	31	0.0	18
		BUN	CRNN	Na	K	Cl	Ca	P	TP	ALB	
Test article		mg/dL	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL	
Dose											
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	31	0.41	142	3.3	101	11.0	9.9	6.6	3.4	
	S.D.	8	0.07	3	0.5	2	0.3	1.5	0.5	0.3	
Dodecyloxirane 100 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	29	0.37	142	3.1	101	10.9	9.0	6.5	3.4	
	S.D.	8	0.06	1	0.4	1	0.4	1.0	0.3	0.2	
Dodecyloxirane 300 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	31	0.39	140	3.6	102	11.0	9.5	6.4	3.4	
	S.D.	10	0.08	3	0.6	3	0.5	1.1	0.4	0.3	
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	
	Mean	25	0.38	142	3.5	103	11.3	9.5	6.5	3.4	
	S.D.	8	0.05	2	0.4	3	0.5	1.6	0.5	0.2	
Not significantly different from Dodecyloxirane 0 mg/kg											

Table 7 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Blood chemistry
Stage : End of administration (non-mating group)
Sex : Female

Study No. : R-1261

		Species : Rat								
		AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL
		IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL
Test article	Dose									
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	51	27	293	0	24.5	72	34	134	0.1
	S.D.	8	5	112	0	13.9	11	7	16	0.0
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	54	35	267	0	18.2	68	16	117	0.1
	S.D.	4	3	93	0	6.2	13	6	16	0.0
		T2 *				T2 **				
		BUN	CRNN	Na	K	Cl	Ca	P	TP	ALB
		mg/dL	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	15	0.30	144	3.5	107	10.3	5.1	6.6	3.8
	S.D.	3	0.04	0	0.3	1	0.3	0.5	0.2	0.1
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	18	0.31	142	3.6	108	10.2	5.5	6.3	3.7
	S.D.	2	0.01	0	0.1	1	0.2	0.1	0.2	0.2
		T2 **								

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05, ** P<0.01

T2 : Student t-test (two-side)

Table 7 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Blood chemistry
Sex : Male
Stage : End of recovery

Study No. : R-1261

		Species : Rat								
		AST	ALT	ALP	γ -GTP	T-BA	T-CHO	TG	PL	T-BIL
		IU/L	IU/L	IU/L	IU/L	μ mol/L	mg/dL	mg/dL	mg/dL	mg/dL
Test article	Dose									
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	60	32	334	1	43.3	73	63	111	0.1
	S.D.	14	8	33	1	50.8	15	20	18	0.0
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	55	33	320	1	27.2	76	63	114	0.1
	S.D.	8	10	36	1	8.2	12	22	13	0.0
		BUN	CRNN	Na	K	Cl	Ca	P	TP	ALB
		mg/dL	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	20	0.28	145	3.8	107	10.4	5.5	6.5	3.3
	S.D.	4	0.02	1	0.2	1	0.2	0.4	0.3	0.1
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	22	0.28	145	3.9	106	10.5	5.8	6.5	3.2
	S.D.	5	0.02	1	0.3	1	0.2	0.5	0.1	0.1

Not significantly different from Dodecyloxirane 0 mg/kg

Table 7 - 5

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Blood chemistry
Sex : Female
Stage : End of recovery (non-mating group)

Study No. : R-1261

		Species : Rat								
		AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL
		IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL
Test article	Dose									GLU
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	54	30	199	0	57.2	79	61	166	0.1
	S.D.	6	4	90	1	33.8	11	42	24	0.1
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	52	26	173	1	15.4	75	64	152	0.1
	S.D.	4	4	73	0	6.3	16	58	28	0.0
		A2 *								
		BUN	CRNN	Na	K	Cl	Ca	P	TP	ALB
		mg/dL	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	22	0.36	142	3.5	106	10.5	4.5	6.9	3.9
	S.D.	4	0.05	1	0.3	1	0.5	0.6	0.7	0.5
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5
	Mean	20	0.35	143	3.6	108	10.2	4.2	6.7	3.8
	S.D.	2	0.05	1	0.3	1	0.3	0.5	0.5	0.4
		T2 *								

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

A2 : Welch test (two-side), T2 : Student t-test (two-side)

Table 8 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration

Study No. : R-1261

		Body weight		Brain		Thyroid-RL		Thymus		Heart	
Test article				AB	RE	AB	RE	AB	RE	AB	RE
Dose		g		g	g/100g	mg	mg/100g	mg	mg/100g	g	g/100g
Dodecyloxirane	n	7		7	7	7	7	7	7	7	7
0 mg/kg	Mean	509		2.18	0.43	20.9	4.1	307	60	1.45	0.29
	S.D.	33		0.09	0.03	3.6	0.7	86	17	0.12	0.01
Dodecyloxirane	n	12		12	12	12	12	12	12	12	12
100 mg/kg	Mean	522		2.17	0.42	21.4	4.1	338	65	1.40	0.27
	S.D.	42		0.08	0.04	3.7	0.6	63	13	0.09	0.02
Dodecyloxirane	n	12		12	12	12	12	12	12	12	12
300 mg/kg	Mean	513		2.14	0.42	22.7	4.4	357	70	1.42	0.28
	S.D.	42		0.15	0.04	3.9	0.8	107	21	0.15	0.02
Dodecyloxirane	n	7		7	7	7	7	7	7	7	7
1000 mg/kg	Mean	487		2.13	0.44	22.6	4.6	310	64	1.33	0.27
	S.D.	23		0.06	0.03	4.4	0.9	70	14	0.07	0.01
		Liver		Spleen		Kidney-RL		Adrenal -RL		Testis-RL	
Test article		AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
Dose		g	g/100g	g	g/100g	g	g/100g	mg	mg/100g	g	g/100g
Dodecyloxirane	n	7		7	7	7	7	7	7	7	7
0 mg/kg	Mean	13.68	2.68	0.72	0.14	3.23	0.64	55	11	3.26	0.64
	S.D.	1.17	0.10	0.10	0.02	0.25	0.05	7	2	0.26	0.07
Dodecyloxirane	n	12		12	12	12	12	12	12	12	12
100 mg/kg	Mean	14.42	2.76	0.67	0.13	3.31	0.63	53	10	3.43	0.66
	S.D.	1.81	0.20	0.07	0.01	0.35	0.04	7	1	0.22	0.06
Dodecyloxirane	n	12		12	12	12	12	12	12	12	12
300 mg/kg	Mean	14.62	2.84	0.72	0.14	3.33	0.65	54	11	3.29	0.64
	S.D.	2.27	0.21	0.14	0.02	0.44	0.07	5	1	0.42	0.09
Dodecyloxirane	n	7		7	7	7	7	7	7	7	7
1000 mg/kg	Mean	14.41	2.96	0.68	0.14	3.28	0.67	56	12	3.36	0.69
	S.D.	0.91	0.12	0.10	0.02	0.27	0.03	6	1	0.20	0.04

AB : Absolute weight, RE : Relative weight by body weight
Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05
DT : Dunnett test (two-side)

Table 8 - 2

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration

Study No. : R-1261

Sex : Male		Species : Rat									
		Epididymis-RL		Prostate		Seminal vesicle		LABC		Cowper's gland-RL	
Test article		AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
Dose		mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	mg	mg/100g
Dodecyloxirane	n	7	7	7	7	7	7	7	7	7	7
0 mg/kg	Mean	1242	245	1.40	0.28	1.79	0.35	1.63	0.32	123.0	24.3
	S.D.	75	23	0.31	0.07	0.40	0.08	0.17	0.05	33.0	7.3
Dodecyloxirane	n	12	12	12	12	12	12	12	12	12	12
100 mg/kg	Mean	1306	252	1.22	0.24	1.63	0.31	1.65	0.32	121.9	23.5
	S.D.	62	21	0.11	0.03	0.29	0.05	0.23	0.03	35.7	6.5
Dodecyloxirane	n	12	12	12	12	12	12	12	12	12	12
300 mg/kg	Mean	1319	258	1.22	0.24	1.58	0.31	1.51	0.30	123.0	24.1
	S.D.	179	36	0.20	0.05	0.20	0.04	0.16	0.05	26.9	5.5
Dodecyloxirane	n	7	7	7	7	7	7	7	7	7	7
1000 mg/kg	Mean	1278	263	1.24	0.26	1.58	0.33	1.51	0.31	126.7	26.2
	S.D.	38	14	0.19	0.04	0.22	0.04	0.13	0.02	17.7	4.7
Glans penis											
		AB	RE								
		mg	mg/100g								
Test article	n	7	7								
Dose	Mean	148.0	29.2								
	S.D.	26.4	5.8								
Dodecyloxirane	n	12	12								
100 mg/kg	Mean	136.6	26.4								
	S.D.	35.0	7.3								
Dodecyloxirane	n	12	12								
300 mg/kg	Mean	130.2	25.5								
	S.D.	18.7	3.8								
Dodecyloxirane	n	7	7								
1000 mg/kg	Mean	109.8	22.6								
	S.D.	14.5	3.3								
DT *											

AB : Absolute weight, RE : Relative weight by body weight

LABC: Levator ani and bulbocavernosus muscle

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

DT : Dunnett test (two-side)

Table 8 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Female

Stage : End of lactation (F0 dams)

Study No. : R-1261

		Body weight		Brain		Thyroid-RL		Thymus		Heart		Species : Rat
Test article		g		AB	RE	AB	RE	AB	RE	AB	RE	
Dose				g	g/100g	mg	mg/100g	mg	mg/100g	g	g/100g	
Dodecyloxirane	n	12		12	12	12	12	12	12	12	12	
0 mg/kg	Mean	343		2.02	0.59	17.3	5.0	246	72	1.12	0.33	
	S.D.	26		0.08	0.05	2.6	0.7	67	18	0.08	0.02	
Dodecyloxirane	n	10		10	10	10	10	10	10	10	10	
100 mg/kg	Mean	346		2.02	0.59	17.4	5.0	235	69	1.15	0.33	
	S.D.	17		0.08	0.04	2.2	0.6	60	19	0.12	0.04	
Dodecyloxirane	n	10		10	10	10	10	10	10	10	10	
300 mg/kg	Mean	362		2.02	0.56	18.1	5.0	252	69	1.18	0.33	
	S.D.	27		0.07	0.04	3.4	0.8	62	16	0.16	0.04	
Dodecyloxirane	n	10		10	10	10	10	10	10	10	10	
1000 mg/kg	Mean	337		2.05	0.61	17.2	5.1	253	76	1.15	0.34	
	S.D.	24		0.08	0.04	2.7	0.8	65	21	0.09	0.03	
		Liver		Spleen		Kidney-RL		Adrenal-RL		Ovary-RL		
Test article		AB	RE	AB	RE	AB	RE	AB	RE	AB	RE	
Dose		g	g/100g	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	
Dodecyloxirane	n	12		12	12	12	12	12	12	12	12	
0 mg/kg	Mean	11.68		3.42	0.58	0.17	2.27	0.67	73	21	92.1	26.9
	S.D.	0.85		0.22	0.08	0.02	0.18	0.04	8	3	9.0	2.7
Dodecyloxirane	n	10		10	10	10	10	10	10	10	10	
100 mg/kg	Mean	12.14		3.52	0.61	0.18	2.35	0.68	67	19	89.6	25.9
	S.D.	0.51		0.12	0.07	0.02	0.16	0.04	8	2	11.9	3.0
Dodecyloxirane	n	10		10	10	10	10	10	10	10	10	
300 mg/kg	Mean	12.97		3.59	0.67	0.19	2.35	0.65	78	22	92.8	25.8
	S.D.	1.00		0.19	0.09	0.03	0.18	0.07	6	1	12.7	4.1
	DT **				DT *							
Dodecyloxirane	n	10		10	10	10	10	10	10	10	10	
1000 mg/kg	Mean	13.24		3.93	0.58	0.17	2.27	0.67	76	23	91.3	27.1
	S.D.	1.31		0.22	0.10	0.02	0.24	0.06	9	2	11.8	3.2
	DT **			DT **								

AB : Absolute weight, RE : Relative weight by body weight

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05, ** P<0.01

DT : Dunnett test (two-side)

Table 8 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Female

Stage : End of lactation (F0 dams)

Study No. : R-1261

Species : Rat

		Uterus	
Test article Dose		AB	RE
		mg	mg/100g
Dodecyloxirane 0 mg/kg	n	12	12
	Mean	519	152
	S.D.	96	28
Dodecyloxirane 100 mg/kg	n	10	10
	Mean	527	152
	S.D.	95	24
Dodecyloxirane 300 mg/kg	n	10	10
	Mean	577	161
	S.D.	88	31
Dodecyloxirane 1000 mg/kg	n	10	10
	Mean	547	162
	S.D.	120	30

AB : Absolute weight, RE : Relative weight by body weight
Not significantly different from Dodecyloxirane 0 mg/kg

Table 8 - 5

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Female

Stage : End of administration (non-mating group)

Study No. : R-1261

		Body weight		Brain		Thyroid-RL		Thymus		Heart	
Test article		AB		RE		AB		RE		AB	
Dose		g		g/100g		mg		mg/100g		g	
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
0 mg/kg	Mean	302	2.03	0.67	18.5	6.2	343	114	0.96	0.32	
	S.D.	22	0.09	0.05	3.5	1.5	70	26	0.06	0.02	
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
1000 mg/kg	Mean	302	1.97	0.66	16.1	5.3	285	96	0.87	0.29	
	S.D.	28	0.05	0.05	3.3	0.7	69	29	0.07	0.01	
A2 *											
		Liver		Spleen		Kidney-RL		Adrenal-RL		Ovary-RL	
Test article		AB		RE		AB		RE		AB	
Dose		g		g/100g		g		g/100g		mg	
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
0 mg/kg	Mean	8.10	2.68	0.54	0.18	1.94	0.64	63	21	88.3	29.2
	S.D.	0.51	0.04	0.08	0.04	0.16	0.05	4	1	15.0	4.4
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
1000 mg/kg	Mean	9.01	2.99	0.48	0.16	1.97	0.65	64	21	89.2	29.5
	S.D.	0.74	0.15	0.07	0.02	0.12	0.04	4	1	10.4	0.8
A2 **											
		Uterus									
Test article		AB		RE							
Dose		mg		mg/100g							
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
0 mg/kg	Mean	679	226	679	226	679	226	679	226	679	226
	S.D.	190	66	190	66	190	66	190	66	190	66
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
1000 mg/kg	Mean	666	224	666	224	666	224	666	224	666	224
	S.D.	249	94	249	94	249	94	249	94	249	94

AB : Absolute weight, RE : Relative weight by body weight

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05, ** P<0.01

A2 : Welch test (two-side)

Table 8 - 6

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of recovery

Study No. : R-1261

		Body weight		Brain		Thyroid-RL		Thymus		Heart	
Test article		AB		RE		AB		RE		AB	
Dose		g		g/100g		mg		mg/100g		g	
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
0 mg/kg	Mean	550	2.16	0.39	22.5	4.1	330	60	1.55	0.28	
	S.D.	27	0.06	0.02	4.1	0.6	32	8	0.09	0.02	
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
1000 mg/kg	Mean	534	2.20	0.41	24.0	4.5	324	60	1.52	0.29	
	S.D.	45	0.04	0.04	3.8	0.9	64	8	0.18	0.03	
		Liver		Spleen		Kidney-RL		Adrenal -RL		Testis-RL	
Test article		AB		RE		AB		RE		AB	
Dose		g		g/100g		g		mg/100g		g	
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
0 mg/kg	Mean	14.37	2.62	0.77	0.14	3.53	0.64	65	12	3.26	0.59
	S.D.	0.79	0.17	0.11	0.02	0.10	0.04	13	2	0.22	0.03
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
1000 mg/kg	Mean	14.69	2.74	0.77	0.14	3.62	0.68	63	12	3.34	0.63
	S.D.	1.69	0.11	0.09	0.01	0.44	0.05	12	3	0.37	0.10
		Epididymis-RL		Prostate		Seminal vesicle		LABC		Cowper's gland-RL	
Test article		AB		RE		AB		RE		AB	
Dose		mg		mg/100g		g		g/100g		mg	
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
0 mg/kg	Mean	1399	255	1.47	0.27	1.92	0.35	1.71	0.31	154.2	28.0
	S.D.	134	27	0.26	0.04	0.19	0.04	0.13	0.02	19.4	3.3
Dodecyloxirane	n	5	5	5	5	5	5	5	5	5	5
1000 mg/kg	Mean	1359	256	1.35	0.25	1.66	0.32	1.60	0.30	136.5	25.6
	S.D.	54	32	0.16	0.04	0.24	0.06	0.13	0.04	26.6	5.1

AB : Absolute weight, RE : Relative weight by body weight

LABC: Levator ani and bulbocavernosus muscle

Not significantly different from Dodecyloxirane 0 mg/kg

Table 8 - 7

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of recovery

Study No. : R-1261

Species : Rat

Glans penis			
Test article		AB	RE
Dose		mg	mg/100g
Dodecyloxirane	n	5	5
0 mg/kg	Mean	123.7	22.4
	S.D.	26.1	4.2
Dodecyloxirane	n	5	5
1000 mg/kg	Mean	145.8	27.4
	S.D.	31.3	5.8

AB : Absolute weight, RE : Relative weight by body weight
Not significantly different from Dodecyloxirane 0 mg/kg

Table 8 - 8

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Female

Stage : End of recovery (non-mating group)

Study No. : R-1261

		Body weight		Brain		Thyroid-RL		Thymus		Heart		Species : Rat	
				AB	RE	AB	RE	AB	RE	AB	RE		
Test article		g		g	g/100g	mg	mg/100g	mg	mg/100g	g	g/100g		
Dose													
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	5		
	Mean	304	2.01	0.66	15.6	5.1	250	82	1.03	0.34			
	S.D.	19	0.10	0.04	1.3	0.4	41	11	0.10	0.03			
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	5		
	Mean	294	2.04	0.69	20.2	6.9	301	102	0.94	0.32			
	S.D.	15	0.05	0.05	1.7	0.5	80	25	0.02	0.02			
						T2 **	T2 **						
		Liver		Spleen		Kidney-RL		Adrenal -RL		Ovary-RL			
				AB	RE	AB	RE	AB	RE	AB	RE		
Test article		g		g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g		
Dose													
Dodecyloxirane 0 mg/kg	n	5	5	5	5	5	5	5	5	5	5		
	Mean	7.84	2.57	0.46	0.15	1.96	0.65	68	22	86.4	28.8		
	S.D.	1.14	0.25	0.05	0.01	0.22	0.05	2	1	21.2	8.1		
Dodecyloxirane 1000 mg/kg	n	5	5	5	5	5	5	5	5	5	5		
	Mean	7.25	2.46	0.46	0.16	2.01	0.68	65	22	84.9	28.8		
	S.D.	0.54	0.09	0.04	0.02	0.13	0.01	7	2	12.7	3.3		
		Uterus											
				AB	RE								
Test article		mg		mg	mg/100g								
Dose													
Dodecyloxirane 0 mg/kg	n	5	5										
	Mean	619	203										
	S.D.	139	42										
Dodecyloxirane 1000 mg/kg	n	5	5										
	Mean	686	232										
	S.D.	185	58										

AB : Absolute weight, RE : Relative weight by body weight
Significantly different from Dodecyloxirane 0 mg/kg : ** P<0.01
T2 : Student t-test (two-side)

Table 9-1 Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats
Gross pathological findings (End of dosing)

Organs	Sex:	M	M	M	M
	Dose (mg/kg) :	0	100	300	1000
Findings	Number:	7	12	12	7
All tissues					
Not remarkable		7	12	12	7

M : Male

Table 9-2 Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats
Gross pathological findings (End of dosing)

Organs	Sex:	F	F
	Dose (mg/kg) :	0	1000
Findings	Number:	5	5
All tissues			
Not remarkable		5	5

F : Female

Table 9-3 Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats
Gross pathological findings (End of dosing,dams)

Organs	Sex:	F	F	F	F
	Dose (mg/kg) :	0	100	300	1000
Findings	Number:	12	10	10	10
Stomach					
Focus, dark red, glandular stomach		1	2	2	0

F : Female

Table 9-4 Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats
Gross pathological findings (End of recovery)

Organs	Sex:	M	M
	Dose (mg/kg) :	0	1000
Findings	Number:	5	5
All tissues			
Not remarkable		5	5

M : Male

Table 9-5 Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats
Gross pathological findings (End of recovery)

Organs	Sex:	F	F
	Dose (mg/kg) :	0	1000
Findings	Number:	5	5
All tissues			
Not remarkable		5	5

F : Female

Table 9-6 Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats
Gross pathological findings (Undelivered)

Organs	Sex:	F	F	F
	Dose (mg/kg) :	100	300	1000
Findings	Number:	2	2	2
All tissues				
Not remarkable		2	2	2

F : Female

Table 10-1 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of dosing)

Organs	Sex:	M	M	M	M
Findings	Dose (mg/kg) : Number:	0 7	100 12	300 12	1000 7
Adrenal					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Bone, femur					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Bone marrow, femur					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Brain (cerebrum, cerebellum, pons)					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Coagulating gland					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Epididymis					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Eye					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Heart					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Intestine, cecum					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Intestine, colon					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Intestine, duodenum					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Intestine, ileum					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Intestine, jejunum					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Intestine, rectum					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Kidney					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Liver					
Number examined		5	2	2	5
Not remarkable		5	2	2	5

M : Male

Table 10-2 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of dosing)

Organs	Sex:	M	M	M	M
Findings	Dose (mg/kg) : Number:	0 7	100 12	300 12	1000 7
Lung (bronchus)					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Lymph node, mesenteric					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Lymph node, submandibular					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Mammary gland, inguinal					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Medulla oblongata					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Muscle, quadriceps femoris					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Optic nerve					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Pancreas					
Number examined		5	2	2	5
Not remarkable		4	2	2	5
Focus, basophilic		1	0	0	0
minimal		1	0	0	0
Parathyroid					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Peyer's patch, ileal					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Pituitary					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Prostate					
Number examined		5	2	2	5
Not remarkable		4	1	1	2
Infiltrate, inflammatory cell		1	1	1	3
minimal		1	1	1	3
Salivary gland, submandibular					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Sciatic nerve					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Seminal vesicle					
Number examined		5	2	2	5
Not remarkable		5	2	2	5

M : Male

Table 10-3 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of dosing)

Organs	Sex:	M	M	M	M
Findings	Dose (mg/kg) : Number:	0 7	100 12	300 12	1000 7
Skin, inguinal					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Spinal cord, thoracic					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Spleen					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Stomach					
Number examined		5	2	2	5
Not remarkable		5	2	2	4
Erosion/ulcer, glandular stomach		0	0	0	1
minimal		0	0	0	1
Testis					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Thymus					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Thyroid					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Trachea					
Number examined		5	2	2	5
Not remarkable		5	2	2	5
Urinary bladder					
Number examined		5	2	2	5
Not remarkable		5	2	2	5

M : Male

Table 10-4 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of dosing)

Organs	Sex:	F	F
Findings	Dose (mg/kg) : Number:	0 5	1000 5
Adrenal			
Number examined		5	5
Not remarkable		4	5
Mineralization		1	0
minimal		1	0
Bone, femur			
Number examined		5	5
Not remarkable		5	5
Bone marrow, femur			
Number examined		5	5
Not remarkable		5	5
Brain (cerebrum, cerebellum, pons)			
Number examined		5	5
Not remarkable		5	5
Eye			
Number examined		5	5
Not remarkable		5	5
Heart			
Number examined		5	5
Not remarkable		5	4
Infiltrate/Fibrosis		0	1
minimal		0	1
Intestine, cecum			
Number examined		5	5
Not remarkable		5	5
Intestine, colon			
Number examined		5	5
Not remarkable		5	5
Intestine, duodenum			
Number examined		5	5
Not remarkable		5	5
Intestine, ileum			
Number examined		5	5
Not remarkable		5	5
Intestine, jejunum			
Number examined		5	5
Not remarkable		5	5
Intestine, rectum			
Number examined		5	5
Not remarkable		5	5
Kidney			
Number examined		5	5
Not remarkable		5	5
Liver			
Number examined		5	5
Not remarkable		5	5
Lung (bronchus)			
Number examined		5	5
Not remarkable		5	5

F : Female

Table 10-5 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of dosing)

Organs	Sex:	F	F
Findings	Dose (mg/kg) : Number:	0 5	1000 5
Lymph node, mesenteric			
Number examined		5	5
Not remarkable		5	5
Lymph node, submandibular			
Number examined		5	5
Not remarkable		5	5
Mammary gland, inguinal			
Number examined		5	5
Not remarkable		5	5
Medulla oblongata			
Number examined		5	5
Not remarkable		5	5
Muscle, quadriceps femoris			
Number examined		5	5
Not remarkable		5	5
Optic nerve			
Number examined		5	5
Not remarkable		5	5
Ovary			
Number examined		5	5
Not remarkable		5	5
Pancreas			
Number examined		5	5
Not remarkable		4	5
Focus, basophilic		1	0
minimal		1	0
Parathyroid			
Number examined		5	5
Not remarkable		5	5
Peyer's patch, ileal			
Number examined		5	5
Not remarkable		5	5
Pituitary			
Number examined		5	5
Not remarkable		5	5
Salivary gland, submandibular			
Number examined		5	5
Not remarkable		5	5
Sciatic nerve			
Number examined		5	5
Not remarkable		5	5
Skin, inguinal			
Number examined		5	5
Not remarkable		5	5
Spinal cord, thoracic			
Number examined		5	5
Not remarkable		5	5

F : Female

Table 10-6 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of dosing)

Organs	Sex:	F	F
Findings	Dose (mg/kg) : Number:	0 5	1000 5
Spleen			
Number examined		5	5
Not remarkable		5	5
Stomach			
Number examined		5	5
Not remarkable		5	5
Thymus			
Number examined		5	5
Not remarkable		5	5
Thyroid			
Number examined		5	5
Not remarkable		5	5
Trachea			
Number examined		5	5
Not remarkable		5	5
Urinary bladder			
Number examined		5	5
Not remarkable		5	5
Uterus			
Number examined		5	5
Not remarkable		5	5
Vagina			
Number examined		5	5
Not remarkable		5	5

F : Female

Table 10-7 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of dosing,Dams)

Organs	Sex:	F	F	F	F
Findings	Dose (mg/kg) : Number:	0 12	100 10	300 10	1000 10
Adrenal					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Bone, femur					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Bone marrow, femur					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Brain (cerebrum, cerebellum, pons)					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Eye					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Heart					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Intestine, cecum					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Intestine, colon					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Intestine, duodenum					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Intestine, ileum					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Intestine, jejunum					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Intestine, rectum					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Kidney					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Liver					
Number examined		5	0	0	5
Not remarkable		5	0	0	4
Necrosis, focal		0	0	0	1
minimal		0	0	0	1
Lung (bronchus)					
Number examined		5	0	0	5
Not remarkable		5	0	0	5

F : Female

Table 10-8 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of dosing,Dams)

Organs	Sex:	F	F	F	F
Findings	Dose (mg/kg) : Number:	0 12	100 10	300 10	1000 10
Lymph node,mesenteric					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Lymph node,submandibular					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Mammary gland,inguinal					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Medulla oblongata					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Muscle,quadriceps femoris					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Optic nerve					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Ovary					
Number examined		5	0	0	5
Not remarkable		4	0	0	5
Cyst,luteal		1	0	0	0
minimal		1	0	0	0
Pancreas					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Parathyroid					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Peyer's patch,ileal					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Pituitary					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Salivary gland,submandibular					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Sciatic nerve					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Skin,inguinal					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Spinal cord,thoracic					
Number examined		5	0	0	5
Not remarkable		5	0	0	5

F : Female

Table 10-9 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of dosing,Dams)

Organs	Sex:	F	F	F	F
Findings	Dose (mg/kg) : Number:	0 12	100 10	300 10	1000 10
Spleen					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Stomach					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Thymus					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Thyroid					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Trachea					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Urinary bladder					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Uterus					
Number examined		5	0	0	5
Not remarkable		5	0	0	5
Vagina					
Number examined		5	0	0	5
Not remarkable		5	0	0	5

F : Female

Table 10-10 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of recovery)

Organs	Sex:	M	M
Findings	Dose (mg/kg) : Number:	0 5	1000 5
Adrenal			
Number examined		5	5
Not remarkable		5	5
Bone, femur			
Number examined		5	5
Not remarkable		5	5
Bone marrow, femur			
Number examined		5	5
Not remarkable		5	5
Brain (cerebrum, cerebellum, pons)			
Number examined		5	5
Not remarkable		5	5
Coagulating gland			
Number examined		5	5
Not remarkable		5	5
Epididymis			
Number examined		5	5
Not remarkable		5	5
Eye			
Number examined		5	5
Not remarkable		4	5
Retinal rosette		1	0
minimal		1	0
Heart			
Number examined		5	5
Not remarkable		4	3
Infiltrate/Fibrosis		1	2
minimal		1	2
Intestine, cecum			
Number examined		5	5
Not remarkable		5	5
Intestine, colon			
Number examined		5	5
Not remarkable		5	5
Intestine, duodenum			
Number examined		5	5
Not remarkable		5	5
Intestine, ileum			
Number examined		5	5
Not remarkable		5	5
Intestine, jejunum			
Number examined		5	5
Not remarkable		5	5
Intestine, rectum			
Number examined		5	5
Not remarkable		5	5
Kidney			
Number examined		5	5
Not remarkable		5	5

M : Male

Table 10-11 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of recovery)

Organs	Sex:	M	M
Findings	Dose (mg/kg) : Number:	0 5	1000 5
Liver			
Number examined		5	5
Not remarkable		5	5
Lung (bronchus)			
Number examined		5	5
Not remarkable		5	5
Lymph node, mesenteric			
Number examined		5	5
Not remarkable		5	5
Lymph node, submandibular			
Number examined		5	5
Not remarkable		5	5
Mammary gland, inguinal			
Number examined		5	5
Not remarkable		5	5
Medulla oblongata			
Number examined		5	5
Not remarkable		5	5
Muscle, quadriceps femoris			
Number examined		5	5
Not remarkable		5	5
Optic nerve			
Number examined		5	5
Not remarkable		5	5
Pancreas			
Number examined		5	5
Not remarkable		5	5
Parathyroid			
Number examined		4	5
Not remarkable		4	5
No sample		1	0
Peyer's patch, ileal			
Number examined		5	5
Not remarkable		5	5
Pituitary			
Number examined		5	5
Not remarkable		5	5
Prostate			
Number examined		5	5
Not remarkable		1	2
Infiltrate, inflammatory cell		4	3
minimal		3	3
mild		1	0
Salivary gland, submandibular			
Number examined		5	5
Not remarkable		5	5
Sciatic nerve			
Number examined		5	5
Not remarkable		5	5

M : Male

Table 10-12 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of recovery)

Organs	Sex:	M	M
Findings	Dose (mg/kg) : Number:	0 5	1000 5
Seminal vesicle			
Number examined		5	5
Not remarkable		5	5
Skin, inguinal			
Number examined		5	5
Not remarkable		5	5
Spinal cord, thoracic			
Number examined		5	5
Not remarkable		5	5
Spleen			
Number examined		5	5
Not remarkable		5	5
Stomach			
Number examined		5	5
Not remarkable		5	5
Testis			
Number examined		5	5
Not remarkable		5	3
Atrophy, tubule		0	2
minimal		0	2
Thymus			
Number examined		5	5
Not remarkable		5	5
Thyroid			
Number examined		5	5
Not remarkable		5	5
Trachea			
Number examined		5	5
Not remarkable		5	5
Urinary bladder			
Number examined		5	5
Not remarkable		5	5

M : Male

Table 10-13 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of recovery)

Organs	Sex:	F	F
Findings	Dose (mg/kg) : Number:	0 5	1000 5
Adrenal			
Number examined		5	5
Not remarkable		5	5
Bone, femur			
Number examined		5	5
Not remarkable		5	5
Bone marrow, femur			
Number examined		5	5
Not remarkable		5	5
Brain (cerebrum, cerebellum, pons)			
Number examined		5	5
Not remarkable		5	5
Eye			
Number examined		5	5
Not remarkable		5	4
Retinal fold		0	1
minimal		0	1
Heart			
Number examined		5	5
Not remarkable		5	5
Intestine, cecum			
Number examined		5	5
Not remarkable		5	5
Intestine, colon			
Number examined		5	5
Not remarkable		5	5
Intestine, duodenum			
Number examined		5	5
Not remarkable		5	5
Intestine, ileum			
Number examined		5	5
Not remarkable		5	5
Intestine, jejunum			
Number examined		5	5
Not remarkable		5	5
Intestine, rectum			
Number examined		5	5
Not remarkable		5	5
Kidney			
Number examined		5	5
Not remarkable		3	5
Urinary cast, hyaline		2	0
minimal		2	0
Fibrosis, interstitium		1	0
minimal		1	0
Liver			
Number examined		5	5
Not remarkable		5	5

F : Female

Table 10-14 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of recovery)

Organs	Sex:	F	F
Findings	Dose (mg/kg) : Number:	0 5	1000 5
Lung (bronchus)			
Number examined		5	5
Not remarkable		5	5
Lymph node, mesenteric			
Number examined		5	5
Not remarkable		5	5
Lymph node, submandibular			
Number examined		5	5
Not remarkable		5	5
Mammary gland, inguinal			
Number examined		5	5
Not remarkable		5	5
Medulla oblongata			
Number examined		5	5
Not remarkable		5	5
Muscle, quadriceps femoris			
Number examined		5	5
Not remarkable		5	5
Optic nerve			
Number examined		5	5
Not remarkable		5	5
Ovary			
Number examined		5	5
Not remarkable		4	5
Age-related atrophy		1	0
mild		1	0
Pancreas			
Number examined		5	5
Not remarkable		5	5
Parathyroid			
Number examined		5	5
Not remarkable		5	5
Peyer's patch, ileal			
Number examined		5	5
Not remarkable		5	5
Pituitary			
Number examined		5	5
Not remarkable		5	5
Salivary gland, submandibular			
Number examined		5	5
Not remarkable		5	5
Sciatic nerve			
Number examined		5	5
Not remarkable		5	5
Skin, inguinal			
Number examined		5	5
Not remarkable		5	5

F : Female

Table 10-15 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (End of recovery)

Organs	Sex:	F	F
Findings	Dose (mg/kg) :	0	1000
	Number:	5	5
Spinal cord, thoracic			
Number examined		5	5
Not remarkable		5	5
Spleen			
Number examined		5	5
Not remarkable		5	5
Stomach			
Number examined		5	5
Not remarkable		5	5
Thymus			
Number examined		5	5
Not remarkable		5	5
Thyroid			
Number examined		5	5
Not remarkable		5	5
Trachea			
Number examined		5	5
Not remarkable		5	5
Urinary bladder			
Number examined		5	5
Not remarkable		5	5
Uterus			
Number examined		5	5
Not remarkable		5	5
Vagina			
Number examined		5	5
Not remarkable		5	5

F : Female

Table 10-16 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (Undelivered)

Organs	Sex:	F	F	F
Findings	Dose (mg/kg) : Number:	100 2	300 2	1000 2
Adrenal				
Number examined		2	2	2
Not remarkable		2	2	2
Bone, femur				
Number examined		2	2	2
Not remarkable		2	2	2
Bone marrow, femur				
Number examined		2	2	2
Not remarkable		2	2	2
Brain (cerebrum, cerebellum, pons)				
Number examined		2	2	2
Not remarkable		2	2	2
Eye				
Number examined		2	2	2
Not remarkable		2	2	2
Heart				
Number examined		2	2	2
Not remarkable		2	2	2
Intestine, cecum				
Number examined		2	2	2
Not remarkable		2	2	2
Intestine, colon				
Number examined		2	2	2
Not remarkable		2	2	2
Intestine, duodenum				
Number examined		2	2	2
Not remarkable		2	2	2
Intestine, ileum				
Number examined		2	2	2
Not remarkable		2	2	2
Intestine, jejunum				
Number examined		2	2	2
Not remarkable		2	2	2
Intestine, rectum				
Number examined		2	2	2
Not remarkable		2	2	2
Kidney				
Number examined		2	2	2
Not remarkable		1	2	2
Mineralization, pelvis		1	0	0
minimal		1	0	0
Liver				
Number examined		2	2	2
Not remarkable		2	2	2
Lung (bronchus)				
Number examined		2	2	2
Not remarkable		2	2	2

F : Female

Table 10-17 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (Undelivered)

Organs	Sex:	F	F	F
Findings	Dose (mg/kg) : Number:	100 2	300 2	1000 2
Lymph node,mesenteric				
Number examined		2	2	2
Not remarkable		2	2	2
Lymph node,submandibular				
Number examined		2	2	2
Not remarkable		2	2	2
Mammary gland,inguinal				
Number examined		2	2	2
Not remarkable		2	2	2
Medulla oblongata				
Number examined		2	2	2
Not remarkable		2	2	2
Muscle,quadriceps femoris				
Number examined		2	2	2
Not remarkable		2	2	2
Optic nerve				
Number examined		2	2	2
Not remarkable		2	2	2
Ovary				
Number examined		2	2	2
Not remarkable		2	2	2
Pancreas				
Number examined		2	2	2
Not remarkable		1	2	2
Focus,basophilic		1	0	0
minimal		1	0	0
Parathyroid				
Number examined		2	2	2
Not remarkable		2	2	2
Peyer's patch,ileal				
Number examined		2	2	2
Not remarkable		2	2	2
Pituitary				
Number examined		2	2	2
Not remarkable		2	2	2
Salivary gland,submandibular				
Number examined		2	2	2
Not remarkable		2	2	2
Sciatic nerve				
Number examined		2	2	2
Not remarkable		2	2	2
Skin,inguinal				
Number examined		2	2	2
Not remarkable		2	2	2
Spinal cord,thoracic				
Number examined		2	2	2
Not remarkable		2	2	2

F : Female

Table 10-18 Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats
Histopathological findings (Undelivered)

Organs	Sex:	F	F	F
Findings	Dose (mg/kg) : Number:	100 2	300 2	1000 2
Spleen				
Number examined		2	2	2
Not remarkable		2	2	2
Stomach				
Number examined		2	2	2
Not remarkable		2	2	2
Thymus				
Number examined		2	2	2
Not remarkable		2	2	2
Thyroid				
Number examined		2	2	2
Not remarkable		2	2	2
Trachea				
Number examined		2	2	2
Not remarkable		2	2	2
Urinary bladder				
Number examined		2	2	2
Not remarkable		2	2	2
Uterus				
Number examined		2	2	2
Not remarkable		2	2	2
Vagina				
Number examined		2	2	2
Not remarkable		2	2	2

F : Female

Table 11

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Estrus cycles
Generation : F0

Study No. : R-1261

Species : Rat

Test article Dose		/Before mating			
		Mean estrous cycle (Days)	Number of estrus	Number of animals with abnormal estrous cycle	Abnormal estrous cycle index (%)
Dodecyloxirane 0 mg/kg	n	12	12	12	0.0
	Mean	4.2	3.8	(0)	
	S.D.	0.3	0.5		
Dodecyloxirane 100 mg/kg	n	12	12	12	8.3
	Mean	4.3	3.6	(1)	
	S.D.	0.5	0.5		
Dodecyloxirane 300 mg/kg	n	12	12	12	0.0
	Mean	4.0	3.7	(0)	
	S.D.	0.0	0.5		
Dodecyloxirane 1000 mg/kg	n	12	12	12	0.0
	Mean	4.2	3.8	(0)	
	S.D.	0.3	0.5		

() : Values in brackets represent number of animals with abnormal estrous cycle.

Abnormal estrous cycle index = (No. of animals with abnormal estrous cycle / No. of animals examined)×100

Not significantly different from Dodecyloxirane 0 mg/kg

Table 12 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Reproductive performance
Generation : F0 Sex : Male

Study No. : R-1261

						Species : Rat
		Number of pairs	Elapsed day of conceiving	1st mating		
Test article Dose				Copulation index (%) a)	Fertility index (%) b)	
Dodecyloxirane 0 mg/kg	n	12	12	(12/12)	(12/12)	
	Mean		3.3	100.0	100.0	
	S.D.		2.8			
Dodecyloxirane 100 mg/kg	n	12	11	(11/12)	(10/11)	
	Mean		2.5	91.7	90.9	
	S.D.		1.2			
Dodecyloxirane 300 mg/kg	n	12	11	(11/12)	(10/11)	
	Mean		3.3	91.7	90.9	
	S.D.		3.8			
Dodecyloxirane 1000 mg/kg	n	12	12	(12/12)	(11/12)	
	Mean		3.3	100.0	91.7	
	S.D.		3.6			

Not significantly different from Dodecyloxirane 0 mg/kg

a): (Number of copulated animals / Number of mated animals)x 100

b): (Number of males which impregnated females / Number of copulated males)x 100

Table 12 - 2

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Reproductive performance
Generation : F0 Sex : Female

Study No. : R-1261

		Species : Rat			
		1st mating			
Test article		Number of	Elapsed day	Copulation	Fertility
Dose		pairs	of conceiving	index (%) a)	index (%) b)
Dodecyloxirane	n	12	12	(12/12)	(12/12)
0 mg/kg	Mean		3.3	100.0	100.0
	S.D.		2.8		
Dodecyloxirane	n	12	11	(11/12)	(10/11)
100 mg/kg	Mean		2.5	91.7	90.9
	S.D.		1.2		
Dodecyloxirane	n	12	11	(11/12)	(10/11)
300 mg/kg	Mean		3.3	91.7	90.9
	S.D.		3.8		
Dodecyloxirane	n	12	12	(12/12)	(11/12)
1000 mg/kg	Mean		3.3	100.0	91.7
	S.D.		3.6		

Not significantly different from Dodecyloxirane 0 mg/kg

a): (Number of copulated animals / Number of mated animals)x 100

b): (Number of pregnant females / Number of copulated females)x 100

Table 13

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Delivery data
Generation : F0

Study No. : R-1261

		Species : Rat										
Test article		Gestation period (day)	Gestation index (%) a)	Number of implantation	Delivery index (%) b)	Live birth index (%) c)	Number of offspring	Number of live newborns			Sex ratio (%) d)	Number of dead newborns
Dose								M	F	Total		
Dodecyloxirane	n	12	12/12	12	12	12	12	12	12	12	91/156	12
0 mg/kg	Mean	22.1	100.0	15.6	85.4	98.6	13.2	7.6	5.4	13.0	58.3	0.2
	S.D.	0.4		2.5	22.0	3.5	3.9	2.9	2.2	3.9		0.4
Dodecyloxirane	n	10	10/10	10	10	10	10	10	10	10	69/140	10
100 mg/kg	Mean	22.1	100.0	15.7	91.4	97.9	14.3	6.9	7.1	14.0	49.3	0.3
	S.D.	0.5		1.6	8.0	6.8	1.5	1.9	1.8	1.8		0.9
Dodecyloxirane	n	10	10/10	10	10	10	10	10	10	10	85/146	10
300 mg/kg	Mean	21.9	100.0	15.9	91.9	100.0	14.6	8.5	6.1	14.6	58.2	0.0
	S.D.	0.3		1.8	11.0	0.0	2.3	1.9	1.7	2.3		0.0
Dodecyloxirane	n	10	10/11	10	10	10	10	10	10	10	69/137	10
1000 mg/kg	Mean	22.3	90.9	14.9	92.1	100.0	13.7	6.9	6.8	13.7	50.4	0.0
	S.D.	0.3		2.5	7.9	0.0	2.4	2.3	2.1	2.4		0.0

M : Male, F : Female

Not significantly different from Dodecyloxirane 0 mg/kg

a): (Number of females delivered live newborns / Number of pregnant females) x 100

b): (Number of delivered pups / Number of implantation) x 100

c): (Number of liveborns / Number of delivered pups) x 100

d): (Number of live males / Number of liveborns) x 100

Table 14

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
External examination of offspring
Generation : F0

Study No. : R-1261

Days after birth : 0
Species : Rat

	Dodecyloxi rane	Dodecyloxi rane	Dodecyloxi rane	Dodecyloxi rane
Test article	0	100	300	1000
Dose	mg/kg	mg/kg	mg/kg	mg/kg
Dose unit	12	10	10	10
Number of dams	158	143	146	137
Number of offspring	0(0.0)	0(0.0)	0(0.0)	0(0.0)
Number of dams with anomalous offspring (incidence %) a)	0(0.0)	0(0.0)	0(0.0)	0(0.0)
Number of offspring with any anomaly (incidence %) b)	0(0.0)	0(0.0)	0(0.0)	0(0.0)

The group mean is the same and unbiased variance 0 for all groups. All data of all groups are same.

a): (Number of dams bearing pups with external anomalies / Number of dams) x 100

b): (Number of pups with external anomalies / Number of delivered pups) x 100

Table 15

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Anogenital of offspring
Generation : F0

Study No. : R-1261

					Unit : mm	Species : Rat
Sex		/Days after birth				
		0				
Dose		Normalized			a)	
Test article		AGD				
Dodecyloxirane 0 mg/kg	Male	n	12	12		
		Mean	2.5	1.3		
		S.D.	0.1	0.1		
	Female	n	12	12		
		Mean	0.9	0.5		
		S.D.	0.1	0.1		
Dodecyloxirane 100 mg/kg	Male	n	10	10		
		Mean	2.5	1.3		
		S.D.	0.2	0.1		
	Female	n	10	10		
		Mean	0.8	0.5		
		S.D.	0.1	0.1		
Dodecyloxirane 300 mg/kg	Male	n	10	10		
		Mean	2.5	1.3		
		S.D.	0.1	0.1		
	Female	n	10	10		
		Mean	0.8	0.4		
		S.D.	0.1	0.0		
Dodecyloxirane 1000 mg/kg	Male	n	10	10		
		Mean	2.5	1.3		
		S.D.	0.1	0.1		
	Female	n	10	10		
		Mean	0.9	0.5		
		S.D.	0.2	0.1		

AGD : Anogenital Distance

Not significantly different from Dodecyloxirane 0 mg/kg

a): AGD (mm) / Cube root of body weight (g)

Table 16

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Litter size and viability index of offspring

Study No. : R-1261

Generation : F0

Species : Rat

		/Days after birth					
Test article Dose		0	4	4	4	13	13
			Pre-culled	Viability index (%) a)	Culled		Viability index (%) b)
Dodecyloxirane 0 mg/kg	Total	156	155		93	93	
	n	12	12	12	12	12	12
	Mean		12.9	99.4		7.8	100.0
	S.D.		3.9	1.9		0.9	0.0
Dodecyloxirane 100 mg/kg	Total	140	137		80	80	
	n	10	10	10	10	10	10
	Mean		13.7	97.8		8.0	100.0
	S.D.		1.9	3.6		0.0	0.0
Dodecyloxirane 300 mg/kg	Total	146	145		80	80	
	n	10	10	10	10	10	10
	Mean		14.5	99.4		8.0	100.0
	S.D.		2.2	1.8		0.0	0.0
Dodecyloxirane 1000 mg/kg	Total	137	134		80	80	
	n	10	10	10	10	10	10
	Mean		13.4	98.1		8.0	100.0
	S.D.		2.2	3.1		0.0	0.0

Not significantly different from Dodecyloxirane 0 mg/kg

a): (Number of pups alive on day 4 after birth / Number of live born pups) × 100

b): (Number of pups alive on day 13 after birth / Number of pups after culling on day 4 after birth) × 100

Table 17

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Sex ratio

Study No. : R-1261

Generation : F0

Species : Rat

Test article Dose	Number of dams		day 4 after birth		
			Number of males	Number of females	Sex ratio (%) a)
Dodecyloxirane 0 mg/kg	12	Total (%)	90	65	90/155 58.1
Dodecyloxirane 100 mg/kg	10	Total (%)	68	69	68/137 49.6
Dodecyloxirane 300 mg/kg	10	Total (%)	84	61	84/145 57.9
Dodecyloxirane 1000 mg/kg	10	Total (%)	69	65	69/134 51.5

Not significantly different from Dodecyloxirane 0 mg/kg

a): (Number of male pups alive on day 4 after birth / Number of pups alive on day 4 after birth) × 100

Table 18 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Body weight of offspring
Generation : F0

Study No. : R-1261

Body weight of offspring			Unit : g					Species : Rat
Test article			/Days after birth					
Dose			0	4/Pre-culled	4/Culled	7	13	
Dodecyloxirane 0 mg/kg	Male	n	12	12	12	12	12	
		Mean	6.9	11.4	11.7	18.6	35.0	
		S.D.	0.7	1.5	1.3	1.4	2.5	
	Female	n	12	12	12	12	12	
		Mean	6.5	10.8	11.0	17.6	33.4	
		S.D.	0.7	1.5	1.4	1.6	2.1	
Dodecyloxirane 100 mg/kg	Male	n	10	10	10	10	10	
		Mean	6.7	11.0	11.2	18.6	35.3	
		S.D.	0.3	0.9	1.0	1.5	2.0	
	Female	n	10	10	10	10	10	
		Mean	6.3	10.5	10.7	17.6	33.9	
		S.D.	0.3	0.9	0.9	1.4	2.1	
Dodecyloxirane 300 mg/kg	Male	n	10	10	10	10	10	
		Mean	6.6	10.6	10.9	17.9	34.8	
		S.D.	0.6	1.2	1.2	2.0	2.4	
	Female	n	10	10	10	10	10	
		Mean	6.1	9.7	10.0	16.4	32.6	
		S.D.	0.4	1.0	0.9	1.5	2.2	
Dodecyloxirane 1000 mg/kg	Male	n	10	10	10	10	10	
		Mean	6.7	10.7	10.9	17.6	31.7	
		S.D.	0.6	1.2	1.1	1.7	3.4	
	Female	n	10	10	10	10	10	
		Mean	6.4	10.4	10.6	17.1	30.9	
		S.D.	0.5	1.0	0.9	1.4	3.0	
					DT *			

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

DT : Dunnett test (two-side)

Table 18 - 2

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Body weight gain of offspring
Generation : F0

Study No. : R-1261

Base : Day 4 after birth
Unit : g

Species : Rat

Test article Dose			/Days after birth		
			/Before culling /After culling		
			0-4	4-7	4-13
Dodecyloxirane 0 mg/kg	Male	n	12	12	12
		Mean	4.5	7.0	23.3
		S.D.	0.9	0.3	1.8
	Female	n	12	12	12
		Mean	4.3	6.7	22.5
		S.D.	0.9	0.5	1.4
Dodecyloxirane 100 mg/kg	Male	n	10	10	10
		Mean	4.3	7.3	24.1
		S.D.	0.7	0.6	1.3
	Female	n	10	10	10
		Mean	4.2	7.0	23.2
		S.D.	0.7	0.8	1.8
Dodecyloxirane 300 mg/kg	Male	n	10	10	10
		Mean	4.0	7.0	23.9
		S.D.	1.0	1.1	1.6
	Female	n	10	10	10
		Mean	3.7	6.4	22.6
		S.D.	0.9	0.9	1.6
Dodecyloxirane 1000 mg/kg	Male	n	10	10	10
		Mean	4.0	6.7	20.9
		S.D.	0.7	1.1	2.8
	Female	n	10	10	10
		Mean	4.1	6.4	20.3
		S.D.	0.6	1.0	2.6

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05

DT : Dunnett test (two-side)

Table 19

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Externally for gross abnormalities in dead pups

Study No. : R-1261

Species : Rat					
	Test article Dose	Dodecyloxirane 0 mg/kg	Dodecyloxirane 100 mg/kg	Dodecyloxirane 300 mg/kg	Dodecyloxirane 1000 mg/kg
Male					
Number of pups examined		0	0	1	0
Number of pups with abnormal findings		0	0	0	0
Female					
Number of pups examined		0	1	0	1
Number of pups with abnormal findings		0	0	0	0

Table 20

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Nipple retention of male pups

Study No. : R-1261

Applicable retention of male pups			Unit : No.	Species : Rat
Test article	Generation : F0		/Days after birth	
Dose			13	
Dodecyloxirane 0 mg/kg	Male	n	12	
		Mean	0.0	
		S.D.	0.0	
Dodecyloxirane 100 mg/kg	Male	n	10	
		Mean	0.0	
		S.D.	0.0	
Dodecyloxirane 300 mg/kg	Male	n	10	
		Mean	0.0	
		S.D.	0.0	
Dodecyloxirane 1000 mg/kg	Male	n	10	
		Mean	0.0	
		S.D.	0.0	

The group mean is the same and unbiased variance 0 for all groups. All data of all groups are same.
Nipple retention: Number of nipple / areolae

Table 21

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Gross pathological findings in pups on postnatal day 13

Study No. : R-1261

Species : Rat					
	Test article Dose	Dodecyloxirane 0 mg/kg	Dodecyloxirane 100 mg/kg	Dodecyloxirane 300 mg/kg	Dodecyloxirane 1000 mg/kg
Male					
Number of pups examined		48	42	41	39
Number of pups with abnormal findings		0	0	0	0
Female					
Number of pups examined		45	38	39	41
Number of pups with abnormal findings		0	0	0	0

Table 22 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : After culled (F1 males)

Study No. : R-1261

Species : Rat

Test article Dose		Body weight g	Thyroid-RL	
			AB mg	RE mg/100g
Dodecyloxirane 0 mg/kg	n	12	12	12
	Mean	36.0	4.2	11.6
	S.D.	2.6	1.1	2.9
Dodecyloxirane 100 mg/kg	n	10	10	10
	Mean	36.6	4.5	12.3
	S.D.	3.4	0.6	2.1
Dodecyloxirane 300 mg/kg	n	10	10	10
	Mean	35.2	4.7	13.2
	S.D.	2.6	1.3	3.4
Dodecyloxirane 1000 mg/kg	n	10	10	10
	Mean	32.2	5.1	16.0
	S.D.	3.2	1.0	3.6
		DT *		DT **

AB : Absolute weight, RE : Relative weight by body weight

Significantly different from Dodecyloxirane 0 mg/kg : * P<0.05, ** P<0.01

DT : Dunnett test (two-side)

Table 22 - 2

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Female

Stage : After culled (F1 females)

Study No. : R-1261

Species : Rat

Test article Dose		Body weight g	Thyroid-RL	
			AB mg	RE mg/100g
Dodecyloxirane 0 mg/kg	n	12	12	12
	Mean	33.6	4.4	13.4
	S.D.	2.4	1.1	4.0
Dodecyloxirane 100 mg/kg	n	10	10	10
	Mean	34.3	4.9	14.3
	S.D.	2.2	1.0	3.0
Dodecyloxirane 300 mg/kg	n	10	10	10
	Mean	33.1	5.4	16.3
	S.D.	2.5	0.8	2.1
Dodecyloxirane 1000 mg/kg	n	10	10	10
	Mean	32.1	4.3	13.3
	S.D.	2.5	1.0	2.7

AB : Absolute weight, RE : Relative weight by body weight
Not significantly different from Dodecyloxirane 0 mg/kg

Appendix 1 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Male

Study No. : R-1261

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	/Day																				
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21
1001	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1002	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1003	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1004	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1005	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1006	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1007	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1008	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1009	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1010	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

∴ No abnormal findings

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	/Day																				
	22	23	24	25	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	42
1001	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1002	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1003	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1004	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1005	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1006	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1007	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1008	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1009	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1010	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Appendix 1 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Male

Study No. : R-1261

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	/Day 43 a)
1001	-
1002	-
1003	-
1004	-
1005	-
1006	-
1007	-
1008	
1009	
1010	
1011	
1012	

-: No abnormal findings

a): Day of necropsy

Appendix 1 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Male

Study No. : R-1261

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	/Day																				
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21
2001	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2002	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2003	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2004	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2005	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2006	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2007	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2008	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2009	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2010	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

∴ No abnormal findings

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	/Day																				
	22	23	24	25	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	42
2001	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2002	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2003	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2004	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2005	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2006	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2007	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2008	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2009	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2010	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Appendix 1 - 6

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Male

Study No. : R-1261

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	/Day 43 a)
2001	-
2002	-
2003	-
2004	-
2005	-
2006	-
2007	-
2008	-
2009	-
2010	-
2011	-
2012	-

:- No abnormal findings

a): Day of necropsy

Animal No.	/Day																				
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21
3001	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3002	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3003	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3004	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3005	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3006	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3007	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3008	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3009	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3010	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Animal No.	/Day																				
	22	23	24	25	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	42
3001	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3002	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3003	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3004	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3005	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3006	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3007	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3008	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3009	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3010	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Appendix 1 - 9

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Male

Study No. : R-1261

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	/Day 43 a)
3001	-
3002	-
3003	-
3004	-
3005	-
3006	-
3007	-
3008	-
3009	-
3010	-
3011	-
3012	-

:- No abnormal findings

a): Day of necropsy

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	/Day																				
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21
4001	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4002	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4003	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4004	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4005	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4006	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4007	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4008	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4009	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4010	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	/Day																				
	22	23	24	25	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	42
4001	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4002	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4003	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4004	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4005	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4006	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4007	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4008	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4009	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4010	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Appendix 1 - 12

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Male

Study No. : R-1261

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	/Day 43 a)
4001	-
4002	-
4003	-
4004	-
4005	-
4006	-
4007	-
4008	
4009	
4010	
4011	
4012	

:- No abnormal findings

a): Day of necropsy

Animal No.	/Day														
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15
1101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1111	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1112	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Animal No.	/Day														
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15
2101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2111	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2112	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Animal No.	/Day														
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15
3101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3111	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3112	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

∴ No abnormal findings

Animal No.	/Day														
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15
4101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4111	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4112	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Period : F0 gestation Day 0-25

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Sex : Female										Dose : Docetaxel 0.1 mg/kg										Species : Rat	
Animal No.	/Day																					
	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	
1101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1111	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
1112	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	

-: No abnormal findings

Animal No.	/Day				
	21	22	23	24	25
1101	-				
1102	-	-			
1103	-	-			
1104	-				
1105	-				
1106	-	-			
1107	-				
1108	-				
1109	-				
1110	-	-			
1111	-				
1112	-				

∴ No abnormal findings

Period : F0 gestation Day 0-25

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Sex : Female										Dose : Doxycycline 100 mg/kg										Species : Rat	
Animal No.	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	
2101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
2102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
2103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
2104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
2105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
2107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
2108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
2110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
2111	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
2112	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	

-: No abnormal findings

Appendix 1 - 20

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs in dams
Sex : Female

Study No. : R-1261

Period : F0 gestation Day 0-25
Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	/Day				
	21	22	23	24	25
2101	-				
2102	-				
2103	-				
2104	-				
2105	-				
2107	-				
2108	-				
2110	-				
2111	-	-			
2112	-				

-: No abnormal findings

Period : F0 gestation Day 0-25

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Sex : Female		Dose : Dodecylxirane 500 mg/kg																		Species : Rat
Animal No.	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20
3101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3111	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3112	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Appendix 1 - 22

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs in dams
Sex : Female

Study No. : R-1261

Period : F0 gestation Day 0-25
Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	/Day				
	21	22	23	24	25
3101	-				
3102	-				
3103	-				
3104	-				
3105	-				
3106	-				
3107	-				
3110	-				
3111	-				
3112	-				

:- No abnormal findings

Period : F0 gestation Day 0-25

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Sex : Female										Dose : Doxycycline 1000 mg/kg										Species : Rat	
Animal No.	/Day																					
	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	
4102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4111	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4112	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	

-: No abnormal findings

Appendix 1 - 24

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs in dams
Sex : Female

Study No. : R-1261

Period : F0 gestation Day 0-25
Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	/Day				
	21	22	23	24	25
4102	-				
4103	-				
4104	-				
4105	-				
4106	-	-	-	-	-
4107	-				
4108	-	-			
4109	-				
4110	-				
4111	-				
4112	-				

∴ No abnormal findings

	Sex : Female										Dose : Doxycycline 5 mg/kg					Species : Rat
Animal No.	/Day	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14 a)
1101		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1102		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1103		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1104		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1105		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1106		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1107		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1108		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1109		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1110		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1111		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1112		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

a): Day of necropsy

Animal No.	/Day													
	0	1	2	3	4	5	6	7	8	9	10	11	12	13 14 a)
2101	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2102	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2103	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2104	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2105	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2107	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2108	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2110	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2111	-	-	-	-	-	-	-	-	-	-	-	-	-	-
2112	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

a): Day of necropsy

	Sex : Female										Dose : Doxycycline 500 mg/kg					Species : Rat
Animal No.	/Day	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14 a)
3101		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3102		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3103			-	-	-	-	-	-	-	-	-	-	-	-	-	-
3104			-	-	-	-	-	-	-	-	-	-	-	-	-	-
3105		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3106		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3107		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3110		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
3111			-	-	-	-	-	-	-	-	-	-	-	-	-	-
3112		-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

a): Day of necropsy

	Sex : Female										Dose : Doxycycline 1000 mg/kg					Species : Rat
	/Day															
Animal No.	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14 a)	
4102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4111	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	
4112	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	

-: No abnormal findings

a): Day of necropsy

Period : Administration day 1-43(non-mating group)

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	/Day																				
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21
5101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

	/Day																				
Animal No.	22	23	24	25	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	42
5101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Appendix 1 - 30

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Clinical signs
Sex : Female

Study No. : R-1261

Period : Administration day 1-43(non-mating group)

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	/Day 43 a)
5101	-
5102	-
5103	-
5104	-
5105	-
5106	
5107	
5108	
5109	
5110	

-: No abnormal findings

a): Day of necropsy

Period : Administration day 1-43(non-mating group)

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	/Day																				
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21
6101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

	/Day																				
Animal No.	22	23	24	25	26	27	28	29	30	31	32	33	34	35	36	37	38	39	40	41	42
6101	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6102	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6103	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6104	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6105	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

Animal No.	/Day 43 a)
6101	-
6102	-
6103	-
6104	-
6105	-
6106	
6107	
6108	
6109	
6110	

-: No abnormal findings

a): Day of necropsy

Animal No.	/Day														
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15 a)
1008	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1009	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1010	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
1012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

a): Day of necropsy

Period : Recovery Day 1-15

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	/Day														
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15 a)
4008	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4009	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4010	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4011	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
4012	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

a): Day of necropsy

Animal No.	/Day														
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15 a)
5106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
5110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

a): Day of necropsy

Animal No.	/Day														
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15 a)
6106	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6107	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6108	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6109	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
6110	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

-: No abnormal findings

a): Day of necropsy

Appendix 2 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 1

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
1001	N	0	0
1002	N	0	0
1003	N	0	0
1004	N	0	0
1005	N	0	0
1006	N	0	0
1007	N	0	0
1008	N	0	0
1009	N	0	0
1010	N	0	0
1011	N	0	0
1012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 1

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2001	N	0	0
2002	N	0	0
2003	N	0	0
2004	N	0	0
2005	N	0	0
2006	N	0	0
2007	N	0	0
2008	N	0	0
2009	N	0	0
2010	N	0	0
2011	N	0	0
2012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 1

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3001	N	0	0
3002	N	0	0
3003	N	0	0
3004	N	0	0
3005	N	0	0
3006	N	0	0
3007	N	0	0
3008	N	0	0
3009	N	0	0
3010	N	0	0
3011	N	0	0
3012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Appendix 2 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4001	N	0	0
4002	N	0	0
4003	N	0	0
4004	N	0	0
4005	N	0	0
4006	N	0	0
4007	N	0	0
4008	N	0	0
4009	N	0	0
4010	N	0	0
4011	N	0	0
4012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
1001	N	0	0
1002	N	0	0
1003	N	0	0
1004	N	0	0
1005	N	0	0
1006	N	0	0
1007	N	0	0
1008	N	0	0
1009	N	0	0
1010	N	0	0
1011	N	0	0
1012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2001	N	0	0
2002	N	0	0
2003	N	0	0
2004	N	0	0
2005	N	0	0
2006	N	0	0
2007	N	0	0
2008	N	0	0
2009	N	0	0
2010	N	0	0
2011	N	0	0
2012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Appendix 2 - 7

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3001	N	0	0
3002	N	0	0
3003	N	0	0
3004	N	0	0
3005	N	0	0
3006	N	0	0
3007	N	0	0
3008	N	0	0
3009	N	0	0
3010	N	0	0
3011	N	0	0
3012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4001	N	0	0
4002	N	0	0
4003	N	0	0
4004	N	0	0
4005	N	0	0
4006	N	0	0
4007	N	0	0
4008	N	0	0
4009	N	0	0
4010	N	0	0
4011	N	0	0
4012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 3

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
1001	N	0	0
1002	N	0	0
1003	N	0	0
1004	N	0	0
1005	N	0	0
1006	N	0	0
1007	N	0	0
1008	N	0	0
1009	N	0	0
1010	N	0	0
1011	N	0	0
1012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Appendix 2 - 10

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 3

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2001	N	0	0
2002	N	0	0
2003	N	0	0
2004	N	0	0
2005	N	0	0
2006	N	0	0
2007	N	0	0
2008	N	0	0
2009	N	0	0
2010	N	0	0
2011	N	0	0
2012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 3

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3001	N	0	0
3002	N	0	0
3003	N	0	0
3004	N	0	0
3005	N	0	0
3006	N	0	0
3007	N	0	0
3008	N	0	0
3009	N	0	0
3010	N	0	0
3011	N	0	0
3012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 3

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4001	N	0	0
4002	N	0	0
4003	N	0	0
4004	N	0	0
4005	N	0	0
4006	N	0	0
4007	N	0	0
4008	N	0	0
4009	N	0	0
4010	N	0	0
4011	N	0	0
4012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 4

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Posture	Convulsion	Abnormal behavior
1001	N	0	0
1002	N	0	0
1003	N	0	0
1004	N	0	0
1005	N	0	0
1006	N	0	0
1007	N	0	0
1008	N	0	0
1009	N	0	0
1010	N	0	0
1011	N	0	0
1012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 4

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2001	N	0	0
2002	N	0	0
2003	N	0	0
2004	N	0	0
2005	N	0	0
2006	N	0	0
2007	N	0	0
2008	N	0	0
2009	N	0	0
2010	N	0	0
2011	N	0	0
2012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 4

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3001	N	0	0
3002	N	0	0
3003	N	0	0
3004	N	0	0
3005	N	0	0
3006	N	0	0
3007	N	0	0
3008	N	0	0
3009	N	0	0
3010	N	0	0
3011	N	0	0
3012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 4

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4001	N	0	0
4002	N	0	0
4003	N	0	0
4004	N	0	0
4005	N	0	0
4006	N	0	0
4007	N	0	0
4008	N	0	0
4009	N	0	0
4010	N	0	0
4011	N	0	0
4012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 5

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Posture	Convulsion	Abnormal behavior
1001	N	0	0
1002	N	0	0
1003	N	0	0
1004	N	0	0
1005	N	0	0
1006	N	0	0
1007	N	0	0
1008	N	0	0
1009	N	0	0
1010	N	0	0
1011	N	0	0
1012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 5

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2001	N	0	0
2002	N	0	0
2003	N	0	0
2004	N	0	0
2005	N	0	0
2006	N	0	0
2007	N	0	0
2008	N	0	0
2009	N	0	0
2010	N	0	0
2011	N	0	0
2012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 5

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3001	N	0	0
3002	N	0	0
3003	N	0	0
3004	N	0	0
3005	N	0	0
3006	N	0	0
3007	N	0	0
3008	N	0	0
3009	N	0	0
3010	N	0	0
3011	N	0	0
3012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 5

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4001	N	0	0
4002	N	0	0
4003	N	0	0
4004	N	0	0
4005	N	0	0
4006	N	0	0
4007	N	0	0
4008	N	0	0
4009	N	0	0
4010	N	0	0
4011	N	0	0
4012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
1001	N	0	0
1002	N	0	0
1003	N	0	0
1004	N	0	0
1005	N	0	0
1006	N	0	0
1007	N	0	0
1008	N	0	0
1009	N	0	0
1010	N	0	0
1011	N	0	0
1012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2001	N	0	0
2002	N	0	0
2003	N	0	0
2004	N	0	0
2005	N	0	0
2006	N	0	0
2007	N	0	0
2008	N	0	0
2009	N	0	0
2010	N	0	0
2011	N	0	0
2012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3001	N	0	0
3002	N	0	0
3003	N	0	0
3004	N	0	0
3005	N	0	0
3006	N	0	0
3007	N	0	0
3008	N	0	0
3009	N	0	0
3010	N	0	0
3011	N	0	0
3012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4001	N	0	0
4002	N	0	0
4003	N	0	0
4004	N	0	0
4005	N	0	0
4006	N	0	0
4007	N	0	0
4008	N	0	0
4009	N	0	0
4010	N	0	0
4011	N	0	0
4012	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Posture	Convulsion	Abnormal behavior
1101	N	0	0
1102	N	0	0
1103	N	0	0
1104	N	0	0
1105	N	0	0
1106	N	0	0
1107	N	0	0
1108	N	0	0
1109	N	0	0
1110	N	0	0
1111	N	0	0
1112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2101	N	0	0
2102	N	0	0
2103	N	0	0
2104	N	0	0
2105	N	0	0
2106	N	0	0
2107	N	0	0
2108	N	0	0
2109	N	0	0
2110	N	0	0
2111	N	0	0
2112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3101	N	0	0
3102	N	0	0
3103	N	0	0
3104	N	0	0
3105	N	0	0
3106	N	0	0
3107	N	0	0
3108	N	0	0
3109	N	0	0
3110	N	0	0
3111	N	0	0
3112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4101	N	0	0
4102	N	0	0
4103	N	0	0
4104	N	0	0
4105	N	0	0
4106	N	0	0
4107	N	0	0
4108	N	0	0
4109	N	0	0
4110	N	0	0
4111	N	0	0
4112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Posture	Convulsion	Abnormal behavior
1101	N	0	0
1102	N	0	0
1103	N	0	0
1104	N	0	0
1105	N	0	0
1106	N	0	0
1107	N	0	0
1108	N	0	0
1109	N	0	0
1110	N	0	0
1111	N	0	0
1112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Appendix 2 - 30

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2101	N	0	0
2102	N	0	0
2103	N	0	0
2104	N	0	0
2105	N	0	0
2106	N	0	0
2107	N	0	0
2108	N	0	0
2109	N	0	0
2110	N	0	0
2111	N	0	0
2112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3101	N	0	0
3102	N	0	0
3103	N	0	0
3104	N	0	0
3105	N	0	0
3106	N	0	0
3107	N	0	0
3108	N	0	0
3109	N	0	0
3110	N	0	0
3111	N	0	0
3112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Posture	Convulsion	Abnormal behavior
4101	N	0	0
4102	N	0	0
4103	N	0	0
4104	N	0	0
4105	N	0	0
4106	N	0	0
4107	N	0	0
4108	N	0	0
4109	N	0	0
4110	N	0	0
4111	N	0	0
4112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0

Sex : Female

Day : 1

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
1101	N	0	0
1102	N	0	0
1103	N	0	0
1104	N	0	0
1105	N	0	0
1106	N	0	0
1107	N	0	0
1108	N	0	0
1109	N	0	0
1110	N	0	0
1111	N	0	0
1112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0

Sex : Female

Day : 1

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2101	N	0	0
2102	N	0	0
2103	N	0	0
2104	N	0	0
2105	N	0	0
2107	N	0	0
2108	N	0	0
2110	N	0	0
2111	N	0	0
2112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 1 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3101	N	0	0
3102	N	0	0
3103	N	0	0
3104	N	0	0
3105	N	0	0
3106	N	0	0
3107	N	0	0
3110	N	0	0
3111	N	0	0
3112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 1 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4102	N	0	0
4103	N	0	0
4104	N	0	0
4105	N	0	0
4106	N	0	0
4107	N	0	0
4108	N	0	0
4109	N	0	0
4110	N	0	0
4111	N	0	0
4112	N	0	0
n	11	11	11

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0

Sex : Female

Day : 7

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Posture	Convulsion	Abnormal behavior
1101	N	0	0
1102	N	0	0
1103	N	0	0
1104	N	0	0
1105	N	0	0
1106	N	0	0
1107	N	0	0
1108	N	0	0
1109	N	0	0
1110	N	0	0
1111	N	0	0
1112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2101	N	0	0
2102	N	0	0
2103	N	0	0
2104	N	0	0
2105	N	0	0
2107	N	0	0
2108	N	0	0
2110	N	0	0
2111	N	0	0
2112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3101	N	0	0
3102	N	0	0
3103	N	0	0
3104	N	0	0
3105	N	0	0
3106	N	0	0
3107	N	0	0
3110	N	0	0
3111	N	0	0
3112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4102	N	0	0
4103	N	0	0
4104	N	0	0
4105	N	0	0
4106	N	0	0
4107	N	0	0
4108	N	0	0
4109	N	0	0
4110	N	0	0
4111	N	0	0
4112	N	0	0
n	11	11	11

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0

Sex : Female

Day : 14

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Posture	Convulsion	Abnormal behavior
1101	N	0	0
1102	N	0	0
1103	N	0	0
1104	N	0	0
1105	N	0	0
1106	N	0	0
1107	N	0	0
1108	N	0	0
1109	N	0	0
1110	N	0	0
1111	N	0	0
1112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2101	N	0	0
2102	N	0	0
2103	N	0	0
2104	N	0	0
2105	N	0	0
2107	N	0	0
2108	N	0	0
2110	N	0	0
2111	N	0	0
2112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3101	N	0	0
3102	N	0	0
3103	N	0	0
3104	N	0	0
3105	N	0	0
3106	N	0	0
3107	N	0	0
3110	N	0	0
3111	N	0	0
3112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4102	N	0	0
4103	N	0	0
4104	N	0	0
4105	N	0	0
4106	N	0	0
4107	N	0	0
4108	N	0	0
4109	N	0	0
4110	N	0	0
4111	N	0	0
4112	N	0	0
n	11	11	11

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0

Sex : Female

Day : 20

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
1101	N	0	0
1102	N	0	0
1103	N	0	0
1104	N	0	0
1105	N	0	0
1106	N	0	0
1107	N	0	0
1108	N	0	0
1109	N	0	0
1110	N	0	0
1111	N	0	0
1112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2101	N	0	0
2102	N	0	0
2103	N	0	0
2104	N	0	0
2105	N	0	0
2107	N	0	0
2108	N	0	0
2110	N	0	0
2111	N	0	0
2112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Appendix 2 - 47

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3101	N	0	0
3102	N	0	0
3103	N	0	0
3104	N	0	0
3105	N	0	0
3106	N	0	0
3107	N	0	0
3110	N	0	0
3111	N	0	0
3112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Appendix 2 - 48

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Home cage observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4102	N	0	0
4103	N	0	0
4104	N	0	0
4105	N	0	0
4106	N	0	0
4107	N	0	0
4108	N	0	0
4109	N	0	0
4110	N	0	0
4111	N	0	0
4112	N	0	0
n	11	11	11

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 lactation

Generation : F0

Sex : Female

Day : 6

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Posture	Convulsion	Abnormal behavior
1101	N	0	0
1102	N	0	0
1103	N	0	0
1104	N	0	0
1105	N	0	0
1106	N	0	0
1107	N	0	0
1108	N	0	0
1109	N	0	0
1110	N	0	0
1111	N	0	0
1112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 lactation

Generation : F0

Sex : Female

Day : 6

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2101	N	0	0
2102	N	0	0
2103	N	0	0
2104	N	0	0
2105	N	0	0
2107	N	0	0
2108	N	0	0
2110	N	0	0
2111	N	0	0
2112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 lactation

Generation : F0

Sex : Female

Day : 6

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3101	N	0	0
3102	N	0	0
3103	N	0	0
3104	N	0	0
3105	N	0	0
3106	N	0	0
3107	N	0	0
3110	N	0	0
3111	N	0	0
3112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 lactation

Generation : F0

Sex : Female

Day : 6

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4102	N	0	0
4103	N	0	0
4104	N	0	0
4105	N	0	0
4107	N	0	0
4108	N	0	0
4109	N	0	0
4110	N	0	0
4111	N	0	0
4112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 lactation

Generation : F0

Sex : Female

Day : 13

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Posture	Convulsion	Abnormal behavior
1101	N	0	0
1102	N	0	0
1103	N	0	0
1104	N	0	0
1105	N	0	0
1106	N	0	0
1107	N	0	0
1108	N	0	0
1109	N	0	0
1110	N	0	0
1111	N	0	0
1112	N	0	0
n	12	12	12

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 lactation

Generation : F0

Sex : Female

Day : 13

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
2101	N	0	0
2102	N	0	0
2103	N	0	0
2104	N	0	0
2105	N	0	0
2107	N	0	0
2108	N	0	0
2110	N	0	0
2111	N	0	0
2112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 lactation

Generation : F0

Sex : Female

Day : 13

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
3101	N	0	0
3102	N	0	0
3103	N	0	0
3104	N	0	0
3105	N	0	0
3106	N	0	0
3107	N	0	0
3110	N	0	0
3111	N	0	0
3112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : F0 lactation

Generation : F0

Sex : Female

Day : 13

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4102	N	0	0
4103	N	0	0
4104	N	0	0
4105	N	0	0
4107	N	0	0
4108	N	0	0
4109	N	0	0
4110	N	0	0
4111	N	0	0
4112	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
5101	N	0	0
5102	N	0	0
5103	N	0	0
5104	N	0	0
5105	N	0	0
5106	N	0	0
5107	N	0	0
5108	N	0	0
5109	N	0	0
5110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
6101	N	0	0
6102	N	0	0
6103	N	0	0
6104	N	0	0
6105	N	0	0
6106	N	0	0
6107	N	0	0
6108	N	0	0
6109	N	0	0
6110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
5101	N	0	0
5102	N	0	0
5103	N	0	0
5104	N	0	0
5105	N	0	0
5106	N	0	0
5107	N	0	0
5108	N	0	0
5109	N	0	0
5110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
6101	N	0	0
6102	N	0	0
6103	N	0	0
6104	N	0	0
6105	N	0	0
6106	N	0	0
6107	N	0	0
6108	N	0	0
6109	N	0	0
6110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 3

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
5101	N	0	0
5102	N	0	0
5103	N	0	0
5104	N	0	0
5105	N	0	0
5106	N	0	0
5107	N	0	0
5108	N	0	0
5109	N	0	0
5110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 3

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
6101	N	0	0
6102	N	0	0
6103	N	0	0
6104	N	0	0
6105	N	0	0
6106	N	0	0
6107	N	0	0
6108	N	0	0
6109	N	0	0
6110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 4

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
5101	N	0	0
5102	N	0	0
5103	N	0	0
5104	N	0	0
5105	N	0	0
5106	N	0	0
5107	N	0	0
5108	N	0	0
5109	N	0	0
5110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 4

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
6101	N	0	0
6102	N	0	0
6103	N	0	0
6104	N	0	0
6105	N	0	0
6106	N	0	0
6107	N	0	0
6108	N	0	0
6109	N	0	0
6110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 5

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
5101	N	0	0
5102	N	0	0
5103	N	0	0
5104	N	0	0
5105	N	0	0
5106	N	0	0
5107	N	0	0
5108	N	0	0
5109	N	0	0
5110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 5

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
6101	N	0	0
6102	N	0	0
6103	N	0	0
6104	N	0	0
6105	N	0	0
6106	N	0	0
6107	N	0	0
6108	N	0	0
6109	N	0	0
6110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
5101	N	0	0
5102	N	0	0
5103	N	0	0
5104	N	0	0
5105	N	0	0
5106	N	0	0
5107	N	0	0
5108	N	0	0
5109	N	0	0
5110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
6101	N	0	0
6102	N	0	0
6103	N	0	0
6104	N	0	0
6105	N	0	0
6106	N	0	0
6107	N	0	0
6108	N	0	0
6109	N	0	0
6110	N	0	0
n	10	10	10

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Recovery

Generation : F0

Sex : Male

Week : 1

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
1008	N	0	0
1009	N	0	0
1010	N	0	0
1011	N	0	0
1012	N	0	0
n	5	5	5

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Recovery

Generation : F0

Sex : Male

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4008	N	0	0
4009	N	0	0
4010	N	0	0
4011	N	0	0
4012	N	0	0
n	5	5	5

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Recovery

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
1008	N	0	0
1009	N	0	0
1010	N	0	0
1011	N	0	0
1012	N	0	0
n	5	5	5

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Recovery

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
4008	N	0	0
4009	N	0	0
4010	N	0	0
4011	N	0	0
4012	N	0	0
n	5	5	5

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
5106	N	0	0
5107	N	0	0
5108	N	0	0
5109	N	0	0
5110	N	0	0
n	5	5	5

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
6106	N	0	0
6107	N	0	0
6108	N	0	0
6109	N	0	0
6110	N	0	0
n	5	5	5

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
5106	N	0	0
5107	N	0	0
5108	N	0	0
5109	N	0	0
5110	N	0	0
n	5	5	5

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Home cage observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Posture	Convulsion	Abnormal behavior
Animal No.			
6106	N	0	0
6107	N	0	0
6108	N	0	0
6109	N	0	0
6110	N	0	0
n	5	5	5

Posture) N: Normal, F: Flattened, H: Hunched

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

In-the-hand observation				Period : F0 before mating			Species : Rat	
Generation : F0	Sex : Male	Week : 1	Dose : Dodecyloxirane 0 mg/kg					
	Ease of removal from cage	Fur condition	Skin	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
1001	0	0	0	0	0	0	0	0
1002	0	0	0	0	0	0	0	0
1003	0	0	0	0	0	0	0	0
1004	0	0	0	0	0	0	0	0
1005	0	0	0	0	0	0	0	0
1006	0	0	0	0	0	0	0	0
1007	0	0	0	0	0	0	0	0
1008	0	0	0	0	0	0	0	0
1009	0	0	0	0	0	0	0	0
1010	0	0	0	0	0	0	0	0
1011	0	0	0	0	0	0	0	0
1012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
1001	0	0	0	0	0	0
1002	0	0	0	0	0	0
1003	0	0	0	0	0	0
1004	0	0	0	0	0	0
1005	0	0	0	0	0	0
1006	0	0	0	0	0	0
1007	0	0	0	0	0	0
1008	0	0	0	0	0	0
1009	0	0	0	0	0	0
1010	0	0	0	0	0	0
1011	0	0	0	0	0	0
1012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

With the reproduction/ developmental toxicity screening test in rats								
In-the-hand observation		Period : F0 before mating			Species : Rat			
Generation : F0	Sex : Male	Week : 1	Dose : Dodecyloxirane 100 mg/kg					
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation	
Animal No.								
2001	0	0	0	0	0	0	0	0
2002	0	0	0	0	0	0	0	0
2003	0	0	0	0	0	0	0	0
2004	0	0	0	0	0	0	0	0
2005	0	0	0	0	0	0	0	0
2006	0	0	0	0	0	0	0	0
2007	0	0	0	0	0	0	0	0
2008	0	0	0	0	0	0	0	0
2009	0	0	0	0	0	0	0	0
2010	0	0	0	0	0	0	0	0
2011	0	0	0	0	0	0	0	0
2012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 before mating		Dose : Dodecyloxirane 300 mg/kg		Species : Rat			
Generation : F0	Sex : Male	Week : 1							
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation		
Animal No.									
3001	0	0	0	0	0	0	0	0	0
3002	0	0	0	0	0	0	0	0	0
3003	0	0	0	0	0	0	0	0	0
3004	0	0	0	0	0	0	0	0	0
3005	0	0	0	0	0	0	0	0	0
3006	0	0	0	0	0	0	0	0	0
3007	0	0	0	0	0	0	0	0	0
3008	0	0	0	0	0	0	0	0	0
3009	0	0	0	0	0	0	0	0	0
3010	0	0	0	0	0	0	0	0	0
3011	0	0	0	0	0	0	0	0	0
3012	0	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
3001	0	0	0	0	0
3002	0	0	0	0	0
3003	0	0	0	0	0
3004	0	0	0	0	0
3005	0	0	0	0	0
3006	0	0	0	0	0
3007	0	0	0	0	0
3008	0	0	0	0	0
3009	0	0	0	0	0
3010	0	0	0	0	0
3011	0	0	0	0	0
3012	0	0	0	0	0
n	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

With the reproduction/ developmental toxicity screening test in rats				Period : F0 before mating		Species : Rat		
In-the-hand observation	Sex : Male	Week : 1	Dose : Dodecyloxirane 1000 mg/kg					
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
4001	0	0	0	0	0	0	0	0
4002	0	0	0	0	0	0	0	0
4003	0	0	0	0	0	0	0	0
4004	0	0	0	0	0	0	0	0
4005	0	0	0	0	0	0	0	0
4006	0	0	0	0	0	0	0	0
4007	0	0	0	0	0	0	0	0
4008	0	0	0	0	0	0	0	0
4009	0	0	0	0	0	0	0	0
4010	0	0	0	0	0	0	0	0
4011	0	0	0	0	0	0	0	0
4012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
4001	0	0	0	0	0
4002	0	0	0	0	0
4003	0	0	0	0	0
4004	0	0	0	0	0
4005	0	0	0	0	0
4006	0	0	0	0	0
4007	0	0	0	0	0
4008	0	0	0	0	0
4009	0	0	0	0	0
4010	0	0	0	0	0
4011	0	0	0	0	0
4012	0	0	0	0	0
n	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 before mating		Species : Rat				
Generation : F0	Sex : Male	Week : 2	Dose : Dodecyloxirane 0 mg/kg					
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
1001	0	0	0	0	0	0	0	0
1002	0	0	0	0	0	0	0	0
1003	0	0	0	0	0	0	0	0
1004	0	0	0	0	0	0	0	0
1005	0	0	0	0	0	0	0	0
1006	0	0	0	0	0	0	0	0
1007	0	0	0	0	0	0	0	0
1008	0	0	0	0	0	0	0	0
1009	0	0	0	0	0	0	0	0
1010	0	0	0	0	0	0	0	0
1011	0	0	0	0	0	0	0	0
1012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
1001	0	0	0	0	0	0
1002	0	0	0	0	0	0
1003	0	0	0	0	0	0
1004	0	0	0	0	0	0
1005	0	0	0	0	0	0
1006	0	0	0	0	0	0
1007	0	0	0	0	0	0
1008	0	0	0	0	0	0
1009	0	0	0	0	0	0
1010	0	0	0	0	0	0
1011	0	0	0	0	0	0
1012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 before mating		Species : Rat				
Generation : F0		Sex : Male	Week : 2	Dose : Dodecyloxirane 100 mg/kg				
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
2001	0	0	0	0	0	0	0	0
2002	0	0	0	0	0	0	0	0
2003	0	0	0	0	0	0	0	0
2004	0	0	0	0	0	0	0	0
2005	0	0	0	0	0	0	0	0
2006	0	0	0	0	0	0	0	0
2007	0	0	0	0	0	0	0	0
2008	0	0	0	0	0	0	0	0
2009	0	0	0	0	0	0	0	0
2010	0	0	0	0	0	0	0	0
2011	0	0	0	0	0	0	0	0
2012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
2001	0	0	0	0	0	0
2002	0	0	0	0	0	0
2003	0	0	0	0	0	0
2004	0	0	0	0	0	0
2005	0	0	0	0	0	0
2006	0	0	0	0	0	0
2007	0	0	0	0	0	0
2008	0	0	0	0	0	0
2009	0	0	0	0	0	0
2010	0	0	0	0	0	0
2011	0	0	0	0	0	0
2012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 before mating		Dose : Dodecyloxirane 300 mg/kg		Species : Rat			
Generation : F0	Sex : Male	Week : 2							
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation		
Animal No.									
3001	0	0	0	0	0	0	0	0	0
3002	0	0	0	0	0	0	0	0	0
3003	0	0	0	0	0	0	0	0	0
3004	0	0	0	0	0	0	0	0	0
3005	0	0	0	0	0	0	0	0	0
3006	0	0	0	0	0	0	0	0	0
3007	0	0	0	0	0	0	0	0	0
3008	0	0	0	0	0	0	0	0	0
3009	0	0	0	0	0	0	0	0	0
3010	0	0	0	0	0	0	0	0	0
3011	0	0	0	0	0	0	0	0	0
3012	0	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling		
Animal No.							
3001	0	0	0	0	0		
3002	0	0	0	0	0		
3003	0	0	0	0	0		
3004	0	0	0	0	0		
3005	0	0	0	0	0		
3006	0	0	0	0	0		
3007	0	0	0	0	0		
3008	0	0	0	0	0		
3009	0	0	0	0	0		
3010	0	0	0	0	0		
3011	0	0	0	0	0		
3012	0	0	0	0	0		
n	12	12	12	12	12		

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 before mating		Dose : Dodecyloxirane 1000 mg/kg		Species : Rat			
Generation : F0	Sex : Male	Week : 2							
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation		
Animal No.									
4001	0	0	0	0	0	0	0	0	0
4002	0	0	0	0	0	0	0	0	0
4003	0	0	0	0	0	0	0	0	0
4004	0	0	0	0	0	0	0	0	0
4005	0	0	0	0	0	0	0	0	0
4006	0	0	0	0	0	0	0	0	0
4007	0	0	0	0	0	0	0	0	0
4008	0	0	0	0	0	0	0	0	0
4009	0	0	0	0	0	0	0	0	0
4010	0	0	0	0	0	0	0	0	0
4011	0	0	0	0	0	0	0	0	0
4012	0	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
4001	0	0	0	0	0
4002	0	0	0	0	0
4003	0	0	0	0	0
4004	0	0	0	0	0
4005	0	0	0	0	0
4006	0	0	0	0	0
4007	0	0	0	0	0
4008	0	0	0	0	0
4009	0	0	0	0	0
4010	0	0	0	0	0
4011	0	0	0	0	0
4012	0	0	0	0	0
n	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation			Period : F0 mating				
	Generation : F0	Sex : Male	Week : 3	Dose : Dodecyloxirane 0 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
1001	0	0	0	0	0	0	0	0
1002	0	0	0	0	0	0	0	0
1003	0	0	0	0	0	0	0	0
1004	0	0	0	0	0	0	0	0
1005	0	0	0	0	0	0	0	0
1006	0	0	0	0	0	0	0	0
1007	0	0	0	0	0	0	0	0
1008	0	0	0	0	0	0	0	0
1009	0	0	0	0	0	0	0	0
1010	0	0	0	0	0	0	0	0
1011	0	0	0	0	0	0	0	0
1012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
1001	0	0	0	0	0	0
1002	0	0	0	0	0	0
1003	0	0	0	0	0	0
1004	0	0	0	0	0	0
1005	0	0	0	0	0	0
1006	0	0	0	0	0	0
1007	0	0	0	0	0	0
1008	0	0	0	0	0	0
1009	0	0	0	0	0	0
1010	0	0	0	0	0	0
1011	0	0	0	0	0	0
1012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation Generation : F0	Sex : Male	Week : 3	Period : F0 mating Dose : Dodecyloxirane 100 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
2001	0	0	0	0	0	0	0	0
2002	0	0	0	0	0	0	0	0
2003	0	0	0	0	0	0	0	0
2004	0	0	0	0	0	0	0	0
2005	0	0	0	0	0	0	0	0
2006	0	0	0	0	0	0	0	0
2007	0	0	0	0	0	0	0	0
2008	0	0	0	0	0	0	0	0
2009	0	0	0	0	0	0	0	0
2010	0	0	0	0	0	0	0	0
2011	0	0	0	0	0	0	0	0
2012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
2001	0	0	0	0	0	0
2002	0	0	0	0	0	0
2003	0	0	0	0	0	0
2004	0	0	0	0	0	0
2005	0	0	0	0	0	0
2006	0	0	0	0	0	0
2007	0	0	0	0	0	0
2008	0	0	0	0	0	0
2009	0	0	0	0	0	0
2010	0	0	0	0	0	0
2011	0	0	0	0	0	0
2012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 mating		Dose : Dodecyloxirane 300 mg/kg		Species : Rat			
Generation : F0	Sex : Male	Week : 3							
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation		
Animal No.									
3001	0	0	0	0	0	0	0	0	0
3002	0	0	0	0	0	0	0	0	0
3003	0	0	0	0	0	0	0	0	0
3004	0	0	0	0	0	0	0	0	0
3005	0	0	0	0	0	0	0	0	0
3006	0	0	0	0	0	0	0	0	0
3007	0	0	0	0	0	0	0	0	0
3008	0	0	0	0	0	0	0	0	0
3009	0	0	0	0	0	0	0	0	0
3010	0	0	0	0	0	0	0	0	0
3011	0	0	0	0	0	0	0	0	0
3012	0	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling		
Animal No.							
3001	0	0	0	0	0		
3002	0	0	0	0	0		
3003	0	0	0	0	0		
3004	0	0	0	0	0		
3005	0	0	0	0	0		
3006	0	0	0	0	0		
3007	0	0	0	0	0		
3008	0	0	0	0	0		
3009	0	0	0	0	0		
3010	0	0	0	0	0		
3011	0	0	0	0	0		
3012	0	0	0	0	0		
n	12	12	12	12	12		

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation Generation : F0	Sex : Male	Week : 3	Period : F0 mating Dose : Dodecyloxirane 1000 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
4001	0	0	0	0	0	0	0	0
4002	0	0	0	0	0	0	0	0
4003	0	0	0	0	0	0	0	0
4004	0	0	0	0	0	0	0	0
4005	0	0	0	0	0	0	0	0
4006	0	0	0	0	0	0	0	0
4007	0	0	0	0	0	0	0	0
4008	0	0	0	0	0	0	0	0
4009	0	0	0	0	0	0	0	0
4010	0	0	0	0	0	0	0	0
4011	0	0	0	0	0	0	0	0
4012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
4001	0	0	0	0	0	0
4002	0	0	0	0	0	0
4003	0	0	0	0	0	0
4004	0	0	0	0	0	0
4005	0	0	0	0	0	0
4006	0	0	0	0	0	0
4007	0	0	0	0	0	0
4008	0	0	0	0	0	0
4009	0	0	0	0	0	0
4010	0	0	0	0	0	0
4011	0	0	0	0	0	0
4012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 mating		Dose : Dodecyloxirane 0 mg/kg		Species : Rat			
Generation : F0	Sex : Male	Week : 4							
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation	
1001	0	0	0	0	0	0	0	0	
1002	0	0	0	0	0	0	0	0	
1003	0	0	0	0	0	0	0	0	
1004	0	0	0	0	0	0	0	0	
1005	0	0	0	0	0	0	0	0	
1006	0	0	0	0	0	0	0	0	
1007	0	0	0	0	0	0	0	0	
1008	0	0	0	0	0	0	0	0	
1009	0	0	0	0	0	0	0	0	
1010	0	0	0	0	0	0	0	0	
1011	0	0	0	0	0	0	0	0	
1012	0	0	0	0	0	0	0	0	
n	12	12	12	12	12	12	12	12	

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling	
1001	0	0	0	0	0	0	
1002	0	0	0	0	0	0	
1003	0	0	0	0	0	0	
1004	0	0	0	0	0	0	
1005	0	0	0	0	0	0	
1006	0	0	0	0	0	0	
1007	0	0	0	0	0	0	
1008	0	0	0	0	0	0	
1009	0	0	0	0	0	0	
1010	0	0	0	0	0	0	
1011	0	0	0	0	0	0	
1012	0	0	0	0	0	0	
n	12	12	12	12	12	12	

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation			Period : F0 mating				
	Generation : F0	Sex : Male	Week : 4	Dose : Dodecyloxirane 100 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
2001	0	0	0	0	0	0	0	0
2002	0	0	0	0	0	0	0	0
2003	0	0	0	0	0	0	0	0
2004	0	0	0	0	0	0	0	0
2005	0	0	0	0	0	0	0	0
2006	0	0	0	0	0	0	0	0
2007	0	0	0	0	0	0	0	0
2008	0	0	0	0	0	0	0	0
2009	0	0	0	0	0	0	0	0
2010	0	0	0	0	0	0	0	0
2011	0	0	0	0	0	0	0	0
2012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
2001	0	0	0	0	0	0
2002	0	0	0	0	0	0
2003	0	0	0	0	0	0
2004	0	0	0	0	0	0
2005	0	0	0	0	0	0
2006	0	0	0	0	0	0
2007	0	0	0	0	0	0
2008	0	0	0	0	0	0
2009	0	0	0	0	0	0
2010	0	0	0	0	0	0
2011	0	0	0	0	0	0
2012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation			Period : F0 mating				
	Generation : F0	Sex : Male	Week : 4	Dose : Dodecyloxirane 300 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
3001	0	0	0	0	0	0	0	0
3002	0	0	0	0	0	0	0	0
3003	0	0	0	0	0	0	0	0
3004	0	0	0	0	0	0	0	0
3005	0	0	0	0	0	0	0	0
3006	0	0	0	0	0	0	0	0
3007	0	0	0	0	0	0	0	0
3008	0	0	0	0	0	0	0	0
3009	0	0	0	0	0	0	0	0
3010	0	0	0	0	0	0	0	0
3011	0	0	0	0	0	0	0	0
3012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
3001	0	0	0	0	0	0
3002	0	0	0	0	0	0
3003	0	0	0	0	0	0
3004	0	0	0	0	0	0
3005	0	0	0	0	0	0
3006	0	0	0	0	0	0
3007	0	0	0	0	0	0
3008	0	0	0	0	0	0
3009	0	0	0	0	0	0
3010	0	0	0	0	0	0
3011	0	0	0	0	0	0
3012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 mating		Dose : Dodecyloxirane 1000 mg/kg		Species : Rat			
Generation : F0	Sex : Male	Week : 4							
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation		
Animal No.									
4001	0	0	0	0	0	0	0	0	0
4002	0	0	0	0	0	0	0	0	0
4003	0	0	0	0	0	0	0	0	0
4004	0	0	0	0	0	0	0	0	0
4005	0	0	0	0	0	0	0	0	0
4006	0	0	0	0	0	0	0	0	0
4007	0	0	0	0	0	0	0	0	0
4008	0	0	0	0	0	0	0	0	0
4009	0	0	0	0	0	0	0	0	0
4010	0	0	0	0	0	0	0	0	0
4011	0	0	0	0	0	0	0	0	0
4012	0	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling		
Animal No.							
4001	0	0	0	0	0		
4002	0	0	0	0	0		
4003	0	0	0	0	0		
4004	0	0	0	0	0		
4005	0	0	0	0	0		
4006	0	0	0	0	0		
4007	0	0	0	0	0		
4008	0	0	0	0	0		
4009	0	0	0	0	0		
4010	0	0	0	0	0		
4011	0	0	0	0	0		
4012	0	0	0	0	0		
n	12	12	12	12	12		

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 mating		Dose : Dodecyloxirane 0 mg/kg		Species : Rat			
Generation : F0		Sex : Male		Week : 5					
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation	
1001	0	0	0	0	0	0	0	0	
1002	0	0	0	0	0	0	0	0	
1003	0	0	0	0	0	0	0	0	
1004	0	0	0	0	0	0	0	0	
1005	0	0	0	0	0	0	0	0	
1006	0	0	0	0	0	0	0	0	
1007	0	0	0	0	0	0	0	0	
1008	0	0	0	0	0	0	0	0	
1009	0	0	0	0	0	0	0	0	
1010	0	0	0	0	0	0	0	0	
1011	0	0	0	0	0	0	0	0	
1012	0	0	0	0	0	0	0	0	
n	12	12	12	12	12	12	12	12	

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling	
1001	0	0	0	0	0	0	
1002	0	0	0	0	0	0	
1003	0	0	0	0	0	0	
1004	0	0	0	0	0	0	
1005	0	0	0	0	0	0	
1006	0	0	0	0	0	0	
1007	0	0	0	0	0	0	
1008	0	0	0	0	0	0	
1009	0	0	0	0	0	0	
1010	0	0	0	0	0	0	
1011	0	0	0	0	0	0	
1012	0	0	0	0	0	0	
n	12	12	12	12	12	12	

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation			Period : F0 mating				
	Generation : F0	Sex : Male	Week : 5	Dose : Dodecyloxirane 100 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
2001	0	0	0	0	0	0	0	0
2002	0	0	0	0	0	0	0	0
2003	0	0	0	0	0	0	0	0
2004	0	0	0	0	0	0	0	0
2005	0	0	0	0	0	0	0	0
2006	0	0	0	0	0	0	0	0
2007	0	0	0	0	0	0	0	0
2008	0	0	0	0	0	0	0	0
2009	0	0	0	0	0	0	0	0
2010	0	0	0	0	0	0	0	0
2011	0	0	0	0	0	0	0	0
2012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
2001	0	0	0	0	0	0
2002	0	0	0	0	0	0
2003	0	0	0	0	0	0
2004	0	0	0	0	0	0
2005	0	0	0	0	0	0
2006	0	0	0	0	0	0
2007	0	0	0	0	0	0
2008	0	0	0	0	0	0
2009	0	0	0	0	0	0
2010	0	0	0	0	0	0
2011	0	0	0	0	0	0
2012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 mating		Dose : Dodecyloxirane 300 mg/kg		Species : Rat			
Generation : F0		Sex : Male		Week : 5					
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation	
3001	0	0	0	0	0	0	0	0	
3002	0	0	0	0	0	0	0	0	
3003	0	0	0	0	0	0	0	0	
3004	0	0	0	0	0	0	0	0	
3005	0	0	0	0	0	0	0	0	
3006	0	0	0	0	0	0	0	0	
3007	0	0	0	0	0	0	0	0	
3008	0	0	0	0	0	0	0	0	
3009	0	0	0	0	0	0	0	0	
3010	0	0	0	0	0	0	0	0	
3011	0	0	0	0	0	0	0	0	
3012	0	0	0	0	0	0	0	0	
n	12	12	12	12	12	12	12	12	

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling	
3001	0	0	0	0	0	0	
3002	0	0	0	0	0	0	
3003	0	0	0	0	0	0	
3004	0	0	0	0	0	0	
3005	0	0	0	0	0	0	
3006	0	0	0	0	0	0	
3007	0	0	0	0	0	0	
3008	0	0	0	0	0	0	
3009	0	0	0	0	0	0	
3010	0	0	0	0	0	0	
3011	0	0	0	0	0	0	
3012	0	0	0	0	0	0	
n	12	12	12	12	12	12	

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation Generation : F0	Sex : Male	Week : 5	Period : F0 mating Dose : Dodecyloxirane 1000 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
4001	0	0	0	0	0	0	0	0
4002	0	0	0	0	0	0	0	0
4003	0	0	0	0	0	0	0	0
4004	0	0	0	0	0	0	0	0
4005	0	0	0	0	0	0	0	0
4006	0	0	0	0	0	0	0	0
4007	0	0	0	0	0	0	0	0
4008	0	0	0	0	0	0	0	0
4009	0	0	0	0	0	0	0	0
4010	0	0	0	0	0	0	0	0
4011	0	0	0	0	0	0	0	0
4012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
4001	0	0	0	0	0	0
4002	0	0	0	0	0	0
4003	0	0	0	0	0	0
4004	0	0	0	0	0	0
4005	0	0	0	0	0	0
4006	0	0	0	0	0	0
4007	0	0	0	0	0	0
4008	0	0	0	0	0	0
4009	0	0	0	0	0	0
4010	0	0	0	0	0	0
4011	0	0	0	0	0	0
4012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation Generation : F0	Sex : Male	Week : 6	Period : F0 mating Dose : Dodecyloxirane 0 mg/kg			Species : Rat	
	Ease of remov- al from cage	Fur condition	Skin	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
1001	0	0	0	0	0	0	0	0
1002	0	0	0	0	0	0	0	0
1003	0	0	0	0	0	0	0	0
1004	0	0	0	0	0	0	0	0
1005	0	0	0	0	0	0	0	0
1006	0	0	0	0	0	0	0	0
1007	0	0	0	0	0	0	0	0
1008	0	0	0	0	0	0	0	0
1009	0	0	0	0	0	0	0	0
1010	0	0	0	0	0	0	0	0
1011	0	0	0	0	0	0	0	0
1012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
1001	0	0	0	0	0	0
1002	0	0	0	0	0	0
1003	0	0	0	0	0	0
1004	0	0	0	0	0	0
1005	0	0	0	0	0	0
1006	0	0	0	0	0	0
1007	0	0	0	0	0	0
1008	0	0	0	0	0	0
1009	0	0	0	0	0	0
1010	0	0	0	0	0	0
1011	0	0	0	0	0	0
1012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation Generation : F0	Sex : Male	Week : 6	Period : F0 mating Dose : Dodecyloxirane 100 mg/kg	Species : Rat			
	Ease of removal from cage	Fur condition	Skin	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
2001	0	0	0	0	0	0	0	0
2002	0	0	0	0	0	0	0	0
2003	0	0	0	0	0	0	0	0
2004	0	0	0	0	0	0	0	0
2005	0	0	0	0	0	0	0	0
2006	0	0	0	0	0	0	0	0
2007	0	0	0	0	0	0	0	0
2008	0	0	0	0	0	0	0	0
2009	0	0	0	0	0	0	0	0
2010	0	0	0	0	0	0	0	0
2011	0	0	0	0	0	0	0	0
2012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
2001	0	0	0	0	0	0
2002	0	0	0	0	0	0
2003	0	0	0	0	0	0
2004	0	0	0	0	0	0
2005	0	0	0	0	0	0
2006	0	0	0	0	0	0
2007	0	0	0	0	0	0
2008	0	0	0	0	0	0
2009	0	0	0	0	0	0
2010	0	0	0	0	0	0
2011	0	0	0	0	0	0
2012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation			Period : F0 mating				
	Generation : F0	Sex : Male	Week : 6	Dose : Dodecyloxirane 300 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
3001	0	0	0	0	0	0	0	0
3002	0	0	0	0	0	0	0	0
3003	0	0	0	0	0	0	0	0
3004	0	0	0	0	0	0	0	0
3005	0	0	0	0	0	0	0	0
3006	0	0	0	0	0	0	0	0
3007	0	0	0	0	0	0	0	0
3008	0	0	0	0	0	0	0	0
3009	0	0	0	0	0	0	0	0
3010	0	0	0	0	0	0	0	0
3011	0	0	0	0	0	0	0	0
3012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
3001	0	0	0	0	0	0
3002	0	0	0	0	0	0
3003	0	0	0	0	0	0
3004	0	0	0	0	0	0
3005	0	0	0	0	0	0
3006	0	0	0	0	0	0
3007	0	0	0	0	0	0
3008	0	0	0	0	0	0
3009	0	0	0	0	0	0
3010	0	0	0	0	0	0
3011	0	0	0	0	0	0
3012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation Generation : F0	Sex : Male	Week : 6	Period : F0 mating Dose : Dodecyloxirane 1000 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
4001	0	0	0	0	0	0	0	0
4002	0	0	0	0	0	0	0	0
4003	0	0	0	0	0	0	0	0
4004	0	0	0	0	0	0	0	0
4005	0	0	0	0	0	0	0	0
4006	0	0	0	0	0	0	0	0
4007	0	0	0	0	0	0	0	0
4008	0	0	0	0	0	0	0	0
4009	0	0	0	0	0	0	0	0
4010	0	0	0	0	0	0	0	0
4011	0	0	0	0	0	0	0	0
4012	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
4001	0	0	0	0	0	0
4002	0	0	0	0	0	0
4003	0	0	0	0	0	0
4004	0	0	0	0	0	0
4005	0	0	0	0	0	0
4006	0	0	0	0	0	0
4007	0	0	0	0	0	0
4008	0	0	0	0	0	0
4009	0	0	0	0	0	0
4010	0	0	0	0	0	0
4011	0	0	0	0	0	0
4012	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation Generation : F0	Sex : Female	Week : 1	Period : F0 before mating Dose : Dodecyloxirane 0 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
1101	0	0	0	0	0	0	0	0
1102	0	0	0	0	0	0	0	0
1103	0	0	0	0	0	0	0	0
1104	0	0	0	0	0	0	0	0
1105	0	0	0	0	0	0	0	0
1106	0	0	0	0	0	0	0	0
1107	0	0	0	0	0	0	0	0
1108	0	0	0	0	0	0	0	0
1109	0	0	0	0	0	0	0	0
1110	0	0	0	0	0	0	0	0
1111	0	0	0	0	0	0	0	0
1112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
1101	0	0	0	0	0	0
1102	0	0	0	0	0	0
1103	0	0	0	0	0	0
1104	0	0	0	0	0	0
1105	0	0	0	0	0	0
1106	0	0	0	0	0	0
1107	0	0	0	0	0	0
1108	0	0	0	0	0	0
1109	0	0	0	0	0	0
1110	0	0	0	0	0	0
1111	0	0	0	0	0	0
1112	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 before mating		Dose : Dodecyloxirane 100 mg/kg		Species : Rat			
Generation : F0	Sex : Female	Week : 1							
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation		
Animal No.									
2101	0	0	0	0	0	0	0	0	0
2102	0	0	0	0	0	0	0	0	0
2103	0	0	0	0	0	0	0	0	0
2104	0	0	0	0	0	0	0	0	0
2105	0	0	0	0	0	0	0	0	0
2106	0	0	0	0	0	0	0	0	0
2107	0	0	0	0	0	0	0	0	0
2108	0	0	0	0	0	0	0	0	0
2109	0	0	0	0	0	0	0	0	0
2110	0	0	0	0	0	0	0	0	0
2111	0	0	0	0	0	0	0	0	0
2112	0	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
2101	0	0	0	0	0
2102	0	0	0	0	0
2103	0	0	0	0	0
2104	0	0	0	0	0
2105	0	0	0	0	0
2106	0	0	0	0	0
2107	0	0	0	0	0
2108	0	0	0	0	0
2109	0	0	0	0	0
2110	0	0	0	0	0
2111	0	0	0	0	0
2112	0	0	0	0	0
n	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Appendix 2 - 103

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

In-the-hand observation				Period : F0 before mating		Species : Rat		
Generation : F0	Sex : Female	Week : 1	Dose : Dodecyloxirane 300 mg/kg	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Ease of remov- al from cage	Fur condition	Skin						
Animal No.								
3101	0	0	0	0	0	0	0	0
3102	0	0	0	0	0	0	0	0
3103	0	0	0	0	0	0	0	0
3104	0	0	0	0	0	0	0	0
3105	0	0	0	0	0	0	0	0
3106	0	0	0	0	0	0	0	0
3107	0	0	0	0	0	0	0	0
3108	0	0	0	0	0	0	0	0
3109	0	0	0	0	0	0	0	0
3110	0	0	0	0	0	0	0	0
3111	0	0	0	0	0	0	0	0
3112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
3101	0	0	0	0	0
3102	0	0	0	0	0
3103	0	0	0	0	0
3104	0	0	0	0	0
3105	0	0	0	0	0
3106	0	0	0	0	0
3107	0	0	0	0	0
3108	0	0	0	0	0
3109	0	0	0	0	0
3110	0	0	0	0	0
3111	0	0	0	0	0
3112	0	0	0	0	0
n	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation				Period : F0 before mating		Species : Rat		
Generation : F0	Sex : Female	Week : 1	Dose : Dodecyloxirane 1000 mg/kg	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Ease of remov- al from cage	Fur condition	Skin						
Animal No.								
4101	0	0	0	0	0	0	0	0
4102	0	0	0	0	0	0	0	0
4103	0	0	0	0	0	0	0	0
4104	0	0	0	0	0	0	0	0
4105	0	0	0	0	0	0	0	0
4106	0	0	0	0	0	0	0	0
4107	0	0	0	0	0	0	0	0
4108	0	0	0	0	0	0	0	0
4109	0	0	0	0	0	0	0	0
4110	0	0	0	0	0	0	0	0
4111	0	0	0	0	0	0	0	0
4112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
4101	0	0	0	0	0
4102	0	0	0	0	0
4103	0	0	0	0	0
4104	0	0	0	0	0
4105	0	0	0	0	0
4106	0	0	0	0	0
4107	0	0	0	0	0
4108	0	0	0	0	0
4109	0	0	0	0	0
4110	0	0	0	0	0
4111	0	0	0	0	0
4112	0	0	0	0	0
n	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation			Period : F0 before mating				
	Generation : F0	Sex : Female	Week : 2	Dose : Dodecyloxirane 0 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
1101	0	0	0	0	0	0	0	0
1102	0	0	0	0	0	0	0	0
1103	0	0	0	0	0	0	0	0
1104	0	0	0	0	0	0	0	0
1105	0	0	0	0	0	0	0	0
1106	0	0	0	0	0	0	0	0
1107	0	0	0	0	0	0	0	0
1108	0	0	0	0	0	0	0	0
1109	0	0	0	0	0	0	0	0
1110	0	0	0	0	0	0	0	0
1111	0	0	0	0	0	0	0	0
1112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
1101	0	0	0	0	0	0
1102	0	0	0	0	0	0
1103	0	0	0	0	0	0
1104	0	0	0	0	0	0
1105	0	0	0	0	0	0
1106	0	0	0	0	0	0
1107	0	0	0	0	0	0
1108	0	0	0	0	0	0
1109	0	0	0	0	0	0
1110	0	0	0	0	0	0
1111	0	0	0	0	0	0
1112	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 before mating		Dose : Dodecyloxirane 100 mg/kg		Species : Rat			
Generation : F0	Sex : Female	Week : 2							
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation		
Animal No.									
2101	0	0	0	0	0	0	0	0	0
2102	0	0	0	0	0	0	0	0	0
2103	0	0	0	0	0	0	0	0	0
2104	0	0	0	0	0	0	0	0	0
2105	0	0	0	0	0	0	0	0	0
2106	0	0	0	0	0	0	0	0	0
2107	0	0	0	0	0	0	0	0	0
2108	0	0	0	0	0	0	0	0	0
2109	0	0	0	0	0	0	0	0	0
2110	0	0	0	0	0	0	0	0	0
2111	0	0	0	0	0	0	0	0	0
2112	0	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
2101	0	0	0	0	0
2102	0	0	0	0	0
2103	0	0	0	0	0
2104	0	0	0	0	0
2105	0	0	0	0	0
2106	0	0	0	0	0
2107	0	0	0	0	0
2108	0	0	0	0	0
2109	0	0	0	0	0
2110	0	0	0	0	0
2111	0	0	0	0	0
2112	0	0	0	0	0
n	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Appendix 2 - 107

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation

Study No. : R-1261

Generation : F0		Sex : Female	Week : 2	Period : F0 before mating Dose : Dodecyloxirane 300 mg/kg		Species : Rat		
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
3101	0	0	0	0	0	0	0	0
3102	0	0	0	0	0	0	0	0
3103	0	0	0	0	0	0	0	0
3104	0	0	0	0	0	0	0	0
3105	0	0	0	0	0	0	0	0
3106	0	0	0	0	0	0	0	0
3107	0	0	0	0	0	0	0	0
3108	0	0	0	0	0	0	0	0
3109	0	0	0	0	0	0	0	0
3110	0	0	0	0	0	0	0	0
3111	0	0	0	0	0	0	0	0
3112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
3101	0	0	0	0	0	0
3102	0	0	0	0	0	0
3103	0	0	0	0	0	0
3104	0	0	0	0	0	0
3105	0	0	0	0	0	0
3106	0	0	0	0	0	0
3107	0	0	0	0	0	0
3108	0	0	0	0	0	0
3109	0	0	0	0	0	0
3110	0	0	0	0	0	0
3111	0	0	0	0	0	0
3112	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Appendix 2 - 108

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

In-the-hand observation				Period : F0 before mating		Species : Rat		
Generation : F0	Sex : Female	Week : 2	Dose : Dodecyloxirane 1000 mg/kg	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Ease of remov- al from cage	Fur condition	Skin						
Animal No.								
4101	0	0	0	0	0	0	0	0
4102	0	0	0	0	0	0	0	0
4103	0	0	0	0	0	0	0	0
4104	0	0	0	0	0	0	0	0
4105	0	0	0	0	0	0	0	0
4106	0	0	0	0	0	0	0	0
4107	0	0	0	0	0	0	0	0
4108	0	0	0	0	0	0	0	0
4109	0	0	0	0	0	0	0	0
4110	0	0	0	0	0	0	0	0
4111	0	0	0	0	0	0	0	0
4112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
4101	0	0	0	0	0
4102	0	0	0	0	0
4103	0	0	0	0	0
4104	0	0	0	0	0
4105	0	0	0	0	0
4106	0	0	0	0	0
4107	0	0	0	0	0
4108	0	0	0	0	0
4109	0	0	0	0	0
4110	0	0	0	0	0
4111	0	0	0	0	0
4112	0	0	0	0	0
n	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Generation : F0		Sex : Female	Day : 1	Period : F0 gestation Dose : Dodecyloxirane 0 mg/kg		Species : Rat		
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
1101	0	0	0	0	0	0	0	0
1102	0	0	0	0	0	0	0	0
1103	0	0	0	0	0	0	0	0
1104	0	0	0	0	0	0	0	0
1105	0	0	0	0	0	0	0	0
1106	0	0	0	0	0	0	0	0
1107	0	0	0	0	0	0	0	0
1108	0	0	0	0	0	0	0	0
1109	0	0	0	0	0	0	0	0
1110	0	0	0	0	0	0	0	0
1111	0	0	0	0	0	0	0	0
1112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
1101	0	0	0	0	0	0
1102	0	0	0	0	0	0
1103	0	0	0	0	0	0
1104	0	0	0	0	0	0
1105	0	0	0	0	0	0
1106	0	0	0	0	0	0
1107	0	0	0	0	0	0
1108	0	0	0	0	0	0
1109	0	0	0	0	0	0
1110	0	0	0	0	0	0
1111	0	0	0	0	0	0
1112	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Appendix 2 - 110

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation

Study No. : R-1261

Generation : F0		Sex : Female	Day : 1	Period : F0 gestation Dose : Dodecyloxirane 100 mg/kg		Species : Rat		
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
2101	0	0	0	0	0	0	0	0
2102	0	0	0	0	0	0	0	0
2103	0	0	0	0	0	0	0	0
2104	0	0	0	0	0	0	0	0
2105	0	0	0	0	0	0	0	0
2107	0	0	0	0	0	0	0	0
2108	0	0	0	0	0	0	0	0
2110	0	0	0	0	0	0	0	0
2111	0	0	0	0	0	0	0	0
2112	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
2101	0	0	0	0	0	0
2102	0	0	0	0	0	0
2103	0	0	0	0	0	0
2104	0	0	0	0	0	0
2105	0	0	0	0	0	0
2107	0	0	0	0	0	0
2108	0	0	0	0	0	0
2110	0	0	0	0	0	0
2111	0	0	0	0	0	0
2112	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Generation : F0		Sex : Female	Day : 1	Period : F0 gestation Dose : Dodecyloxirane 300 mg/kg		Species : Rat		
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
3101	0	0	0	0	0	0	0	0
3102	0	0	0	0	0	0	0	0
3103	0	0	0	0	0	0	0	0
3104	0	0	0	0	0	0	0	0
3105	0	0	0	0	0	0	0	0
3106	0	0	0	0	0	0	0	0
3107	0	0	0	0	0	0	0	0
3110	0	0	0	0	0	0	0	0
3111	0	0	0	0	0	0	0	0
3112	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
3101	0	0	0	0	0	0
3102	0	0	0	0	0	0
3103	0	0	0	0	0	0
3104	0	0	0	0	0	0
3105	0	0	0	0	0	0
3106	0	0	0	0	0	0
3107	0	0	0	0	0	0
3110	0	0	0	0	0	0
3111	0	0	0	0	0	0
3112	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Appendix 2 - 112

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation

Study No. : R-1261

Generation : F0		Sex : Female	Day : 1	Period : F0 gestation Dose : Dodecyloxirane 1000 mg/kg		Species : Rat		
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
4102	0	0	0	0	0	0	0	0
4103	0	0	0	0	0	0	0	0
4104	0	0	0	0	0	0	0	0
4105	0	0	0	0	0	0	0	0
4106	0	0	0	0	0	0	0	0
4107	0	0	0	0	0	0	0	0
4108	0	0	0	0	0	0	0	0
4109	0	0	0	0	0	0	0	0
4110	0	0	0	0	0	0	0	0
4111	0	0	0	0	0	0	0	0
4112	0	0	0	0	0	0	0	0
n	11	11	11	11	11	11	11	11

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
4102	0	0	0	0	0	0
4103	0	0	0	0	0	0
4104	0	0	0	0	0	0
4105	0	0	0	0	0	0
4106	0	0	0	0	0	0
4107	0	0	0	0	0	0
4108	0	0	0	0	0	0
4109	0	0	0	0	0	0
4110	0	0	0	0	0	0
4111	0	0	0	0	0	0
4112	0	0	0	0	0	0
n	11	11	11	11	11	11

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : F0 gestation

Generation : F0

Sex : Female

Day : 7

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
1101	0	0	0	0	0	0	0	0
1102	0	0	0	0	0	0	0	0
1103	0	0	0	0	0	0	0	0
1104	0	0	0	0	0	0	0	0
1105	0	0	0	0	0	0	0	0
1106	0	0	0	0	0	0	0	0
1107	0	0	0	0	0	0	0	0
1108	0	0	0	0	0	0	0	0
1109	0	0	0	0	0	0	0	0
1110	0	0	0	0	0	0	0	0
1111	0	0	0	0	0	0	0	0
1112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
1101	0	0	0	0	0	0
1102	0	0	0	0	0	0
1103	0	0	0	0	0	0
1104	0	0	0	0	0	0
1105	0	0	0	0	0	0
1106	0	0	0	0	0	0
1107	0	0	0	0	0	0
1108	0	0	0	0	0	0
1109	0	0	0	0	0	0
1110	0	0	0	0	0	0
1111	0	0	0	0	0	0
1112	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

With the reproduction/ developmental toxicity screening test in rats				Period : F0 gestation			Species : Rat	
In-the-hand observation		Sex : Female	Day : 7	Dose : Dodecyloxirane 100 mg/kg				
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
2101	0	0	0	0	0	0	0	0
2102	0	0	0	0	0	0	0	0
2103	0	0	0	0	0	0	0	0
2104	0	0	0	0	0	0	0	0
2105	0	0	0	0	0	0	0	0
2107	0	0	0	0	0	0	0	0
2108	0	0	0	0	0	0	0	0
2110	0	0	0	0	0	0	0	0
2111	0	0	0	0	0	0	0	0
2112	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Piloerection		Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
2101	0	0	0	0	0	0
2102	0	0	0	0	0	0
2103	0	0	0	0	0	0
2104	0	0	0	0	0	0
2105	0	0	0	0	0	0
2107	0	0	0	0	0	0
2108	0	0	0	0	0	0
2110	0	0	0	0	0	0
2111	0	0	0	0	0	0
2112	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Appendix 2 - 115

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation

Study No. : R-1261

Generation : F0		Sex : Female	Day : 7	Period : F0 gestation Dose : Dodecyloxirane 300 mg/kg		Species : Rat		
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
3101	0	0	0	0	0	0	0	0
3102	0	0	0	0	0	0	0	0
3103	0	0	0	0	0	0	0	0
3104	0	0	0	0	0	0	0	0
3105	0	0	0	0	0	0	0	0
3106	0	0	0	0	0	0	0	0
3107	0	0	0	0	0	0	0	0
3110	0	0	0	0	0	0	0	0
3111	0	0	0	0	0	0	0	0
3112	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
3101	0	0	0	0	0	0
3102	0	0	0	0	0	0
3103	0	0	0	0	0	0
3104	0	0	0	0	0	0
3105	0	0	0	0	0	0
3106	0	0	0	0	0	0
3107	0	0	0	0	0	0
3110	0	0	0	0	0	0
3111	0	0	0	0	0	0
3112	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Appendix 2 - 116

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

In-the-hand observation			Period : F0 gestation			Species : Rat		
Generation : F0	Sex : Female	Day : 7	Dose : Dodecyloxirane 1000 mg/kg					
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
4102	0	0	0	0	0	0	0	0
4103	0	0	0	0	0	0	0	0
4104	0	0	0	0	0	0	0	0
4105	0	0	0	0	0	0	0	0
4106	0	0	0	0	0	0	0	0
4107	0	0	0	0	0	0	0	0
4108	0	0	0	0	0	0	0	0
4109	0	0	0	0	0	0	0	0
4110	0	0	0	0	0	0	0	0
4111	0	0	0	0	0	0	0	0
4112	0	0	0	0	0	0	0	0
n	11	11	11	11	11	11	11	11
Piloerection			Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling	
Animal No.								
4102	0	0	0	0	0	0		
4103	0	0	0	0	0	0		
4104	0	0	0	0	0	0		
4105	0	0	0	0	0	0		
4106	0	0	0	0	0	0		
4107	0	0	0	0	0	0		
4108	0	0	0	0	0	0		
4109	0	0	0	0	0	0		
4110	0	0	0	0	0	0		
4111	0	0	0	0	0	0		
4112	0	0	0	0	0	0		
n	11	11	11	11	11	11		

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 gestation		Dose : Dodecyloxirane 0 mg/kg		Species : Rat			
Generation : F0	Sex : Female	Day : 14							
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation		
Animal No.									
1101	0	0	0	0	0	0	0	0	0
1102	0	0	0	0	0	0	0	0	0
1103	0	0	0	0	0	0	0	0	0
1104	0	0	0	0	0	0	0	0	0
1105	0	0	0	0	0	0	0	0	0
1106	0	0	0	0	0	0	0	0	0
1107	0	0	0	0	0	0	0	0	0
1108	0	0	0	0	0	0	0	0	0
1109	0	0	0	0	0	0	0	0	0
1110	0	0	0	0	0	0	0	0	0
1111	0	0	0	0	0	0	0	0	0
1112	0	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
1101	0	0	0	0	0
1102	0	0	0	0	0
1103	0	0	0	0	0
1104	0	0	0	0	0
1105	0	0	0	0	0
1106	0	0	0	0	0
1107	0	0	0	0	0
1108	0	0	0	0	0
1109	0	0	0	0	0
1110	0	0	0	0	0
1111	0	0	0	0	0
1112	0	0	0	0	0
n	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Generation : F0		Sex : Female	Day : 14	Period : F0 gestation Dose : Dodecyloxirane 100 mg/kg		Species : Rat		
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
2101	0	0	0	0	0	0	0	0
2102	0	0	0	0	0	0	0	0
2103	0	0	0	0	0	0	0	0
2104	0	0	0	0	0	0	0	0
2105	0	0	0	0	0	0	0	0
2107	0	0	0	0	0	0	0	0
2108	0	0	0	0	0	0	0	0
2110	0	0	0	0	0	0	0	0
2111	0	0	0	0	0	0	0	0
2112	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
2101	0	0	0	0	0	0
2102	0	0	0	0	0	0
2103	0	0	0	0	0	0
2104	0	0	0	0	0	0
2105	0	0	0	0	0	0
2107	0	0	0	0	0	0
2108	0	0	0	0	0	0
2110	0	0	0	0	0	0
2111	0	0	0	0	0	0
2112	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Generation : F0		Sex : Female	Day : 14	Period : F0 gestation Dose : Dodecyloxirane 300 mg/kg		Species : Rat		
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
3101	0	0	0	0	0	0	0	0
3102	0	0	0	0	0	0	0	0
3103	0	0	0	0	0	0	0	0
3104	0	0	0	0	0	0	0	0
3105	0	0	0	0	0	0	0	0
3106	0	0	0	0	0	0	0	0
3107	0	0	0	0	0	0	0	0
3110	0	0	0	0	0	0	0	0
3111	0	0	0	0	0	0	0	0
3112	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
3101	0	0	0	0	0	0
3102	0	0	0	0	0	0
3103	0	0	0	0	0	0
3104	0	0	0	0	0	0
3105	0	0	0	0	0	0
3106	0	0	0	0	0	0
3107	0	0	0	0	0	0
3110	0	0	0	0	0	0
3111	0	0	0	0	0	0
3112	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 gestation		Dose : Dodecyloxirane 1000 mg/kg		Species : Rat			
Generation : F0	Sex : Female	Day : 14							
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation		
Animal No.									
4102	0	0	0	0	0	0	0	0	0
4103	0	0	0	0	0	0	0	0	0
4104	0	0	0	0	0	0	0	0	0
4105	0	0	0	0	0	0	0	0	0
4106	0	0	0	0	0	0	0	0	0
4107	0	0	0	0	0	0	0	0	0
4108	0	0	0	0	0	0	0	0	0
4109	0	0	0	0	0	0	0	0	0
4110	0	0	0	0	0	0	0	0	0
4111	0	0	0	0	0	0	0	0	0
4112	0	0	0	0	0	0	0	0	0
n	11	11	11	11	11	11	11	11	11

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
4102	0	0	0	0	0
4103	0	0	0	0	0
4104	0	0	0	0	0
4105	0	0	0	0	0
4106	0	0	0	0	0
4107	0	0	0	0	0
4108	0	0	0	0	0
4109	0	0	0	0	0
4110	0	0	0	0	0
4111	0	0	0	0	0
4112	0	0	0	0	0
n	11	11	11	11	11

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

	In-the-hand observation			Period : F0 gestation				
	Generation : F0	Sex : Female	Day : 20	Dose : Dodecyloxirane 0 mg/kg			Species : Rat	
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
1101	0	0	0	0	0	0	0	0
1102	0	0	0	0	0	0	0	0
1103	0	0	0	0	0	0	0	0
1104	0	0	0	0	0	0	0	0
1105	0	0	0	0	0	0	0	0
1106	0	0	0	0	0	0	0	0
1107	0	0	0	0	0	0	0	0
1108	0	0	0	0	0	0	0	0
1109	0	0	0	0	0	0	0	0
1110	0	0	0	0	0	0	0	0
1111	0	0	0	0	0	0	0	0
1112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.						
1101	0	0	0	0	0	0
1102	0	0	0	0	0	0
1103	0	0	0	0	0	0
1104	0	0	0	0	0	0
1105	0	0	0	0	0	0
1106	0	0	0	0	0	0
1107	0	0	0	0	0	0
1108	0	0	0	0	0	0
1109	0	0	0	0	0	0
1110	0	0	0	0	0	0
1111	0	0	0	0	0	0
1112	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Generation : F0		Sex : Female	Day : 20	Period : F0 gestation Dose : Dodecyloxirane 100 mg/kg		Species : Rat		
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
2101	0	0	0	0	0	0	0	0
2102	0	0	0	0	0	0	0	0
2103	0	0	0	0	0	0	0	0
2104	0	0	0	0	0	0	0	0
2105	0	0	0	0	0	0	0	0
2107	0	0	0	0	0	0	0	0
2108	0	0	0	0	0	0	0	0
2110	0	0	0	0	0	0	0	0
2111	0	0	0	0	0	0	0	0
2112	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
2101	0	0	0	0	0	0
2102	0	0	0	0	0	0
2103	0	0	0	0	0	0
2104	0	0	0	0	0	0
2105	0	0	0	0	0	0
2107	0	0	0	0	0	0
2108	0	0	0	0	0	0
2110	0	0	0	0	0	0
2111	0	0	0	0	0	0
2112	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 gestation		Species : Rat				
Generation : F0		Sex : Female	Day : 20	Dose : Dodecyloxirane 300 mg/kg				
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
3101	0	0	0	0	0	0	0	0
3102	0	0	0	0	0	0	0	0
3103	0	0	0	0	0	0	0	0
3104	0	0	0	0	0	0	0	0
3105	0	0	0	0	0	0	0	0
3106	0	0	0	0	0	0	0	0
3107	0	0	0	0	0	0	0	0
3110	0	0	0	0	0	0	0	0
3111	0	0	0	0	0	0	0	0
3112	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
3101	0	0	0	0	0	0
3102	0	0	0	0	0	0
3103	0	0	0	0	0	0
3104	0	0	0	0	0	0
3105	0	0	0	0	0	0
3106	0	0	0	0	0	0
3107	0	0	0	0	0	0
3110	0	0	0	0	0	0
3111	0	0	0	0	0	0
3112	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 gestation		Dose : Dodecyloxirane 1000 mg/kg		Species : Rat			
Generation : F0	Sex : Female	Day : 20							
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation		
Animal No.									
4102	0	0	0	0	0	0	0	0	0
4103	0	0	0	0	0	0	0	0	0
4104	0	0	0	0	0	0	0	0	0
4105	0	0	0	0	0	0	0	0	0
4106	0	0	0	0	0	0	0	0	0
4107	0	0	0	0	0	0	0	0	0
4108	0	0	0	0	0	0	0	0	0
4109	0	0	0	0	0	0	0	0	0
4110	0	0	0	0	0	0	0	0	0
4111	0	0	0	0	0	0	0	0	0
4112	0	0	0	0	0	0	0	0	0
n	11	11	11	11	11	11	11	11	11

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
4102	0	0	0	0	0
4103	0	0	0	0	0
4104	0	0	0	0	0
4105	0	0	0	0	0
4106	0	0	0	0	0
4107	0	0	0	0	0
4108	0	0	0	0	0
4109	0	0	0	0	0
4110	0	0	0	0	0
4111	0	0	0	0	0
4112	0	0	0	0	0
n	11	11	11	11	11

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation
Generation : F0 Sex : Female Day : 6 Period : F0 lactation
Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
1101	0	0	0	0	0	0	0	0
1102	0	0	0	0	0	0	0	0
1103	0	0	0	0	0	0	0	0
1104	0	0	0	0	0	0	0	0
1105	0	0	0	0	0	0	0	0
1106	0	0	0	0	0	0	0	0
1107	0	0	0	0	0	0	0	0
1108	0	0	0	0	0	0	0	0
1109	0	0	0	0	0	0	0	0
1110	0	0	0	0	0	0	0	0
1111	0	0	0	0	0	0	0	0
1112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
1101	0	0	0	0	0	0
1102	0	0	0	0	0	0
1103	0	0	0	0	0	0
1104	0	0	0	0	0	0
1105	0	0	0	0	0	0
1106	0	0	0	0	0	0
1107	0	0	0	0	0	0
1108	0	0	0	0	0	0
1109	0	0	0	0	0	0
1110	0	0	0	0	0	0
1111	0	0	0	0	0	0
1112	0	0	0	0	0	0
n	12	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Appendix 2 - 126

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
In-the-hand observation

Study No. : R-1261

Generation : F0		Sex : Female	Day : 6	Period : F0 lactation Dose : Dodecyloxirane 100 mg/kg		Species : Rat		
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
2101	0	0	0	0	0	0	0	0
2102	0	0	0	0	0	0	0	0
2103	0	0	0	0	0	0	0	0
2104	0	0	0	0	0	0	0	0
2105	0	0	0	0	0	0	0	0
2107	0	0	0	0	0	0	0	0
2108	0	0	0	0	0	0	0	0
2110	0	0	0	0	0	0	0	0
2111	0	0	0	0	0	0	0	0
2112	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
2101	0	0	0	0	0	0
2102	0	0	0	0	0	0
2103	0	0	0	0	0	0
2104	0	0	0	0	0	0
2105	0	0	0	0	0	0
2107	0	0	0	0	0	0
2108	0	0	0	0	0	0
2110	0	0	0	0	0	0
2111	0	0	0	0	0	0
2112	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 lactation		Dose : Dodecyloxirane 300 mg/kg		Species : Rat			
Generation : F0		Sex : Female		Day : 6					
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation	
3101	0	0	0	0	0	0	0	0	
3102	0	0	0	0	0	0	0	0	
3103	0	0	0	0	0	0	0	0	
3104	0	0	0	0	0	0	0	0	
3105	0	0	0	0	0	0	0	0	
3106	0	0	0	0	0	0	0	0	
3107	0	0	0	0	0	0	0	0	
3110	0	0	0	0	0	0	0	0	
3111	0	0	0	0	0	0	0	0	
3112	0	0	0	0	0	0	0	0	
n	10	10	10	10	10	10	10	10	

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling	
3101	0	0	0	0	0	0	
3102	0	0	0	0	0	0	
3103	0	0	0	0	0	0	
3104	0	0	0	0	0	0	
3105	0	0	0	0	0	0	
3106	0	0	0	0	0	0	
3107	0	0	0	0	0	0	
3110	0	0	0	0	0	0	
3111	0	0	0	0	0	0	
3112	0	0	0	0	0	0	
n	10	10	10	10	10	10	

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 lactation		Dose : Dodecyloxirane 1000 mg/kg		Species : Rat			
Generation : F0		Sex : Female		Day : 6					
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation	
4102	0	0	0	0	0	0	0	0	
4103	0	0	0	0	0	0	0	0	
4104	0	0	0	0	0	0	0	0	
4105	0	0	0	0	0	0	0	0	
4107	0	0	0	0	0	0	0	0	
4108	0	0	0	0	0	0	0	0	
4109	0	0	0	0	0	0	0	0	
4110	0	0	0	0	0	0	0	0	
4111	0	0	0	0	0	0	0	0	
4112	0	0	0	0	0	0	0	0	
n	10	10	10	10	10	10	10	10	

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling	
4102	0	0	0	0	0	0	
4103	0	0	0	0	0	0	
4104	0	0	0	0	0	0	
4105	0	0	0	0	0	0	
4107	0	0	0	0	0	0	
4108	0	0	0	0	0	0	
4109	0	0	0	0	0	0	
4110	0	0	0	0	0	0	
4111	0	0	0	0	0	0	
4112	0	0	0	0	0	0	
n	10	10	10	10	10	10	

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

With the reproduction/ developmental toxicity screening test in rats								
In-the-hand observation		Period : F0 lactation		Species : Rat				
Generation : F0	Sex : Female	Day : 13	Dose : Dodecylloxirane 0 mg/kg					
Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation	
Animal No.								
1101	0	0	0	0	0	0	0	0
1102	0	0	0	0	0	0	0	0
1103	0	0	0	0	0	0	0	0
1104	0	0	0	0	0	0	0	0
1105	0	0	0	0	0	0	0	0
1106	0	0	0	0	0	0	0	0
1107	0	0	0	0	0	0	0	0
1108	0	0	0	0	0	0	0	0
1109	0	0	0	0	0	0	0	0
1110	0	0	0	0	0	0	0	0
1111	0	0	0	0	0	0	0	0
1112	0	0	0	0	0	0	0	0
n	12	12	12	12	12	12	12	12

Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
Animal No.					
1101	0	0	0	0	0
1102	0	0	0	0	0
1103	0	0	0	0	0
1104	0	0	0	0	0
1105	0	0	0	0	0
1106	0	0	0	0	0
1107	0	0	0	0	0
1108	0	0	0	0	0
1109	0	0	0	0	0
1110	0	0	0	0	0
1111	0	0	0	0	0
1112	0	0	0	0	0
n	12	12	12	12	12

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 lactation		Species : Rat				
Generation : F0		Sex : Female	Day : 13	Dose : Dodecyloxirane 100 mg/kg				
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
2101	0	0	0	0	0	0	0	0
2102	0	0	0	0	0	0	0	0
2103	0	0	0	0	0	0	0	0
2104	0	0	0	0	0	0	0	0
2105	0	0	0	0	0	0	0	0
2107	0	0	0	0	0	0	0	0
2108	0	0	0	0	0	0	0	0
2110	0	0	0	0	0	0	0	0
2111	0	0	0	0	0	0	0	0
2112	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
2101	0	0	0	0	0	0
2102	0	0	0	0	0	0
2103	0	0	0	0	0	0
2104	0	0	0	0	0	0
2105	0	0	0	0	0	0
2107	0	0	0	0	0	0
2108	0	0	0	0	0	0
2110	0	0	0	0	0	0
2111	0	0	0	0	0	0
2112	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 lactation		Dose : Dodecyloxirane 300 mg/kg		Species : Rat			
Generation : F0		Sex : Female		Day : 13					
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation	
3101	0	0	0	0	0	0	0	0	
3102	0	0	0	0	0	0	0	0	
3103	0	0	0	0	0	0	0	0	
3104	0	0	0	0	0	0	0	0	
3105	0	0	0	0	0	0	0	0	
3106	0	0	0	0	0	0	0	0	
3107	0	0	0	0	0	0	0	0	
3110	0	0	0	0	0	0	0	0	
3111	0	0	0	0	0	0	0	0	
3112	0	0	0	0	0	0	0	0	
n	10	10	10	10	10	10	10	10	

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling	
3101	0	0	0	0	0	0	
3102	0	0	0	0	0	0	
3103	0	0	0	0	0	0	
3104	0	0	0	0	0	0	
3105	0	0	0	0	0	0	
3106	0	0	0	0	0	0	
3107	0	0	0	0	0	0	
3110	0	0	0	0	0	0	
3111	0	0	0	0	0	0	
3112	0	0	0	0	0	0	
n	10	10	10	10	10	10	

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation		Period : F0 lactation		Dose : Dodecyloxirane 1000 mg/kg		Species : Rat			
Generation : F0		Sex : Female		Day : 13					
Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation	
4102	0	0	0	0	0	0	0	0	
4103	0	0	0	0	0	0	0	0	
4104	0	0	0	0	0	0	0	0	
4105	0	0	0	0	0	0	0	0	
4107	0	0	0	0	0	0	0	0	
4108	0	0	0	0	0	0	0	0	
4109	0	0	0	0	0	0	0	0	
4110	0	0	0	0	0	0	0	0	
4111	0	0	0	0	0	0	0	0	
4112	0	0	0	0	0	0	0	0	
n	10	10	10	10	10	10	10	10	

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling	
4102	0	0	0	0	0	0	
4103	0	0	0	0	0	0	
4104	0	0	0	0	0	0	
4105	0	0	0	0	0	0	
4107	0	0	0	0	0	0	
4108	0	0	0	0	0	0	
4109	0	0	0	0	0	0	
4110	0	0	0	0	0	0	
4111	0	0	0	0	0	0	
4112	0	0	0	0	0	0	
n	10	10	10	10	10	10	

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
5101	0	0	0	0	0	0	0	0
5102	0	0	0	0	0	0	0	0
5103	0	0	0	0	0	0	0	0
5104	0	0	0	0	0	0	0	0
5105	0	0	0	0	0	0	0	0
5106	0	0	0	0	0	0	0	0
5107	0	0	0	0	0	0	0	0
5108	0	0	0	0	0	0	0	0
5109	0	0	0	0	0	0	0	0
5110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
5101	0	0	0	0	0	0
5102	0	0	0	0	0	0
5103	0	0	0	0	0	0
5104	0	0	0	0	0	0
5105	0	0	0	0	0	0
5106	0	0	0	0	0	0
5107	0	0	0	0	0	0
5108	0	0	0	0	0	0
5109	0	0	0	0	0	0
5110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
6101	0	0	0	0	0	0	0	0
6102	0	0	0	0	0	0	0	0
6103	0	0	0	0	0	0	0	0
6104	0	0	0	0	0	0	0	0
6105	0	0	0	0	0	0	0	0
6106	0	0	0	0	0	0	0	0
6107	0	0	0	0	0	0	0	0
6108	0	0	0	0	0	0	0	0
6109	0	0	0	0	0	0	0	0
6110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
6101	0	0	0	0	0	0
6102	0	0	0	0	0	0
6103	0	0	0	0	0	0
6104	0	0	0	0	0	0
6105	0	0	0	0	0	0
6106	0	0	0	0	0	0
6107	0	0	0	0	0	0
6108	0	0	0	0	0	0
6109	0	0	0	0	0	0
6110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
5101	0	0	0	0	0	0	0	0
5102	0	0	0	0	0	0	0	0
5103	0	0	0	0	0	0	0	0
5104	0	0	0	0	0	0	0	0
5105	0	0	0	0	0	0	0	0
5106	0	0	0	0	0	0	0	0
5107	0	0	0	0	0	0	0	0
5108	0	0	0	0	0	0	0	0
5109	0	0	0	0	0	0	0	0
5110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
5101	0	0	0	0	0	0
5102	0	0	0	0	0	0
5103	0	0	0	0	0	0
5104	0	0	0	0	0	0
5105	0	0	0	0	0	0
5106	0	0	0	0	0	0
5107	0	0	0	0	0	0
5108	0	0	0	0	0	0
5109	0	0	0	0	0	0
5110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecylxirane 1000 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
6101	0	0	0	0	0	0	0	0
6102	0	0	0	0	0	0	0	0
6103	0	0	0	0	0	0	0	0
6104	0	0	0	0	0	0	0	0
6105	0	0	0	0	0	0	0	0
6106	0	0	0	0	0	0	0	0
6107	0	0	0	0	0	0	0	0
6108	0	0	0	0	0	0	0	0
6109	0	0	0	0	0	0	0	0
6110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
6101	0	0	0	0	0	0
6102	0	0	0	0	0	0
6103	0	0	0	0	0	0
6104	0	0	0	0	0	0
6105	0	0	0	0	0	0
6106	0	0	0	0	0	0
6107	0	0	0	0	0	0
6108	0	0	0	0	0	0
6109	0	0	0	0	0	0
6110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 3

Dose : Dodecylxirane 0 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
5101	0	0	0	0	0	0	0	0
5102	0	0	0	0	0	0	0	0
5103	0	0	0	0	0	0	0	0
5104	0	0	0	0	0	0	0	0
5105	0	0	0	0	0	0	0	0
5106	0	0	0	0	0	0	0	0
5107	0	0	0	0	0	0	0	0
5108	0	0	0	0	0	0	0	0
5109	0	0	0	0	0	0	0	0
5110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
5101	0	0	0	0	0	0
5102	0	0	0	0	0	0
5103	0	0	0	0	0	0
5104	0	0	0	0	0	0
5105	0	0	0	0	0	0
5106	0	0	0	0	0	0
5107	0	0	0	0	0	0
5108	0	0	0	0	0	0
5109	0	0	0	0	0	0
5110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 3

Dose : Dodecylxirane 1000 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
6101	0	0	0	0	0	0	0	0
6102	0	0	0	0	0	0	0	0
6103	0	0	0	0	0	0	0	0
6104	0	0	0	0	0	0	0	0
6105	0	0	0	0	0	0	0	0
6106	0	0	0	0	0	0	0	0
6107	0	0	0	0	0	0	0	0
6108	0	0	0	0	0	0	0	0
6109	0	0	0	0	0	0	0	0
6110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
6101	0	0	0	0	0	0
6102	0	0	0	0	0	0
6103	0	0	0	0	0	0
6104	0	0	0	0	0	0
6105	0	0	0	0	0	0
6106	0	0	0	0	0	0
6107	0	0	0	0	0	0
6108	0	0	0	0	0	0
6109	0	0	0	0	0	0
6110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 4

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
5101	0	0	0	0	0	0	0	0
5102	0	0	0	0	0	0	0	0
5103	0	0	0	0	0	0	0	0
5104	0	0	0	0	0	0	0	0
5105	0	0	0	0	0	0	0	0
5106	0	0	0	0	0	0	0	0
5107	0	0	0	0	0	0	0	0
5108	0	0	0	0	0	0	0	0
5109	0	0	0	0	0	0	0	0
5110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
5101	0	0	0	0	0	0
5102	0	0	0	0	0	0
5103	0	0	0	0	0	0
5104	0	0	0	0	0	0
5105	0	0	0	0	0	0
5106	0	0	0	0	0	0
5107	0	0	0	0	0	0
5108	0	0	0	0	0	0
5109	0	0	0	0	0	0
5110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 4

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
6101	0	0	0	0	0	0	0	0
6102	0	0	0	0	0	0	0	0
6103	0	0	0	0	0	0	0	0
6104	0	0	0	0	0	0	0	0
6105	0	0	0	0	0	0	0	0
6106	0	0	0	0	0	0	0	0
6107	0	0	0	0	0	0	0	0
6108	0	0	0	0	0	0	0	0
6109	0	0	0	0	0	0	0	0
6110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
6101	0	0	0	0	0	0
6102	0	0	0	0	0	0
6103	0	0	0	0	0	0
6104	0	0	0	0	0	0
6105	0	0	0	0	0	0
6106	0	0	0	0	0	0
6107	0	0	0	0	0	0
6108	0	0	0	0	0	0
6109	0	0	0	0	0	0
6110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 5

Dose : Dodecylxirane 0 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
5101	0	0	0	0	0	0	0	0
5102	0	0	0	0	0	0	0	0
5103	0	0	0	0	0	0	0	0
5104	0	0	0	0	0	0	0	0
5105	0	0	0	0	0	0	0	0
5106	0	0	0	0	0	0	0	0
5107	0	0	0	0	0	0	0	0
5108	0	0	0	0	0	0	0	0
5109	0	0	0	0	0	0	0	0
5110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
5101	0	0	0	0	0	0
5102	0	0	0	0	0	0
5103	0	0	0	0	0	0
5104	0	0	0	0	0	0
5105	0	0	0	0	0	0
5106	0	0	0	0	0	0
5107	0	0	0	0	0	0
5108	0	0	0	0	0	0
5109	0	0	0	0	0	0
5110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 5

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
6101	0	0	0	0	0	0	0	0
6102	0	0	0	0	0	0	0	0
6103	0	0	0	0	0	0	0	0
6104	0	0	0	0	0	0	0	0
6105	0	0	0	0	0	0	0	0
6106	0	0	0	0	0	0	0	0
6107	0	0	0	0	0	0	0	0
6108	0	0	0	0	0	0	0	0
6109	0	0	0	0	0	0	0	0
6110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
6101	0	0	0	0	0	0
6102	0	0	0	0	0	0
6103	0	0	0	0	0	0
6104	0	0	0	0	0	0
6105	0	0	0	0	0	0
6106	0	0	0	0	0	0
6107	0	0	0	0	0	0
6108	0	0	0	0	0	0
6109	0	0	0	0	0	0
6110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
5101	0	0	0	0	0	0	0	0
5102	0	0	0	0	0	0	0	0
5103	0	0	0	0	0	0	0	0
5104	0	0	0	0	0	0	0	0
5105	0	0	0	0	0	0	0	0
5106	0	0	0	0	0	0	0	0
5107	0	0	0	0	0	0	0	0
5108	0	0	0	0	0	0	0	0
5109	0	0	0	0	0	0	0	0
5110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
5101	0	0	0	0	0	0
5102	0	0	0	0	0	0
5103	0	0	0	0	0	0
5104	0	0	0	0	0	0
5105	0	0	0	0	0	0
5106	0	0	0	0	0	0
5107	0	0	0	0	0	0
5108	0	0	0	0	0	0
5109	0	0	0	0	0	0
5110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecylxirane 1000 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
6101	0	0	0	0	0	0	0	0
6102	0	0	0	0	0	0	0	0
6103	0	0	0	0	0	0	0	0
6104	0	0	0	0	0	0	0	0
6105	0	0	0	0	0	0	0	0
6106	0	0	0	0	0	0	0	0
6107	0	0	0	0	0	0	0	0
6108	0	0	0	0	0	0	0	0
6109	0	0	0	0	0	0	0	0
6110	0	0	0	0	0	0	0	0
n	10	10	10	10	10	10	10	10

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
6101	0	0	0	0	0	0
6102	0	0	0	0	0	0
6103	0	0	0	0	0	0
6104	0	0	0	0	0	0
6105	0	0	0	0	0	0
6106	0	0	0	0	0	0
6107	0	0	0	0	0	0
6108	0	0	0	0	0	0
6109	0	0	0	0	0	0
6110	0	0	0	0	0	0
n	10	10	10	10	10	10

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation				Period : Recovery					
Generation : F0		Sex : Male	Week : 1	Dose : Dodecyloxirane 0 mg/kg			Species : Rat		
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation	
Animal No.									
1008	0	0	0	0	0	0	0	0	
1009	0	0	0	0	0	0	0	0	
1010	0	0	0	0	0	0	0	0	
1011	0	0	0	0	0	0	0	0	
1012	0	0	0	0	0	0	0	0	
n	5	5	5	5	5	5	5	5	

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
1008	0	0	0	0	0	0
1009	0	0	0	0	0	0
1010	0	0	0	0	0	0
1011	0	0	0	0	0	0
1012	0	0	0	0	0	0
n	5	5	5	5	5	5

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation			Period : Recovery					
Generation : F0	Sex : Male	Week : 1	Dose : Dodecyloxirane 1000 mg/kg			Species : Rat		
	Ease of removal from cage	Fur condition	Skin	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
4008	0	0	0	0	0	0	0	0
4009	0	0	0	0	0	0	0	0
4010	0	0	0	0	0	0	0	0
4011	0	0	0	0	0	0	0	0
4012	0	0	0	0	0	0	0	0
n	5	5	5	5	5	5	5	5

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
4008	0	0	0	0	0	0
4009	0	0	0	0	0	0
4010	0	0	0	0	0	0
4011	0	0	0	0	0	0
4012	0	0	0	0	0	0
n	5	5	5	5	5	5

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation			Period : Recovery					
Generation : F0		Sex : Male	Week : 2	Dose : Dodecyloxirane 0 mg/kg		Species : Rat		
	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
1008	0	0	0	0	0	0	0	0
1009	0	0	0	0	0	0	0	0
1010	0	0	0	0	0	0	0	0
1011	0	0	0	0	0	0	0	0
1012	0	0	0	0	0	0	0	0
n	5	5	5	5	5	5	5	5

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
1008	0	0	0	0	0	0
1009	0	0	0	0	0	0
1010	0	0	0	0	0	0
1011	0	0	0	0	0	0
1012	0	0	0	0	0	0
n	5	5	5	5	5	5

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation			Period : Recovery					
Generation : F0	Sex : Male	Week : 2	Dose : Dodecyloxirane 1000 mg/kg			Species : Rat		
	Ease of removal from cage	Fur condition	Skin	Secretions- Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
Animal No.								
4008	0	0	0	0	0	0	0	0
4009	0	0	0	0	0	0	0	0
4010	0	0	0	0	0	0	0	0
4011	0	0	0	0	0	0	0	0
4012	0	0	0	0	0	0	0	0
n	5	5	5	5	5	5	5	5

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
4008	0	0	0	0	0	0
4009	0	0	0	0	0	0
4010	0	0	0	0	0	0
4011	0	0	0	0	0	0
4012	0	0	0	0	0	0
n	5	5	5	5	5	5

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
5106	0	0	0	0	0	0	0	0
5107	0	0	0	0	0	0	0	0
5108	0	0	0	0	0	0	0	0
5109	0	0	0	0	0	0	0	0
5110	0	0	0	0	0	0	0	0
n	5	5	5	5	5	5	5	5

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
5106	0	0	0	0	0	0
5107	0	0	0	0	0	0
5108	0	0	0	0	0	0
5109	0	0	0	0	0	0
5110	0	0	0	0	0	0
n	5	5	5	5	5	5

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
6106	0	0	0	0	0	0	0	0
6107	0	0	0	0	0	0	0	0
6108	0	0	0	0	0	0	0	0
6109	0	0	0	0	0	0	0	0
6110	0	0	0	0	0	0	0	0
n	5	5	5	5	5	5	5	5

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
6106	0	0	0	0	0	0
6107	0	0	0	0	0	0
6108	0	0	0	0	0	0
6109	0	0	0	0	0	0
6110	0	0	0	0	0	0
n	5	5	5	5	5	5

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
5106	0	0	0	0	0	0	0	0
5107	0	0	0	0	0	0	0	0
5108	0	0	0	0	0	0	0	0
5109	0	0	0	0	0	0	0	0
5110	0	0	0	0	0	0	0	0
n	5	5	5	5	5	5	5	5

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
5106	0	0	0	0	0	0
5107	0	0	0	0	0	0
5108	0	0	0	0	0	0
5109	0	0	0	0	0	0
5110	0	0	0	0	0	0
n	5	5	5	5	5	5

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

In-the-hand observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Ease of removal from cage	Fur condition	Skin	Secretions-Eye, Nose	Exophthalmos	Palpebral closure	Mucosal membranes	Lacrimation
6106	0	0	0	0	0	0	0	0
6107	0	0	0	0	0	0	0	0
6108	0	0	0	0	0	0	0	0
6109	0	0	0	0	0	0	0	0
6110	0	0	0	0	0	0	0	0
n	5	5	5	5	5	5	5	5

Animal No.	Piloerection	Pupil size	Salivation	Abnormal respiration	Vocalization	Reactivity to handling
6106	0	0	0	0	0	0
6107	0	0	0	0	0	0
6108	0	0	0	0	0	0
6109	0	0	0	0	0	0
6110	0	0	0	0	0	0
n	5	5	5	5	5	5

Ease of removal from cage) -1: Atypically docile, 0: Easy, 1: Some resistance/avoidance, 2: Difficult, 3: Very difficult

Fur condition) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Skin) 0: Normal, 1: Slight, 2: Moderate, 3: Severe

Secretions-Eye, Nose) 0: Absent, 1: Present

Exophthalmos) 0: Absent, 1: Present

Palpebral closure) 0: Normal, 1: Slightly closed, 2: Half closed, 3: Completely closed

Mucosal membranes) 0: Normal, 1: Slight, 2: Moderate, 3: Marked

Lacrimation) 0: Absent, 1: Present

Piloerection) 0: Absent, 1: Present

Pupil size) -1: Miosis, 0: Normal, 1: Half opened pupil, 2: Mydriasis

Salivation) 0: None, 1: Slight, 2: Moderate, 3: Marked

Abnormal respiration) 0: None, 1: Slight, 2: Moderate, 3: Severe

Vocalization) 0: None, 1: Soft, 2: Moderate, 3: Loud/aggressive

Reactivity to handling) -1: Atypically docile, 0: Easy, 1: Slightly awkward, 2: Awkward, 3: Very Awkward

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1001	0	0	0	0	0	N	0	0
1002	0	0	0	0	0	N	0	0
1003	0	0	0	0	0	N	0	0
1004	0	0	0	0	0	N	0	0
1005	0	0	0	0	0	N	0	1
1006	0	0	0	0	0	N	0	0
1007	0	0	0	0	0	N	0	0
1008	0	0	0	0	0	N	0	1
1009	0	0	0	0	0	N	0	0
1010	0	0	0	0	0	N	0	0
1011	0	0	0	0	0	N	0	0
1012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1001	2	0
1002	0	0
1003	5	0
1004	3	0
1005	7	0
1006	0	0
1007	3	0
1008	4	0
1009	5	0
1010	2	0
1011	4	0
1012	2	0
n	12	12
Mean	3	0
S.D.	2	0

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2001	0	0	0	0	0	N	0	0
2002	0	0	0	0	0	N	0	1
2003	0	0	0	0	0	N	0	0
2004	0	0	0	0	0	N	0	0
2005	0	0	0	0	0	N	0	0
2006	0	0	0	0	0	N	0	0
2007	0	0	0	0	0	N	0	0
2008	0	0	0	0	0	N	0	0
2009	0	0	0	0	0	N	0	0
2010	0	0	0	0	0	N	0	0
2011	0	0	0	0	0	N	0	0
2012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2001	4	0
2002	3	0
2003	3	0
2004	5	0
2005	0	0
2006	7	0
2007	2	0
2008	4	0
2009	1	0
2010	0	0
2011	4	0
2012	2	0
n	12	12
Mean	3	0
S.D.	2	0

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3001	0	0	0	0	0	N	0	0
3002	0	0	0	0	0	N	0	0
3003	0	0	0	0	0	N	0	0
3004	0	0	0	0	0	N	0	0
3005	0	0	0	0	0	N	0	0
3006	0	0	0	0	0	N	0	0
3007	0	0	0	0	0	N	0	0
3008	0	0	0	0	0	N	0	0
3009	0	0	0	0	0	N	0	0
3010	0	0	0	0	0	N	0	0
3011	0	0	0	0	0	N	0	0
3012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3001	3	0
3002	2	0
3003	5	0
3004	5	0
3005	7	0
3006	8	0
3007	10	0
3008	3	0
3009	4	0
3010	5	0
3011	2	0
3012	1	0
n	12	12
Mean	5	0
S.D.	3	0

Open field observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4001	0	0	0	0	0	N	0	0
4002	0	0	0	0	0	N	0	0
4003	0	0	0	0	0	N	0	0
4004	0	0	0	0	0	N	0	1
4005	0	0	0	0	0	N	0	0
4006	0	0	0	0	0	N	0	0
4007	0	0	0	0	0	N	0	1
4008	0	0	0	0	0	N	0	0
4009	0	0	0	0	0	N	0	0
4010	0	0	0	0	0	N	0	0
4011	0	0	0	0	0	N	0	0
4012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4001	3	0
4002	9	0
4003	2	0
4004	3	0
4005	5	0
4006	6	0
4007	0	0
4008	3	0
4009	4	0
4010	4	0
4011	0	0
4012	2	0
n	12	12
Mean	3	0
S.D.	3	0

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 2 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
1001	0	0	0	0	0	N	0	0
1002	0	0	0	0	0	N	0	0
1003	0	0	0	0	0	N	0	0
1004	0	0	0	0	0	N	0	0
1005	0	0	0	0	0	N	0	0
1006	0	0	0	0	0	N	0	0
1007	0	0	0	0	0	N	0	0
1008	0	0	0	0	0	N	0	0
1009	0	0	0	0	0	N	0	0
1010	0	0	0	0	0	N	0	0
1011	0	0	0	0	0	N	0	0
1012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 2 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1001	4	0
1002	0	0
1003	8	0
1004	6	0
1005	3	0
1006	4	0
1007	2	0
1008	1	0
1009	3	0
1010	2	0
1011	2	0
1012	2	0
n	12	12
Mean	3	0
S.D.	2	0

Open field observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2001	0	0	0	0	0	N	0	0
2002	0	0	0	0	0	N	0	1
2003	0	0	0	0	0	N	0	0
2004	0	0	0	0	0	N	0	0
2005	0	0	0	0	0	N	0	0
2006	0	0	0	0	0	N	0	0
2007	0	0	0	0	0	N	0	0
2008	0	0	0	0	0	N	0	0
2009	0	0	0	0	0	N	0	0
2010	0	0	0	0	0	N	0	0
2011	0	0	0	0	0	N	0	0
2012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 2 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2001	3	0
2002	5	0
2003	5	0
2004	5	0
2005	0	0
2006	4	0
2007	6	0
2008	4	0
2009	2	0
2010	2	0
2011	3	0
2012	4	0
n	12	12
Mean	4	0
S.D.	2	0

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 2 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3001	0	0	0	0	0	N	0	0
3002	0	0	0	0	0	N	0	0
3003	0	0	0	0	0	N	0	0
3004	0	0	0	0	0	N	0	0
3005	0	0	0	0	0	N	0	0
3006	0	0	0	0	0	N	0	0
3007	0	0	0	0	0	N	0	0
3008	0	0	0	0	0	N	0	0
3009	0	0	0	0	0	N	0	0
3010	0	0	0	0	0	N	0	0
3011	0	0	0	0	0	N	0	0
3012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 2 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3001	5	0
3002	2	1
3003	9	0
3004	3	0
3005	7	0
3006	6	0
3007	8	0
3008	4	0
3009	5	0
3010	4	0
3011	5	0
3012	1	0
n	12	12
Mean	5	0
S.D.	2	0

Open field observation

Period : F0 before mating

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4001	0	0	0	0	0	N	0	0
4002	0	0	0	0	0	N	0	0
4003	0	0	0	0	0	N	0	0
4004	0	0	0	0	0	N	0	1
4005	0	0	0	0	0	N	0	0
4006	0	0	0	0	0	N	0	0
4007	0	0	0	0	0	N	0	0
4008	0	0	0	0	0	N	0	0
4009	0	0	0	0	0	N	0	0
4010	0	0	0	0	0	N	0	0
4011	0	0	0	0	0	N	0	0
4012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Male Week : 2 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4001	4	0
4002	7	0
4003	5	0
4004	4	0
4005	8	0
4006	7	0
4007	0	0
4008	6	0
4009	4	0
4010	3	0
4011	2	0
4012	3	0
n	12	12
Mean	4	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 3 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1001	0	0	0	0	0	N	0	0
1002	0	0	0	0	0	N	0	0
1003	0	0	0	0	0	N	0	0
1004	0	0	0	0	0	N	0	0
1005	0	0	0	0	0	N	0	0
1006	0	0	0	0	0	N	0	0
1007	0	0	0	0	0	N	0	0
1008	0	0	0	0	0	N	0	0
1009	0	0	0	0	0	N	0	0
1010	0	0	0	0	0	N	0	0
1011	0	0	0	0	0	N	0	0
1012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 3 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1001	2	0
1002	5	0
1003	8	0
1004	7	0
1005	2	0
1006	2	0
1007	2	0
1008	2	0
1009	2	0
1010	5	0
1011	1	0
1012	5	0
n	12	12
Mean	4	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 3 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2001	0	0	0	0	0	N	0	0
2002	0	0	0	0	0	N	0	1
2003	0	0	0	0	0	N	0	0
2004	0	0	0	0	0	N	0	0
2005	0	0	0	0	0	N	0	0
2006	0	0	0	0	0	N	0	0
2007	0	0	0	0	0	N	0	0
2008	0	0	0	0	0	N	0	0
2009	0	0	0	0	0	N	0	0
2010	0	0	0	0	0	N	0	0
2011	0	0	0	0	0	N	0	0
2012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 3 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2001	3	0
2002	3	0
2003	2	0
2004	6	0
2005	1	0
2006	6	0
2007	5	0
2008	4	0
2009	1	0
2010	0	0
2011	1	0
2012	5	0
n	12	12
Mean	3	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 3 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3001	0	0	0	0	0	N	0	0
3002	0	0	0	0	0	N	0	0
3003	0	0	0	0	0	N	0	0
3004	0	0	0	0	0	N	0	0
3005	0	0	0	0	0	N	0	0
3006	0	0	0	0	0	N	0	0
3007	0	0	0	0	0	N	0	0
3008	0	0	0	0	0	N	0	0
3009	0	0	0	0	0	N	0	0
3010	0	0	0	0	0	N	0	0
3011	0	0	0	0	0	N	0	0
3012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 3 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3001	3	0
3002	1	0
3003	5	0
3004	4	0
3005	8	0
3006	4	0
3007	7	0
3008	3	0
3009	7	0
3010	3	0
3011	5	0
3012	4	0
n	12	12
Mean	5	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 3 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4001	0	0	0	0	0	N	0	0
4002	0	0	0	0	0	N	0	0
4003	0	0	0	0	0	N	0	0
4004	0	0	0	0	0	N	0	1
4005	0	0	0	0	0	N	0	0
4006	0	0	0	0	0	N	0	0
4007	0	0	0	0	0	N	0	0
4008	0	0	0	0	0	N	0	0
4009	0	0	0	0	0	N	0	0
4010	0	0	0	0	0	N	0	0
4011	0	0	0	0	0	N	0	0
4012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 3 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4001	3	0
4002	7	0
4003	5	0
4004	2	0
4005	1	0
4006	2	0
4007	2	0
4008	5	0
4009	2	0
4010	6	0
4011	2	0
4012	1	0
n	12	12
Mean	3	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 4 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1001	0	0	0	0	0	N	0	0
1002	0	0	0	0	0	N	0	0
1003	0	0	0	0	0	N	0	0
1004	0	0	0	0	0	N	0	0
1005	0	0	0	0	0	N	0	0
1006	0	0	0	0	0	N	0	0
1007	0	0	0	0	0	N	0	0
1008	0	0	0	0	0	N	0	0
1009	0	0	0	0	0	N	0	0
1010	0	0	0	0	0	N	0	0
1011	0	0	0	0	0	N	0	0
1012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 4 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1001	3	0
1002	4	0
1003	6	0
1004	4	0
1005	1	0
1006	1	0
1007	2	0
1008	3	0
1009	2	0
1010	3	0
1011	3	0
1012	2	0
n	12	12
Mean	3	0
S.D.	1	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 4 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2001	0	0	0	0	0	N	0	1
2002	0	0	0	0	0	N	0	1
2003	0	0	0	0	0	N	0	0
2004	0	0	0	0	0	N	0	0
2005	0	0	0	0	0	N	0	0
2006	0	0	0	0	0	N	0	0
2007	0	0	0	0	0	N	0	0
2008	0	0	0	0	0	N	0	0
2009	0	0	0	0	0	N	0	0
2010	0	0	0	0	0	N	0	0
2011	0	0	0	0	0	N	0	0
2012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 4 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2001	3	0
2002	2	0
2003	5	0
2004	7	0
2005	2	0
2006	5	0
2007	5	0
2008	6	0
2009	2	0
2010	3	0
2011	5	0
2012	7	0
n	12	12
Mean	4	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 4 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3001	0	0	0	0	0	N	0	0
3002	0	0	0	0	0	N	0	0
3003	0	0	0	0	0	N	0	0
3004	0	0	0	0	0	N	0	0
3005	0	0	0	0	0	N	0	0
3006	0	0	0	0	0	N	0	0
3007	0	0	0	0	0	N	0	0
3008	0	0	0	0	0	N	0	0
3009	0	0	0	0	0	N	0	0
3010	0	0	0	0	0	N	0	0
3011	0	0	0	0	0	N	0	0
3012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 4 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3001	2	0
3002	2	0
3003	5	0
3004	5	0
3005	5	0
3006	1	0
3007	4	0
3008	4	0
3009	3	0
3010	4	0
3011	5	0
3012	7	0
n	12	12
Mean	4	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 4 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4001	0	0	0	0	0	N	0	0
4002	0	0	0	0	0	N	0	0
4003	0	0	0	0	0	N	0	1
4004	0	0	0	0	0	N	0	1
4005	0	0	0	0	0	N	0	0
4006	0	0	0	0	0	N	0	0
4007	0	0	0	0	0	N	0	0
4008	0	0	0	0	0	N	0	0
4009	0	0	0	0	0	N	0	0
4010	0	0	0	0	0	N	0	0
4011	0	0	0	0	0	N	0	0
4012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 4 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4001	4	0
4002	4	0
4003	5	0
4004	4	0
4005	3	0
4006	3	0
4007	2	0
4008	3	0
4009	7	0
4010	4	0
4011	2	0
4012	0	0
n	12	12
Mean	3	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 5 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1001	0	0	0	0	0	N	0	0
1002	0	0	0	0	0	N	0	0
1003	0	0	0	0	0	N	0	0
1004	0	0	0	0	0	N	0	0
1005	0	0	0	0	0	N	0	0
1006	0	0	0	0	0	N	0	0
1007	0	0	0	0	0	N	0	0
1008	0	0	0	0	0	N	0	0
1009	0	0	0	0	0	N	0	0
1010	0	0	0	0	0	N	0	0
1011	0	0	0	0	0	N	0	0
1012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 5 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1001	4	0
1002	5	0
1003	6	0
1004	4	0
1005	0	0
1006	1	0
1007	2	0
1008	5	0
1009	3	0
1010	4	0
1011	2	0
1012	4	0
n	12	12
Mean	3	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 5 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2001	0	0	0	0	0	N	0	0
2002	0	0	0	0	0	N	0	1
2003	0	0	0	0	0	N	0	0
2004	0	0	0	0	0	N	0	0
2005	0	0	0	0	0	N	0	0
2006	0	0	0	0	0	N	0	0
2007	0	0	0	0	0	N	0	0
2008	0	0	0	0	0	N	0	0
2009	0	0	0	0	0	N	0	1
2010	0	0	0	0	0	N	0	0
2011	0	0	0	0	0	N	0	0
2012	0	0	0	0	0	N	0	1
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 5 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2001	5	0
2002	3	0
2003	3	0
2004	8	0
2005	3	0
2006	6	0
2007	5	0
2008	6	0
2009	1	0
2010	1	0
2011	2	0
2012	4	0
n	12	12
Mean	4	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 5 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3001	0	0	0	0	0	N	0	0
3002	0	0	0	0	0	N	0	0
3003	0	0	0	0	0	N	0	0
3004	0	0	0	0	0	N	0	0
3005	0	0	0	0	0	N	0	0
3006	0	0	0	0	0	N	0	0
3007	0	0	0	0	0	N	0	0
3008	0	0	0	0	0	N	0	0
3009	0	0	0	0	0	N	0	0
3010	0	0	0	0	0	N	0	0
3011	0	0	0	0	0	N	0	0
3012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 5 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3001	4	0
3002	2	0
3003	7	0
3004	4	0
3005	5	0
3006	5	0
3007	5	0
3008	5	0
3009	4	0
3010	3	0
3011	3	0
3012	5	0
n	12	12
Mean	4	0
S.D.	1	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 5 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4001	0	0	0	0	0	N	0	0
4002	0	0	0	0	0	N	0	0
4003	0	0	0	0	0	N	0	0
4004	0	0	0	0	0	N	0	0
4005	0	0	0	0	0	N	0	0
4006	0	0	0	0	0	N	0	0
4007	0	0	0	0	0	N	0	0
4008	0	0	0	0	0	N	0	0
4009	0	0	0	0	0	N	0	0
4010	0	0	0	0	0	N	0	1
4011	0	0	0	0	0	N	0	0
4012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 5 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4001	2	0
4002	6	0
4003	4	0
4004	4	0
4005	2	0
4006	3	0
4007	0	0
4008	4	0
4009	7	0
4010	3	0
4011	2	0
4012	3	0
n	12	12
Mean	3	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 6 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1001	0	0	0	0	0	N	0	0
1002	0	0	0	0	0	N	0	0
1003	0	0	0	0	0	N	0	0
1004	0	0	0	0	0	N	0	0
1005	0	0	0	0	0	N	0	0
1006	0	0	0	0	0	N	0	1
1007	0	0	0	0	0	N	0	0
1008	0	0	0	0	0	N	0	0
1009	0	0	0	0	0	N	0	0
1010	0	0	0	0	0	N	0	0
1011	0	0	0	0	0	N	0	0
1012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 6 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1001	5	0
1002	4	0
1003	6	0
1004	5	0
1005	1	0
1006	3	0
1007	3	0
1008	2	0
1009	3	0
1010	6	0
1011	6	0
1012	5	0
n	12	12
Mean	4	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 6 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2001	0	0	0	0	0	N	0	1
2002	0	0	0	0	0	N	0	1
2003	0	0	0	0	0	N	0	0
2004	0	0	0	0	0	N	0	0
2005	0	0	0	0	0	N	0	0
2006	0	0	0	0	0	N	0	0
2007	0	0	0	0	0	N	0	0
2008	0	0	0	0	0	N	0	0
2009	0	0	0	0	0	N	0	1
2010	0	0	0	0	0	N	0	0
2011	0	0	0	0	0	N	0	0
2012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 6 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2001	3	0
2002	4	0
2003	5	0
2004	4	0
2005	3	0
2006	7	0
2007	5	0
2008	6	0
2009	4	0
2010	3	0
2011	3	0
2012	8	0
n	12	12
Mean	5	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 6 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3001	0	0	0	0	0	N	0	0
3002	0	0	0	0	0	N	0	0
3003	0	0	0	0	0	N	0	0
3004	0	0	0	0	0	N	0	0
3005	0	0	0	0	0	N	0	0
3006	0	0	0	0	0	N	0	0
3007	0	0	0	0	0	N	0	0
3008	0	0	0	0	0	N	0	0
3009	0	0	0	0	0	N	0	0
3010	0	0	0	0	0	N	0	0
3011	0	0	0	0	0	N	0	0
3012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 6 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3001	6	0
3002	1	0
3003	4	0
3004	5	0
3005	9	0
3006	4	0
3007	7	0
3008	5	0
3009	2	0
3010	5	0
3011	6	0
3012	6	0
n	12	12
Mean	5	0
S.D.	2	0

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 6 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4001	0	0	0	0	0	N	0	0
4002	0	0	0	0	0	N	0	0
4003	0	0	0	0	0	N	0	0
4004	0	0	0	0	0	N	0	1
4005	0	0	0	0	0	N	0	0
4006	0	0	0	0	0	N	0	0
4007	0	0	0	0	0	N	0	0
4008	0	0	0	0	0	N	0	0
4009	0	0	0	0	0	N	0	0
4010	0	0	0	0	0	N	0	0
4011	0	0	0	0	0	N	0	0
4012	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 mating

Generation : F0 Sex : Male Week : 6 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4001	5	0
4002	5	0
4003	6	0
4004	3	0
4005	4	0
4006	3	0
4007	4	0
4008	7	0
4009	3	0
4010	6	0
4011	3	0
4012	2	0
n	12	12
Mean	4	0
S.D.	2	0

Open field observation

Generation : F0 Sex : Female Week : 1 Period : F0 before mating

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1101	0	0	0	0	0	N	0	0
1102	0	0	0	0	0	N	0	0
1103	0	0	0	0	0	N	0	0
1104	0	0	0	0	0	N	0	1
1105	0	0	0	0	0	N	0	0
1106	0	0	0	0	0	N	0	0
1107	0	0	0	0	0	N	0	0
1108	0	0	0	0	0	N	0	0
1109	0	0	0	0	0	N	0	0
1110	0	0	0	0	0	N	0	0
1111	0	0	0	0	0	N	0	0
1112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

With the reproduction/ developmental toxicity screening test in rats						
Open field observation		Period : F0 before mating		Species : Rat		
Generation : F0	Sex : Female	Week : 1	Dose : Dodecyloxirane 0 mg/kg			
Rearing		Defecation count				
Animal No.						
1101	8	0				
1102	7	0				
1103	5	0				
1104	6	0				
1105	4	0				
1106	9	0				
1107	3	0				
1108	10	0				
1109	5	0				
1110	2	0				
1111	5	0				
1112	1	0				
n	12	12				
Mean	5	0				
S.D.	3	0				

Open field observation

Generation : F0 Sex : Female Week : 1 Period : F0 before mating

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2101	0	0	0	0	0	N	0	0
2102	0	0	0	0	0	N	0	0
2103	0	0	0	0	0	N	0	0
2104	0	0	0	0	0	N	0	0
2105	0	0	0	0	0	N	0	0
2106	0	0	0	0	0	N	0	0
2107	0	0	0	0	0	N	0	0
2108	0	0	0	0	0	N	0	0
2109	0	0	0	0	0	N	0	0
2110	0	0	0	0	0	N	0	0
2111	0	0	0	0	0	N	0	0
2112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2101	4	0
2102	6	0
2103	9	0
2104	4	0
2105	5	0
2106	7	0
2107	10	0
2108	4	0
2109	8	0
2110	5	0
2111	6	0
2112	2	0
n	12	12
Mean	6	0
S.D.	2	0

Open field observation

Generation : F0 Sex : Female Week : 1 Period : F0 before mating

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3101	0	0	0	0	0	N	0	0
3102	0	0	0	0	0	N	0	0
3103	0	0	0	0	0	N	0	0
3104	0	0	0	0	0	N	0	0
3105	0	0	0	0	0	N	0	0
3106	0	0	0	0	0	N	0	0
3107	0	0	0	0	0	N	0	0
3108	0	0	0	0	0	N	0	0
3109	0	0	0	0	0	N	0	0
3110	0	0	0	0	0	N	0	0
3111	0	0	0	0	0	N	0	0
3112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Female Week : 1 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3101	6	0
3102	5	0
3103	8	0
3104	6	0
3105	7	0
3106	7	0
3107	11	0
3108	5	0
3109	8	0
3110	6	0
3111	4	0
3112	5	0
n	12	12
Mean	7	0
S.D.	2	0

Open field observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4101	0	0	0	0	0	N	0	0
4102	0	0	0	0	0	N	0	0
4103	0	0	0	0	0	N	0	0
4104	0	0	0	0	0	N	0	0
4105	0	0	0	0	0	N	0	0
4106	0	0	0	0	0	N	0	0
4107	0	0	0	0	0	N	0	0
4108	0	0	0	0	0	N	0	0
4109	0	0	0	0	0	N	0	0
4110	0	0	0	0	0	N	0	0
4111	0	0	0	0	0	N	0	0
4112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Female Week : 1 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4101	4	0
4102	2	0
4103	6	0
4104	7	0
4105	7	0
4106	5	0
4107	5	0
4108	2	0
4109	3	0
4110	4	0
4111	6	0
4112	8	0
n	12	12
Mean	5	0
S.D.	2	0

Open field observation

Generation : F0 Sex : Female Week : 2 Period : F0 before mating

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1101	0	0	0	0	0	N	0	0
1102	0	0	0	0	0	N	0	0
1103	0	0	0	0	0	N	0	0
1104	0	0	0	0	0	N	0	0
1105	0	0	0	0	0	N	0	0
1106	0	0	0	0	0	N	0	0
1107	0	0	0	0	0	N	0	0
1108	0	0	0	0	0	N	0	0
1109	0	0	0	0	0	N	0	0
1110	0	0	0	0	0	N	0	0
1111	0	0	0	0	0	N	0	0
1112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Female Week : 2 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1101	7	0
1102	2	0
1103	9	0
1104	5	0
1105	5	0
1106	8	0
1107	6	0
1108	10	0
1109	8	0
1110	4	0
1111	4	0
1112	3	0
n	12	12
Mean	6	0
S.D.	3	0

Open field observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2101	0	0	0	0	0	N	0	0
2102	0	0	0	0	0	N	0	0
2103	0	0	0	0	0	N	0	0
2104	0	0	0	0	0	N	0	0
2105	0	0	0	0	0	N	0	0
2106	0	0	0	0	0	N	0	0
2107	0	0	0	0	0	N	0	0
2108	0	0	0	0	0	N	0	0
2109	0	0	0	0	0	N	0	0
2110	0	0	0	0	0	N	0	0
2111	0	0	0	0	0	N	0	0
2112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Appendix 2 - 212

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Open field observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2101	7	0
2102	5	0
2103	7	0
2104	7	0
2105	6	0
2106	6	0
2107	9	0
2108	8	0
2109	8	0
2110	8	0
2111	6	0
2112	7	0
n	12	12
Mean	7	0
S.D.	1	0

Open field observation

Generation : F0 Sex : Female Week : 2 Period : F0 before mating

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3101	0	0	0	0	0	N	0	0
3102	0	0	0	0	0	N	0	0
3103	0	0	0	0	0	N	0	0
3104	0	0	0	0	0	N	0	0
3105	0	0	0	0	0	N	0	0
3106	0	0	0	0	0	N	0	0
3107	0	0	0	0	0	N	0	0
3108	0	0	0	0	0	N	0	0
3109	0	0	0	0	0	N	0	0
3110	0	0	0	0	0	N	0	0
3111	0	0	0	0	0	N	0	0
3112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0 Sex : Female Week : 2 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3101	8	0
3102	5	0
3103	7	0
3104	7	0
3105	7	0
3106	8	0
3107	6	0
3108	9	0
3109	7	0
3110	8	0
3111	7	0
3112	5	0
n	12	12
Mean	7	0
S.D.	1	0

Open field observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4101	0	0	0	0	0	N	0	0
4102	0	0	0	0	0	N	0	0
4103	0	0	0	0	0	N	0	0
4104	0	0	0	0	0	N	0	0
4105	0	0	0	0	0	N	0	0
4106	0	0	0	0	0	N	0	0
4107	0	0	0	0	0	N	0	0
4108	0	0	0	0	0	N	0	0
4109	0	0	0	0	0	N	0	0
4110	0	0	0	0	0	N	0	0
4111	0	0	0	0	0	N	0	0
4112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 before mating

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4101	9	0
4102	8	0
4103	5	0
4104	8	0
4105	7	0
4106	7	0
4107	7	0
4108	9	0
4109	8	0
4110	7	0
4111	7	0
4112	9	0
n	12	12
Mean	8	0
S.D.	1	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 1 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1101	0	0	0	0	0	N	0	0
1102	0	0	0	0	0	N	0	0
1103	0	0	0	0	0	N	0	0
1104	0	0	0	0	0	N	0	0
1105	0	0	0	0	0	N	0	0
1106	0	0	0	0	0	N	0	0
1107	0	0	0	0	0	N	0	0
1108	0	0	0	0	0	N	0	0
1109	0	0	0	0	0	N	0	0
1110	0	0	0	0	0	N	0	0
1111	0	0	0	0	0	N	0	0
1112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 1 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1101	6	0
1102	4	0
1103	6	0
1104	7	0
1105	5	0
1106	9	0
1107	4	0
1108	9	0
1109	10	0
1110	5	0
1111	7	0
1112	6	0
n	12	12
Mean	7	0
S.D.	2	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 1 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2101	0	0	0	0	0	N	0	0
2102	0	0	0	0	0	N	0	0
2103	0	0	0	0	0	N	0	0
2104	0	0	0	0	0	N	0	0
2105	0	0	0	0	0	N	0	0
2107	0	0	0	0	0	N	0	0
2108	0	0	0	0	0	N	0	0
2110	0	0	0	0	0	N	0	0
2111	0	0	0	0	0	N	0	0
2112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 1 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2101	8	0
2102	5	0
2103	6	0
2104	8	0
2105	5	0
2107	8	0
2108	6	0
2110	6	0
2111	7	0
2112	9	0
n	10	10
Mean	7	0
S.D.	1	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 1 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3101	0	0	0	0	0	N	0	0
3102	0	0	0	0	0	N	0	0
3103	0	0	0	0	0	N	0	0
3104	0	0	0	0	0	N	0	0
3105	0	0	0	0	0	N	0	0
3106	0	0	0	0	0	N	0	0
3107	0	0	0	0	0	N	0	0
3110	0	0	0	0	0	N	0	0
3111	0	0	0	0	0	N	0	0
3112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 1 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3101	5	0
3102	5	0
3103	8	0
3104	10	0
3105	6	0
3106	8	0
3107	8	0
3110	4	0
3111	3	0
3112	8	0
n	10	10
Mean	7	0
S.D.	2	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 1 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4102	0	0	0	0	0	N	0	0
4103	0	0	0	0	0	N	0	0
4104	0	0	0	0	0	N	0	0
4105	0	0	0	0	0	N	0	0
4106	0	0	0	0	0	N	0	0
4107	0	0	0	0	0	N	0	0
4108	0	0	0	0	0	N	0	0
4109	0	0	0	0	0	N	0	0
4110	0	0	0	0	0	N	0	0
4111	0	0	0	0	0	N	0	0
4112	0	0	0	0	0	N	0	0
n	11	11	11	11	11	11	11	11

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 1 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4102	6	0
4103	3	0
4104	7	0
4105	5	0
4106	6	0
4107	8	0
4108	6	0
4109	8	0
4110	6	0
4111	7	0
4112	6	0
n	11	11
Mean	6	0
S.D.	1	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1101	0	0	0	0	0	N	0	0
1102	0	0	0	0	0	N	0	0
1103	0	0	0	0	0	N	0	0
1104	0	0	0	0	0	N	0	0
1105	0	0	0	0	0	N	0	0
1106	0	0	0	0	0	N	0	0
1107	0	0	0	0	0	N	0	0
1108	0	0	0	0	0	N	0	0
1109	0	0	0	0	0	N	0	0
1110	0	0	0	0	0	N	0	0
1111	0	0	0	0	0	N	0	0
1112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1101	8	0
1102	3	0
1103	8	0
1104	4	0
1105	3	0
1106	11	0
1107	4	0
1108	10	0
1109	11	0
1110	8	0
1111	8	0
1112	4	0
n	12	12
Mean	7	0
S.D.	3	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2101	0	0	0	0	0	N	0	0
2102	0	0	0	0	0	N	0	0
2103	0	0	0	0	0	N	0	0
2104	0	0	0	0	0	N	0	0
2105	0	0	0	0	0	N	0	0
2107	0	0	0	0	0	N	0	0
2108	0	0	0	0	0	N	0	0
2110	0	0	0	0	0	N	0	0
2111	0	0	0	0	0	N	0	0
2112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2101	7	0
2102	6	0
2103	6	0
2104	7	0
2105	4	0
2107	7	0
2108	7	0
2110	10	0
2111	8	0
2112	7	0
n	10	10
Mean	7	0
S.D.	2	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3101	0	0	0	0	0	N	0	0
3102	0	0	0	0	0	N	0	0
3103	0	0	0	0	0	N	0	0
3104	0	0	0	0	0	N	0	0
3105	0	0	0	0	0	N	0	0
3106	0	0	0	0	0	N	0	0
3107	0	0	0	0	0	N	0	0
3110	0	0	0	0	0	N	0	0
3111	0	0	0	0	0	N	0	0
3112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3101	6	0
3102	2	0
3103	8	0
3104	6	0
3105	7	0
3106	8	0
3107	9	0
3110	5	0
3111	4	0
3112	7	0
n	10	10
Mean	6	0
S.D.	2	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4102	0	0	0	0	0	N	0	0
4103	0	0	0	0	0	N	0	0
4104	0	0	0	0	0	N	0	0
4105	0	0	0	0	0	N	0	0
4106	0	0	0	0	0	N	0	0
4107	0	0	0	0	0	N	0	0
4108	0	0	0	0	0	N	0	0
4109	0	0	0	0	0	N	0	0
4110	0	0	0	0	0	N	0	0
4111	0	0	0	0	0	N	0	0
4112	0	0	0	0	0	N	0	0
n	11	11	11	11	11	11	11	11

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 7 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4102	5	0
4103	4	0
4104	7	0
4105	4	0
4106	4	0
4107	7	0
4108	7	0
4109	5	0
4110	5	0
4111	8	0
4112	5	0
n	11	11
Mean	6	0
S.D.	1	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1101	0	0	0	0	0	N	0	0
1102	0	0	0	0	0	N	0	0
1103	0	0	0	0	0	N	0	0
1104	0	0	0	0	0	N	0	0
1105	0	0	0	0	0	N	0	0
1106	0	0	0	0	0	N	0	0
1107	0	0	0	0	0	N	0	0
1108	0	0	0	0	0	N	0	0
1109	0	0	0	0	0	N	0	0
1110	0	0	0	0	0	N	0	0
1111	0	0	0	0	0	N	0	0
1112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1101	7	0
1102	3	0
1103	4	0
1104	3	0
1105	5	0
1106	8	0
1107	5	0
1108	10	0
1109	8	0
1110	7	0
1111	5	0
1112	6	0
n	12	12
Mean	6	0
S.D.	2	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2101	0	0	0	0	0	N	0	0
2102	0	0	0	0	0	N	0	0
2103	0	0	0	0	0	N	0	0
2104	0	0	0	0	0	N	0	0
2105	0	0	0	0	0	N	0	0
2107	0	0	0	0	0	N	0	0
2108	0	0	0	0	0	N	0	0
2110	0	0	0	0	0	N	0	0
2111	0	0	0	0	0	N	0	0
2112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2101	7	0
2102	4	0
2103	5	0
2104	4	0
2105	5	0
2107	4	0
2108	5	0
2110	4	0
2111	5	0
2112	6	0
n	10	10
Mean	5	0
S.D.	1	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3101	0	0	0	0	0	N	0	0
3102	0	0	0	0	0	N	0	0
3103	0	0	0	0	0	N	0	0
3104	0	0	0	0	0	N	0	0
3105	0	0	0	0	0	N	0	0
3106	0	0	0	0	0	N	0	0
3107	0	0	0	0	0	N	0	0
3110	0	0	0	0	0	N	0	0
3111	0	0	0	0	0	N	0	0
3112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3101	6	0
3102	3	0
3103	5	0
3104	7	0
3105	6	0
3106	5	0
3107	5	0
3110	4	0
3111	5	0
3112	8	0
n	10	10
Mean	5	0
S.D.	1	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4102	0	0	0	0	0	N	0	0
4103	0	0	0	0	0	N	0	0
4104	0	0	0	0	0	N	0	0
4105	0	0	0	0	0	N	0	0
4106	0	0	0	0	0	N	0	0
4107	0	0	0	0	0	N	0	0
4108	0	0	0	0	0	N	0	0
4109	0	0	0	0	0	N	0	0
4110	0	0	0	0	0	N	0	0
4111	0	0	0	0	0	N	0	0
4112	0	0	0	0	0	N	0	1
n	11	11	11	11	11	11	11	11

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 14 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4102	6	0
4103	4	0
4104	6	0
4105	1	0
4106	4	0
4107	4	0
4108	3	0
4109	4	0
4110	4	0
4111	7	0
4112	3	0
n	11	11
Mean	4	0
S.D.	2	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1101	0	0	0	0	0	N	0	0
1102	0	0	0	0	0	N	0	0
1103	0	0	0	0	0	N	0	0
1104	0	0	0	0	0	N	0	0
1105	0	0	0	0	0	N	0	0
1106	0	0	0	0	0	N	0	0
1107	0	0	0	0	0	N	0	0
1108	0	0	0	0	0	N	0	0
1109	0	0	0	0	0	N	0	0
1110	0	0	0	0	0	N	0	0
1111	0	0	0	0	0	N	0	0
1112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1101	6	0
1102	2	0
1103	3	0
1104	2	0
1105	4	0
1106	4	0
1107	4	0
1108	6	0
1109	3	0
1110	1	0
1111	5	0
1112	0	0
n	12	12
Mean	3	0
S.D.	2	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2101	0	0	0	0	0	N	0	0
2102	0	0	0	0	0	N	0	0
2103	0	0	0	0	0	N	0	0
2104	0	0	0	0	0	N	0	0
2105	0	0	0	0	0	N	0	0
2107	0	0	0	0	0	N	0	0
2108	0	0	0	0	0	N	0	0
2110	0	0	0	0	0	N	0	0
2111	0	0	0	0	0	N	0	0
2112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2101	3	0
2102	5	0
2103	4	0
2104	3	0
2105	6	0
2107	1	0
2108	6	0
2110	3	0
2111	4	0
2112	6	0
n	10	10
Mean	4	0
S.D.	2	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3101	0	0	0	0	0	N	0	0
3102	0	0	0	0	0	N	0	0
3103	0	0	0	0	0	N	0	0
3104	0	0	0	0	0	N	0	0
3105	0	0	0	0	0	N	0	0
3106	0	0	0	0	0	N	0	1
3107	0	0	0	0	0	N	0	0
3110	0	0	0	0	0	N	0	0
3111	0	0	0	0	0	N	0	0
3112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3101	4	0
3102	2	0
3103	4	0
3104	3	0
3105	1	0
3106	4	0
3107	4	0
3110	6	0
3111	4	0
3112	4	0
n	10	10
Mean	4	0
S.D.	1	0

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4102	0	0	0	0	0	N	0	0
4103	0	0	0	0	0	N	0	0
4104	0	0	0	0	0	N	0	0
4105	0	0	0	0	0	N	0	0
4106	0	0	0	0	0	N	0	0
4107	0	0	0	0	0	N	0	0
4108	0	0	0	0	0	N	0	0
4109	0	0	0	0	0	N	0	0
4110	0	0	0	0	0	N	0	0
4111	0	0	0	0	0	N	0	0
4112	0	0	0	0	0	N	0	0
n	11	11	11	11	11	11	11	11

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 gestation

Generation : F0 Sex : Female Day : 20 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4102	5	0
4103	5	0
4104	4	0
4105	1	1
4106	4	0
4107	4	0
4108	4	0
4109	4	0
4110	1	0
4111	5	0
4112	5	0
n	11	11
Mean	4	0
S.D.	1	0

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 6 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1101	0	0	0	0	0	N	0	0
1102	0	0	0	0	0	N	0	0
1103	0	0	0	0	0	N	0	0
1104	0	0	0	0	0	N	0	0
1105	0	0	0	0	0	N	0	0
1106	0	0	0	0	0	N	0	0
1107	0	0	0	0	0	N	0	0
1108	0	0	0	0	0	N	0	0
1109	0	0	0	0	0	N	0	0
1110	0	0	0	0	0	N	0	0
1111	0	0	0	0	0	N	0	0
1112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 6 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1101	5	0
1102	6	0
1103	6	0
1104	3	0
1105	5	0
1106	8	0
1107	3	0
1108	3	0
1109	8	0
1110	7	0
1111	4	0
1112	5	0
n	12	12
Mean	5	0
S.D.	2	0

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 6 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2101	0	0	0	0	0	N	0	0
2102	0	0	0	0	0	N	0	0
2103	0	0	0	0	0	N	0	0
2104	0	0	0	0	0	N	0	0
2105	0	0	0	0	0	N	0	0
2107	0	0	0	0	0	N	0	0
2108	0	0	0	0	0	N	0	0
2110	0	0	0	0	0	N	0	0
2111	0	0	0	0	0	N	0	0
2112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 6 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2101	6	0
2102	7	0
2103	7	0
2104	5	0
2105	7	0
2107	3	0
2108	4	0
2110	7	0
2111	4	0
2112	7	0
n	10	10
Mean	6	0
S.D.	2	0

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 6 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3101	0	0	0	0	0	N	0	0
3102	0	0	0	0	0	N	0	0
3103	0	0	0	0	0	N	0	0
3104	0	0	0	0	0	N	0	0
3105	0	0	0	0	0	N	0	0
3106	0	0	0	0	0	N	0	0
3107	0	0	0	0	0	N	0	0
3110	0	0	0	0	0	N	0	0
3111	0	0	0	0	0	N	0	0
3112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 6 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3101	4	0
3102	4	0
3103	3	0
3104	5	0
3105	5	0
3106	12	0
3107	1	0
3110	2	0
3111	1	0
3112	10	0
n	10	10
Mean	5	0
S.D.	4	0

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 6 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4102	0	0	0	0	0	N	0	0
4103	0	0	0	0	0	N	0	0
4104	0	0	0	0	0	N	0	0
4105	0	0	0	0	0	N	0	0
4107	0	0	0	0	0	N	0	0
4108	0	0	0	0	0	N	0	0
4109	0	0	0	0	0	N	0	0
4110	0	0	0	0	0	N	0	0
4111	0	0	0	0	0	N	0	0
4112	0	0	0	0	0	N	0	1
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 6 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4102	5	0
4103	6	0
4104	5	0
4105	1	0
4107	6	0
4108	5	0
4109	4	0
4110	4	0
4111	3	0
4112	5	0
n	10	10
Mean	4	0
S.D.	2	0

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1101	0	0	0	0	0	N	0	0
1102	0	0	0	0	0	N	0	0
1103	0	0	0	0	0	N	0	0
1104	0	0	0	0	0	N	0	0
1105	0	0	0	0	0	N	0	0
1106	0	0	0	0	0	N	0	0
1107	0	0	0	0	0	N	0	0
1108	0	0	0	0	0	N	0	0
1109	0	0	0	0	0	N	0	0
1110	0	0	0	0	0	N	0	0
1111	0	0	0	0	0	N	0	0
1112	0	0	0	0	0	N	0	0
n	12	12	12	12	12	12	12	12

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1101	7	0
1102	5	0
1103	6	0
1104	7	0
1105	5	0
1106	7	0
1107	2	0
1108	6	0
1109	7	0
1110	6	0
1111	6	0
1112	2	0
n	12	12
Mean	6	0
S.D.	2	0

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
2101	0	0	0	0	0	N	0	0
2102	0	0	0	0	0	N	0	0
2103	0	0	0	0	0	N	0	0
2104	0	0	0	0	0	N	0	0
2105	0	0	0	0	0	N	0	0
2107	0	0	0	0	0	N	0	0
2108	0	0	0	0	0	N	0	0
2110	0	0	0	0	0	N	0	0
2111	0	0	0	0	0	N	0	0
2112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
2101	7	0
2102	9	0
2103	5	0
2104	5	0
2105	9	0
2107	8	0
2108	8	0
2110	6	0
2111	8	0
2112	8	0
n	10	10
Mean	7	0
S.D.	1	0

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
3101	0	0	0	0	0	N	0	0
3102	0	0	0	0	0	N	0	0
3103	0	0	0	0	0	N	0	0
3104	0	0	0	0	0	N	0	0
3105	0	0	0	0	0	N	0	0
3106	0	0	0	0	0	N	0	0
3107	0	0	0	0	0	N	0	0
3110	0	0	0	0	0	N	0	0
3111	0	0	0	0	0	N	0	0
3112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
3101	6	0
3102	5	0
3103	8	0
3104	5	0
3105	7	0
3106	8	0
3107	4	0
3110	7	0
3111	4	0
3112	5	0
n	10	10
Mean	6	0
S.D.	2	0

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4102	0	0	0	0	0	N	0	0
4103	0	0	0	0	0	N	0	0
4104	0	0	0	0	0	N	0	0
4105	0	0	0	0	0	N	0	0
4107	0	0	0	0	0	N	0	0
4108	0	0	0	0	0	N	0	0
4109	0	0	0	0	0	N	0	0
4110	0	0	0	0	0	N	0	0
4111	0	0	0	0	0	N	0	0
4112	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4102	8	0
4103	5	0
4104	7	0
4105	3	0
4107	6	0
4108	3	0
4109	8	0
4110	4	0
4111	6	0
4112	7	0
n	10	10
Mean	6	0
S.D.	2	0

Open field observation

Period : Administration (non-mating group)

Generation : F0 Sex : Female Week : 1 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
5101	0	0	0	0	0	N	0	0
5102	0	0	0	0	0	N	0	0
5103	0	0	0	0	0	N	0	0
5104	0	0	0	0	0	N	0	0
5105	0	0	0	0	0	N	0	0
5106	0	0	0	0	0	N	0	0
5107	0	0	0	0	0	N	0	0
5108	0	0	0	0	0	N	0	0
5109	0	0	0	0	0	N	0	0
5110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
5101	6	0
5102	10	0
5103	6	0
5104	7	0
5105	7	0
5106	5	0
5107	6	0
5108	5	0
5109	5	0
5110	2	0
n	10	10
Mean	6	0
S.D.	2	0

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
6101	0	0	0	0	0	N	0	0
6102	0	0	0	0	0	N	0	0
6103	0	0	0	0	0	N	0	0
6104	0	0	0	0	0	N	0	0
6105	0	0	0	0	0	N	0	0
6106	0	0	0	0	0	N	0	0
6107	0	0	0	0	0	N	0	0
6108	0	0	0	0	0	N	0	0
6109	0	0	0	0	0	N	0	0
6110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
6101	5	0
6102	6	0
6103	8	0
6104	7	0
6105	9	0
6106	4	0
6107	5	0
6108	4	0
6109	3	0
6110	6	0
n	10	10
Mean	6	0
S.D.	2	0

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
5101	0	0	0	0	0	N	0	0
5102	0	0	0	0	0	N	0	0
5103	0	0	0	0	0	N	0	0
5104	0	0	0	0	0	N	0	0
5105	0	0	0	0	0	N	0	0
5106	0	0	0	0	0	N	0	0
5107	0	0	0	0	0	N	0	0
5108	0	0	0	0	0	N	0	0
5109	0	0	0	0	0	N	0	0
5110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
5101	10	0
5102	5	0
5103	5	0
5104	10	0
5105	9	0
5106	11	0
5107	8	0
5108	6	0
5109	6	0
5110	3	0
n	10	10
Mean	7	0
S.D.	3	0

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
6101	0	0	0	0	0	N	0	0
6102	0	0	0	0	0	N	0	0
6103	0	0	0	0	0	N	0	0
6104	0	0	0	0	0	N	0	0
6105	0	0	0	0	0	N	0	0
6106	0	0	0	0	0	N	0	0
6107	0	0	0	0	0	N	0	0
6108	0	0	0	0	0	N	0	0
6109	0	0	0	0	0	N	0	0
6110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
6101	7	0
6102	7	0
6103	7	0
6104	8	0
6105	9	0
6106	6	0
6107	8	0
6108	5	0
6109	5	0
6110	7	0
n	10	10
Mean	7	0
S.D.	1	0

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 3

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
5101	0	0	0	0	0	N	0	0
5102	0	0	0	0	0	N	0	0
5103	0	0	0	0	0	N	0	0
5104	0	0	0	0	0	N	0	0
5105	0	0	0	0	0	N	0	0
5106	0	0	0	0	0	N	0	0
5107	0	0	0	0	0	N	0	0
5108	0	0	0	0	0	N	0	0
5109	0	0	0	0	0	N	0	0
5110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 3

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
5101	11	0
5102	9	0
5103	9	0
5104	11	0
5105	11	0
5106	11	0
5107	8	0
5108	5	0
5109	7	0
5110	6	0
n	10	10
Mean	9	0
S.D.	2	0

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 3

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
6101	0	0	0	0	0	N	0	0
6102	0	0	0	0	0	N	0	0
6103	0	0	0	0	0	N	0	0
6104	0	0	0	0	0	N	0	0
6105	0	0	0	0	0	N	0	0
6106	0	0	0	0	0	N	0	0
6107	0	0	0	0	0	N	0	0
6108	0	0	0	0	0	N	0	0
6109	0	0	0	0	0	N	0	0
6110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 3

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
6101	5	0
6102	7	0
6103	8	0
6104	9	0
6105	4	0
6106	6	0
6107	5	0
6108	9	0
6109	5	0
6110	9	0
n	10	10
Mean	7	0
S.D.	2	0

Open field observation

Period : Administration (non-mating group)

Generation : F0 Sex : Female Week : 4 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
5101	0	0	0	0	0	N	0	0
5102	0	0	0	0	0	N	0	0
5103	0	0	0	0	0	N	0	0
5104	0	0	0	0	0	N	0	0
5105	0	0	0	0	0	N	0	0
5106	0	0	0	0	0	N	0	0
5107	0	0	0	0	0	N	0	0
5108	0	0	0	0	0	N	0	0
5109	0	0	0	0	0	N	0	0
5110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 4

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
5101	8	0
5102	8	0
5103	4	0
5104	8	0
5105	4	0
5106	12	0
5107	5	0
5108	4	0
5109	7	0
5110	7	0
n	10	10
Mean	7	0
S.D.	3	0

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 4

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
6101	0	0	0	0	0	N	0	0
6102	0	0	0	0	0	N	0	0
6103	0	0	0	0	0	N	0	0
6104	0	0	0	0	0	N	0	0
6105	0	0	0	0	0	N	0	0
6106	0	0	0	0	0	N	0	0
6107	0	0	0	0	0	N	0	0
6108	0	0	0	0	0	N	0	0
6109	0	0	0	0	0	N	0	0
6110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 4

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
6101	4	0
6102	6	0
6103	4	0
6104	7	0
6105	4	0
6106	6	0
6107	8	0
6108	4	0
6109	5	0
6110	8	0
n	10	10
Mean	6	0
S.D.	2	0

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 5

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
5101	0	0	0	0	0	N	0	0
5102	0	0	0	0	0	N	0	0
5103	0	0	0	0	0	N	0	0
5104	0	0	0	0	0	N	0	0
5105	0	0	0	0	0	N	0	0
5106	0	0	0	0	0	N	0	0
5107	0	0	0	0	0	N	0	0
5108	0	0	0	0	0	N	0	0
5109	0	0	0	0	0	N	0	0
5110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 5

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
5101	8	0
5102	6	0
5103	6	0
5104	7	0
5105	7	0
5106	10	0
5107	8	0
5108	4	0
5109	9	0
5110	8	0
n	10	10
Mean	7	0
S.D.	2	0

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 5

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
6101	0	0	0	0	0	N	0	0
6102	0	0	0	0	0	N	0	0
6103	0	0	0	0	0	N	0	0
6104	0	0	0	0	0	N	0	0
6105	0	0	0	0	0	N	0	0
6106	0	0	0	0	0	N	0	0
6107	0	0	0	0	0	N	0	0
6108	0	0	0	0	0	N	0	0
6109	0	0	0	0	0	N	0	0
6110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 5

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
6101	8	0
6102	6	0
6103	8	0
6104	10	0
6105	9	0
6106	5	0
6107	9	0
6108	11	0
6109	5	0
6110	7	0
n	10	10
Mean	8	0
S.D.	2	0

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
5101	0	0	0	0	0	N	0	0
5102	0	0	0	0	0	N	0	0
5103	0	0	0	0	0	N	0	0
5104	0	0	0	0	0	N	0	0
5105	0	0	0	0	0	N	0	0
5106	0	0	0	0	0	N	0	0
5107	0	0	0	0	0	N	0	0
5108	0	0	0	0	0	N	0	0
5109	0	0	0	0	0	N	0	0
5110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
5101	10	0
5102	10	0
5103	7	0
5104	10	0
5105	8	0
5106	9	0
5107	6	0
5108	7	0
5109	8	0
5110	7	0
n	10	10
Mean	8	0
S.D.	1	0

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
6101	0	0	0	0	0	N	0	0
6102	0	0	0	0	0	N	0	0
6103	0	0	0	0	0	N	0	0
6104	0	0	0	0	0	N	0	0
6105	0	0	0	0	0	N	0	0
6106	0	0	0	0	0	N	0	0
6107	0	0	0	0	0	N	0	0
6108	0	0	0	0	0	N	0	0
6109	0	0	0	0	0	N	0	0
6110	0	0	0	0	0	N	0	0
n	10	10	10	10	10	10	10	10

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
6101	8	0
6102	5	0
6103	9	0
6104	11	0
6105	9	0
6106	6	0
6107	8	0
6108	5	0
6109	7	0
6110	10	0
n	10	10
Mean	8	0
S.D.	2	0

Open field observation

Period : Recovery

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1008	0	0	0	0	0	N	0	0
1009	0	0	0	0	0	N	0	1
1010	0	0	0	0	0	N	0	1
1011	0	0	0	0	0	N	0	0
1012	0	0	0	0	0	N	0	0
n	5	5	5	5	5	5	5	5

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Recovery

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1008	3	0
1009	4	0
1010	8	0
1011	5	0
1012	2	0
n	5	5
Mean	4	0
S.D.	2	0

Open field observation

Period : Recovery

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4008	0	0	0	0	0	N	0	0
4009	0	0	0	0	0	N	0	0
4010	0	0	0	0	0	N	0	1
4011	0	0	0	0	0	N	0	0
4012	0	0	0	0	0	N	0	0
n	5	5	5	5	5	5	5	5

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Recovery

Generation : F0 Sex : Male Week : 1 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4008	4	0
4009	4	0
4010	3	0
4011	0	0
4012	5	0
n	5	5
Mean	3	0
S.D.	2	0

Open field observation

Period : Recovery

Generation : F0 Sex : Male Week : 2 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
1008	0	0	0	0	0	N	0	0
1009	0	0	0	0	0	N	0	0
1010	0	0	0	0	0	N	0	1
1011	0	0	0	0	0	N	0	0
1012	0	0	0	0	0	N	0	0
n	5	5	5	5	5	5	5	5

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Recovery

Generation : F0 Sex : Male Week : 2 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
1008	5	0
1009	6	0
1010	6	0
1011	7	0
1012	7	0
n	5	5
Mean	6	0
S.D.	1	0

Open field observation

Period : Recovery

Generation : F0 Sex : Male Week : 2 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
4008	0	0	0	0	0	N	0	0
4009	0	0	0	0	0	N	0	0
4010	0	0	0	0	0	N	0	1
4011	0	0	0	0	0	N	0	0
4012	0	0	0	0	0	N	0	0
n	5	5	5	5	5	5	5	5

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Recovery

Generation : F0 Sex : Male Week : 2 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
4008	6	0
4009	3	0
4010	3	0
4011	4	0
4012	2	0
n	5	5
Mean	4	0
S.D.	2	0

Open field observation

Generation : F0

Sex : Female

Week : 1

Period : Recovery (non-mating group)

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
5106	0	0	0	0	0	N	0	0
5107	0	0	0	0	0	N	0	0
5108	0	0	0	0	0	N	0	0
5109	0	0	0	0	0	N	0	0
5110	0	0	0	0	0	N	0	0
n	5	5	5	5	5	5	5	5

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
5106	5	0
5107	8	0
5108	3	0
5109	6	0
5110	7	0
n	5	5
Mean	6	0
S.D.	2	0

Open field observation

Generation : F0

Sex : Female

Week : 1

Period : Recovery (non-mating group)

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
6106	0	0	0	0	0	N	0	0
6107	0	0	0	0	0	N	0	0
6108	0	0	0	0	0	N	0	0
6109	0	0	0	0	0	N	0	0
6110	0	0	0	0	0	N	0	0
n	5	5	5	5	5	5	5	5

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 1

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
6106	5	0
6107	2	0
6108	4	0
6109	4	0
6110	5	0
n	5	5
Mean	4	0
S.D.	1	0

Open field observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
5106	0	0	0	0	0	N	0	0
5107	0	0	0	0	0	N	0	0
5108	0	0	0	0	0	N	0	0
5109	0	0	0	0	0	N	0	0
5110	0	0	0	0	0	N	0	0
n	5	5	5	5	5	5	5	5

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
5106	9	0
5107	5	0
5108	5	0
5109	8	0
5110	7	0
n	5	5
Mean	7	0
S.D.	2	0

Open field observation

Generation : F0

Sex : Female

Week : 2

Period : Recovery (non-mating group)

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Arousal	Convulsion	Abnormal behavior	Stereotypy	Gait	Posture	Grooming	Urination
Animal No.								
6106	0	0	0	0	0	N	0	0
6107	0	0	0	0	0	N	0	0
6108	0	0	0	0	0	N	0	0
6109	0	0	0	0	0	N	0	0
6110	0	0	0	0	0	N	0	0
n	5	5	5	5	5	5	5	5

Arousal) -2: Unconscious/semi-conscious, -1: Reduced awareness, 0: Normal, 1: Increased alertness, 2: Markedly alert

Convulsion) 0: None, 1: Minor, 2: Moderate, 3: Severe

Abnormal behavior) 0: None, 1: Minor, 2: Moderate, 3: Severe

Stereotypy) 0: None, 1: Minor, 2: Moderate, 3: Severe

Gait) U: No/minimal location, 0: Normal, 1: Slight, 2: Moderate, 3: Markedly

Posture) N: Normal, F: Flattened, H: Hunched

Grooming) 0: None, 1: Occasional bouts (up to four), 2: Numerous (more than four)

Urination) 0: None, 1: Small amount, 2: Moderate amount, 3: Large/excessive amount

Open field observation

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	Rearing	Defecation count
Animal No.		
6106	8	0
6107	8	0
6108	7	0
6109	6	0
6110	8	0
n	5	5
Mean	7	0
S.D.	1	0

Manipulative test

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
1008	0	0	0	0	P
1009	0	0	0	0	P
1010	0	0	0	0	P
1011	0	0	0	0	P
1012	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial righting reflex-1	Aerial righting reflex-2	Aerial righting reflex-3	Aerial righting reflex-total
1008	0	0	0	0
1009	0	0	0	0
1010	0	0	0	0
1011	0	0	0	0
1012	0	0	0	0
n	5	5	5	5

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
2008	0	0	0	0	P
2009	0	0	0	0	P
2010	0	0	0	0	P
2011	0	0	0	0	P
2012	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial righting reflex-1	Aerial righting reflex-2	Aerial righting reflex-3	Aerial righting reflex-total
2008	0	0	0	0
2009	0	0	0	0
2010	0	0	0	0
2011	0	0	0	0
2012	0	0	0	0
n	5	5	5	5
	0	0	0	0
	0	0	0	0

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
3008	0	0	0	0	P
3009	0	0	0	0	P
3010	0	0	0	0	P
3011	0	0	0	0	P
3012	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial righting reflex-1	Aerial righting reflex-2	Aerial righting reflex-3	Aerial righting reflex-total
3008	0	0	0	0
3009	0	0	0	0
3010	0	0	0	0
3011	0	0	0	0
3012	0	0	0	0
n	5	5	5	5
	0	0	0	0
	0	0	0	0

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
4008	0	0	0	0	P
4009	0	0	0	0	P
4010	0	0	0	0	P
4011	0	0	0	0	P
4012	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial righting reflex-1	Aerial righting reflex-2	Aerial righting reflex-3	Aerial righting reflex-total
4008	0	0	0	0
4009	0	0	0	0
4010	0	0	0	0
4011	0	0	0	0
4012	0	0	0	0
n	5	5	5	5
	0	0	0	0
	0	0	0	0

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Generation : F0

Sex : Female

Day : 13

Period : F0 lactation

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
1101	0	0	0	0	P
1105	0	0	0	0	P
1107	0	0	0	0	P
1108	0	0	0	0	P
1111	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial right-ing reflex-1	Aerial right-ing reflex-2	Aerial right-ing reflex-3	Aerial right-ing reflex-total
1101	0	0	0	0
1105	0	0	0	0
1107	0	0	0	0
1108	0	0	0	0
1111	0	0	0	0
n	5	5	5	5

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Generation : F0

Sex : Female

Day : 13

Period : F0 lactation

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
2104	0	0	0	0	P
2105	0	0	0	0	P
2107	0	0	0	0	P
2108	0	0	0	0	P
2112	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial right-ing reflex-1	Aerial right-ing reflex-2	Aerial right-ing reflex-3	Aerial right-ing reflex-total
2104	0	0	0	0
2105	0	0	0	0
2107	0	0	0	0
2108	0	0	0	0
2112	0	0	0	0
n	5	5	5	5

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Generation : F0

Sex : Female

Day : 13

Period : F0 lactation

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
3101	0	0	0	0	P
3102	0	0	0	0	P
3104	0	0	0	0	P
3110	0	0	0	0	P
3111	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial right-ing reflex-1	Aerial right-ing reflex-2	Aerial right-ing reflex-3	Aerial right-ing reflex-total
3101	0	0	0	0
3102	0	0	0	0
3104	0	0	0	0
3110	0	0	0	0
3111	0	0	0	0
n	5	5	5	5

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Generation : F0

Sex : Female

Day : 13

Period : F0 lactation

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
4102	0	0	0	0	P
4103	0	0	0	0	P
4107	0	0	0	0	P
4109	0	0	0	0	P
4111	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial right-ing reflex-1	Aerial right-ing reflex-2	Aerial right-ing reflex-3	Aerial right-ing reflex-total
4102	0	0	0	0
4103	0	0	0	0
4107	0	0	0	0
4109	0	0	0	0
4111	0	0	0	0
n	5	5	5	5

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
5106	0	0	0	0	P
5107	0	0	0	0	P
5108	0	0	0	0	P
5109	0	0	0	0	P
5110	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial righting reflex-1	Aerial righting reflex-2	Aerial righting reflex-3	Aerial righting reflex-total
5106	0	0	0	0
5107	0	0	0	0
5108	0	0	0	0
5109	0	0	0	0
5110	0	0	0	0
n	5	5	5	5

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
6106	0	0	0	0	P
6107	0	0	0	0	P
6108	0	0	0	0	P
6109	0	0	0	0	P
6110	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial righting reflex-1	Aerial righting reflex-2	Aerial righting reflex-3	Aerial righting reflex-total
6106	0	0	0	0
6107	0	0	0	0
6108	0	0	0	0
6109	0	0	0	0
6110	0	0	0	0
n	5	5	5	5

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Period : Recovery

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
1008	0	0	0	0	P
1009	0	0	0	0	P
1010	0	0	0	0	P
1011	0	0	0	0	P
1012	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial righting reflex-1	Aerial righting reflex-2	Aerial righting reflex-3	Aerial righting reflex-total
1008	0	0	0	0
1009	0	0	0	0
1010	0	0	0	0
1011	0	0	0	0
1012	0	0	0	0
n	5	5	5	5

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Period : Recovery

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
4008	0	0	0	0	P
4009	0	0	0	0	P
4010	0	0	0	0	P
4011	0	0	0	0	P
4012	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial righting reflex-1	Aerial righting reflex-2	Aerial righting reflex-3	Aerial righting reflex-total
4008	0	0	0	0
4009	0	0	0	0
4010	0	0	0	0
4011	0	0	0	0
4012	0	0	0	0
n	5	5	5	5

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
5106	0	0	0	0	P
5107	0	0	0	0	P
5108	0	0	0	0	P
5109	0	0	0	0	P
5110	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial righting reflex-1	Aerial righting reflex-2	Aerial righting reflex-3	Aerial righting reflex-total
5106	0	0	0	0
5107	0	0	0	0
5108	0	0	0	0
5109	0	0	0	0
5110	0	0	0	0
n	5	5	5	5

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Manipulative test

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Auditory response	Approach response	Touch response	Tail pinch response	Pupillary reflex
6106	0	0	0	0	P
6107	0	0	0	0	P
6108	0	0	0	0	P
6109	0	0	0	0	P
6110	0	0	0	0	P
n	5	5	5	5	5

Animal No.	Aerial righting reflex-1	Aerial righting reflex-2	Aerial righting reflex-3	Aerial righting reflex-total
6106	0	0	0	0
6107	0	0	0	0
6108	0	0	0	0
6109	0	0	0	0
6110	0	0	0	0
n	5	5	5	5
	0	0	0	0
	0	0	0	0

Auditory response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Approach response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Touch response) -1: No reaction/ignores, 0: Normal, 1: Abnormality fearful/aggressive reaction

Tail pinch response) -2: None, -1: Weak, 0: Normal, 1: Exaggerate

Pupillary reflex) P: Pass, both, F: Failed, neither, L: Left pupil responds, R: Right pupil responds

Aerial righting reflex-1 to -3) 0: Normal, 1: Failure, 2:None

Aerial righting reflex-total) Total score of 3 times

Grip strength

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
1008	1200	1119	1160	653	478	566
1009	1578	1661	1620	761	725	743
1010	1524	1507	1516	764	605	685
1011	1508	1596	1552	976	967	972
1012	1234	1258	1246	522	426	474
n	5	5	5	5	5	5
Mean	1409	1428	1419	735	640	688
S.D.	177	231	203	167	216	190

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
2008	1693	1615	1654	738	532	635
2009	1269	1234	1252	843	770	807
2010	1407	1053	1230	694	621	658
2011	1364	1595	1480	820	630	725
2012	1142	1235	1189	718	655	687
n	5	5	5	5	5	5
Mean	1375	1346	1361	763	642	702
S.D.	205	248	199	65	85	67

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
3008	1516	1489	1503	790	727	759
3009	1210	1387	1299	552	512	532
3010	1389	1232	1311	687	599	643
3011	1320	1241	1281	588	520	554
3012	1297	1015	1156	652	557	605
n	5	5	5	5	5	5
Mean	1346	1273	1310	654	583	619
S.D.	114	180	124	93	88	90

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : F0 mating

Generation : F0

Sex : Male

Week : 6

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
4008	1200	1519	1360	792	752	772
4009	1135	1327	1231	620	459	540
4010	1510	1659	1585	763	556	660
4011	1542	1596	1569	839	765	802
4012	1510	1422	1466	669	764	717
n	5	5	5	5	5	5
Mean	1379	1505	1442	737	659	698
S.D.	195	133	149	90	143	104

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : F0 lactation

Generation : F0

Sex : Female

Day : 13

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
1101	1128	1174	1151	502	567	535
1105	1260	1467	1364	567	602	585
1107	849	872	861	403	341	372
1108	1239	1362	1301	626	646	636
1111	823	740	782	367	323	345
n	5	5	5	5	5	5
Mean	1060	1123	1092	493	496	495
S.D.	211	311	260	109	152	130

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : F0 lactation

Generation : F0

Sex : Female

Day : 13

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
2104	880	836	858	367	405	386
2105	1065	880	973	491	498	495
2107	714	809	762	353	344	349
2108	911	1119	1015	355	316	336
2112	851	774	813	427	378	403
n	5	5	5	5	5	5
Mean	884	884	884	399	388	394
S.D.	126	137	107	60	70	63

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : F0 lactation

Generation : F0

Sex : Female

Day : 13

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
3101	1200	1141	1171	858	753	806
3102	837	968	903	452	458	455
3104	1584	1639	1612	785	815	800
3110	1104	1087	1096	462	450	456
3111	1040	990	1015	503	524	514
n	5	5	5	5	5	5
Mean	1153	1165	1159	612	600	606
S.D.	275	274	272	194	172	181

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : F0 lactation

Generation : F0

Sex : Female

Day : 13

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
4102	1408	1371	1390	484	487	486
4103	1220	1124	1172	635	696	666
4107	955	991	973	593	642	618
4109	801	795	798	352	302	327
4111	859	952	906	433	423	428
n	5	5	5	5	5	5
Mean	1049	1047	1048	499	510	505
S.D.	257	216	235	116	161	138

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
5106	1151	1393	1272	481	598	540
5107	1038	1001	1020	422	644	533
5108	1625	1706	1666	871	789	830
5109	886	964	925	720	766	743
5110	1051	991	1021	560	425	493
n	5	5	5	5	5	5
Mean	1150	1211	1181	611	644	628
S.D.	282	329	300	184	147	149

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : Administration (non-mating group)

Generation : F0

Sex : Female

Week : 6

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
6106	1371	1241	1306	757	705	731
6107	1013	999	1006	714	704	709
6108	1069	1205	1137	670	552	611
6109	1580	1359	1470	634	705	670
6110	1173	1414	1294	739	753	746
n	5	5	5	5	5	5
Mean	1241	1244	1243	703	684	693
S.D.	233	161	177	50	77	54

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : Recovery

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
1008	1900	1869	1885	1050	906	978
1009	2119	1898	2009	1156	970	1063
1010	1558	1570	1564	1108	965	1037
1011	1923	1750	1837	959	1043	1001
1012	1779	1655	1717	555	670	613
n	5	5	5	5	5	5
Mean	1856	1748	1802	966	911	938
S.D.	206	139	169	241	143	185

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : Recovery

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
4008	1792	1696	1744	802	789	796
4009	1302	1303	1303	546	705	626
4010	1648	1652	1650	531	568	550
4011	1896	1698	1797	930	833	882
4012	1652	1688	1670	1037	1091	1064
n	5	5	5	5	5	5
Mean	1658	1607	1633	769	797	784
S.D.	224	171	194	226	193	205

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
5106	1302	1414	1358	528	559	544
5107	1184	1090	1137	846	803	825
5108	1266	1474	1370	824	704	764
5109	1502	1530	1516	863	763	813
5110	1147	1036	1092	774	593	684
n	5	5	5	5	5	5
Mean	1280	1309	1295	767	684	726
S.D.	139	229	176	138	106	116

Grip strength of fore/hind limb-mean) mean value of 2 times

Grip strength

Period : Recovery (non-mating group)

Generation : F0

Sex : Female

Week : 2

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Grip strength of fore limb-1 g	Grip strength of fore limb-2 g	Grip strength of fore limb-mean g	Grip strength of hind limb-1 g	Grip strength of hind limb-2 g	Grip strength of hind limb-mean g
6106	1678	1434	1556	662	652	657
6107	1144	1103	1124	542	466	504
6108	1327	1528	1428	693	717	705
6109	1587	1569	1578	606	680	643
6110	1495	1439	1467	592	587	590
n	5	5	5	5	5	5
Mean	1446	1415	1431	619	620	620
S.D.	213	184	182	59	99	77

Grip strength of fore/hind limb-mean) mean value of 2 times

Motor activity		Period : F0 mating		Dose : Dodecyloxirane 0 mg/kg		Species : Rat	
Generation : F0		Sex : Male	Week : 6				
	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
1008	466	390	206	175	0	54	1291
1009	358	141	69	30	9	37	644
1010	403	395	186	156	51	6	1197
1011	446	374	357	233	96	8	1514
1012	385	336	245	220	181	94	1461
n	5	5	5	5	5	5	5
Mean	412	327	213	163	67	40	1221
S.D.	44	107	104	81	74	36	347

Motor activity		Period : F0 mating		Dose : Dodecyloxirane 100 mg/kg		Species : Rat	
Generation : F0		Sex : Male	Week : 6				
	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
2008	434	359	353	311	211	13	1681
2009	427	345	74	64	0	5	915
2010	435	384	252	192	276	265	1804
2011	378	325	296	24	275	278	1576
2012	449	291	354	214	339	118	1765
n	5	5	5	5	5	5	5
Mean	425	341	266	161	220	136	1548
S.D.	27	35	115	117	131	132	365

Motor activity		Period : F0 mating		Dose : Dodecyloxirane 300 mg/kg		Species : Rat	
Generation : F0		Sex : Male	Week : 6				
	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
3008	393	222	138	20	3	4	780
3009	225	232	134	9	0	3	603
3010	480	435	120	20	4	119	1178
3011	441	329	177	97	12	87	1143
3012	431	371	395	263	198	85	1743
n	5	5	5	5	5	5	5
Mean	394	318	193	82	43	60	1089
S.D.	99	91	115	107	87	53	439

Motor activity		Period : F0 mating		Dose : Dodecyloxirane 1000 mg/kg		Species : Rat	
Generation : F0		Sex : Male	Week : 6				
	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
4008	441	311	72	200	0	76	1100
4009	497	137	10	54	338	175	1211
4010	461	158	65	60	93	17	854
4011	455	450	221	155	6	18	1305
4012	304	368	161	37	2	45	917
n	5	5	5	5	5	5	5
Mean	432	285	106	101	88	66	1077
S.D.	74	135	84	72	145	65	191

Motor activity

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
1101	438	295	229	181	90	296	1529
1105	355	362	339	318	182	231	1787
1107	442	220	370	354	337	262	1985
1108	387	156	196	79	254	54	1126
1111	403	275	186	254	249	276	1643
n	5	5	5	5	5	5	5
Mean	405	262	264	237	222	224	1614
S.D.	36	78	85	110	92	98	322

Motor activity		Period : F0 lactation		Dose : Dodecyloxirane 100 mg/kg		Species : Rat	
Generation : F0		Sex : Female		Day : 13		Total(0'-60')	
	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	
Animal No.	Count	Count	Count	Count	Count	Count	Count
2104	388	163	34	172	82	120	959
2105	454	56	260	131	3	16	920
2107	481	297	53	265	398	233	1727
2108	423	128	45	236	193	35	1060
2112	355	79	24	241	123	243	1065
n	5	5	5	5	5	5	5
Mean	420	145	83	209	160	129	1146
S.D.	50	95	99	56	150	107	331

Motor activity

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
3101	440	225	346	179	406	243	1839
3102	337	176	188	12	4	2	719
3104	321	69	178	9	145	23	745
3110	384	122	36	5	28	7	582
3111	398	252	249	281	223	235	1638
n	5	5	5	5	5	5	5
Mean	376	169	199	97	161	102	1105
S.D.	48	75	113	127	163	125	586

Motor activity

Period : F0 lactation

Generation : F0 Sex : Female Day : 13 Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
4102	382	204	99	1	0	12	698
4103	262	21	175	88	21	35	602
4107	448	332	128	0	0	0	908
4109	378	367	320	304	221	365	1955
4111	315	20	156	37	4	15	547
n	5	5	5	5	5	5	5
Mean	357	189	176	86	49	85	942
S.D.	71	165	86	127	96	157	583

Motor activity		Period : Administration (non-mating group)		Species : Rat			
Generation : F0		Sex : Female		Week : 6		Dose : Dodecyloxirane 0 mg/kg	
	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
5106	423	326	335	410	422	360	2276
5107	346	408	293	286	80	293	1706
5108	240	187	167	112	22	23	751
5109	158	206	152	21	19	27	583
5110	388	179	92	6	17	2	684
n	5	5	5	5	5	5	5
Mean	311	261	208	167	112	141	1200
S.D.	110	101	102	176	175	171	752

with the reproduction/ developmental toxicity screening test in rats							
Motor activity		Period : Administration (non-mating group)					
Generation : F0	Sex : Female	Week : 6	Dose : Dodecyloxirane 1000 mg/kg			Species : Rat	
0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')	
Animal No.	Count	Count	Count	Count	Count	Count	Count
6106	342	224	250	87	116	212	1231
6107	359	235	159	6	122	308	1189
6108	332	308	182	223	95	306	1446
6109	316	310	213	0	147	294	1280
6110	286	87	163	0	73	202	811
n	5	5	5	5	5	5	5
Mean	327	233	193	63	111	264	1191
S.D.	28	91	38	97	28	53	234

Motor activity

Period : Recovery

Generation : F0

Sex : Male

Week : 2

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
1008	423	318	328	5	11	249	1334
1009	283	180	9	2	7	0	481
1010	325	243	77	32	2	0	679
1011	378	199	112	1	2	11	703
1012	445	347	217	165	37	32	1243
n	5	5	5	5	5	5	5
Mean	371	257	149	41	12	58	888
S.D.	67	73	125	70	15	107	377

Motor activity		Period : Recovery		Dose : Dodecyloxirane 1000 mg/kg		Species : Rat	
Generation : F0		Sex : Male	Week : 2				
	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
4008	368	263	202	25	67	25	950
4009	483	292	258	71	47	69	1220
4010	414	343	81	88	10	10	946
4011	444	377	359	202	222	83	1687
4012	342	285	215	232	75	77	1226
n	5	5	5	5	5	5	5
Mean	410	312	223	124	84	53	1206
S.D.	57	47	100	89	81	33	302

with the reproduction/ developmental toxicity screening test in rats							
Motor activity		Period : Recovery (non-mating group)					
Generation : F0	Sex : Female	Week : 2	Dose : Dodecyloxirane 0 mg/kg			Species : Rat	
0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')	
Animal No.	Count	Count	Count	Count	Count	Count	
5106	291	311	263	154	269	295	1583
5107	362	332	255	44	8	15	1016
5108	303	219	72	2	6	11	613
5109	267	16	87	29	0	0	399
5110	372	166	244	13	0	144	939
n	5	5	5	5	5	5	5
Mean	319	209	184	48	57	93	910
S.D.	46	127	96	61	119	127	451

Motor activity		Period : Recovery (non-mating group)		Species : Rat			
Generation : F0		Sex : Female		Week : 2		Dose : Dodecyloxirane 1000 mg/kg	
	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	Total(0'-60')
Animal No.	Count	Count	Count	Count	Count	Count	Count
6106	371	318	286	221	291	51	1538
6107	248	21	6	285	44	24	628
6108	367	518	39	159	115	1	1199
6109	445	329	501	267	267	258	2067
6110	279	279	229	43	4	1	835
n	5	5	5	5	5	5	5
Mean	342	293	212	195	144	67	1253
S.D.	79	178	201	98	130	109	573

Body weight

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day							Body weight
	1	8	15	22	29	36	42	gain
1001	382	432	468	480	513	531	538	156
1002	376	393	404	429	457	462	474	98
1003	400	436	471	484	514	528	535	135
1004	351	384	413	430	457	477	493	142
1005	383	406	421	443	469	491	511	128
1006	402	441	479	494	530	562	586	184
1007	354	404	432	455	480	502	509	155
1008	373	411	436	453	484	509	503	130
1009	424	447	468	483	511	538	545	121
1010	408	446	472	498	523	549	554	146
1011	398	425	464	487	532	561	576	178
1012	358	401	442	462	492	513	537	179
n	12	12	12	12	12	12	12	12
Mean	384	419	448	467	497	519	530	146
S.D.	23	22	26	24	27	32	33	26

Body weight

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Sex : Male

Dose : Dodecyloxirane 100 mg/kg

Unit : g

Species : Rat

Animal No.	/Day							Body weight gain
	1	8	15	22	29	36	42	
2001	398	440	477	505	532	561	577	179
2002	404	450	492	511	548	575	597	193
2003	393	423	453	470	500	513	517	124
2004	415	461	501	531	560	588	582	167
2005	381	410	434	454	479	501	511	130
2006	355	386	408	431	451	469	489	134
2007	366	401	423	440	470	482	497	131
2008	385	426	458	479	500	521	530	145
2009	375	402	432	454	474	497	508	133
2010	357	381	399	402	423	437	448	91
2011	378	425	462	480	512	516	560	182
2012	387	431	469	494	521	542	554	167
n	12	12	12	12	12	12	12	12
Mean	383	420	451	471	498	517	531	148
S.D.	18	25	32	37	40	44	44	30

Body weight

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Sex : Male

Dose : Dodecyloxirane 300 mg/kg

Unit : g

Species : Rat

Animal No.	/Day							Body weight gain
	1	8	15	22	29	36	42	
3001	381	430	476	506	550	588	609	228
3002	359	387	403	426	444	459	466	107
3003	388	423	454	460	479	486	499	111
3004	398	431	451	467	493	516	537	139
3005	396	433	469	488	509	536	555	159
3006	352	383	410	434	458	474	489	137
3007	367	392	414	433	453	468	476	109
3008	414	462	496	501	525	558	567	153
3009	377	410	440	454	479	501	507	130
3010	367	402	432	444	468	476	491	124
3011	408	444	477	502	527	554	565	157
3012	379	421	454	477	499	518	530	151
n	12	12	12	12	12	12	12	12
Mean	382	418	448	466	490	511	524	142
S.D.	19	24	29	29	33	41	44	33

Body weight

Period : F0 before mating Day 1-15, F0 mating Day 15-43

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day							Body weight
	1	8	15	22	29	36	42	gain
4001	379	404	429	448	474	496	509	130
4002	360	376	402	426	456	483	488	128
4003	383	405	424	433	451	454	470	87
4004	371	410	433	449	472	490	506	135
4005	352	382	410	425	457	476	494	142
4006	410	438	463	489	519	531	546	136
4007	361	396	418	438	469	489	509	148
4008	397	441	470	486	517	540	557	160
4009	397	440	482	488	517	531	556	159
4010	358	393	413	419	435	450	446	88
4011	394	422	448	466	500	520	534	140
4012	414	435	453	476	497	516	536	122
n	12	12	12	12	12	12	12	12
Mean	381	412	437	454	480	498	513	131
S.D.	21	23	26	26	29	30	35	23

Body weight

Period : F0 before mating Day 1-15

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Body weight gain
	1	8	15	
1101	247	267	273	26
1102	236	239	260	24
1103	259	276	286	27
1104	270	289	291	21
1105	248	255	262	14
1106	266	275	286	20
1107	246	247	261	15
1108	263	277	287	24
1109	246	250	250	4
1110	235	242	243	8
1111	211	228	236	25
1112	269	273	280	11
n	12	12	12	12
Mean	250	260	268	18
S.D.	17	19	19	8

Appendix 3 - 6

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Body weight
Sex : Female
Period : F0 before mating Day 1-15
Dose : Dodecyloxirane 100 mg/kg

Study No. : R-1261

Animal No.	/Day			Body weight gain	Unit : g	Species : Rat
	1	8	15			
2101	253	269	289	36		
2102	242	248	253	11		
2103	256	263	270	14		
2104	238	258	267	29		
2105	239	248	266	27		
2106	268	277	281	13		
2107	248	262	267	19		
2108	242	251	269	27		
2109	274	288	285	11		
2110	268	283	286	18		
2111	229	236	241	12		
2112	263	290	308	45		
n	12	12	12	12		
Mean	252	264	274	22		
S.D.	14	17	18	11		

Appendix 3 - 7

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Body weight
Sex : Female
Period : F0 before mating Day 1-15
Dose : Dodecyloxirane 300 mg/kg

Study No. : R-1261

Animal No.	/Day			Body weight gain	Unit : g	Species : Rat
	1	8	15			
3101	260	273	290	30		
3102	240	249	268	28		
3103	253	261	271	18		
3104	254	264	276	22		
3105	273	277	281	8		
3106	248	252	254	6		
3107	237	243	249	12		
3108	252	262	259	7		
3109	238	241	245	7		
3110	274	284	303	29		
3111	245	247	270	25		
3112	229	251	271	42		
n	12	12	12	12		
Mean	250	259	270	20		
S.D.	14	14	17	12		

Appendix 3 - 8

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Body weight

Period : F0 before mating Day 1-15

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Body weight gain
	1	8	15	
4101	242	267	274	32
4102	265	278	286	21
4103	243	255	272	29
4104	232	239	253	21
4105	232	239	243	11
4106	259	271	297	38
4107	238	253	271	33
4108	261	283	294	33
4109	269	283	294	25
4110	270	286	283	13
4111	226	230	242	16
4112	280	287	293	13
n	12	12	12	12
Mean	251	264	275	24
S.D.	18	20	20	9

Body weight

Period : F0 gestation Day 0-25

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Body weight gain
	0	7	14	20	
1101	278	313	359	413	135
1102	300	323	350	380	80
1103	297	334	367	467	170
1104	307	354	397	492	185
1105	267	322	372	473	206
1106	298	338	379	439	141
1107	267	300	342	412	145
1108	290	320	354	438	148
1109	272	300	335	406	134
1110	251	276	304	379	128
1111	244	292	320	416	172
1112	304	356	393	492	188
n	12	12	12	12	12
Mean	281	319	356	434	153
S.D.	21	24	28	40	34

Appendix 3 - 10

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Body weight

Period : F0 gestation Day 0-25

Sex : Female

Dose : Dodecyloxirane 100 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Body weight gain
	0	7	14	20	
2101	285	322	353	441	156
2102	257	274	320	408	151
2103	290	332	364	465	175
2104	275	308	344	440	165
2105	278	298	328	406	128
2107	274	307	342	427	153
2108	279	325	366	451	172
2110	317	367	407	511	194
2111	267	300	341	434	167
2112	309	335	373	455	146
n	10	10	10	10	10
Mean	283	317	354	444	161
S.D.	18	25	25	30	18

Appendix 3 - 11

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Body weight

Period : F0 gestation Day 0-25

Sex : Female

Dose : Dodecyloxirane 300 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Body weight gain
	0	7	14	20	
3101	292	332	374	458	166
3102	271	311	336	420	149
3103	280	321	351	418	138
3104	283	324	370	465	182
3105	297	351	400	506	209
3106	276	323	370	453	177
3107	327	354	392	488	161
3110	313	368	423	511	198
3111	277	331	378	491	214
3112	270	310	352	429	159
n	10	10	10	10	10
Mean	289	333	375	464	175
S.D.	19	19	26	35	25

Body weight

Period : F0 gestation Day 0-25

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Body weight gain
	0	7	14	20	
4102	287	321	368	453	166
4103	275	310	358	437	162
4104	268	297	328	398	130
4105	254	287	322	404	150
4106	300	340	368	344	44
4107	278	313	349	456	178
4108	303	336	371	425	122
4109	305	346	381	485	180
4110	295	332	368	446	151
4111	244	284	326	402	158
4112	353	375	403	484	131
n	11	11	11	11	11
Mean	287	322	358	430	143
S.D.	29	27	25	42	38

Body weight

Period : F0 lactation Day 0-14

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Body weight gain
	0	4	7	13	
1101	353	350	351	355	2
1102	356	351	354	372	16
1103	367	355	361	368	1
1104	377	387	392	396	19
1105	361	384	365	394	33
1106	378	373	377	393	15
1107	333	338	338	364	31
1108	341	352	349	360	19
1109	321	333	325	350	29
1110	306	300	299	318	12
1111	309	336	327	337	28
1112	397	376	379	403	6
n	12	12	12	12	12
Mean	350	353	351	368	18
S.D.	29	25	26	26	11

Appendix 3 - 14

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Body weight

Period : F0 lactation Day 0-14

Sex : Female

Dose : Dodecyloxirane 100 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Body weight gain
	0	4	7	13	
2101	341	355	362	357	16
2102	318	338	340	369	51
2103	348	365	355	384	36
2104	364	350	349	365	1
2105	340	342	342	354	14
2107	344	323	325	340	-4
2108	364	373	369	381	17
2110	405	402	402	415	10
2111	332	340	341	359	27
2112	373	373	364	386	13
n	10	10	10	10	10
Mean	353	356	355	371	18
S.D.	25	23	21	21	16

Body weight

Period : F0 lactation Day 0-14

Sex : Female

Dose : Dodecyloxirane 300 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Body weight gain
	0	4	7	13	
3101	359	381	371	388	29
3102	334	345	346	355	21
3103	364	355	355	361	-3
3104	374	357	353	380	6
3105	407	407	413	420	13
3106	354	375	386	401	47
3107	399	412	402	398	-1
3110	419	439	439	446	27
3111	386	365	357	374	-12
3112	360	344	345	358	-2
n	10	10	10	10	10
Mean	376	378	377	388	13
S.D.	27	32	32	29	18

Body weight

Period : F0 lactation Day 0-14

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Body weight gain
	0	4	7	13	
4102	370	361	355	366	-4
4103	325	350	339	357	32
4104	333	330	338	354	21
4105	302	317	306	324	22
4107	337	350	346	361	24
4108	373	376	377	382	9
4109	384	388	388	388	4
4110	335	360	360	375	40
4111	323	332	333	338	15
4112	392	371	380	400	8
n	10	10	10	10	10
Mean	347	354	352	365	17
S.D.	30	22	25	23	13

Body weight

Period : Administration (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day							Body weight
	1	8	15	22	29	36	42	gain
5101	264	278	290	307	312	323	325	61
5102	265	281	286	313	334	334	341	76
5103	250	255	258	258	269	280	279	29
5104	251	263	276	288	298	299	308	57
5105	239	249	264	271	278	277	288	49
5106	249	270	275	273	294	293	299	50
5107	273	279	282	304	325	334	335	62
5108	234	250	264	284	298	303	304	70
5109	240	250	260	275	284	293	297	57
5110	230	253	263	267	278	291	295	65
n	10	10	10	10	10	10	10	10
Mean	250	263	272	284	297	303	307	58
S.D.	14	13	12	19	21	21	20	13

Body weight

Period : Administration (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day							Body weight
	1	8	15	22	29	36	42	gain
6101	267	286	301	312	327	348	340	73
6102	243	258	261	277	281	292	297	54
6103	264	280	301	310	312	328	332	68
6104	260	263	277	289	301	307	316	56
6105	224	237	244	253	266	261	276	52
6106	272	287	293	304	316	313	320	48
6107	238	245	253	264	276	275	272	34
6108	252	274	267	279	297	304	307	55
6109	252	263	276	278	305	310	309	57
6110	260	279	280	284	303	313	308	48
n	10	10	10	10	10	10	10	10
Mean	253	267	275	285	298	305	308	55
S.D.	15	17	19	19	19	25	22	11

Body weight

Period : Recovery Day 1-15

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Body weight gain
	1	8	14	
1008	509	515	528	19
1009	544	562	577	33
1010	559	571	585	26
1011	577	597	595	18
1012	540	556	577	37
n	5	5	5	5
Mean	546	560	572	27
S.D.	25	30	26	8

Body weight

Period : Recovery Day 1-15

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Body weight gain
	1	8	14	
4008	556	579	591	35
4009	551	576	598	47
4010	449	466	479	30
4011	532	545	566	34
4012	536	558	562	26
n	5	5	5	5
Mean	525	545	559	34
S.D.	44	46	47	8

Body weight

Period : Recovery Day 1-15 (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Body weight gain
	1	8	14	
5106	299	314	306	7
5107	339	347	349	10
5108	298	314	316	18
5109	300	299	301	1
5110	298	304	310	12
n	5	5	5	5
Mean	307	316	316	10
S.D.	18	19	19	6

Body weight

Period : Recovery Day 1-15 (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Body weight gain
	1	8	14	
6106	311	325	325	14
6107	270	281	287	17
6108	297	310	314	17
6109	317	308	322	5
6110	313	307	314	1
n	5	5	5	5
Mean	302	306	312	11
S.D.	19	16	15	7

Appendix 4 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption
Sex : Male
Period : F0 before mating Day 1-15
Dose : Dodecyloxirane 0 mg/kg

Study No. : R-1261

Animal No.	/Day			Total food consumption	Unit : g	Species : Rat
	2	8	15			
1001	24	27	27	78		
1002	23	20	16	59		
1003	22	21	21	64		
1004	17	21	21	59		
1005	19	17	17	53		
1006	21	24	24	69		
1007	19	23	19	61		
1008	19	24	23	66		
1009	18	23	21	62		
1010	24	20	22	66		
1011	19	22	25	66		
1012	18	22	28	68		
n	12	12	12	12		
Mean	20	22	22	64		
S.D.	2	3	4	6		

Appendix 4 - 2

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : F0 before mating Day 1-15

Sex : Male

Dose : Dodecyloxirane 100 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Total food consumption
	2	8	15	
2001	20	19	23	62
2002	24	23	23	70
2003	20	21	20	61
2004	21	24	26	71
2005	20	20	21	61
2006	18	19	17	54
2007	20	22	18	60
2008	20	23	23	66
2009	21	18	21	60
2010	16	16	16	48
2011	23	26	25	74
2012	24	28	28	80
n	12	12	12	12
Mean	21	22	22	64
S.D.	2	3	4	9

Appendix 4 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption
Sex : Male
Period : F0 before mating Day 1-15
Dose : Dodecyloxirane 300 mg/kg

Study No. : R-1261

Animal No.	/Day			Total food consumption	Unit : g	Species : Rat
	2	8	15			
3001	24	27	28	79		
3002	19	15	15	49		
3003	20	22	20	62		
3004	20	24	18	62		
3005	20	20	19	59		
3006	19	19	20	58		
3007	22	20	20	62		
3008	22	24	25	71		
3009	21	22	20	63		
3010	21	20	20	61		
3011	25	25	25	75		
3012	19	21	21	61		
n	12	12	12	12		
Mean	21	22	21	64		
S.D.	2	3	4	8		

Appendix 4 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : F0 before mating Day 1-15

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Total food consumption
	2	8	15	
4001	19	19	20	58
4002	19	18	22	59
4003	22	19	18	59
4004	16	20	22	58
4005	18	19	22	59
4006	22	20	21	63
4007	21	22	21	64
4008	21	24	26	71
4009	21	23	25	69
4010	25	19	21	65
4011	23	23	27	73
4012	20	23	22	65
n	12	12	12	12
Mean	21	21	22	64
S.D.	2	2	3	5

Food consumption

Period : F0 mating Day 15-43

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day		
	30	36	42
1001	21	21	19
1002	20	21	23
1003	20	21	19
1004	17	22	22
1005	21	22	22
1006	24	25	25
1007	20	20	19
1008	22	25	18
1009	18	21	19
1010	17	24	22
1011	22	28	25
1012	21	22	23
n	12	12	12
Mean	20	23	21
S.D.	2	2	2

Appendix 4 - 6

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : F0 mating Day 15-43

Sex : Male

Dose : Dodecyloxirane 100 mg/kg

Unit : g

Species : Rat

Animal No.	/Day		
	30	36	42
2001	22	26	23
2002	24	26	25
2003	21	19	16
2004	24	23	22
2005	20	20	16
2006	20	23	24
2007	19	19	19
2008	19	22	19
2009	16	23	21
2010	13	15	15
2011	24	23	24
2012	24	26	21
n	12	12	12
Mean	21	22	20
S.D.	3	3	3

Appendix 4 - 7

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : F0 mating Day 15-43

Sex : Male

Dose : Dodecyloxirane 300 mg/kg

Unit : g

Species : Rat

Animal No.	/Day		
	30	36	42
3001	24	29	27
3002	15	17	17
3003	16	16	19
3004	22	23	18
3005	20	22	23
3006	20	22	22
3007	19	20	21
3008	20	26	24
3009	21	19	16
3010	17	21	18
3011	21	26	23
3012	19	20	20
n	12	12	12
Mean	20	22	21
S.D.	3	4	3

Food consumption

Period : F0 mating Day 15-43

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day		
	30	36	42
4001	22	20	22
4002	21	26	22
4003	16	18	18
4004	24	24	22
4005	23	20	22
4006	21	22	22
4007	22	23	21
4008	22	21	20
4009	18	20	24
4010	18	21	16
4011	24	26	24
4012	21	21	23
n	12	12	12
Mean	21	22	21
S.D.	2	2	2

Appendix 4 - 9

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption
Sex : Female
Period : F0 before mating Day 1-15
Dose : Dodecyloxirane 0 mg/kg

Study No. : R-1261

Animal No.	/Day			Total food consumption	Unit : g	Species : Rat
	2	8	15			
1101	12	14	12	38		
1102	8	13	13	34		
1103	13	16	18	47		
1104	15	19	11	45		
1105	11	16	11	38		
1106	10	16	16	42		
1107	14	10	16	40		
1108	16	14	16	46		
1109	12	15	12	39		
1110	10	11	10	31		
1111	10	13	14	37		
1112	16	14	14	44		
n	12	12	12	12		
Mean	12	14	14	40		
S.D.	3	2	3	5		

Appendix 4 - 10

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption
Sex : Female
Period : F0 before mating Day 1-15
Dose : Dodecyloxirane 100 mg/kg

Study No. : R-1261

Animal No.	/Day			Total food consumption	Unit : g	Species : Rat
	2	8	15			
2101	10	13	16	39		
2102	9	13	13	35		
2103	15	13	12	40		
2104	13	16	16	45		
2105	19	11	16	46		
2106	13	19	13	45		
2107	10	14	16	40		
2108	14	10	19	43		
2109	17	18	11	46		
2110	16	18	15	49		
2111	13	13	13	39		
2112	17	17	18	52		
n	12	12	12	12		
Mean	14	15	15	43		
S.D.	3	3	2	5		

Appendix 4 - 11

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Food consumption
Sex : Female
Period : F0 before mating Day 1-15
Dose : Dodecyloxirane 300 mg/kg

Study No. : R-1261

Animal No.	/Day			Total food consumption	Unit : g	Species : Rat
	2	8	15			
3101	14	15	15	44		
3102	14	10	17	41		
3103	10	13	14	37		
3104	13	15	16	44		
3105	12	15	13	40		
3106	13	13	12	38		
3107	12	12	11	35		
3108	14	15	12	41		
3109	13	13	12	38		
3110	19	16	19	54		
3111	18	12	19	49		
3112	17	16	21	54		
n	12	12	12	12		
Mean	14	14	15	43		
S.D.	3	2	3	6		

Appendix 4 - 12

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : F0 before mating Day 1-15

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Total food consumption
	2	8	15	
4101	17	17	17	51
4102	13	14	14	41
4103	14	13	16	43
4104	13	12	15	40
4105	11	13	10	34
4106	16	14	17	47
4107	14	15	16	45
4108	11	20	20	51
4109	18	21	17	56
4110	10	20	14	44
4111	11	12	13	36
4112	18	17	13	48
n	12	12	12	12
Mean	14	16	15	45
S.D.	3	3	3	6

Food consumption

Period : F0 gestation Day 0-25

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Total food consumption
	1	7	14	20	
1101	16	18	20	22	76
1102	14	14	17	16	61
1103	19	23	23	26	91
1104	18	25	25	23	91
1105	15	22	22	27	86
1106	19	20	25	28	92
1107	18	18	19	22	77
1108	15	14	16	25	70
1109	14	16	19	19	68
1110	7	16	16	18	57
1111	15	24	16	21	76
1112	17	25	22	21	85
n	12	12	12	12	12
Mean	16	20	20	22	78
S.D.	3	4	3	4	12

Appendix 4 - 14

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : F0 gestation Day 0-25

Sex : Female

Dose : Dodecyloxirane 100 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Total food consumption
	1	7	14	20	
2101	13	21	21	24	79
2102	0	16	24	21	61
2103	12	23	25	24	84
2104	17	24	23	25	89
2105	17	15	17	23	72
2107	12	18	24	22	76
2108	24	22	19	25	90
2110	20	25	26	32	103
2111	20	19	19	25	83
2112	14	20	22	26	82
n	10	10	10	10	10
Mean	15	20	22	25	82
S.D.	7	3	3	3	11

Appendix 4 - 15

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : F0 gestation Day 0-25

Sex : Female

Dose : Dodecyloxirane 300 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Total food consumption
	1	7	14	20	
3101	16	19	20	23	78
3102	19	17	20	22	78
3103	18	20	26	25	89
3104	18	21	22	24	85
3105	13	27	33	34	107
3106	18	20	22	24	84
3107	14	23	23	24	84
3110	19	27	29	27	102
3111	24	22	22	35	103
3112	16	21	22	22	81
n	10	10	10	10	10
Mean	18	22	24	26	89
S.D.	3	3	4	5	11

Food consumption

Period : F0 gestation Day 0-25

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Total food consumption
	1	7	14	20	
4102	15	18	22	25	80
4103	19	17	23	21	80
4104	14	18	18	21	71
4105	15	21	23	20	79
4106	17	22	22	14	75
4107	19	21	21	27	88
4108	17	18	22	21	78
4109	20	24	21	23	88
4110	17	21	25	23	86
4111	19	17	20	25	81
4112	19	24	21	24	88
n	11	11	11	11	11
Mean	17	20	22	22	81
S.D.	2	3	2	3	6

Food consumption

Period : F0 lactation Day 0-14

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Total food consumption
	2	4	7	13	
1101	19	35	44	59	157
1102	17	25	26	43	111
1103	20	40	42	57	159
1104	28	40	39	61	168
1105	30	50	35	63	178
1106	22	32	39	56	149
1107	23	40	38	54	155
1108	30	40	38	61	169
1109	20	37	41	58	156
1110	20	32	38	55	145
1111	29	38	41	59	167
1112	22	33	37	63	155
n	12	12	12	12	12
Mean	23	37	38	57	156
S.D.	5	6	5	5	17

Food consumption

Period : F0 lactation Day 0-14

Sex : Female

Dose : Dodecyloxirane 100 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Total food consumption
	2	4	7	13	
2101	29	44	40	54	167
2102	32	43	46	65	186
2103	26	41	40	59	166
2104	25	32	41	61	159
2105	17	34	37	58	146
2107	17	30	40	47	134
2108	24	39	33	58	154
2110	28	43	41	71	183
2111	32	47	46	55	180
2112	24	35	37	49	145
n	10	10	10	10	10
Mean	25	39	40	58	162
S.D.	5	6	4	7	18

Food consumption

Period : F0 lactation Day 0-14

Sex : Female

Dose : Dodecyloxirane 300 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Total food consumption
	2	4	7	13	
3101	25	39	40	59	163
3102	27	40	35	57	159
3103	22	33	41	55	151
3104	20	36	34	57	147
3105	30	46	48	66	190
3106	26	45	44	70	185
3107	32	38	41	56	167
3110	30	41	43	57	171
3111	27	40	33	60	160
3112	23	32	38	52	145
n	10	10	10	10	10
Mean	26	39	40	59	164
S.D.	4	5	5	5	15

Appendix 4 - 20

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : F0 lactation Day 0-14

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day				Total food consumption
	2	4	7	13	
4102	18	32	34	50	134
4103	27	42	41	56	166
4104	24	36	43	55	158
4105	25	40	35	39	139
4107	26	36	37	52	151
4108	22	32	36	53	143
4109	30	49	56	60	195
4110	28	41	42	59	170
4111	25	35	34	51	145
4112	24	31	46	54	155
n	10	10	10	10	10
Mean	25	37	40	53	156
S.D.	3	6	7	6	18

Food consumption

Period : Administration Day 1-15(non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Total food consumption
	2	8	15	
5101	15	10	16	41
5102	12	14	14	40
5103	9	15	14	38
5104	12	13	15	40
5105	13	11	15	39
5106	13	19	15	47
5107	12	15	12	39
5108	11	15	16	42
5109	10	17	18	45
5110	14	15	18	47
n	10	10	10	10
Mean	12	14	15	42
S.D.	2	3	2	3

Food consumption

Period : Administration Day 1-15(non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Total food consumption
	2	8	15	
6101	12	16	14	42
6102	14	14	15	43
6103	14	16	18	48
6104	11	14	14	39
6105	10	16	15	41
6106	15	16	16	47
6107	11	15	12	38
6108	16	19	12	47
6109	13	17	18	48
6110	12	19	16	47
n	10	10	10	10
Mean	13	16	15	44
S.D.	2	2	2	4

Food consumption

Period : Administration Day 15-43(non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day		
	30	36	42
5101	16	16	13
5102	16	16	17
5103	10	17	13
5104	15	13	15
5105	15	14	14
5106	15	17	17
5107	13	17	13
5108	15	14	15
5109	12	18	18
5110	15	17	17
n	10	10	10
Mean	14	16	15
S.D.	2	2	2

Appendix 4 - 24

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : Administration Day 15-43(non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day		
	30	36	42
6101	18	20	13
6102	15	16	16
6103	16	18	15
6104	17	15	16
6105	15	14	17
6106	15	15	18
6107	12	16	14
6108	15	18	15
6109	13	17	15
6110	17	20	11
n	10	10	10
Mean	15	17	15
S.D.	2	2	2

Appendix 4 - 25

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : Recovery Day 1-15

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Total food consumption
	2	8	14	
1008	22	22	23	67
1009	23	24	24	71
1010	20	26	23	69
1011	22	31	28	81
1012	22	28	26	76
n	5	5	5	5
Mean	22	26	25	73
S.D.	1	3	2	6

Appendix 4 - 26

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

Food consumption

Period : Recovery Day 1-15

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Total food consumption
	2	8	14	
4008	24	32	26	82
4009	24	27	29	80
4010	18	26	23	67
4011	26	27	28	81
4012	27	29	26	82
n	5	5	5	5
Mean	24	28	26	78
S.D.	3	2	2	6

Food consumption

Period : Recovery Day 1-15 (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Total food consumption
	2	8	14	
5106	12	21	17	50
5107	16	20	16	52
5108	16	18	16	50
5109	14	18	19	51
5110	22	20	21	63
n	5	5	5	5
Mean	16	19	18	53
S.D.	4	1	2	6

Food consumption

Period : Recovery Day 1-15 (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Unit : g

Species : Rat

Animal No.	/Day			Total food consumption
	2	8	14	
6106	16	22	20	58
6107	13	18	19	50
6108	16	21	16	53
6109	15	17	21	53
6110	17	16	18	51
n	5	5	5	5
Mean	15	19	19	53
S.D.	2	3	2	3

Urinalysis

Stage : Week 6 of administration

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	pH	Protein	Ketones	Glucose	Occult Blood	Bilirubin	Urobilinogen	Color
Animal No.								
1008	8.5	-	1+	-	+/-	-	+/-	Y
1009	>=9.0	1+	1+	-	-	-	+/-	Y
1010	8.5	2+	2+	-	-	1+	+/-	Y
1011	7.0	1+	1+	-	-	-	+/-	Y
1012	8.5	1+	1+	-	-	-	+/-	Y
n	5	5	5	5	5	5	5	5

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Urinalysis

Stage : Week 6 of administration

Sex : Male

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	pH	Protein	Ketones	Glucose	Occult Blood	Bilirubin	Urobilinogen	Color
Animal No.								
2008	8.0	-	+/-	-	+/-	-	+/-	Y
2009	8.5	-	+/-	-	+/-	-	+/-	Y
2010	7.5	2+	2+	-	-	-	+/-	Y
2011	8.5	+/-	1+	-	+/-	-	+/-	Y
2012	8.0	1+	2+	-	-	-	+/-	Y
n	5	5	5	5	5	5	5	5

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Urinalysis

Stage : Week 6 of administration

Sex : Male

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	pH	Protein	Ketones	Glucose	Occult Blood	Bilirubin	Urobilinogen	Color
Animal No.								
3008	7.5	+/-	1+	-	-	-	+/-	Y
3009	8.5	1+	1+	-	-	1+	+/-	Y
3010	8.0	1+	1+	-	+/-	-	+/-	Y
3011	7.0	+/-	1+	-	-	-	+/-	Y
3012	7.5	1+	2+	-	-	1+	+/-	Y
n	5	5	5	5	5	5	5	5

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

	pH	Protein	Ketones	Glucose	Occult Blood	Bilirubin	Urobilinogen	Color
Animal No.								
4008	6.5	2+	3+	1+	-	1+	+/-	Y
4009	8.5	1+	2+	-	+/-	-	+/-	Y
4010	8.5	2+	2+	-	-	-	+/-	Y
4011	8.0	2+	2+	-	-	1+	+/-	Y
4012	>=9.0	2+	2+	1+	-	1+	+/-	Y
n	5	5	5	5	5	5	5	5

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Urinalysis

Stage : Week 6 of administration

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	pH	Protein	Ketones	Glucose	Occult Blood	Bilirubin	Urobilinogen	Color
Animal No.								
5106	6.0	-	-	-	-	-	+/-	Y
5107	6.5	+/-	1+	-	-	-	+/-	Y
5108	6.5	2+	1+	-	-	-	+/-	Y
5109	7.0	+/-	1+	-	-	-	+/-	Y
5110	8.5	+/-	+/-	-	-	-	+/-	Y
n	5	5	5	5	5	5	5	5

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Urinalysis

Stage : Week 6 of administration

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	pH	Protein	Ketones	Glucose	Occult Blood	Bilirubin	Urobilinogen	Color
Animal No.								
6106	6.5	-	2+	-	-	-	+/-	Y
6107	6.5	1+	3+	-	-	-	+/-	Y
6108	6.5	+/-	2+	-	-	-	+/-	Y
6109	6.5	1+	3+	-	-	-	+/-	Y
6110	6.0	1+	2+	-	-	-	+/-	Y
n	5	5	5	5	5	5	5	5

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Animal No.	RBC	WBC	Squamous epithelial cells	Small round epithelial cells	Cast	Crystal phosphate salts	Crystal calcium oxalate
1008	-	-	+/-	-	-	-	-
1009	-	-	+/-	-	-	-	-
1010	-	-	+/-	-	-	+/-	-
1011	-	-	+/-	-	-	-	-
1012	-	-	+/-	-	-	+/-	-
n	5	5	5	5	5	5	5

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Animal No.	RBC	WBC	Squamous epithelial cells	Small round epithelial cells	Cast	Crystal phosphate salts	Crystal calcium oxalate
2008	-	-	+/-	-	-	-	-
2009	-	-	+/-	-	-	+/-	-
2010	-	-	+/-	-	-	-	-
2011	-	-	+/-	-	-	-	-
2012	-	-	+/-	-	-	-	-
n	5	5	5	5	5	5	5

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Animal No.	RBC	WBC	Squamous epithelial cells	Small round epithelial cells	Cast	Crystal phosphate salts	Crystal calcium oxalate
3008	-	-	+/-	-	-	-	-
3009	-	-	+/-	-	-	-	-
3010	-	-	+/-	-	-	-	-
3011	-	-	+/-	-	-	-	-
3012	-	-	+/-	-	-	-	-
n	5	5	5	5	5	5	5

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Urinalysis

Stage : Week 6 of administration

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	RBC	WBC	Squamous epithelial cells	Small round epithelial cells	Cast	Crystal phosphate salts	Crystal calcium oxalate
4008	-	-	+/-	-	-	-	-
4009	-	-	+/-	-	-	-	-
4010	-	-	+/-	-	-	+/-	-
4011	-	-	+/-	-	-	-	-
4012	-	-	+/-	-	-	+/-	-
n	5	5	5	5	5	5	5

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Urinalysis

Stage : Week 6 of administration (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	RBC	WBC	Squamous epithelial cells	Small round epithelial cells	Cast	Crystal phosphate salts	Crystal calcium oxalate
5106	-	-	+/-	-	-	-	-
5107	-	-	+/-	-	-	-	-
5108	-	-	+/-	-	-	-	-
5109	-	-	+/-	-	-	+/-	-
5110	-	-	+/-	-	-	-	-
n	5	5	5	5	5	5	5

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Animal No.	RBC	WBC	Squamous epithelial cells	Small round epithelial cells	Cast	Crystal phosphate salts	Crystal calcium oxalate
6106	-	-	+/-	-	-	-	-
6107	-	-	+/-	-	-	-	-
6108	-	-	+/-	-	-	+/-	-
6109	-	-	+/-	-	-	-	-
6110	-	-	+/-	-	-	-	-
n	5	5	5	5	5	5	5

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Urinalysis

Stage : Week 6 of administration

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
1008	45	17.4	1264	1.8	3.5	2.5
1009	32	8.3	1976	1.2	2.6	1.9
1010	40	9.6	1156	0.7	1.8	1.0
1011	57	19.8	986	1.7	3.2	2.2
1012	35	12.1	1736	1.7	3.5	2.5
n	5	5	5	5	5	5
Mean	42	13.4	1424	1.4	2.9	2.0
S.D.	10	5.0	416	0.5	0.7	0.6

Urinalysis

Stage : Week 6 of administration

Sex : Male

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
2008	57	32.6	864	2.3	4.7	3.3
2009	48	19.7	1346	2.5	4.3	3.1
2010	19	6.1	2128	1.2	2.3	1.7
2011	34	9.2	2052	1.5	3.2	2.4
2012	37	15.3	1350	1.5	3.4	2.4
n	5	5	5	5	5	5
Mean	39	16.6	1548	1.8	3.6	2.6
S.D.	14	10.4	533	0.6	0.9	0.6

Urinalysis

Stage : Week 6 of administration

Sex : Male

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
3008	32	13.2	1988	1.7	4.3	2.5
3009	38	10.7	1404	0.9	2.6	1.5
3010	40	7.8	1540	1.0	1.9	1.4
3011	43	13.1	1928	1.4	4.3	2.6
3012	33	10.6	1856	1.4	3.3	2.2
n	5	5	5	5	5	5
Mean	37	11.1	1743	1.3	3.3	2.0
S.D.	5	2.2	256	0.3	1.1	0.6

Urinalysis

Stage : Week 6 of administration

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
4008	36	10.7	1934	1.2	3.2	1.9
4009	32	7.5	2260	1.2	2.3	1.7
4010	32	5.9	2232	0.9	2.0	1.5
4011	36	9.1	2462	1.5	3.4	2.3
4012	42	6.5	2010	0.7	2.2	1.2
n	5	5	5	5	5	5
Mean	36	7.9	2180	1.1	2.6	1.7
S.D.	4	2.0	211	0.3	0.6	0.4

Urinalysis

Stage : Week 6 of administration (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
5106	44	14.4	1072	1.2	2.3	1.5
5107	29	8.1	1446	0.8	1.8	1.1
5108	21	2.4	3074	0.5	1.1	0.7
5109	27	8.6	1682	1.2	2.2	1.6
5110	43	11.1	1198	0.8	2.2	1.5
n	5	5	5	5	5	5
Mean	33	8.9	1694	0.9	1.9	1.3
S.D.	10	4.4	806	0.3	0.5	0.4

Urinalysis

Stage : Week 6 of administration (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
6106	27	7.3	1992	1.3	2.2	1.7
6107	32	2.4	1928	0.3	0.6	0.3
6108	30	5.4	1906	0.7	1.5	1.1
6109	38	5.8	1950	0.7	1.7	1.1
6110	26	7.7	1938	1.3	1.9	1.5
n	5	5	5	5	5	5
Mean	31	5.7	1943	0.9	1.6	1.1
S.D.	5	2.1	32	0.4	0.6	0.5

Urinalysis

Stage : Week 2 of recovery

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	pH	Protein	Ketones	Glucose	Occult Blood	Bilirubin	Urobilinogen	Color
Animal No.								
1008	8.5	-	1+	-	+/-	-	+/-	Y
1009	>=9.0	+/-	+/-	-	-	1+	+/-	Y
1010	>=9.0	1+	+/-	-	+/-	-	+/-	Y
1011	8.5	-	-	-	1+	-	+/-	Y
1012	8.5	1+	+/-	-	+/-	-	+/-	Y
n	5	5	5	5	5	5	5	5

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Urinalysis

Stage : Week 2 of recovery

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	pH	Protein	Ketones	Glucose	Occult Blood	Bilirubin	Urobilinogen	Color
Animal No.								
4008	>=9.0	1+	1+	-	-	1+	+/-	Y
4009	>=9.0	1+	1+	-	-	-	+/-	Y
4010	>=9.0	1+	1+	-	-	-	+/-	Y
4011	>=9.0	2+	2+	-	-	-	+/-	Y
4012	7.0	1+	2+	-	-	1+	+/-	Y
n	5	5	5	5	5	5	5	5

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

with the reproduction/ developmental toxicity screening test in rats									
Urinalysis					Stage : Week 2 of recovery (non-mating group)				Species : Rat
Sex : Female					Dose : Dodecyloxirane 0 mg/kg				
					Occult Blood	Bilirubin	Urobilinogen	Color	
pH					Protein	Ketones	Glucose		
Animal No.									
5106	7.5	+/-	-	-	-	-	+/-	Y	
5107	6.5	2+	+/-	-	-	-	+/-	Y	
5108	7.5	1+	1+	1+	-	1+	+/-	Y	
5109	7.0	+/-	+/-	-	-	-	+/-	Y	
5110	>=9.0	-	-	-	-	-	+/-	Y	
n	5	5	5	5	5	5	5	5	

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Urinalysis

Stage : Week 2 of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	pH	Protein	Ketones	Glucose	Occult Blood	Bilirubin	Urobilinogen	Color
Animal No.								
6106	8.5	-	-	-	-	-	+/-	Y
6107	8.5	-	-	-	-	-	+/-	Y
6108	>=9.0	1+	+/-	1+	-	1+	+/-	Y
6109	>=9.0	1+	+/-	-	-	-	+/-	Y
6110	8.5	+/-	+/-	-	-	-	+/-	Y
n	5	5	5	5	5	5	5	5

Color LY: Light Yellow Y: Yellow DY: Dark Yellow BR: Brown RE: Red RB: Reddish brown WH: White BL: Blue CL: Colorless OT: Other

Protein) -:Negative , +/-:15 , 1+:30 , 2+:100 , 3+:>=300 mg/dL

Ketones) -:Negative , +/-:5 , 1+:15 , 2+:40 , 3+:80 mg/dL

Glucose) -:Negative , 1+:100 , 2+:250 , 3+:500 , 4+:>=1000 mg/dL

Occult Blood) -:Negative , +/-:0.015 , 1+:0.062 , 2+:0.135 , 3+:0.405 mg/dL

Bilirubin) -:Negative , 1+:0.8 , 2+:1.6 , 3+:3.2 mg/dL

Urobilinogen) +/-:0.1 - 1.0 , 1+:2.0 , 2+:4.0 , 3+:>=8.0 Ehrlich U/dL

Urinalysis

Stage : Week 2 of recovery

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	RBC	WBC	Squamous epithelial cells	Small round epithelial cells	Cast	Crystal phosphate salts	Crystal calcium oxalate
1008	-	-	+/-	-	-	-	-
1009	-	-	+/-	-	-	-	-
1010	-	-	+/-	-	-	-	-
1011	-	-	+/-	-	-	-	-
1012	-	-	+/-	-	-	+/-	-
n	5	5	5	5	5	5	5

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Urinalysis

Stage : Week 2 of recovery

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	RBC	WBC	Squamous epithelial cells	Small round epithelial cells	Cast	Crystal phosphate salts	Crystal calcium oxalate
4008	-	-	+/-	-	-	+/-	-
4009	-	-	+/-	-	-	-	-
4010	-	-	+/-	-	-	-	-
4011	-	-	+/-	-	-	-	-
4012	-	-	+/-	-	-	+/-	-
n	5	5	5	5	5	5	5

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Urinalysis

Stage : Week 2 of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	RBC	WBC	Squamous epithelial cells	Small round epithelial cells	Cast	Crystal phosphate salts	Crystal calcium oxalate
5106	-	-	+/-	-	-	+/-	-
5107	-	-	+/-	-	-	-	-
5108	-	-	+/-	-	-	-	-
5109	-	-	+/-	-	-	-	-
5110	-	-	+/-	-	-	-	-
n	5	5	5	5	5	5	5

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Urinalysis

Stage : Week 2 of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	RBC	WBC	Squamous epithelial cells	Small round epithelial cells	Cast	Crystal phosphate salts	Crystal calcium oxalate
6106	-	-	+/-	-	-	-	-
6107	-	-	+/-	-	-	-	-
6108	-	-	+/-	-	-	+/-	-
6109	-	-	+/-	-	-	-	-
6110	-	-	+/-	-	-	-	-
n	5	5	5	5	5	5	5

RBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

WBC) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Squamous epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Small round epithelial cells) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Cast) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal phosphate salts) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Crystal calcium oxalate) -:Negative , +/-:Slight , 1+:Mild , 2+:Moderate , 3+:Severe

Urinalysis

Stage : Week 2 of recovery

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
1008	50	16.1	1680	2.0	4.2	2.6
1009	48	14.5	2000	2.1	4.6	3.0
1010	46	12.3	1424	1.2	2.9	1.9
1011	91	41.6	888	2.9	5.6	3.7
1012	49	11.1	1936	1.8	3.5	2.2
n	5	5	5	5	5	5
Mean	57	19.1	1586	2.0	4.2	2.7
S.D.	19	12.7	452	0.6	1.0	0.7

Urinalysis

Stage : Week 2 of recovery

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
4008	44	13.8	2104	2.2	4.7	3.0
4009	40	11.7	2132	1.8	4.1	2.7
4010	34	10.0	2290	1.6	3.6	2.5
4011	41	12.9	2520	2.2	5.1	3.2
4012	45	9.1	2434	1.6	3.7	2.2
n	5	5	5	5	5	5
Mean	41	11.5	2296	1.9	4.2	2.7
S.D.	4	2.0	182	0.3	0.6	0.4

Urinalysis

Stage : Week 2 of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
5106	37	8.3	1554	0.9	1.9	1.2
5107	40	8.0	1854	1.0	2.3	1.5
5108	30	4.6	2660	0.8	1.8	1.2
5109	37	8.2	1756	1.1	2.3	1.7
5110	39	11.0	1434	1.2	2.1	1.4
n	5	5	5	5	5	5
Mean	37	8.0	1852	1.0	2.1	1.4
S.D.	4	2.3	481	0.2	0.2	0.2

Urinalysis

Stage : Week 2 of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Water consumption mL/24h	Urine volume mL/24h	Osmotic pressure mOsm/kg	U-Na mmol/24h	U-K mmol/24h	U-Cl mmol/24h
6106	34	10.1	1846	1.3	2.7	1.8
6107	36	6.2	1814	0.9	1.6	1.2
6108	30	7.3	2246	1.2	2.5	1.9
6109	32	8.6	1780	1.2	2.4	1.7
6110	34	5.1	2302	0.8	1.9	1.2
n	5	5	5	5	5	5
Mean	33	7.5	1998	1.1	2.2	1.6
S.D.	2	2.0	254	0.2	0.5	0.3

Hematology

Stage : End of administration

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
1001	886	15.2	44.3	50.0	17.2	34.3	12.1	180.8	97.8	87.2	70.7
1002	875	15.4	45.4	51.8	17.6	33.9	12.4	165.3	100.0	66.3	55.7
1003	857	15.1	43.3	50.6	17.6	34.9	12.1	172.5	96.9	62.4	47.4
1004	853	15.4	44.7	52.5	18.0	34.4	12.9	220.5	121.3	104.6	74.3
1005	847	15.3	45.6	53.8	18.0	33.6	13.0	227.1	101.0	52.8	39.9
n	5	5	5	5	5	5	5	5	5	5	5
Mean	864	15.3	44.7	51.7	17.7	34.2	12.5	193.2	103.4	74.7	57.6
S.D.	16	0.1	0.9	1.5	0.3	0.5	0.4	28.5	10.1	20.9	14.8

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
1001	14.1	0.7	0.0	1.1	0.6	13.6	15.5	345
1002	7.8	1.0	0.0	1.6	0.2	13.0	13.7	291
1003	12.3	0.7	0.0	1.6	0.4	12.9	16.1	299
1004	24.8	2.1	0.2	2.9	0.4	14.5	18.4	333
1005	11.3	0.6	0.1	0.8	0.2	13.1	16.3	331
n	5	5	5	5	5	5	5	5
Mean	14.1	1.0	0.1	1.6	0.4	13.4	16.0	320
S.D.	6.4	0.6	0.1	0.8	0.2	0.7	1.7	23

Hematology

Stage : End of administration

Sex : Male

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
2001	777	14.3	40.8	52.6	18.4	35.0	12.4	165.6	110.1	85.8	72.9
2002	777	14.5	42.5	54.7	18.7	34.2	13.0	206.0	115.7	87.2	70.0
2003	857	15.5	45.2	52.7	18.0	34.2	12.3	166.1	109.0	69.6	56.1
2004	851	14.9	43.0	50.5	17.5	34.6	12.4	151.9	110.9	66.2	54.4
2005	865	14.9	43.3	50.0	17.2	34.3	12.8	182.0	103.2	53.4	34.7
n	5	5	5	5	5	5	5	5	5	5	5
Mean	825	14.8	43.0	52.1	18.0	34.5	12.6	174.3	109.8	72.4	57.6
S.D.	44	0.5	1.6	1.9	0.6	0.3	0.3	20.7	4.5	14.2	15.2

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
2001	9.9	0.9	0.1	1.5	0.4	13.9	17.1	281
2002	13.4	1.4	0.1	1.7	0.5	12.3	12.8	344
2003	10.6	1.3	0.1	1.3	0.2	14.6	15.9	344
2004	9.8	0.5	0.1	1.0	0.5	13.6	12.0	345
2005	16.1	0.7	0.0	1.7	0.1	14.6	17.3	371
n	5	5	5	5	5	5	5	5
Mean	12.0	1.0	0.1	1.4	0.3	13.8	15.0	337
S.D.	2.7	0.4	0.0	0.3	0.2	0.9	2.5	33

Hematology

Stage : End of administration

Sex : Male

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
3001	818	15.0	43.5	53.2	18.4	34.5	12.6	216.9	115.7	82.6	71.1
3002	823	15.4	44.2	53.7	18.6	34.7	12.0	123.7	105.6	80.5	65.8
3003	838	14.9	43.1	51.5	17.8	34.5	12.2	162.5	93.8	79.8	58.5
3004	835	15.1	43.4	51.9	18.1	34.8	12.2	205.7	105.4	125.3	100.5
3005	812	15.7	46.9	57.7	19.3	33.5	12.1	186.9	107.2	98.6	77.3
n	5	5	5	5	5	5	5	5	5	5	5
Mean	825	15.2	44.2	53.6	18.4	34.4	12.2	179.1	105.5	93.4	74.6
S.D.	11	0.3	1.6	2.5	0.6	0.5	0.2	37.2	7.8	19.4	16.0

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
3001	8.4	0.9	0.2	1.2	0.8	12.9	13.2	275
3002	12.4	0.6	0.1	1.2	0.4	12.2	13.5	266
3003	15.8	1.5	0.1	3.4	0.5	14.1	16.8	315
3004	19.8	1.5	0.2	2.8	0.6	12.9	16.8	291
3005	17.1	1.6	0.1	1.9	0.5	12.8	14.6	309
n	5	5	5	5	5	5	5	5
Mean	14.7	1.2	0.1	2.1	0.6	13.0	15.0	291
S.D.	4.4	0.4	0.1	1.0	0.2	0.7	1.7	21

Hematology

Stage : End of administration

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
4001	820	15.2	44.3	54.0	18.6	34.4	12.7	160.1	86.6	82.0	68.8
4002	815	14.9	42.7	52.4	18.3	35.0	12.5	119.3	98.6	70.2	52.8
4003	861	15.0	43.7	50.8	17.5	34.4	12.5	156.3	133.1	64.5	42.9
4004	847	15.5	44.9	53.0	18.3	34.4	12.1	192.8	122.9	78.5	58.8
4005	848	14.9	44.5	52.5	17.6	33.6	13.9	163.0	103.7	64.4	50.1
n	5	5	5	5	5	5	5	5	5	5	5
Mean	838	15.1	44.0	52.5	18.1	34.4	12.7	158.3	109.0	71.9	54.7
S.D.	20	0.3	0.9	1.2	0.5	0.5	0.7	26.2	18.8	8.1	9.7

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
4001	10.4	1.0	0.1	1.4	0.4	12.9	15.4	284
4002	14.1	1.0	0.1	1.9	0.3	14.0	14.1	272
4003	19.3	0.9	0.0	1.0	0.2	13.1	14.8	324
4004	15.3	0.7	0.1	3.3	0.3	14.3	15.1	278
4005	11.4	0.8	0.1	1.8	0.2	14.2	15.6	328
n	5	5	5	5	5	5	5	5
Mean	14.1	0.9	0.1	1.9	0.3	13.7	15.0	297
S.D.	3.5	0.1	0.0	0.9	0.1	0.7	0.6	27

Hematology
Sex : FemaleStage : End of lactation (F0 dams)
Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
1101	730	15.0	41.2	56.4	20.6	36.5	13.4	231.0	130.6	149.3	76.9
1105	717	15.6	43.8	61.1	21.7	35.6	13.5	294.1	112.7	88.9	40.6
1107	736	13.8	40.7	55.3	18.7	33.8	13.8	216.3	138.4	67.0	41.4
1108	756	16.0	45.1	59.6	21.1	35.4	13.4	308.1	126.1	110.6	35.7
1111	752	15.7	45.1	60.0	20.8	34.7	13.9	269.5	115.9	79.7	30.6
n	5	5	5	5	5	5	5	5	5	5	5
Mean	738	15.2	43.2	58.5	20.6	35.2	13.6	263.8	124.7	99.1	45.0
S.D.	16	0.9	2.1	2.5	1.1	1.0	0.2	39.5	10.6	32.3	18.3

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
1101	61.6	1.8	0.3	6.7	2.0	11.6	13.9	466
1105	45.8	0.6	0.1	1.5	0.4	13.9	15.7	690
1107	23.4	0.4	0.1	1.2	0.4	13.1	12.8	527
1108	69.0	0.8	0.1	4.0	1.1	11.3	12.8	534
1111	44.0	0.9	0.0	3.0	1.2	13.3	16.4	359
n	5	5	5	5	5	5	5	5
Mean	48.8	0.9	0.1	3.3	1.0	12.6	14.3	515
S.D.	17.7	0.5	0.1	2.2	0.7	1.1	1.7	120

Hematology
Sex : FemaleStage : End of lactation (F0 dams)
Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
2104	740	15.0	43.5	58.8	20.3	34.5	13.2	227.5	115.9	109.1	55.0
2105	759	15.3	42.4	56.0	20.2	36.2	13.0	289.0	127.0	115.7	58.8
2107	750	15.0	43.4	57.9	20.0	34.6	13.4	259.2	123.7	80.2	34.7
2108	782	15.5	45.3	58.0	19.8	34.2	13.0	321.2	153.7	65.9	30.9
2112	749	14.7	42.3	56.4	19.7	34.9	14.0	236.3	127.5	80.6	38.9
n	5	5	5	5	5	5	5	5	5	5	5
Mean	756	15.1	43.4	57.4	20.0	34.9	13.3	266.6	129.6	90.3	43.7
S.D.	16	0.3	1.2	1.2	0.3	0.8	0.4	38.7	14.3	21.2	12.5

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
2104	46.8	0.8	0.1	4.8	1.7	14.4	17.4	372
2105	52.5	0.6	0.2	2.8	0.8	10.2	11.5	475
2107	41.6	0.6	0.0	2.7	0.5	12.6	12.1	429
2108	30.6	0.9	0.0	3.0	0.5	13.8	16.6	437
2112	36.8	0.7	0.1	3.1	1.1	12.4	12.2	544
n	5	5	5	5	5	5	5	5
Mean	41.7	0.7	0.1	3.3	0.9	12.7	14.0	451
S.D.	8.5	0.1	0.1	0.9	0.5	1.6	2.8	64

Hematology
Sex : FemaleStage : End of lactation (F0 dams)
Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
3101	730	15.7	42.5	58.2	21.5	36.9	12.7	287.6	150.6	128.5	73.9
3102	763	16.1	44.4	58.2	21.1	36.3	13.9	252.2	112.6	126.6	79.9
3104	719	15.3	43.1	60.0	21.3	35.5	13.0	223.4	114.0	59.6	31.8
3110	801	16.2	46.1	57.6	20.3	35.2	14.2	291.3	114.6	74.7	39.3
3111	713	15.2	43.7	61.3	21.4	34.8	12.9	277.5	119.3	58.6	18.2
n	5	5	5	5	5	5	5	5	5	5	5
Mean	745	15.7	44.0	59.1	21.1	35.7	13.3	266.4	122.2	89.6	48.6
S.D.	37	0.5	1.4	1.5	0.5	0.9	0.7	28.5	16.1	35.2	27.0

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
3101	48.0	1.5	0.2	4.3	0.7	11.9	14.6	491
3102	37.8	0.6	0.3	5.9	2.1	12.9	14.1	432
3104	25.7	0.9	0.0	0.9	0.3	11.5	12.1	584
3110	31.1	0.5	0.1	3.3	0.4	12.2	13.7	355
3111	37.7	0.4	0.0	2.0	0.2	13.4	14.8	361
n	5	5	5	5	5	5	5	5
Mean	36.1	0.8	0.1	3.3	0.7	12.4	13.9	445
S.D.	8.4	0.4	0.1	1.9	0.8	0.8	1.1	96

Hematology
Sex : FemaleStage : End of lactation (F0 dams)
Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
4102	775	16.4	44.4	57.3	21.1	36.9	15.3	240.9	124.3	96.0	59.6
4103	759	16.2	44.8	59.0	21.3	36.1	14.3	266.5	122.8	59.9	24.7
4107	745	16.0	43.8	58.8	21.5	36.6	14.4	214.3	107.3	109.1	58.6
4109	696	15.1	42.4	60.9	21.7	35.6	14.6	184.3	104.6	121.6	53.6
4111	704	15.1	42.4	60.3	21.4	35.6	13.3	218.4	122.8	79.9	35.4
n	5	5	5	5	5	5	5	5	5	5	5
Mean	736	15.8	43.6	59.3	21.4	36.2	14.4	224.9	116.4	93.3	46.4
S.D.	34	0.6	1.1	1.4	0.2	0.6	0.7	30.8	9.6	24.3	15.5

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
4102	30.7	0.8	0.2	4.3	0.4	11.8	12.5	322
4103	31.9	0.5	0.1	2.3	0.4	11.2	13.3	398
4107	41.9	1.4	0.1	6.2	0.9	14.1	14.7	286
4109	56.8	1.1	0.1	7.9	2.1	13.0	13.5	481
4111	41.3	0.8	0.0	1.9	0.3	10.1	10.6	445
n	5	5	5	5	5	5	5	5
Mean	40.5	0.9	0.1	4.5	0.8	12.0	12.9	386
S.D.	10.5	0.3	0.1	2.6	0.8	1.6	1.5	82

Hematology
Sex : FemaleStage : End of administration (non-mating group)
Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁶ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ³ /μL	10 ³ /μL	10 ² /μL
5101	784	14.5	40.5	51.6	18.5	35.8	11.5	123.9	101.1	87.1	72.4
5102	818	14.5	41.9	51.3	17.7	34.5	12.5	185.7	123.0	73.6	62.5
5103	802	14.6	41.7	52.0	18.2	35.0	12.2	171.9	112.3	64.6	50.1
5104	761	13.9	40.8	53.6	18.3	34.1	12.1	213.2	118.3	67.2	58.9
5105	801	14.9	43.3	54.1	18.6	34.3	11.4	184.6	97.4	61.7	52.9
n	5	5	5	5	5	5	5	5	5	5	5
Mean	793	14.5	41.6	52.5	18.3	34.7	11.9	175.9	110.4	70.8	59.4
S.D.	22	0.4	1.1	1.3	0.4	0.7	0.5	32.7	11.0	10.1	8.8

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
5101	11.8	0.9	0.1	1.1	0.8	12.7	12.8	253
5102	9.3	0.5	0.1	0.9	0.3	13.7	13.2	211
5103	11.1	1.1	0.1	1.8	0.4	14.5	15.5	205
5104	6.3	0.5	0.1	0.9	0.6	13.1	11.6	200
5105	6.0	1.1	0.0	1.2	0.4	14.4	13.6	213
n	5	5	5	5	5	5	5	5
Mean	8.9	0.8	0.1	1.2	0.5	13.7	13.3	216
S.D.	2.7	0.3	0.0	0.4	0.2	0.8	1.4	21

Hematology
Sex : FemaleStage : End of administration (non-mating group)
Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁶ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ³ /μL	10 ³ /μL	10 ² /μL
6101	730	14.0	41.0	56.1	19.2	34.2	12.1	149.2	106.7	69.9	55.4
6102	780	14.7	42.4	54.4	18.9	34.7	11.5	159.0	107.0	87.0	74.1
6103	806	15.1	43.2	53.6	18.7	34.8	11.4	105.9	128.0	61.7	50.8
6104	744	14.4	42.3	56.9	19.3	33.9	11.5	220.0	111.3	45.3	34.5
6105	761	14.4	41.9	55.0	18.9	34.4	12.3	164.3	108.5	38.8	30.6
n	5	5	5	5	5	5	5	5	5	5	5
Mean	764	14.5	42.2	55.2	19.0	34.4	11.8	159.7	112.3	60.5	49.1
S.D.	30	0.4	0.8	1.3	0.2	0.4	0.4	40.8	9.0	19.3	17.5

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
6101	10.6	0.7	0.0	2.7	0.6	13.0	12.1	221
6102	9.8	0.4	0.1	2.1	0.5	12.6	11.2	215
6103	6.2	0.8	0.1	3.1	0.7	14.0	13.9	207
6104	8.8	0.3	0.0	1.3	0.3	13.8	12.6	198
6105	6.5	0.7	0.0	0.9	0.1	13.9	12.1	192
n	5	5	5	5	5	5	5	5
Mean	8.4	0.6	0.0	2.0	0.4	13.5	12.4	207
S.D.	2.0	0.2	0.1	0.9	0.2	0.6	1.0	12

Hematology

Stage : End of recovery

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
1008	883	16.4	45.1	51.0	18.6	36.4	12.2	129.5	104.6	63.0	49.9
1009	925	16.9	48.2	52.1	18.2	35.0	12.7	183.6	107.6	79.5	63.9
1010	880	15.6	44.4	50.4	17.7	35.2	12.2	155.4	107.8	86.4	64.5
1011	878	15.9	45.2	51.4	18.1	35.2	13.1	140.1	112.4	71.0	56.2
1012	816	14.3	40.5	49.6	17.6	35.4	12.2	157.6	83.3	47.3	30.8
n	5	5	5	5	5	5	5	5	5	5	5
Mean	876	15.8	44.7	50.9	18.0	35.4	12.5	153.2	103.1	69.4	53.1
S.D.	39	1.0	2.8	1.0	0.4	0.6	0.4	20.5	11.4	15.2	13.8

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
1008	9.5	0.9	0.1	2.0	0.7	14.7	15.4	320
1009	12.3	1.2	0.1	1.6	0.4	15.1	19.1	314
1010	16.7	1.7	0.2	2.9	0.4	14.5	17.6	291
1011	11.4	1.1	0.2	1.5	0.6	14.7	15.2	343
1012	13.4	1.1	0.1	1.5	0.4	13.7	13.0	328
n	5	5	5	5	5	5	5	5
Mean	12.7	1.2	0.1	1.9	0.5	14.5	16.1	319
S.D.	2.7	0.3	0.1	0.6	0.1	0.5	2.4	19

Hematology

Stage : End of recovery

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
4008	797	15.2	43.0	53.9	19.1	35.4	13.1	187.3	102.3	75.7	59.2
4009	840	16.0	44.5	53.0	19.1	36.0	12.4	148.9	105.8	83.9	65.0
4010	888	15.9	45.2	50.9	17.9	35.1	12.4	151.2	116.3	74.7	54.5
4011	808	14.5	41.6	51.5	18.0	34.9	13.8	206.0	116.8	83.8	67.1
4012	885	16.0	45.3	51.3	18.1	35.2	12.7	192.3	98.3	67.7	48.5
n	5	5	5	5	5	5	5	5	5	5	5
Mean	844	15.5	43.9	52.1	18.4	35.3	12.9	177.1	107.9	77.2	58.9
S.D.	42	0.7	1.6	1.3	0.6	0.4	0.6	25.7	8.3	6.8	7.6

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
4008	12.2	1.5	0.1	2.5	0.3	13.6	18.2	298
4009	14.4	1.4	0.1	2.7	0.3	13.6	18.5	276
4010	16.4	1.4	0.1	1.8	0.6	14.4	20.0	340
4011	12.8	1.2	0.1	1.9	0.6	14.3	18.6	338
4012	16.0	1.4	0.1	1.5	0.3	14.0	19.6	308
n	5	5	5	5	5	5	5	5
Mean	14.4	1.4	0.1	2.1	0.4	14.0	19.0	312
S.D.	1.9	0.1	0.0	0.5	0.2	0.4	0.8	27

Hematology

Stage : End of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
5106	758	15.5	41.9	55.3	20.4	36.9	11.2	165.8	101.3	65.6	56.8
5107	715	14.5	39.5	55.2	20.3	36.7	12.9	186.8	102.5	96.8	74.6
5108	777	16.3	43.7	56.2	21.0	37.3	10.6	115.7	83.7	54.7	41.5
5109	795	15.5	42.4	53.3	19.4	36.5	11.2	145.9	117.5	33.3	27.8
5110	761	15.1	40.4	53.1	19.9	37.4	11.4	134.6	114.4	46.1	33.0
n	5	5	5	5	5	5	5	5	5	5	5
Mean	761	15.4	41.6	54.6	20.2	37.0	11.5	149.8	103.9	59.3	46.7
S.D.	30	0.7	1.7	1.4	0.6	0.4	0.9	27.5	13.3	24.1	19.1

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
5106	5.9	0.8	0.1	1.2	0.7	13.2	13.4	244
5107	18.1	0.9	0.1	2.7	0.4	13.5	16.3	206
5108	9.3	1.6	0.1	2.0	0.3	15.7	15.2	234
5109	4.0	0.7	0.0	0.6	0.1	16.1	12.6	199
5110	10.2	1.1	0.0	1.5	0.2	13.9	12.3	208
n	5	5	5	5	5	5	5	5
Mean	9.5	1.0	0.1	1.6	0.3	14.5	14.0	218
S.D.	5.4	0.4	0.1	0.8	0.2	1.3	1.7	20

Hematology

Stage : End of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	RBC	HGB	HCT	MCV	MCH	MCHC	RDW	#RETIC	PLT	WBC	#LYMP
Animal No.	10 ⁴ /μL	g/dL	%	fL	pg	g/dL	%	10 ⁹ /L	10 ⁴ /μL	10 ² /μL	10 ² /μL
6106	728	15.2	40.9	56.2	20.9	37.3	12.1	201.0	93.4	49.5	41.7
6107	779	16.1	43.3	55.6	20.7	37.2	11.4	139.1	130.6	32.2	27.8
6108	656	14.6	38.6	58.8	22.3	37.8	13.5	148.9	5.3	23.3	18.8
6109	794	15.4	41.7	52.5	19.4	37.0	11.9	158.8	109.7	32.7	26.4
6110	780	15.3	41.6	53.4	19.7	36.8	11.6	172.7	98.9	45.9	32.9
n	5	5	5	5	5	5	5	5	5	5	5
Mean	747	15.3	41.2	55.3	20.6	37.2	12.1	164.1	87.6	36.7	29.5
S.D.	57	0.5	1.7	2.5	1.1	0.4	0.8	24.1	48.1	10.8	8.5

	#NEUT	#EOS	#BASO	#MONO	#LUC	PT	APTT	FIB
Animal No.	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	10 ² /μL	s	s	mg/dL
6106	5.4	0.5	0.1	1.3	0.4	14.4	17.8	261
6107	3.2	0.4	0.0	0.6	0.1	16.0	12.8	215
6108	3.2	0.4	0.0	0.7	0.2	15.1	13.5	220
6109	4.6	0.9	0.0	0.6	0.1	15.1	12.5	260
6110	9.2	1.1	0.0	2.5	0.2	13.5	13.4	206
n	5	5	5	5	5	5	5	5
Mean	5.1	0.7	0.0	1.1	0.2	14.8	14.0	232
S.D.	2.5	0.3	0.0	0.8	0.1	0.9	2.2	26

Blood chemistry

Stage : End of administration

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
1001	67	48	316	0	51.1	81	18	120	0.1	138	20
1002	54	47	651	0	39.4	61	40	101	0.1	151	14
1003	56	35	560	0	53.8	44	23	81	0.1	161	12
1004	52	33	624	0	21.3	68	42	113	0.1	158	14
1005	70	49	979	0	55.3	71	39	114	0.1	115	17
n	5	5	5	5	5	5	5	5	5	5	5
Mean	60	42	626	0	44.2	65	32	106	0.1	145	15
S.D.	8	8	238	0	14.2	14	11	15	0.0	19	3

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
1001	0.27	143	3.9	104	10.5	5.5	6.7	3.3
1002	0.24	144	3.8	107	10.1	6.3	5.9	3.2
1003	0.29	142	3.7	104	10.3	5.5	6.3	3.3
1004	0.30	144	4.0	106	10.7	5.6	6.6	3.4
1005	0.29	147	3.7	106	10.8	6.1	6.5	3.3
n	5	5	5	5	5	5	5	5
Mean	0.28	144	3.8	105	10.5	5.8	6.4	3.3
S.D.	0.02	2	0.1	1	0.3	0.4	0.3	0.1

Blood chemistry

Stage : End of administration

Sex : Male

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
2001	57	44	587	0	19.5	76	19	117	0.1	167	18
2002	63	52	447	0	20.6	63	65	116	0.1	154	17
2003	58	42	770	0	36.7	65	21	98	0.1	147	19
2004	86	41	451	0	48.8	69	50	110	0.1	179	16
2005	54	28	660	0	9.3	49	62	86	0.1	172	13
n	5	5	5	5	5	5	5	5	5	5	5
Mean	64	41	583	0	27.0	64	43	105	0.1	164	17
S.D.	13	9	139	0	15.7	10	22	13	0.0	13	2

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
2001	0.29	143	3.8	108	10.1	5.6	6.0	3.1
2002	0.30	143	3.8	107	10.5	5.8	6.3	3.2
2003	0.34	144	3.7	107	10.1	5.5	6.0	3.2
2004	0.32	143	3.7	107	10.4	5.0	6.3	3.3
2005	0.26	142	3.9	106	10.2	5.1	6.3	3.2
n	5	5	5	5	5	5	5	5
Mean	0.30	143	3.8	107	10.3	5.4	6.2	3.2
S.D.	0.03	1	0.1	1	0.2	0.3	0.2	0.1

Blood chemistry

Stage : End of administration

Sex : Male

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
3001	56	48	728	0	46.1	69	33	122	0.1	167	18
3002	50	29	387	0	42.4	90	36	131	0.1	141	16
3003	73	40	357	0	61.7	68	77	119	0.1	133	17
3004	57	39	578	0	11.0	72	56	107	0.1	145	13
3005	56	38	565	0	24.8	90	78	127	0.1	169	13
n	5	5	5	5	5	5	5	5	5	5	5
Mean	58	39	523	0	37.2	78	56	121	0.1	151	15
S.D.	9	7	152	0	19.7	11	22	9	0.0	16	2

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
3001	0.29	142	3.9	104	10.4	6.5	6.3	3.3
3002	0.36	143	3.7	106	10.4	5.2	6.4	3.3
3003	0.34	144	3.8	106	10.4	5.7	6.3	3.3
3004	0.26	144	3.7	105	10.5	6.4	6.5	3.3
3005	0.22	144	3.6	104	10.7	5.3	6.6	3.3
n	5	5	5	5	5	5	5	5
Mean	0.29	143	3.7	105	10.5	5.8	6.4	3.3
S.D.	0.06	1	0.1	1	0.1	0.6	0.1	0.0

Blood chemistry

Stage : End of administration

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	AST	ALT	ALP	γ -GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μ mol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
4001	61	63	288	0	8.3	78	23	112	0.1	136	22
4002	74	69	694	0	37.4	80	39	121	0.1	126	18
4003	71	48	671	0	43.4	85	21	122	0.1	132	22
4004	70	65	575	0	20.1	81	30	113	0.1	142	18
4005	67	65	699	0	27.8	85	47	125	0.1	147	19
n	5	5	5	5	5	5	5	5	5	5	5
Mean	69	62	585	0	27.4	82	32	119	0.1	137	20
S.D.	5	8	174	0	13.9	3	11	6	0.0	8	2

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
4001	0.30	143	3.8	108	10.1	5.5	6.2	3.3
4002	0.34	144	3.6	108	10.0	5.9	5.9	3.2
4003	0.35	144	3.8	108	9.8	5.4	5.9	3.2
4004	0.30	143	3.5	106	10.0	5.0	6.2	3.3
4005	0.27	142	3.9	107	10.2	6.2	5.9	3.2
n	5	5	5	5	5	5	5	5
Mean	0.31	143	3.7	107	10.0	5.6	6.0	3.2
S.D.	0.03	1	0.2	1	0.1	0.5	0.2	0.1

Blood chemistry

Stage : End of lactation (F0 dams)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
1101	93	73	404	1	32.1	96	112	185	0.1	124	40
1105	175	70	313	1	57.4	85	77	176	0.1	131	28
1107	151	96	429	1	72.2	111	106	199	0.1	125	25
1108	118	70	496	1	47.5	119	103	226	0.1	137	38
1111	91	68	268	1	46.2	101	81	186	0.0	126	23
n	5	5	5	5	5	5	5	5	5	5	5
Mean	126	75	382	1	51.1	102	96	194	0.1	129	31
S.D.	37	12	91	0	14.9	13	16	19	0.0	5	8

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
1101	0.51	138	3.5	100	10.8	12.0	6.1	3.2
1105	0.38	141	3.7	102	11.1	9.4	6.8	3.3
1107	0.37	144	3.2	103	11.1	8.3	6.0	3.2
1108	0.47	139	3.6	99	11.5	11.0	7.2	3.7
1111	0.34	146	2.5	102	10.7	8.9	6.9	3.7
n	5	5	5	5	5	5	5	5
Mean	0.41	142	3.3	101	11.0	9.9	6.6	3.4
S.D.	0.07	3	0.5	2	0.3	1.5	0.5	0.3

Blood chemistry

Stage : End of lactation (F0 dams)

Sex : Female

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

	AST	ALT	ALP	γ -GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μ mol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
2104	73	49	476	1	68.1	78	90	167	0.1	138	26
2105	151	97	374	1	76.5	94	119	186	0.1	135	41
2107	94	70	421	1	51.0	95	92	175	0.1	128	29
2108	77	45	390	1	17.3	82	115	152	0.0	141	19
2112	89	68	437	1	59.0	116	55	206	0.1	105	29
n	5	5	5	5	5	5	5	5	5	5	5
Mean	97	66	420	1	54.4	93	94	177	0.1	129	29
S.D.	31	21	40	0	22.8	15	26	20	0.0	14	8

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
2104	0.31	143	2.6	101	10.7	8.1	6.2	3.3
2105	0.46	140	3.6	102	11.0	9.8	6.1	3.2
2107	0.38	143	2.8	100	10.8	9.6	6.4	3.3
2108	0.34	142	2.9	101	10.6	7.7	6.8	3.6
2112	0.38	143	3.5	103	11.6	9.9	6.8	3.5
n	5	5	5	5	5	5	5	5
Mean	0.37	142	3.1	101	10.9	9.0	6.5	3.4
S.D.	0.06	1	0.4	1	0.4	1.0	0.3	0.2

Blood chemistry

Stage : End of lactation (F0 dams)

Sex : Female

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

	AST	ALT	ALP	γ -GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μ mol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
3101	78	52	401	1	43.5	96	86	166	0.1	136	34
3102	73	67	346	1	23.4	97	73	167	0.1	124	27
3104	166	111	538	1	104.9	109	138	198	0.1	135	47
3110	121	81	458	1	52.1	107	286	213	0.1	162	26
3111	99	61	258	1	26.3	129	40	211	0.0	153	22
n	5	5	5	5	5	5	5	5	5	5	5
Mean	107	74	400	1	50.0	108	125	191	0.1	142	31
S.D.	38	23	107	0	32.9	13	97	23	0.0	15	10

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
3101	0.43	141	3.4	101	10.5	9.6	5.9	3.1
3102	0.40	140	3.5	102	10.5	8.5	6.0	3.1
3104	0.51	135	4.3	97	11.4	11.3	6.7	3.5
3110	0.33	140	3.8	103	11.1	8.9	6.6	3.5
3111	0.30	144	2.8	106	11.7	9.3	6.9	3.7
n	5	5	5	5	5	5	5	5
Mean	0.39	140	3.6	102	11.0	9.5	6.4	3.4
S.D.	0.08	3	0.6	3	0.5	1.1	0.4	0.3

Blood chemistry

Stage : End of lactation (F0 dams)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
4102	66	47	445	1	58.8	98	165	178	0.1	162	20
4103	90	71	291	1	157.2	142	149	246	0.1	115	20
4107	78	53	478	1	57.5	96	89	173	0.1	142	23
4109	90	56	396	1	13.8	93	70	172	0.1	127	21
4111	166	130	542	1	44.7	107	88	190	0.1	139	39
n	5	5	5	5	5	5	5	5	5	5	5
Mean	98	71	430	1	66.4	107	112	192	0.1	137	25
S.D.	39	34	94	0	53.9	20	42	31	0.0	18	8

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
4102	0.32	143	3.3	103	10.7	7.2	6.5	3.1
4103	0.35	145	3.4	107	11.9	10.3	6.9	3.6
4107	0.39	142	3.4	103	11.1	9.6	5.8	3.4
4109	0.36	141	3.3	101	11.1	8.8	6.5	3.5
4111	0.46	139	4.3	99	11.7	11.4	7.0	3.6
n	5	5	5	5	5	5	5	5
Mean	0.38	142	3.5	103	11.3	9.5	6.5	3.4
S.D.	0.05	2	0.4	3	0.5	1.6	0.5	0.2

Blood chemistry

Stage : End of administration (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
5101	54	27	455	0	34.5	59	31	119	0.1	142	20
5102	51	23	308	0	41.0	84	24	149	0.1	178	15
5103	63	35	215	0	26.2	69	41	126	0.1	122	15
5104	45	24	163	0	11.3	64	33	122	0.1	166	13
5105	43	27	325	0	9.6	83	41	153	0.1	169	11
n	5	5	5	5	5	5	5	5	5	5	5
Mean	51	27	293	0	24.5	72	34	134	0.1	155	15
S.D.	8	5	112	0	13.9	11	7	16	0.0	23	3

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
5101	0.36	144	3.5	106	10.6	5.6	6.7	3.8
5102	0.29	144	3.8	109	10.1	4.5	6.7	3.8
5103	0.31	143	3.6	107	10.3	5.5	6.5	3.7
5104	0.29	144	3.1	108	10.1	4.9	6.3	3.7
5105	0.26	144	3.6	106	10.6	4.8	6.6	3.8
n	5	5	5	5	5	5	5	5
Mean	0.30	144	3.5	107	10.3	5.1	6.6	3.8
S.D.	0.04	0	0.3	1	0.3	0.5	0.2	0.1

Blood chemistry

Stage : End of administration (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	AST	ALT	ALP	γ -GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μ mol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
6101	51	31	324	0	21.3	58	11	106	0.1	174	19
6102	54	33	188	0	7.4	72	16	123	0.1	135	15
6103	57	38	210	0	21.3	52	25	100	0.1	127	21
6104	50	37	209	0	22.7	74	18	115	0.1	144	16
6105	59	35	404	0	18.5	84	9	140	0.1	130	19
n	5	5	5	5	5	5	5	5	5	5	5
Mean	54	35	267	0	18.2	68	16	117	0.1	142	18
S.D.	4	3	93	0	6.2	13	6	16	0.0	19	2

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
6101	0.31	142	3.7	108	10.4	5.6	6.4	3.7
6102	0.30	142	3.8	110	10.3	5.7	6.3	3.7
6103	0.33	143	3.6	107	10.3	5.5	6.4	3.8
6104	0.30	142	3.5	108	9.8	5.4	6.0	3.3
6105	0.31	142	3.6	109	10.3	5.4	6.6	3.9
n	5	5	5	5	5	5	5	5
Mean	0.31	142	3.6	108	10.2	5.5	6.3	3.7
S.D.	0.01	0	0.1	1	0.2	0.1	0.2	0.2

Blood chemistry

Stage : End of recovery

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	AST	ALT	ALP	γ -GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μ mol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
1008	57	35	289	0	19.1	84	58	112	0.1	132	17
1009	84	45	363	1	133.6	63	68	107	0.1	134	27
1010	53	29	362	1	18.3	63	49	92	0.1	144	19
1011	57	29	347	1	14.9	61	44	103	0.1	144	17
1012	47	23	310	0	30.7	93	94	141	0.1	177	18
n	5	5	5	5	5	5	5	5	5	5	5
Mean	60	32	334	1	43.3	73	63	111	0.1	146	20
S.D.	14	8	33	1	50.8	15	20	18	0.0	18	4

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
1008	0.26	145	3.6	107	10.4	5.2	6.6	3.2
1009	0.30	145	3.9	108	10.8	6.0	6.6	3.5
1010	0.29	144	3.8	106	10.5	5.4	6.5	3.2
1011	0.28	145	3.6	107	10.2	5.9	6.0	3.2
1012	0.25	144	4.0	105	10.3	5.1	6.6	3.3
n	5	5	5	5	5	5	5	5
Mean	0.28	145	3.8	107	10.4	5.5	6.5	3.3
S.D.	0.02	1	0.2	1	0.2	0.4	0.3	0.1

Blood chemistry

Stage : End of recovery

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	AST	ALT	ALP	γ -GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μ mol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
4008	53	24	285	1	40.9	59	74	97	0.1	184	17
4009	46	28	312	1	26.5	84	72	123	0.1	152	20
4010	52	31	380	1	19.9	70	43	110	0.1	140	25
4011	68	49	309	0	26.5	90	88	131	0.1	170	29
4012	55	32	313	0	22.3	79	38	107	0.1	153	21
n	5	5	5	5	5	5	5	5	5	5	5
Mean	55	33	320	1	27.2	76	63	114	0.1	160	22
S.D.	8	10	36	1	8.2	12	22	13	0.0	17	5

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
4008	0.28	143	4.2	104	10.2	5.8	6.6	3.2
4009	0.27	144	4.0	105	10.8	5.8	6.4	3.3
4010	0.27	145	3.6	107	10.5	5.9	6.4	3.2
4011	0.31	145	4.1	106	10.5	6.5	6.6	3.3
4012	0.28	146	3.4	106	10.6	5.1	6.7	3.2
n	5	5	5	5	5	5	5	5
Mean	0.28	145	3.9	106	10.5	5.8	6.5	3.2
S.D.	0.02	1	0.3	1	0.2	0.5	0.1	0.1

Blood chemistry

Stage : End of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

	AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
5106	54	27	188	0	79.4	88	43	167	0.2	123	20
5107	55	36	101	0	25.7	90	133	205	0.2	138	22
5108	48	25	255	1	104.5	81	44	160	0.1	132	28
5109	50	30	131	0	29.8	73	60	161	0.1	165	18
5110	64	30	322	1	46.8	63	25	138	0.1	128	20
n	5	5	5	5	5	5	5	5	5	5	5
Mean	54	30	199	0	57.2	79	61	166	0.1	137	22
S.D.	6	4	90	1	33.8	11	42	24	0.1	16	4

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
5106	0.42	140	3.4	106	10.2	5.2	6.6	3.6
5107	0.30	143	3.3	104	11.3	4.2	8.0	4.7
5108	0.40	141	4.0	107	10.3	4.7	6.3	3.5
5109	0.36	143	3.4	107	10.4	3.6	7.0	4.2
5110	0.32	143	3.2	107	10.3	5.0	6.6	3.7
n	5	5	5	5	5	5	5	5
Mean	0.36	142	3.5	106	10.5	4.5	6.9	3.9
S.D.	0.05	1	0.3	1	0.5	0.6	0.7	0.5

Blood chemistry

Stage : End of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

	AST	ALT	ALP	γ-GTP	T-BA	T-CHO	TG	PL	T-BIL	GLU	BUN
Animal No.	IU/L	IU/L	IU/L	IU/L	μmol/L	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL	mg/dL
6106	55	26	237	1	17.2	72	22	153	0.2	154	21
6107	53	24	101	1	25.5	69	35	133	0.1	119	19
6108	48	32	263	1	9.8	55	69	127	0.1	163	24
6109	56	24	145	1	11.6	83	32	147	0.1	140	19
6110	48	23	119	0	13.1	97	162	198	0.1	147	18
n	5	5	5	5	5	5	5	5	5	5	5
Mean	52	26	173	1	15.4	75	64	152	0.1	145	20
S.D.	4	4	73	0	6.3	16	58	28	0.0	17	2

	CRNN	Na	K	Cl	Ca	P	TP	ALB
Animal No.	mg/dL	mmol/L	mmol/L	mmol/L	mg/dL	mg/dL	g/dL	g/dL
6106	0.44	143	3.2	108	10.4	4.2	7.3	4.1
6107	0.35	143	4.0	110	10.0	4.5	6.4	3.5
6108	0.34	141	3.8	108	10.2	4.6	6.5	3.8
6109	0.34	143	3.6	107	10.0	4.3	6.0	3.2
6110	0.29	144	3.6	108	10.6	3.3	7.1	4.2
n	5	5	5	5	5	5	5	5
Mean	0.35	143	3.6	108	10.2	4.2	6.7	3.8
S.D.	0.05	1	0.3	1	0.3	0.5	0.5	0.4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration
Dose : Dodecyloxirane 0 mg/kg

Animal No.	Body weight		Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	g	AB g	RE g/100g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
1001	530		2.29	0.43	9.1	1.7	10.5	2.0	19.6	3.7
1002	468		2.13	0.46	8.8	1.9	10.2	2.2	19.0	4.1
1003	520		2.24	0.43	10.5	2.0	10.9	2.1	21.4	4.1
1004	483		2.11	0.44	7.3	1.5	9.2	1.9	16.5	3.4
1005	499		2.28	0.46	15.3	3.1	12.8	2.6	28.1	5.6
1006	568		2.17	0.38	10.5	1.8	11.1	2.0	21.6	3.8
1007	498		2.05	0.41	9.8	2.0	10.2	2.0	20.0	4.0
n	7		7	7	7	7	7	7	7	7
Mean	509		2.18	0.43	10.2	2.0	10.7	2.1	20.9	4.1
S.D.	33		0.09	0.03	2.5	0.5	1.1	0.2	3.6	0.7

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB mg	RE mg/100g	AB g	RE g/100g	AB g	RE g/100g	AB g	RE g/100g	AB g	RE g/100g
1001	303	57	1.59	0.30	14.62	2.76	0.63	0.12	1.75	0.33
1002	337	72	1.43	0.31	12.39	2.65	0.78	0.17	1.60	0.34
1003	236	45	1.46	0.28	14.04	2.70	0.64	0.12	1.59	0.31
1004	416	86	1.30	0.27	13.57	2.81	0.75	0.16	1.47	0.30
1005	207	41	1.36	0.27	13.13	2.63	0.74	0.15	1.68	0.34
1006	415	73	1.62	0.29	15.59	2.74	0.87	0.15	1.73	0.30
1007	235	47	1.42	0.29	12.45	2.50	0.60	0.12	1.55	0.31
n	7	7	7	7	7	7	7	7	7	7
Mean	307	60	1.45	0.29	13.68	2.68	0.72	0.14	1.62	0.32
S.D.	86	17	0.12	0.01	1.17	0.10	0.10	0.02	0.10	0.02

Animal No.	Kidney-L		Kidney-RL		Adrenal -R		Adrenal -L		Adrenal -RL	
	AB g	RE g/100g	AB g	RE g/100g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
1001	1.91	0.36	3.66	0.69	25	5	27	5	52	10
1002	1.70	0.36	3.30	0.71	35	7	32	7	67	14
1003	1.56	0.30	3.15	0.61	25	5	28	5	53	10
1004	1.40	0.29	2.87	0.59	28	6	31	6	59	12
1005	1.55	0.31	3.23	0.65	27	5	30	6	57	11
1006	1.63	0.29	3.36	0.59	24	4	25	4	49	9
1007	1.50	0.30	3.05	0.61	21	4	25	5	46	9
n	7	7	7	7	7	7	7	7	7	7
Mean	1.61	0.32	3.23	0.64	26	5	28	5	55	11
S.D.	0.16	0.03	0.25	0.05	4	1	3	1	7	2

AB : Absolute weight, RE : Relative weight by body weight

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration
Dose : Dodecyloxirane 0 mg/kg

Animal No.	Testis-R		Testis-L		Testis-RL		Epididymis-R		Epididymis-L	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g
1001	1.74	0.33	1.74	0.33	3.48	0.66	670	126	661	125
1002	1.76	0.38	1.66	0.35	3.42	0.73	679	145	631	135
1003	1.70	0.33	1.65	0.32	3.35	0.64	585	113	583	112
1004	1.42	0.29	1.41	0.29	2.83	0.59	607	126	555	115
1005	1.79	0.36	1.75	0.35	3.54	0.71	673	135	647	130
1006	1.58	0.28	1.51	0.27	3.09	0.54	628	111	564	99
1007	1.53	0.31	1.57	0.32	3.10	0.62	614	123	595	119
n	7	7	7	7	7	7	7	7	7	7
Mean	1.65	0.33	1.61	0.32	3.26	0.64	637	126	605	119
S.D.	0.14	0.04	0.12	0.03	0.26	0.07	37	12	42	12

Animal No.	Epididymis-RL		Prostate		Seminal vesicle		LABC		Cowper's gland-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	mg	mg/100g
1001	1331	251	0.90	0.17	1.26	0.24	1.38	0.26	50.1	9.5
1002	1310	280	1.81	0.39	2.05	0.44	1.92	0.41	76.1	16.3
1003	1168	225	1.42	0.27	1.65	0.32	1.66	0.32	58.2	11.2
1004	1162	241	1.18	0.24	1.28	0.27	1.48	0.31	48.6	10.1
1005	1320	265	1.33	0.27	2.30	0.46	1.63	0.33	76.4	15.3
1006	1192	210	1.75	0.31	1.94	0.34	1.72	0.30	74.5	13.1
1007	1209	243	1.38	0.28	2.02	0.41	1.64	0.33	40.2	8.1
n	7	7	7	7	7	7	7	7	7	7
Mean	1242	245	1.40	0.28	1.79	0.35	1.63	0.32	60.6	11.9
S.D.	75	23	0.31	0.07	0.40	0.08	0.17	0.05	15.1	3.1

Animal No.	Cowper's gland-L		Cowper's gland-RL		Glans penis	
	AB	RE	AB	RE	AB	RE
	mg	mg/100g	mg	mg/100g	mg	mg/100g
1001	49.0	9.2	99.1	18.7	129.0	24.3
1002	100.6	21.5	176.7	37.8	130.9	28.0
1003	59.3	11.4	117.5	22.6	177.1	34.1
1004	47.5	9.8	96.1	19.9	140.3	29.0
1005	71.8	14.4	148.2	29.7	150.1	30.1
1006	64.0	11.3	138.5	24.4	118.7	20.9
1007	44.9	9.0	85.1	17.1	189.9	38.1
n	7	7	7	7	7	7
Mean	62.4	12.4	123.0	24.3	148.0	29.2
S.D.	19.4	4.4	33.0	7.3	26.4	5.8

AB : Absolute weight, RE : Relative weight by body weight
LABC: Levator ani and bulbocavernosus muscle

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration
Dose : Dodecyloxirane 100 mg/kg

Study No. : R-1261

Animal No.	Body weight		Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	g	AB g	RE g/100g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
2001	566	2.21	0.39	11.5	2.0	6.7	1.2	18.2	3.2	
2002	583	2.21	0.38	13.1	2.2	11.3	1.9	24.4	4.2	
2003	508	2.09	0.41	10.1	2.0	10.6	2.1	20.7	4.1	
2004	574	2.27	0.40	15.0	2.6	14.9	2.6	29.9	5.2	
2005	504	2.11	0.42	11.1	2.2	11.1	2.2	22.2	4.4	
2006	487	2.15	0.44	10.4	2.1	10.7	2.2	21.1	4.3	
2007	484	2.30	0.48	11.4	2.4	7.2	1.5	18.6	3.8	
2008	521	2.14	0.41	10.0	1.9	7.5	1.4	17.5	3.4	
2009	497	2.26	0.45	12.6	2.5	10.2	2.1	22.8	4.6	
2010	444	2.16	0.49	8.9	2.0	9.2	2.1	18.1	4.1	
2011	544	2.10	0.39	13.4	2.5	11.7	2.2	25.1	4.6	
2012	546	2.06	0.38	9.7	1.8	8.4	1.5	18.1	3.3	
n	12	12	12	12	12	12	12	12	12	12
Mean	522	2.17	0.42	11.4	2.2	10.0	1.9	21.4	4.1	
S.D.	42	0.08	0.04	1.8	0.3	2.3	0.4	3.7	0.6	

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB mg	RE mg/100g	AB g	RE g/100g	AB g	RE g/100g	AB g	RE g/100g	AB g	RE g/100g
2001	267	47	1.45	0.26	15.68	2.77	0.70	0.12	1.78	0.31
2002	409	70	1.43	0.25	15.63	2.68	0.67	0.11	1.86	0.32
2003	276	54	1.41	0.28	13.13	2.58	0.77	0.15	1.41	0.28
2004	418	73	1.36	0.24	17.50	3.05	0.75	0.13	1.68	0.29
2005	296	59	1.39	0.28	13.00	2.58	0.51	0.10	1.47	0.29
2006	265	54	1.32	0.27	15.13	3.11	0.63	0.13	1.49	0.31
2007	407	84	1.54	0.32	12.72	2.63	0.74	0.15	1.52	0.31
2008	300	58	1.32	0.25	14.35	2.75	0.70	0.13	1.77	0.34
2009	302	61	1.48	0.30	12.74	2.56	0.60	0.12	1.68	0.34
2010	414	93	1.18	0.27	11.42	2.57	0.60	0.14	1.31	0.30
2011	314	58	1.42	0.26	16.06	2.95	0.65	0.12	1.82	0.33
2012	389	71	1.44	0.26	15.73	2.88	0.68	0.12	1.67	0.31
n	12	12	12	12	12	12	12	12	12	12
Mean	338	65	1.40	0.27	14.42	2.76	0.67	0.13	1.62	0.31
S.D.	63	13	0.09	0.02	1.81	0.20	0.07	0.01	0.18	0.02

AB : Absolute weight, RE : Relative weight by body weight

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration
Dose : Dodecyloxirane 100 mg/kg

Animal No.	Kidney-L		Kidney-RL		Adrenal -R		Adrenal -L		Adrenal -RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
2001	1.93	0.34	3.71	0.66	25	4	24	4	49	9
2002	1.91	0.33	3.77	0.65	30	5	30	5	60	10
2003	1.51	0.30	2.92	0.57	24	5	24	5	48	9
2004	1.81	0.32	3.49	0.61	34	6	31	5	65	11
2005	1.56	0.31	3.03	0.60	22	4	26	5	48	10
2006	1.60	0.33	3.09	0.63	27	6	29	6	56	11
2007	1.54	0.32	3.06	0.63	25	5	28	6	53	11
2008	1.82	0.35	3.59	0.69	22	4	26	5	48	9
2009	1.64	0.33	3.32	0.67	22	4	23	5	45	9
2010	1.38	0.31	2.69	0.61	21	5	22	5	43	10
2011	1.88	0.35	3.70	0.68	29	5	31	6	60	11
2012	1.62	0.30	3.29	0.60	30	5	31	6	61	11
n	12	12	12	12	12	12	12	12	12	12
Mean	1.68	0.32	3.31	0.63	26	5	27	5	53	10
S.D.	0.18	0.02	0.35	0.04	4	1	3	1	7	1

Animal No.	Testis-R		Testis-L		Testis-RL		Epididymis-R		Epididymis-L	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g
2001	1.73	0.31	1.65	0.29	3.38	0.60	656	116	643	114
2002	1.60	0.27	1.64	0.28	3.24	0.56	685	117	665	114
2003	1.82	0.36	1.80	0.35	3.62	0.71	690	136	687	135
2004	1.80	0.31	1.90	0.33	3.70	0.64	686	120	684	119
2005	1.88	0.37	1.79	0.36	3.67	0.73	629	125	625	124
2006	1.68	0.34	1.61	0.33	3.29	0.68	684	140	675	139
2007	1.60	0.33	1.60	0.33	3.20	0.66	589	122	598	124
2008	1.70	0.33	1.76	0.34	3.46	0.66	630	121	670	129
2009	1.89	0.38	1.88	0.38	3.77	0.76	691	139	677	136
2010	1.54	0.35	1.57	0.35	3.11	0.70	660	149	613	138
2011	1.68	0.31	1.72	0.32	3.40	0.63	609	112	617	113
2012	1.68	0.31	1.60	0.29	3.28	0.60	639	117	667	122
n	12	12	12	12	12	12	12	12	12	12
Mean	1.72	0.33	1.71	0.33	3.43	0.66	654	126	652	126
S.D.	0.11	0.03	0.11	0.03	0.22	0.06	35	12	31	10

AB : Absolute weight, RE : Relative weight by body weight

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration
Dose : Dodecyloxirane 100 mg/kg

Animal No.	Epididymis-RL		Prostate		Seminal vesicle		LABC		Cowper's gland-R	
	AB mg	RE mg/100g	AB g	RE g/100g	AB g	RE g/100g	AB g	RE g/100g	AB mg	RE mg/100g
2001	1299	230	1.16	0.20	1.76	0.31	1.97	0.35	105.8	18.7
2002	1350	232	1.25	0.21	1.65	0.28	1.82	0.31	34.7	6.0
2003	1377	271	1.15	0.23	1.62	0.32	1.56	0.31	47.4	9.3
2004	1370	239	1.05	0.18	1.94	0.34	1.83	0.32	54.2	9.4
2005	1254	249	1.22	0.24	1.23	0.24	1.66	0.33	48.3	9.6
2006	1359	279	1.15	0.24	1.71	0.35	1.61	0.33	56.7	11.6
2007	1187	245	1.32	0.27	2.10	0.43	1.69	0.35	76.6	15.8
2008	1300	250	1.46	0.28	1.38	0.26	1.82	0.35	57.8	11.1
2009	1368	275	1.34	0.27	1.26	0.25	1.15	0.23	55.0	11.1
2010	1273	287	1.18	0.27	1.39	0.31	1.32	0.30	49.9	11.2
2011	1226	225	1.25	0.23	1.98	0.36	1.78	0.33	55.9	10.3
2012	1306	239	1.16	0.21	1.49	0.27	1.61	0.29	67.6	12.4
n	12	12	12	12	12	12	12	12	12	12
Mean	1306	252	1.22	0.24	1.63	0.31	1.65	0.32	59.2	11.4
S.D.	62	21	0.11	0.03	0.29	0.05	0.23	0.03	18.0	3.2

Animal No.	Cowper's gland-L		Cowper's gland-RL		Glans penis	
	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
2001	109.8	19.4	215.6	38.1	93.8	16.6
2002	41.6	7.1	76.3	13.1	139.6	23.9
2003	52.5	10.3	99.9	19.7	106.1	20.9
2004	59.4	10.3	113.6	19.8	114.6	20.0
2005	54.3	10.8	102.6	20.4	125.9	25.0
2006	60.1	12.3	116.8	24.0	97.1	19.9
2007	83.0	17.1	159.6	33.0	194.7	40.2
2008	63.5	12.2	121.3	23.3	128.5	24.7
2009	66.7	13.4	121.7	24.5	188.2	37.9
2010	47.3	10.7	97.2	21.9	116.0	26.1
2011	54.9	10.1	110.8	20.4	159.6	29.3
2012	59.4	10.9	127.0	23.3	175.5	32.1
n	12	12	12	12	12	12
Mean	62.7	12.1	121.9	23.5	136.6	26.4
S.D.	18.1	3.3	35.7	6.5	35.0	7.3

AB : Absolute weight, RE : Relative weight by body weight
LABC: Levator ani and bulbocavernosus muscle

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration
Dose : Dodecyloxirane 300 mg/kg

Animal No.	Body weight		Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	g	AB g	RE g/100g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
3001	598		1.97	0.33	11.9	2.0	9.1	1.5	21.0	3.5
3002	460		1.90	0.41	7.2	1.6	8.8	1.9	16.0	3.5
3003	480		2.09	0.44	10.2	2.1	8.5	1.8	18.7	3.9
3004	525		2.49	0.47	13.8	2.6	14.4	2.7	28.2	5.4
3005	540		2.15	0.40	8.5	1.6	12.2	2.3	20.7	3.8
3006	480		2.09	0.44	15.5	3.2	12.6	2.6	28.1	5.9
3007	470		2.32	0.49	8.7	1.9	10.3	2.2	19.0	4.0
3008	557		2.22	0.40	12.2	2.2	12.7	2.3	24.9	4.5
3009	495		2.10	0.42	10.6	2.1	11.2	2.3	21.8	4.4
3010	480		2.16	0.45	13.7	2.9	13.6	2.8	27.3	5.7
3011	551		2.09	0.38	13.2	2.4	10.0	1.8	23.2	4.2
3012	519		2.14	0.41	10.8	2.1	12.6	2.4	23.4	4.5
n	12		12	12	12	12	12	12	12	12
Mean	513		2.14	0.42	11.4	2.2	11.3	2.2	22.7	4.4
S.D.	42		0.15	0.04	2.5	0.5	2.0	0.4	3.9	0.8

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB mg	RE mg/100g	AB g	RE g/100g	AB g	RE g/100g	AB g	RE g/100g	AB g	RE g/100g
3001	393	66	1.62	0.27	19.34	3.23	0.66	0.11	1.70	0.28
3002	270	59	1.17	0.25	12.92	2.81	0.66	0.14	1.37	0.30
3003	305	64	1.31	0.27	13.12	2.73	0.58	0.12	1.51	0.31
3004	555	106	1.45	0.28	14.68	2.80	0.91	0.17	1.72	0.33
3005	427	79	1.50	0.28	16.81	3.11	0.99	0.18	1.93	0.36
3006	367	76	1.29	0.27	13.01	2.71	0.54	0.11	1.38	0.29
3007	245	52	1.32	0.28	12.08	2.57	0.67	0.14	1.47	0.31
3008	235	42	1.60	0.29	16.70	3.00	0.72	0.13	1.69	0.30
3009	485	98	1.27	0.26	12.85	2.60	0.69	0.14	1.58	0.32
3010	446	93	1.47	0.31	12.86	2.68	0.66	0.14	1.87	0.39
3011	221	40	1.59	0.29	16.82	3.05	0.91	0.17	1.81	0.33
3012	339	65	1.39	0.27	14.26	2.75	0.69	0.13	1.53	0.29
n	12	12	12	12	12	12	12	12	12	12
Mean	357	70	1.42	0.28	14.62	2.84	0.72	0.14	1.63	0.32
S.D.	107	21	0.15	0.02	2.27	0.21	0.14	0.02	0.19	0.03

AB : Absolute weight, RE : Relative weight by body weight

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration
Dose : Dodecyloxirane 300 mg/kg

Animal No.	Kidney-L		Kidney-RL		Adrenal -R		Adrenal -L		Adrenal -RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
3001	1.76	0.29	3.46	0.58	24	4	28	5	52	9
3002	1.37	0.30	2.74	0.60	27	6	30	7	57	12
3003	1.53	0.32	3.04	0.63	20	4	23	5	43	9
3004	1.88	0.36	3.60	0.69	25	5	29	6	54	10
3005	2.04	0.38	3.97	0.74	25	5	29	5	54	10
3006	1.34	0.28	2.72	0.57	27	6	30	6	57	12
3007	1.50	0.32	2.97	0.63	28	6	30	6	58	12
3008	1.76	0.32	3.45	0.62	27	5	30	5	57	10
3009	1.61	0.33	3.19	0.64	23	5	26	5	49	10
3010	2.13	0.44	4.00	0.83	30	6	33	7	63	13
3011	1.89	0.34	3.70	0.67	25	5	32	6	57	10
3012	1.61	0.31	3.14	0.61	23	4	28	5	51	10
n	12	12	12	12	12	12	12	12	12	12
Mean	1.70	0.33	3.33	0.65	25	5	29	6	54	11
S.D.	0.25	0.04	0.44	0.07	3	1	3	1	5	1

Animal No.	Testis-R		Testis-L		Testis-RL		Epididymis-R		Epididymis-L	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g
3001	1.55	0.26	1.62	0.27	3.17	0.53	647	108	639	107
3002	1.59	0.35	1.53	0.33	3.12	0.68	625	136	596	130
3003	1.15	0.24	1.04	0.22	2.19	0.46	556	116	471	98
3004	1.85	0.35	1.80	0.34	3.65	0.70	798	152	862	164
3005	1.68	0.31	1.69	0.31	3.37	0.62	619	115	589	109
3006	1.44	0.30	1.45	0.30	2.89	0.60	601	125	554	115
3007	1.83	0.39	1.85	0.39	3.68	0.78	707	150	699	149
3008	1.61	0.29	1.61	0.29	3.22	0.58	609	109	576	103
3009	1.78	0.36	1.83	0.37	3.61	0.73	745	151	713	144
3010	1.78	0.37	1.78	0.37	3.56	0.74	658	137	651	136
3011	1.74	0.32	1.77	0.32	3.51	0.64	802	146	751	136
3012	1.74	0.34	1.74	0.34	3.48	0.67	688	133	673	130
n	12	12	12	12	12	12	12	12	12	12
Mean	1.65	0.32	1.64	0.32	3.29	0.64	671	132	648	127
S.D.	0.20	0.05	0.23	0.05	0.42	0.09	79	17	102	20

AB : Absolute weight, RE : Relative weight by body weight

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration
Dose : Dodecyloxirane 300 mg/kg

Animal No.	Epididymis-RL		Prostate		Seminal vesicle		LABC		Cowper's gland-R	
	AB mg	RE mg/100g	AB g	RE g/100g	AB g	RE g/100g	AB g	RE g/100g	AB mg	RE mg/100g
3001	1286	215	1.27	0.21	1.63	0.27	1.44	0.24	59.3	9.9
3002	1221	265	1.46	0.32	1.26	0.27	1.65	0.36	74.4	16.2
3003	1027	214	1.14	0.24	1.55	0.32	1.38	0.29	43.9	9.1
3004	1660	316	1.37	0.26	1.78	0.34	1.49	0.28	68.0	13.0
3005	1208	224	1.15	0.21	1.54	0.29	1.32	0.24	60.7	11.2
3006	1155	241	1.26	0.26	1.36	0.28	1.45	0.30	50.1	10.4
3007	1406	299	1.34	0.29	1.54	0.33	1.78	0.38	53.2	11.3
3008	1185	213	0.91	0.16	1.67	0.30	1.52	0.27	38.1	6.8
3009	1458	295	1.03	0.21	1.42	0.29	1.26	0.25	49.0	9.9
3010	1309	273	1.27	0.26	1.98	0.41	1.63	0.34	68.1	14.2
3011	1553	282	0.93	0.17	1.75	0.32	1.72	0.31	68.8	12.5
3012	1361	262	1.55	0.30	1.46	0.28	1.48	0.29	78.1	15.0
n	12	12	12	12	12	12	12	12	12	12
Mean	1319	258	1.22	0.24	1.58	0.31	1.51	0.30	59.3	11.6
S.D.	179	36	0.20	0.05	0.20	0.04	0.16	0.05	12.6	2.7

Animal No.	Cowper's gland-L		Cowper's gland-RL		Glans penis	
	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
3001	54.0	9.0	113.3	18.9	122.1	20.4
3002	59.1	12.8	133.5	29.0	116.9	25.4
3003	49.4	10.3	93.3	19.4	131.2	27.3
3004	84.3	16.1	152.3	29.0	130.9	24.9
3005	80.3	14.9	141.0	26.1	169.3	31.4
3006	56.2	11.7	106.3	22.1	156.7	32.6
3007	62.2	13.2	115.4	24.6	128.9	27.4
3008	40.2	7.2	78.3	14.1	129.5	23.2
3009	41.3	8.3	90.3	18.2	105.0	21.2
3010	81.7	17.0	149.8	31.2	104.6	21.8
3011	81.5	14.8	150.3	27.3	128.5	23.3
3012	74.0	14.3	152.1	29.3	138.5	26.7
n	12	12	12	12	12	12
Mean	63.7	12.5	123.0	24.1	130.2	25.5
S.D.	16.2	3.2	26.9	5.5	18.7	3.8

AB : Absolute weight, RE : Relative weight by body weight
LABC: Levator ani and bulbocavernosus muscle

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Male

Stage : End of administration
Dose : Dodecyloxirane 1000 mg/kg

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
		g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
4001	487	2.08	0.43	9.6	2.0	10.3	2.1	19.9	4.1
4002	478	2.16	0.45	6.6	1.4	10.8	2.3	17.4	3.6
4003	449	2.14	0.48	11.5	2.6	11.0	2.4	22.5	5.0
4004	495	2.23	0.45	13.9	2.8	15.5	3.1	29.4	5.9
4005	481	2.13	0.44	12.5	2.6	10.7	2.2	23.2	4.8
4006	528	2.07	0.39	8.5	1.6	10.2	1.9	18.7	3.5
4007	488	2.08	0.43	13.3	2.7	13.7	2.8	27.0	5.5
n	7	7	7	7	7	7	7	7	7
Mean	487	2.13	0.44	10.8	2.2	11.7	2.4	22.6	4.6
S.D.	23	0.06	0.03	2.7	0.6	2.0	0.4	4.4	0.9

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
4001	301	62	1.32	0.27	14.77	3.03	0.69	0.14	1.64	0.34
4002	224	47	1.37	0.29	13.73	2.87	0.54	0.11	1.46	0.31
4003	333	74	1.25	0.28	12.63	2.81	0.55	0.12	1.42	0.32
4004	360	73	1.30	0.26	14.99	3.03	0.82	0.17	1.50	0.30
4005	212	44	1.29	0.27	15.07	3.13	0.66	0.14	1.61	0.33
4006	342	65	1.48	0.28	15.07	2.85	0.73	0.14	1.82	0.34
4007	400	82	1.31	0.27	14.60	2.99	0.76	0.16	1.65	0.34
n	7	7	7	7	7	7	7	7	7	7
Mean	310	64	1.33	0.27	14.41	2.96	0.68	0.14	1.59	0.33
S.D.	70	14	0.07	0.01	0.91	0.12	0.10	0.02	0.14	0.02

Animal No.	Kidney-L		Kidney-RL		Adrenal -R		Adrenal -L		Adrenal -RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
4001	1.70	0.35	3.34	0.69	28	6	29	6	57	12
4002	1.74	0.36	3.20	0.67	24	5	25	5	49	10
4003	1.48	0.33	2.90	0.65	32	7	30	7	62	14
4004	1.66	0.34	3.16	0.64	25	5	28	6	53	11
4005	1.60	0.33	3.21	0.67	25	5	23	5	48	10
4006	1.96	0.37	3.78	0.72	31	6	32	6	63	12
4007	1.69	0.35	3.34	0.68	29	6	31	6	60	12
n	7	7	7	7	7	7	7	7	7	7
Mean	1.69	0.35	3.28	0.67	28	6	28	6	56	12
S.D.	0.15	0.01	0.27	0.03	3	1	3	1	6	1

AB : Absolute weight, RE : Relative weight by body weight

Animal No.	Testis-R		Testis-L		Testis-RL		Epididymis-R		Epididymis-L	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g
4001	1.57	0.32	1.55	0.32	3.12	0.64	600	123	628	129
4002	1.69	0.35	1.64	0.34	3.33	0.70	683	143	609	127
4003	1.60	0.36	1.59	0.35	3.19	0.71	669	149	612	136
4004	1.80	0.36	1.84	0.37	3.64	0.74	657	133	690	139
4005	1.75	0.36	1.78	0.37	3.53	0.73	642	133	614	128
4006	1.78	0.34	1.75	0.33	3.53	0.67	641	121	650	123
4007	1.60	0.33	1.60	0.33	3.20	0.66	615	126	636	130
n	7	7	7	7	7	7	7	7	7	7
Mean	1.68	0.35	1.68	0.34	3.36	0.69	644	133	634	130
S.D.	0.10	0.02	0.11	0.02	0.20	0.04	29	10	29	5

Animal No.	Epididymis-RL		Prostate		Seminal vesicle		LABC		Cowper's gland-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	mg	mg/100g
4001	1228	252	1.29	0.26	1.93	0.40	1.56	0.32	65.8	13.5
4002	1292	270	1.52	0.32	1.50	0.31	1.61	0.34	52.2	10.9
4003	1281	285	1.07	0.24	1.29	0.29	1.39	0.31	80.2	17.9
4004	1347	272	1.14	0.23	1.60	0.32	1.48	0.30	70.4	14.2
4005	1256	261	1.00	0.21	1.40	0.29	1.31	0.27	64.6	13.4
4006	1291	245	1.21	0.23	1.56	0.30	1.67	0.32	44.3	8.4
4007	1251	256	1.44	0.30	1.80	0.37	1.54	0.32	64.6	13.2
n	7	7	7	7	7	7	7	7	7	7
Mean	1278	263	1.24	0.26	1.58	0.33	1.51	0.31	63.2	13.1
S.D.	38	14	0.19	0.04	0.22	0.04	0.13	0.02	11.8	2.9

Animal No.	Cowper's gland-L		Cowper's gland-RL		Glans penis	
	AB	RE	AB	RE	AB	RE
	mg	mg/100g	mg	mg/100g	mg	mg/100g
4001	65.2	13.4	131.0	26.9	99.4	20.4
4002	62.3	13.0	114.5	24.0	138.4	29.0
4003	76.7	17.1	156.9	34.9	97.4	21.7
4004	69.7	14.1	140.1	28.3	105.7	21.4
4005	49.2	10.2	113.8	23.7	111.0	23.1
4006	61.1	11.6	105.4	20.0	99.3	18.8
4007	60.5	12.4	125.1	25.6	117.3	24.0
n	7	7	7	7	7	7
Mean	63.5	13.1	126.7	26.2	109.8	22.6
S.D.	8.5	2.2	17.7	4.7	14.5	3.3

AB : Absolute weight, RE : Relative weight by body weight
LABC: Levator ani and bulbocavernosus muscle

Organ weight

Stage : End of lactation (F0 dams)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
	g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
1101	352	2.00	0.57	8.4	2.4	8.1	2.3	16.5	4.7
1102	332	1.92	0.58	7.8	2.3	6.6	2.0	14.4	4.3
1103	355	2.03	0.57	7.9	2.2	7.9	2.2	15.8	4.5
1104	374	2.08	0.56	9.7	2.6	8.3	2.2	18.0	4.8
1105	364	1.97	0.54	11.5	3.2	12.0	3.3	23.5	6.5
1106	363	1.95	0.54	8.9	2.5	9.5	2.6	18.4	5.1
1107	330	1.98	0.60	8.4	2.5	8.8	2.7	17.2	5.2
1108	341	2.17	0.64	10.9	3.2	8.1	2.4	19.0	5.6
1109	333	2.15	0.65	10.7	3.2	8.1	2.4	18.8	5.6
1110	300	1.98	0.66	7.9	2.6	8.1	2.7	16.0	5.3
1111	297	2.01	0.68	7.0	2.4	7.2	2.4	14.2	4.8
1112	373	2.01	0.54	8.0	2.1	7.2	1.9	15.2	4.1
n	12	12	12	12	12	12	12	12	12
Mean	343	2.02	0.59	8.9	2.6	8.3	2.4	17.3	5.0
S.D.	26	0.08	0.05	1.4	0.4	1.4	0.4	2.6	0.7

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
1101	156	44	1.19	0.34	11.81	3.36	0.53	0.15	1.20	0.34
1102	251	76	1.06	0.32	10.61	3.20	0.64	0.19	1.17	0.35
1103	148	42	1.15	0.32	12.59	3.55	0.58	0.16	1.14	0.32
1104	291	78	1.15	0.31	12.75	3.41	0.52	0.14	1.18	0.32
1105	284	78	1.20	0.33	10.91	3.00	0.48	0.13	1.22	0.34
1106	365	101	1.20	0.33	12.50	3.44	0.66	0.18	1.17	0.32
1107	225	68	1.10	0.33	11.14	3.38	0.50	0.15	0.97	0.29
1108	282	83	1.15	0.34	11.51	3.38	0.50	0.15	1.03	0.30
1109	330	99	1.01	0.30	10.90	3.27	0.64	0.19	1.20	0.36
1110	186	62	0.96	0.32	10.77	3.59	0.55	0.18	1.09	0.36
1111	203	68	1.10	0.37	11.63	3.92	0.57	0.19	1.04	0.35
1112	230	62	1.17	0.31	12.99	3.48	0.75	0.20	1.22	0.33
n	12	12	12	12	12	12	12	12	12	12
Mean	246	72	1.12	0.33	11.68	3.42	0.58	0.17	1.14	0.33
S.D.	67	18	0.08	0.02	0.85	0.22	0.08	0.02	0.08	0.02

AB : Absolute weight, RE : Relative weight by body weight

Animal No.	Kidney-L		Kidney-RL		Adrenal-R		Adrenal-L		Adrenal-RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
1101	1.24	0.35	2.44	0.69	34	10	39	11	73	21
1102	1.11	0.33	2.28	0.69	37	11	40	12	77	23
1103	1.09	0.31	2.23	0.63	32	9	33	9	65	18
1104	1.16	0.31	2.34	0.63	31	8	32	9	63	17
1105	1.23	0.34	2.45	0.67	32	9	33	9	65	18
1106	1.27	0.35	2.44	0.67	42	12	49	13	91	25
1107	1.03	0.31	2.00	0.61	34	10	35	11	69	21
1108	1.05	0.31	2.08	0.61	40	12	42	12	82	24
1109	1.25	0.38	2.45	0.74	34	10	35	11	69	21
1110	1.06	0.35	2.15	0.72	34	11	38	13	72	24
1111	0.97	0.33	2.01	0.68	31	10	34	11	65	22
1112	1.18	0.32	2.40	0.64	38	10	42	11	80	21
n	12	12	12	12	12	12	12	12	12	12
Mean	1.14	0.33	2.27	0.67	35	10	38	11	73	21
S.D.	0.10	0.02	0.18	0.04	4	1	5	1	8	3

Animal No.	Ovary-R		Ovary-L		Ovary-RL		Uterus	
	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
1101	44.2	12.6	47.7	13.6	91.9	26.1	546	155
1102	47.1	14.2	48.3	14.5	95.4	28.7	576	173
1103	56.7	16.0	40.5	11.4	97.2	27.4	458	129
1104	38.8	10.4	47.6	12.7	86.4	23.1	443	118
1105	46.1	12.7	41.4	11.4	87.5	24.0	483	133
1106	47.1	13.0	60.7	16.7	107.8	29.7	507	140
1107	38.8	11.8	40.5	12.3	79.3	24.0	551	167
1108	54.5	16.0	38.7	11.3	93.2	27.3	569	167
1109	56.4	16.9	47.7	14.3	104.1	31.3	640	192
1110	40.0	13.3	36.9	12.3	76.9	25.6	288	96
1111	43.1	14.5	47.9	16.1	91.0	30.6	523	176
1112	42.6	11.4	51.4	13.8	94.0	25.2	643	172
n	12	12	12	12	12	12	12	12
Mean	46.3	13.6	45.8	13.4	92.1	26.9	519	152
S.D.	6.5	2.0	6.6	1.8	9.0	2.7	96	28

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : End of lactation (F0 dams)

Sex : Female

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
	g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
2101	344	2.09	0.61	9.3	2.7	6.2	1.8	15.5	4.5
2102	337	2.01	0.60	7.1	2.1	8.0	2.4	15.1	4.5
2103	348	2.01	0.58	7.1	2.0	13.0	3.7	20.1	5.8
2104	340	2.06	0.61	9.3	2.7	7.4	2.2	16.7	4.9
2105	333	1.89	0.57	7.2	2.2	7.5	2.3	14.7	4.4
2107	328	2.07	0.63	9.8	3.0	9.8	3.0	19.6	6.0
2108	358	2.14	0.60	10.1	2.8	7.8	2.2	17.9	5.0
2110	381	1.91	0.50	11.2	2.9	9.3	2.4	20.5	5.4
2111	328	2.08	0.63	9.3	2.8	8.8	2.7	18.1	5.5
2112	358	1.95	0.54	7.4	2.1	7.9	2.2	15.3	4.3
n	10	10	10	10	10	10	10	10	10
Mean	346	2.02	0.59	8.8	2.5	8.6	2.5	17.4	5.0
S.D.	17	0.08	0.04	1.5	0.4	1.9	0.5	2.2	0.6

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
2101	219	64	1.12	0.33	12.34	3.59	0.61	0.18	1.17	0.34
2102	296	88	1.30	0.39	12.09	3.59	0.71	0.21	1.31	0.39
2103	168	48	1.07	0.31	12.32	3.54	0.61	0.18	1.19	0.34
2104	244	72	1.38	0.41	12.34	3.63	0.74	0.22	1.19	0.35
2105	172	52	1.04	0.31	11.57	3.47	0.56	0.17	1.06	0.32
2107	277	84	1.18	0.36	11.76	3.59	0.59	0.18	1.06	0.32
2108	308	86	1.22	0.34	12.53	3.50	0.61	0.17	1.08	0.30
2110	187	49	1.03	0.27	13.17	3.46	0.61	0.16	1.36	0.36
2111	312	95	1.05	0.32	11.73	3.58	0.52	0.16	1.31	0.40
2112	168	47	1.06	0.30	11.56	3.23	0.54	0.15	1.12	0.31
n	10	10	10	10	10	10	10	10	10	10
Mean	235	69	1.15	0.33	12.14	3.52	0.61	0.18	1.19	0.34
S.D.	60	19	0.12	0.04	0.51	0.12	0.07	0.02	0.11	0.03

AB : Absolute weight, RE : Relative weight by body weight

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Female

Stage : End of lactation (F0 dams)
Dose : Dodecyloxirane 100 mg/kg

Animal No.	Kidney-L		Kidney-RL		Adrenal-R		Adrenal-L		Adrenal-RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
2101	1.19	0.35	2.36	0.69	37	11	39	11	76	22
2102	1.12	0.33	2.43	0.72	38	11	41	12	79	23
2103	1.17	0.34	2.36	0.68	30	9	33	9	63	18
2104	1.17	0.34	2.36	0.69	33	10	26	8	59	17
2105	1.09	0.33	2.15	0.65	27	8	26	8	53	16
2107	1.13	0.34	2.19	0.67	31	9	37	11	68	21
2108	1.13	0.32	2.21	0.62	37	10	39	11	76	21
2110	1.32	0.35	2.68	0.70	48	13	20	5	68	18
2111	1.19	0.36	2.50	0.76	34	10	36	11	70	21
2112	1.12	0.31	2.24	0.63	29	8	31	9	60	17
n	10	10	10	10	10	10	10	10	10	10
Mean	1.16	0.34	2.35	0.68	34	10	33	10	67	19
S.D.	0.06	0.01	0.16	0.04	6	2	7	2	8	2

Animal No.	Ovary-R		Ovary-L		Ovary-RL		Uterus	
	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
2101	34.4	10.0	51.2	14.9	85.6	24.9	495	144
2102	49.7	14.7	40.7	12.1	90.4	26.8	484	144
2103	44.2	12.7	59.1	17.0	103.3	29.7	435	125
2104	47.0	13.8	41.3	12.1	88.3	26.0	628	185
2105	40.5	12.2	32.4	9.7	72.9	21.9	510	153
2107	31.5	9.6	48.3	14.7	79.8	24.3	477	145
2108	38.3	10.7	39.7	11.1	78.0	21.8	592	165
2110	54.2	14.2	57.3	15.0	111.5	29.3	729	191
2111	41.4	12.6	57.3	17.5	98.7	30.1	502	153
2112	54.0	15.1	33.3	9.3	87.3	24.4	417	116
n	10	10	10	10	10	10	10	10
Mean	43.5	12.6	46.1	13.3	89.6	25.9	527	152
S.D.	7.8	1.9	10.0	2.9	11.9	3.0	95	24

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : End of lactation (F0 dams)

Sex : Female

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
	g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
3101	367	2.07	0.56	10.7	2.9	8.5	2.3	19.2	5.2
3102	334	1.97	0.59	8.6	2.6	7.0	2.1	15.6	4.7
3103	336	2.09	0.62	5.8	1.7	6.3	1.9	12.1	3.6
3104	370	1.96	0.53	7.9	2.1	7.8	2.1	15.7	4.2
3105	393	2.08	0.53	10.5	2.7	10.9	2.8	21.4	5.4
3106	361	2.16	0.60	8.0	2.2	7.0	1.9	15.0	4.2
3107	388	1.97	0.51	10.0	2.6	11.2	2.9	21.2	5.5
3110	405	1.97	0.49	11.9	2.9	11.3	2.8	23.2	5.7
3111	330	1.96	0.59	10.0	3.0	9.0	2.7	19.0	5.8
3112	335	1.98	0.59	9.7	2.9	8.8	2.6	18.5	5.5
n	10	10	10	10	10	10	10	10	10
Mean	362	2.02	0.56	9.3	2.6	8.8	2.4	18.1	5.0
S.D.	27	0.07	0.04	1.8	0.4	1.8	0.4	3.4	0.8

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
3101	313	85	1.12	0.31	12.51	3.41	0.74	0.20	1.10	0.30
3102	288	86	1.11	0.33	11.59	3.47	0.82	0.25	1.04	0.31
3103	245	73	1.23	0.37	13.00	3.87	0.68	0.20	1.19	0.35
3104	156	42	1.02	0.28	12.39	3.35	0.59	0.16	1.18	0.32
3105	355	90	1.58	0.40	14.70	3.74	0.69	0.18	1.26	0.32
3106	276	76	1.20	0.33	13.35	3.70	0.74	0.20	1.26	0.35
3107	257	66	1.10	0.28	13.02	3.36	0.53	0.14	1.22	0.31
3110	263	65	1.26	0.31	14.49	3.58	0.67	0.17	1.13	0.28
3111	189	57	1.21	0.37	12.57	3.81	0.62	0.19	1.27	0.38
3112	179	53	1.00	0.30	12.04	3.59	0.59	0.18	1.11	0.33
n	10	10	10	10	10	10	10	10	10	10
Mean	252	69	1.18	0.33	12.97	3.59	0.67	0.19	1.18	0.33
S.D.	62	16	0.16	0.04	1.00	0.19	0.09	0.03	0.08	0.03

AB : Absolute weight, RE : Relative weight by body weight

Animal No.	Kidney-L		Kidney-RL		Adrenal-R		Adrenal-L		Adrenal-RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
3101	1.06	0.29	2.16	0.59	34	9	41	11	75	20
3102	1.01	0.30	2.05	0.61	32	10	41	12	73	22
3103	1.14	0.34	2.33	0.69	34	10	35	10	69	21
3104	1.20	0.32	2.38	0.64	40	11	43	12	83	22
3105	1.30	0.33	2.56	0.65	38	10	43	11	81	21
3106	1.16	0.32	2.42	0.67	39	11	42	12	81	22
3107	1.20	0.31	2.42	0.62	38	10	42	11	80	21
3110	1.14	0.28	2.27	0.56	43	11	43	11	86	21
3111	1.37	0.42	2.64	0.80	40	12	42	13	82	25
3112	1.14	0.34	2.25	0.67	30	9	37	11	67	20
n	10	10	10	10	10	10	10	10	10	10
Mean	1.17	0.33	2.35	0.65	37	10	41	11	78	22
S.D.	0.10	0.04	0.18	0.07	4	1	3	1	6	1

Animal No.	Ovary-R		Ovary-L		Ovary-RL		Uterus	
	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
3101	44.5	12.1	48.9	13.3	93.4	25.4	631	172
3102	36.0	10.8	50.8	15.2	86.8	26.0	604	181
3103	39.6	11.8	45.8	13.6	85.4	25.4	691	206
3104	28.2	7.6	47.1	12.7	75.3	20.4	573	155
3105	54.2	13.8	60.3	15.3	114.5	29.1	640	163
3106	58.0	16.1	42.7	11.8	100.7	27.9	458	127
3107	36.0	9.3	42.8	11.0	78.8	20.3	435	112
3110	52.0	12.8	37.0	9.1	89.0	22.0	520	128
3111	56.3	17.1	54.1	16.4	110.4	33.5	545	165
3112	47.8	14.3	45.9	13.7	93.7	28.0	676	202
n	10	10	10	10	10	10	10	10
Mean	45.3	12.6	47.5	13.2	92.8	25.8	577	161
S.D.	10.1	2.9	6.5	2.2	12.7	4.1	88	31

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : End of lactation (F0 dams)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
	g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
4102	335	2.04	0.61	6.1	1.8	7.7	2.3	13.8	4.1
4103	302	2.07	0.69	9.9	3.3	9.8	3.2	19.7	6.5
4104	333	2.07	0.62	8.3	2.5	8.4	2.5	16.7	5.0
4105	312	2.09	0.67	10.0	3.2	8.5	2.7	18.5	5.9
4107	324	1.97	0.61	7.2	2.2	7.9	2.4	15.1	4.7
4108	362	2.01	0.56	11.0	3.0	9.3	2.6	20.3	5.6
4109	367	2.07	0.56	11.9	3.2	8.3	2.3	20.2	5.5
4110	343	2.16	0.63	7.3	2.1	5.5	1.6	12.8	3.7
4111	320	1.87	0.58	8.3	2.6	7.2	2.3	15.5	4.8
4112	371	2.13	0.57	10.2	2.7	8.7	2.3	18.9	5.1
n	10	10	10	10	10	10	10	10	10
Mean	337	2.05	0.61	9.0	2.7	8.1	2.4	17.2	5.1
S.D.	24	0.08	0.04	1.9	0.5	1.2	0.4	2.7	0.8

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
4102	325	97	1.03	0.31	13.13	3.92	0.51	0.15	1.03	0.31
4103	245	81	1.12	0.37	11.87	3.93	0.48	0.16	1.07	0.35
4104	117	35	1.08	0.32	13.12	3.94	0.50	0.15	1.00	0.30
4105	306	98	1.28	0.41	11.28	3.62	0.54	0.17	1.09	0.35
4107	301	93	1.11	0.34	12.40	3.83	0.56	0.17	1.02	0.31
4108	309	85	1.11	0.31	13.45	3.72	0.74	0.20	1.18	0.33
4109	221	60	1.23	0.34	13.77	3.75	0.56	0.15	1.15	0.31
4110	226	66	1.28	0.37	15.03	4.38	0.77	0.22	1.36	0.40
4111	284	89	1.06	0.33	12.79	4.00	0.50	0.16	1.01	0.32
4112	193	52	1.19	0.32	15.51	4.18	0.59	0.16	1.25	0.34
n	10	10	10	10	10	10	10	10	10	10
Mean	253	76	1.15	0.34	13.24	3.93	0.58	0.17	1.12	0.33
S.D.	65	21	0.09	0.03	1.31	0.22	0.10	0.02	0.12	0.03

AB : Absolute weight, RE : Relative weight by body weight

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Organ weight
Sex : Female

Stage : End of lactation (F0 dams)
Dose : Dodecyloxirane 1000 mg/kg

Animal No.	Kidney-L		Kidney-RL		Adrenal-R		Adrenal-L		Adrenal-RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
4102	1.11	0.33	2.14	0.64	36	11	35	10	71	21
4103	1.09	0.36	2.16	0.72	36	12	42	14	78	26
4104	1.01	0.30	2.01	0.60	35	11	31	9	66	20
4105	1.13	0.36	2.22	0.71	36	12	40	13	76	24
4107	0.98	0.30	2.00	0.62	31	10	34	10	65	20
4108	1.22	0.34	2.40	0.66	38	10	41	11	79	22
4109	1.13	0.31	2.28	0.62	46	13	45	12	91	25
4110	1.39	0.41	2.75	0.80	42	12	44	13	86	25
4111	1.13	0.35	2.14	0.67	31	10	33	10	64	20
4112	1.30	0.35	2.55	0.69	40	11	44	12	84	23
n	10	10	10	10	10	10	10	10	10	10
Mean	1.15	0.34	2.27	0.67	37	11	39	11	76	23
S.D.	0.12	0.03	0.24	0.06	5	1	5	2	9	2

Animal No.	Ovary-R		Ovary-L		Ovary-RL		Uterus	
	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
4102	28.8	8.6	48.9	14.6	77.7	23.2	496	148
4103	47.7	15.8	43.2	14.3	90.9	30.1	460	152
4104	39.7	11.9	35.0	10.5	74.7	22.4	463	139
4105	34.4	11.0	41.1	13.2	75.5	24.2	470	151
4107	37.0	11.4	52.3	16.1	89.3	27.6	410	127
4108	47.3	13.1	53.3	14.7	100.6	27.8	674	186
4109	54.5	14.9	51.1	13.9	105.6	28.8	611	166
4110	50.7	14.8	45.7	13.3	96.4	28.1	804	234
4111	53.9	16.8	51.3	16.0	105.2	32.9	516	161
4112	52.4	14.1	44.5	12.0	96.9	26.1	565	152
n	10	10	10	10	10	10	10	10
Mean	44.6	13.2	46.6	13.9	91.3	27.1	547	162
S.D.	9.0	2.5	5.8	1.7	11.8	3.2	120	30

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : End of administration (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
		g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
5101	319	1.96	0.61	8.3	2.6	9.4	2.9	17.7	5.5
5102	329	2.18	0.66	7.6	2.3	7.7	2.3	15.3	4.7
5103	279	2.07	0.74	6.7	2.4	11.6	4.2	18.3	6.6
5104	302	1.95	0.65	8.7	2.9	8.1	2.7	16.8	5.6
5105	282	2.01	0.71	10.3	3.7	14.2	5.0	24.5	8.7
n	5	5	5	5	5	5	5	5	5
Mean	302	2.03	0.67	8.3	2.8	10.2	3.4	18.5	6.2
S.D.	22	0.09	0.05	1.3	0.6	2.7	1.1	3.5	1.5

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
5101	314	98	0.98	0.31	8.40	2.63	0.49	0.15	0.86	0.27
5102	281	85	1.02	0.31	8.76	2.66	0.49	0.15	1.06	0.32
5103	352	126	0.90	0.32	7.49	2.68	0.67	0.24	1.01	0.36
5104	460	152	0.90	0.30	8.09	2.68	0.53	0.18	0.99	0.33
5105	310	110	1.00	0.35	7.74	2.74	0.51	0.18	0.90	0.32
n	5	5	5	5	5	5	5	5	5	5
Mean	343	114	0.96	0.32	8.10	2.68	0.54	0.18	0.96	0.32
S.D.	70	26	0.06	0.02	0.51	0.04	0.08	0.04	0.08	0.03

Animal No.	Kidney-L		Kidney-RL		Adrenal-R		Adrenal-L		Adrenal-RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
5101	0.91	0.29	1.77	0.55	34	11	32	10	66	21
5102	1.14	0.35	2.20	0.67	34	10	31	9	65	20
5103	0.91	0.33	1.92	0.69	36	13	21	8	57	20
5104	0.97	0.32	1.96	0.65	32	11	34	11	66	22
5105	0.96	0.34	1.86	0.66	32	11	29	10	61	22
n	5	5	5	5	5	5	5	5	5	5
Mean	0.98	0.33	1.94	0.64	34	11	29	10	63	21
S.D.	0.09	0.02	0.16	0.05	2	1	5	1	4	1

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : End of administration (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Ovary-R		Ovary-L		Ovary-RL		Uterus	
	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
5101	50.9	16.0	46.4	14.5	97.3	30.5	528	166
5102	55.0	16.7	49.6	15.1	104.6	31.8	628	191
5103	41.1	14.7	49.2	17.6	90.3	32.4	520	186
5104	35.0	11.6	30.1	10.0	65.1	21.6	977	324
5105	43.7	15.5	40.6	14.4	84.3	29.9	744	264
n	5	5	5	5	5	5	5	5
Mean	45.1	14.9	43.2	14.3	88.3	29.2	679	226
S.D.	7.9	2.0	8.1	2.7	15.0	4.4	190	66

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : End of administration (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
		g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
6101	332	1.99	0.60	11.0	3.3	10.7	3.2	21.7	6.5
6102	281	1.95	0.69	7.3	2.6	6.1	2.2	13.4	4.8
6103	321	1.98	0.62	8.3	2.6	8.0	2.5	16.3	5.1
6104	310	2.04	0.66	8.6	2.8	6.3	2.0	14.9	4.8
6105	266	1.90	0.71	5.9	2.2	8.4	3.2	14.3	5.4
n	5	5	5	5	5	5	5	5	5
Mean	302	1.97	0.66	8.2	2.7	7.9	2.6	16.1	5.3
S.D.	28	0.05	0.05	1.9	0.4	1.9	0.6	3.3	0.7

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
6101	269	81	0.92	0.28	9.83	2.96	0.51	0.15	1.02	0.31
6102	405	144	0.84	0.30	8.61	3.06	0.41	0.15	0.89	0.32
6103	226	70	0.94	0.29	8.83	2.75	0.47	0.15	1.06	0.33
6104	260	84	0.89	0.29	9.69	3.13	0.59	0.19	0.96	0.31
6105	267	100	0.77	0.29	8.09	3.04	0.42	0.16	0.95	0.36
n	5	5	5	5	5	5	5	5	5	5
Mean	285	96	0.87	0.29	9.01	2.99	0.48	0.16	0.98	0.33
S.D.	69	29	0.07	0.01	0.74	0.15	0.07	0.02	0.07	0.02

Animal No.	Kidney-L		Kidney-RL		Adrenal-R		Adrenal-L		Adrenal-RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
6101	0.97	0.29	1.99	0.60	35	11	35	11	70	21
6102	0.90	0.32	1.79	0.64	30	11	31	11	61	22
6103	1.06	0.33	2.12	0.66	30	9	35	11	65	20
6104	1.06	0.34	2.02	0.65	29	9	36	12	65	21
6105	0.97	0.36	1.92	0.72	30	11	31	12	61	23
n	5	5	5	5	5	5	5	5	5	5
Mean	0.99	0.33	1.97	0.65	31	10	34	11	64	21
S.D.	0.07	0.03	0.12	0.04	2	1	2	1	4	1

AB : Absolute weight, RE : Relative weight by body weight

Stage : End of administration (non-mating group)

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Ovary-R		Ovary-L		Ovary-RL		Uterus	
	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
6101	53.6	16.1	48.1	14.5	101.7	30.6	478	144
6102	45.9	16.3	35.1	12.5	81.0	28.8	1036	369
6103	45.4	14.1	49.8	15.5	95.2	29.7	472	147
6104	45.7	14.7	45.9	14.8	91.6	29.5	811	262
6105	40.8	15.3	35.5	13.3	76.3	28.7	533	200
n	5	5	5	5	5	5	5	5
Mean	46.3	15.3	42.9	14.1	89.2	29.5	666	224
S.D.	4.6	0.9	7.1	1.2	10.4	0.8	249	94

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : End of recovery

Sex : Male

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
	g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
1008	505	2.10	0.42	9.2	1.8	8.6	1.7	17.8	3.5
1009	556	2.13	0.38	12.7	2.3	11.3	2.0	24.0	4.3
1010	559	2.24	0.40	13.7	2.5	7.7	1.4	21.4	3.8
1011	579	2.12	0.37	14.2	2.5	14.5	2.5	28.7	5.0
1012	552	2.20	0.40	11.4	2.1	9.2	1.7	20.6	3.7
n	5	5	5	5	5	5	5	5	5
Mean	550	2.16	0.39	12.2	2.2	10.3	1.9	22.5	4.1
S.D.	27	0.06	0.02	2.0	0.3	2.7	0.4	4.1	0.6

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
1008	361	71	1.49	0.30	14.01	2.77	0.81	0.16	1.75	0.35
1009	340	61	1.63	0.29	15.46	2.78	0.76	0.14	1.73	0.31
1010	324	58	1.67	0.30	14.74	2.64	0.91	0.16	1.80	0.32
1011	347	60	1.47	0.25	14.31	2.47	0.77	0.13	1.77	0.31
1012	277	50	1.48	0.27	13.34	2.42	0.61	0.11	1.62	0.29
n	5	5	5	5	5	5	5	5	5	5
Mean	330	60	1.55	0.28	14.37	2.62	0.77	0.14	1.73	0.32
S.D.	32	8	0.09	0.02	0.79	0.17	0.11	0.02	0.07	0.02

Animal No.	Kidney-L		Kidney-RL		Adrenal -R		Adrenal -L		Adrenal -RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
1008	1.82	0.36	3.57	0.71	26	5	27	5	53	10
1009	1.71	0.31	3.44	0.62	33	6	33	6	66	12
1010	1.84	0.33	3.64	0.65	29	5	30	5	59	11
1011	1.84	0.32	3.61	0.62	31	5	32	6	63	11
1012	1.79	0.32	3.41	0.62	42	8	44	8	86	16
n	5	5	5	5	5	5	5	5	5	5
Mean	1.80	0.33	3.53	0.64	32	6	33	6	65	12
S.D.	0.05	0.02	0.10	0.04	6	1	6	1	13	2

AB : Absolute weight, RE : Relative weight by body weight

Animal No.	Testis-R		Testis-L		Testis-RL		Epididymis-R		Epididymis-L	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g
1008	1.55	0.31	1.52	0.30	3.07	0.61	747	148	707	140
1009	1.57	0.28	1.62	0.29	3.19	0.57	645	116	644	116
1010	1.57	0.28	1.50	0.27	3.07	0.55	647	116	599	107
1011	1.77	0.31	1.79	0.31	3.56	0.61	827	143	753	130
1012	1.65	0.30	1.76	0.32	3.41	0.62	690	125	738	134
n	5	5	5	5	5	5	5	5	5	5
Mean	1.62	0.30	1.64	0.30	3.26	0.59	711	130	688	125
S.D.	0.09	0.02	0.13	0.02	0.22	0.03	77	15	65	14
Animal No.	Epididymis-RL		Prostate		Seminal vesicle		LABC		Cowper's gland-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	mg	mg/100g
1008	1454	288	1.35	0.27	1.97	0.39	1.59	0.31	73.8	14.6
1009	1289	232	1.26	0.23	2.04	0.37	1.66	0.30	73.0	13.1
1010	1246	223	1.24	0.22	1.87	0.33	1.91	0.34	88.9	15.9
1011	1580	273	1.83	0.32	2.09	0.36	1.74	0.30	84.4	14.6
1012	1428	259	1.65	0.30	1.62	0.29	1.64	0.30	66.7	12.1
n	5	5	5	5	5	5	5	5	5	5
Mean	1399	255	1.47	0.27	1.92	0.35	1.71	0.31	77.4	14.1
S.D.	134	27	0.26	0.04	0.19	0.04	0.13	0.02	9.1	1.5
Animal No.	Cowper's gland-L		Cowper's gland-RL		Glans penis					
	AB	RE	AB	RE	AB	RE				
	mg	mg/100g	mg	mg/100g	mg	mg/100g				
1008	72.0	14.3	145.8	28.9	95.8	19.0				
1009	66.6	12.0	139.6	25.1	162.9	29.3				
1010	96.1	17.2	185.0	33.1	126.1	22.6				
1011	77.1	13.3	161.5	27.9	129.4	22.3				
1012	72.6	13.2	139.3	25.2	104.2	18.9				
n	5	5	5	5	5	5				
Mean	76.9	14.0	154.2	28.0	123.7	22.4				
S.D.	11.4	2.0	19.4	3.3	26.1	4.2				

AB : Absolute weight, RE : Relative weight by body weight
LABC: Levator ani and bulbocavernosus muscle

Organ weight

Stage : End of recovery

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
		g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
4008	562	2.20	0.39	13.7	2.4	12.7	2.3	26.4	4.7
4009	572	2.16	0.38	8.0	1.4	9.4	1.6	17.4	3.0
4010	458	2.17	0.47	14.0	3.1	10.7	2.3	24.7	5.4
4011	537	2.25	0.42	15.5	2.9	11.5	2.1	27.0	5.0
4012	542	2.24	0.41	14.3	2.6	10.2	1.9	24.5	4.5
n	5	5	5	5	5	5	5	5	5
Mean	534	2.20	0.41	13.1	2.5	10.9	2.0	24.0	4.5
S.D.	45	0.04	0.04	2.9	0.7	1.3	0.3	3.8	0.9

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
4008	393	70	1.40	0.25	15.37	2.73	0.84	0.15	1.73	0.31
4009	359	63	1.48	0.26	16.15	2.82	0.78	0.14	2.03	0.35
4010	224	49	1.32	0.29	11.84	2.59	0.61	0.13	1.48	0.32
4011	312	58	1.68	0.31	14.54	2.71	0.80	0.15	1.84	0.34
4012	332	61	1.73	0.32	15.54	2.87	0.83	0.15	1.72	0.32
n	5	5	5	5	5	5	5	5	5	5
Mean	324	60	1.52	0.29	14.69	2.74	0.77	0.14	1.76	0.33
S.D.	64	8	0.18	0.03	1.69	0.11	0.09	0.01	0.20	0.02

Animal No.	Kidney-L		Kidney-RL		Adrenal -R		Adrenal -L		Adrenal -RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
4008	1.71	0.30	3.44	0.61	27	5	29	5	56	10
4009	2.14	0.37	4.17	0.73	28	5	25	4	53	9
4010	1.55	0.34	3.03	0.66	29	6	29	6	58	13
4011	2.07	0.39	3.91	0.73	43	8	41	8	84	16
4012	1.85	0.34	3.57	0.66	31	6	31	6	62	11
n	5	5	5	5	5	5	5	5	5	5
Mean	1.86	0.35	3.62	0.68	32	6	31	6	63	12
S.D.	0.25	0.03	0.44	0.05	7	1	6	1	12	3

AB : Absolute weight, RE : Relative weight by body weight

	Testis-R		Testis-L		Testis-RL		Epididymis-R		Epididymis-L	
Animal No.	AB g	RE g/100g	AB g	RE g/100g	AB g	RE g/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
4008	1.40	0.25	1.44	0.26	2.84	0.51	677	120	628	112
4009	1.74	0.30	1.71	0.30	3.45	0.60	693	121	684	120
4010	1.74	0.38	1.74	0.38	3.48	0.76	732	160	691	151
4011	1.52	0.28	1.60	0.30	3.12	0.58	635	118	665	124
4012	1.90	0.35	1.91	0.35	3.81	0.70	678	125	711	131
n	5	5	5	5	5	5	5	5	5	5
Mean	1.66	0.31	1.68	0.32	3.34	0.63	683	129	676	128
S.D.	0.20	0.05	0.17	0.05	0.37	0.10	35	18	31	15

	Epididymis-RL		Prostate		Seminal vesicle		LABC		Cowper's gland-R	
Animal No.	AB mg	RE mg/100g	AB g	RE g/100g	AB g	RE g/100g	AB g	RE g/100g	AB mg	RE mg/100g
4008	1305	232	1.09	0.19	1.38	0.25	1.42	0.25	57.9	10.3
4009	1377	241	1.52	0.27	1.78	0.31	1.54	0.27	57.2	10.0
4010	1423	311	1.39	0.30	1.90	0.41	1.62	0.35	65.0	14.2
4011	1300	242	1.35	0.25	1.44	0.27	1.67	0.31	84.3	15.7
4012	1389	256	1.39	0.26	1.82	0.34	1.75	0.32	68.9	12.7
n	5	5	5	5	5	5	5	5	5	5
Mean	1359	256	1.35	0.25	1.66	0.32	1.60	0.30	66.7	12.6
S.D.	54	32	0.16	0.04	0.24	0.06	0.13	0.04	11.0	2.5

	Cowper's gland-L		Cowper's gland-RL		Glans penis	
Animal No.	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
4008	66.5	11.8	124.4	22.1	124.3	22.1
4009	61.9	10.8	119.1	20.8	125.9	22.0
4010	53.8	11.7	118.8	25.9	123.1	26.9
4011	97.6	18.2	181.9	33.9	162.0	30.2
4012	69.3	12.8	138.2	25.5	193.7	35.7
n	5	5	5	5	5	5
Mean	69.8	13.1	136.5	25.6	145.8	27.4
S.D.	16.6	3.0	26.6	5.1	31.3	5.8

AB : Absolute weight, RE : Relative weight by body weight
LABC: Levator ani and bulbocavernosus muscle

Organ weight

Stage : End of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
		g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
5106	302	2.04	0.68	7.9	2.6	7.8	2.6	15.7	5.2
5107	337	2.04	0.61	7.8	2.3	8.5	2.5	16.3	4.8
5108	299	2.00	0.67	9.4	3.1	5.8	1.9	15.2	5.1
5109	288	1.85	0.64	6.7	2.3	6.9	2.4	13.6	4.7
5110	293	2.12	0.72	9.5	3.2	7.5	2.6	17.0	5.8
n	5	5	5	5	5	5	5	5	5
Mean	304	2.01	0.66	8.3	2.7	7.3	2.4	15.6	5.1
S.D.	19	0.10	0.04	1.2	0.4	1.0	0.3	1.3	0.4

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
5106	253	84	0.98	0.32	7.32	2.42	0.49	0.16	0.86	0.28
5107	307	91	1.11	0.33	9.72	2.88	0.50	0.15	1.11	0.33
5108	199	67	1.08	0.36	7.03	2.35	0.48	0.16	0.91	0.30
5109	266	92	1.08	0.38	8.08	2.81	0.37	0.13	0.97	0.34
5110	224	76	0.88	0.30	7.04	2.40	0.46	0.16	0.99	0.34
n	5	5	5	5	5	5	5	5	5	5
Mean	250	82	1.03	0.34	7.84	2.57	0.46	0.15	0.97	0.32
S.D.	41	11	0.10	0.03	1.14	0.25	0.05	0.01	0.09	0.03

Animal No.	Kidney-L		Kidney-RL		Adrenal -R		Adrenal -L		Adrenal -RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
5106	0.89	0.29	1.75	0.58	32	11	35	12	67	22
5107	1.20	0.36	2.31	0.69	33	10	36	11	69	20
5108	0.90	0.30	1.81	0.61	33	11	37	12	70	23
5109	0.93	0.32	1.90	0.66	33	11	35	12	68	24
5110	1.04	0.35	2.03	0.69	30	10	34	12	64	22
n	5	5	5	5	5	5	5	5	5	5
Mean	0.99	0.32	1.96	0.65	32	11	35	12	68	22
S.D.	0.13	0.03	0.22	0.05	1	1	1	0	2	1

AB : Absolute weight, RE : Relative weight by body weight

Animal No.	Ovary-R		Ovary-L		Ovary-RL		Uterus	
	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
5106	49.6	16.4	54.4	18.0	104.0	34.4	539	178
5107	22.2	6.6	33.5	9.9	55.7	16.5	787	234
5108	39.3	13.1	33.7	11.3	73.0	24.4	486	163
5109	50.5	17.5	50.6	17.6	101.1	35.1	752	261
5110	45.6	15.6	52.7	18.0	98.3	33.5	531	181
n	5	5	5	5	5	5	5	5
Mean	41.4	13.8	45.0	15.0	86.4	28.8	619	203
S.D.	11.6	4.4	10.5	4.0	21.2	8.1	139	42

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : End of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Body weight	Brain		Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB	RE	AB	RE	AB	RE	AB	RE
	g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
6106	301	2.06	0.68	11.8	3.9	9.4	3.1	21.2	7.0
6107	268	2.07	0.77	10.8	4.0	8.2	3.1	19.0	7.1
6108	305	1.95	0.64	11.8	3.9	10.7	3.5	22.5	7.4
6109	299	2.08	0.70	10.4	3.5	9.6	3.2	20.0	6.7
6110	298	2.03	0.68	10.1	3.4	8.1	2.7	18.2	6.1
n	5	5	5	5	5	5	5	5	5
Mean	294	2.04	0.69	11.0	3.7	9.2	3.1	20.2	6.9
S.D.	15	0.05	0.05	0.8	0.3	1.1	0.3	1.7	0.5

Animal No.	Thymus		Heart		Liver		Spleen		Kidney-R	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	mg	mg/100g	g	g/100g	g	g/100g	g	g/100g	g	g/100g
6106	237	79	0.96	0.32	7.74	2.57	0.47	0.16	1.04	0.35
6107	242	90	0.94	0.35	6.43	2.40	0.48	0.18	0.89	0.33
6108	342	112	0.96	0.31	7.71	2.53	0.39	0.13	1.03	0.34
6109	423	141	0.92	0.31	7.07	2.36	0.51	0.17	0.97	0.32
6110	263	88	0.92	0.31	7.29	2.45	0.45	0.15	1.01	0.34
n	5	5	5	5	5	5	5	5	5	5
Mean	301	102	0.94	0.32	7.25	2.46	0.46	0.16	0.99	0.34
S.D.	80	25	0.02	0.02	0.54	0.09	0.04	0.02	0.06	0.01

Animal No.	Kidney-L		Kidney-RL		Adrenal -R		Adrenal -L		Adrenal -RL	
	AB	RE	AB	RE	AB	RE	AB	RE	AB	RE
	g	g/100g	g	g/100g	mg	mg/100g	mg	mg/100g	mg	mg/100g
6106	1.03	0.34	2.07	0.69	34	11	37	12	71	24
6107	0.89	0.33	1.78	0.66	27	10	30	11	57	21
6108	1.08	0.35	2.11	0.69	33	11	39	13	72	24
6109	1.03	0.34	2.00	0.67	25	8	32	11	57	19
6110	1.06	0.36	2.07	0.69	31	10	35	12	66	22
n	5	5	5	5	5	5	5	5	5	5
Mean	1.02	0.34	2.01	0.68	30	10	35	12	65	22
S.D.	0.07	0.01	0.13	0.01	4	1	4	1	7	2

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : End of recovery (non-mating group)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Ovary-R		Ovary-L		Ovary-RL		Uterus	
	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
6106	56.0	18.6	43.3	14.4	99.3	33.0	684	227
6107	33.5	12.5	32.4	12.1	65.9	24.6	476	178
6108	42.6	14.0	44.3	14.5	86.9	28.5	531	174
6109	43.2	14.4	37.2	12.4	80.4	26.9	844	282
6110	44.8	15.0	47.1	15.8	91.9	30.8	895	300
n	5	5	5	5	5	5	5	5
Mean	44.0	14.9	40.9	13.8	84.9	28.8	686	232
S.D.	8.0	2.3	6.0	1.6	12.7	3.3	185	58

AB : Absolute weight, RE : Relative weight by body weight

Appendix 9-1(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1001 Male 0 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Pancreas Focus,basophilic: minimal

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine,duodenum, Intestine,jejunum
Intestine,ileum, Intestine,cecum, Intestine,colon, Intestine,rectum, Kidney
Lymph node,mesenteric, Lymph node,submandibular, Liver, Lung(bronchus)
Mammary gland,inguinal, Medulla oblongata, Muscle,quadriceps femoris
Optic nerve, Parathyroid, Pituitary, Prostate, Peyer's patch,ileal
Salivary gland,submandibular, Skin,inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord,thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-2 (1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1002 Male 0 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain (cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung (bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Prostate, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-3 (1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1003 Male 0 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Prostate Infiltrate, inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain (cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung (bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Animal No. 1004 Male 0 mg/kg Day 43 End of dosing

All tissues Not remarkable

Following tissues : Not remarkable

881

Animal No. 1005 Male 0 mg/kg Day 43 End of dosing

All tissues Not remarkable

Following tissues : Not remarkable

882

Animal No. 1006 Male 0 mg/kg Day 43 End of dosing

All tissues	Not remarkable
-------------	----------------

No tissues examined

Appendix 9-7(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1007 Male 0 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-8(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2001 Male 100 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Animal No. 2002 Male 100 mg/kg Day 43 End of dosing

All tissues Not remarkable

No tissues examined

Animal No. 2003 Male 100 mg/kg Day 43 End of dosing

All tissues Not remarkable

No tissues examined

Animal No. 2004 Male 100 mg/kg Day 43 End of dosing

All tissues	Not remarkable
-------------	----------------

No tissues examined

Appendix 9-12(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2005 Male 100 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-13(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2006 Male 100 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine,duodenum, Intestine,jejunum
Intestine,ileum, Intestine,cecum, Intestine,colon, Intestine,rectum, Kidney
Lymph node,mesenteric, Lymph node,submandibular, Liver, Lung(bronchus)
Mammary gland,inguinal, Medulla oblongata, Muscle,quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Prostate, Peyer's patch,ileal
Salivary gland,submandibular, Skin,inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord,thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Animal No. 2007 Male 100 mg/kg Day 43 End of dosing

All tissues Not remarkable

No tissues examined

Appendix 9-15(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2008 Male 100 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-16(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2009 Male 100 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Prostate Infiltrate, inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung(bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-17(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2010 Male 100 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-18(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2011 Male 100 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-19(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2012 Male 100 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-20(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3001 Male 300 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Animal No. 3002 Male 300 mg/kg Day 43 End of dosing

All tissues Not remarkable

No tissues examined

Animal No. 3003 Male 300 mg/kg Day 43 End of dosing

All tissues	Not remarkable
-------------	----------------

No tissues examined

Appendix 9-23(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3004 Male 300 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Animal No. 3005 Male 300 mg/kg Day 43 End of dosing

All tissues	Not remarkable
-------------	----------------

No tissues examined

Appendix 9-25(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3006 Male 300 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-26(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3007 Male 300 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-27(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3008 Male 300 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine,duodenum, Intestine,jejunum
Intestine,ileum, Intestine,cecum, Intestine,colon, Intestine,rectum, Kidney
Lymph node,mesenteric, Lymph node,submandibular, Liver, Lung(bronchus)
Mammary gland,inguinal, Medulla oblongata, Muscle,quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Prostate, Peyer's patch,ileal
Salivary gland,submandibular, Skin,inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord,thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-28(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3009 Male 300 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Prostate Infiltrate, inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung(bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-29(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3010 Male 300 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Animal No. 3011 Male 300 mg/kg Day 43 End of dosing

All tissues Not remarkable

No tissues examined

Animal No. 3012 Male 300 mg/kg Day 43 End of dosing

All tissues Not remarkable

No tissues examined

Appendix 9-32(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4001 Male 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Prostate Infiltrate, inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung(bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-33(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4002 Male 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Stomach Erosion/ulcer,glandular stomach: minimal

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine,duodenum, Intestine,jejunum
Intestine,ileum, Intestine,cecum, Intestine,colon, Intestine,rectum, Kidney
Lymph node,mesenteric, Lymph node,submandibular, Liver, Lung(bronchus)
Mammary gland,inguinal, Medulla oblongata, Muscle,quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Prostate, Peyer's patch,ileal
Salivary gland,submandibular, Skin,inguinal, Sciatic nerve, Spleen
Seminal vesicle, Spinal cord,thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-34(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4003 Male 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Prostate Infiltrate, inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung(bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-35(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4004 Male 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Prostate Infiltrate, inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung(bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-36(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4005 Male 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine,duodenum, Intestine,jejunum
Intestine,ileum, Intestine,cecum, Intestine,colon, Intestine,rectum, Kidney
Lymph node,mesenteric, Lymph node,submandibular, Liver, Lung(bronchus)
Mammary gland,inguinal, Medulla oblongata, Muscle,quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Prostate, Peyer's patch,ileal
Salivary gland,submandibular, Skin,inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord,thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Animal No. 4006 Male 1000 mg/kg Day 43 End of dosing

All tissues Not remarkable

No tissues examined

Appendix 9-38(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4007 Male 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-39(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1101 Female 0 mg/kg Day 52 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-40(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1102 Female 0 mg/kg Day 62 End of dosing, Dams

Gross pathology:

Stomach Focus, dark red, glandular stomach: 2 present
each 2x1mm

Other tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-41(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1103 Female 0 mg/kg Day 54 End of dosing, Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-42(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1104 Female 0 mg/kg Day 55 End of dosing, Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-43(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1105 Female 0 mg/kg Day 52 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

Ovary Cyst,luteal: minimal, unilateral

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Parathyroid, Pancreas
Pituitary, Peyer's patch,ileal, Salivary gland,submandibular, Skin,inguinal
Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus, Trachea, Thyroid
Urinary bladder, Uterus, Vagina

Appendix 9-44(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1106 Female 0 mg/kg Day 54 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-45(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1107 Female 0 mg/kg Day 51 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-46(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1108 Female 0 mg/kg Day 52 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-47(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1109 Female 0 mg/kg Day 55 End of dosing, Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-48(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1110 Female 0 mg/kg Day 55 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-49(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1111 Female 0 mg/kg Day 53 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-50(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1112 Female 0 mg/kg Day 54 End of dosing, Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-51(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2101 Female 100 mg/kg Day 54 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-52(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2102 Female 100 mg/kg Day 54 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-53(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2103 Female 100 mg/kg Day 55 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Animal No. 2104 Female 100 mg/kg Day 53 End of dosing, Dams

All tissues	Not remarkable
-------------	----------------

No tissues examined

Appendix 9-55(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2105 Female 100 mg/kg Day 52 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-56(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2106 Female 100 mg/kg Day 41 Undelivered

Gross pathology:

All tissues Not remarkable

Histopathology:

Pancreas Focus,basophilic: minimal

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pituitary, Peyer's patch,ileal, Salivary gland,submandibular, Skin,inguinal
Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus, Trachea, Thyroid
Urinary bladder, Uterus, Vagina

Animal No. 2107 Female 100 mg/kg Day 53 End of dosing, Dams

All tissues	Not remarkable
-------------	----------------

No tissues examined

Appendix 9-58(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2108 Female 100 mg/kg Day 51 End of dosing, Dams

Gross pathology:

Stomach Focus, dark red, glandular stomach: 1 present
3x1mm

Other tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-59(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2109 Female 100 mg/kg Day 54 Undelivered

Gross pathology:

All tissues Not remarkable

Histopathology:

Kidney Mineralization,pelvis: minimal, unilateral

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-60(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2110 Female 100 mg/kg Day 55 End of dosing,Dams

Gross pathology:

Stomach Focus, dark red, glandular stomach: 2 present
each 1x1mm

Other tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-61(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 2111 Female 100 mg/kg Day 56 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Animal No. 2112 Female 100 mg/kg Day 53 End of dosing, Dams

All tissues	Not remarkable
-------------	----------------

No tissues examined

Appendix 9-63(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3101 Female 300 mg/kg Day 52 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Animal No. 3102 Female 300 mg/kg Day 52 End of dosing, Dams

All tissues Not remarkable

No tissues examined

Appendix 9-65(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3103 Female 300 mg/kg Day 53 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-66(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3104 Female 300 mg/kg Day 52 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-67(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3105 Female 300 mg/kg Day 56 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-68(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3106 Female 300 mg/kg Day 55 End of dosing, Dams

Gross pathology:

Stomach Focus, dark red, glandular stomach: 2 present
each 1x1mm

Other tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-69(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3107 Female 300 mg/kg Day 65 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-70(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3108 Female 300 mg/kg Day 54 Undelivered

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-71(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3109 Female 300 mg/kg Day 42 Undelivered

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-72(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3110 Female 300 mg/kg Day 52 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-73(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 3111 Female 300 mg/kg Day 51 End of dosing,Dams

Gross pathology:

Stomach Focus, dark red, glandular stomach: 1 present
1x2mm

Other tissues Not remarkable

Histopathology:

No tissues examined

Animal No. 3112 Female 300 mg/kg Day 53 End of dosing, Dams

All tissues Not remarkable

No tissues examined

Appendix 9-75(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4101 Female 1000 mg/kg Day 42 Undelivered

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-76(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4102 Female 1000 mg/kg Day 52 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-77(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4103 Female 1000 mg/kg Day 52 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Animal No. 4104 Female 1000 mg/kg Day 55 End of dosing, Dams

All tissues	Not remarkable
-------------	----------------

No tissues examined

Appendix 9-79(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4105 Female 1000 mg/kg Day 55 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-80(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4106 Female 1000 mg/kg Day 41 Undelivered

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-81(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4107 Female 1000 mg/kg Day 54 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

Liver Necrosis,focal: minimal

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-82(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4108 Female 1000 mg/kg Day 54 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-83(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4109 Female 1000 mg/kg Day 53 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-84(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with
the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4110 Female 1000 mg/kg Day 55 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

No tissues examined

Appendix 9-85(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4111 Female 1000 mg/kg Day 52 End of dosing,Dams

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Animal No. 4112 Female 1000 mg/kg Day 65 End of dosing, Dams

All tissues Not remarkable

No tissues examined

Appendix 9-87(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 5101 Female 0 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons), Eye
Heart, Intestine, duodenum, Intestine, jejunum, Intestine, ileum, Intestine, cecum
Intestine, colon, Intestine, rectum, Kidney, Lymph node, mesenteric
Lymph node, submandibular, Liver, Lung(bronchus), Mammary gland, inguinal
Medulla oblongata, Muscle, quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch, ileal, Salivary gland, submandibular
Skin, inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord, thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-88(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 5102 Female 0 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons), Eye
Heart, Intestine, duodenum, Intestine, jejunum, Intestine, ileum, Intestine, cecum
Intestine, colon, Intestine, rectum, Kidney, Lymph node, mesenteric
Lymph node, submandibular, Liver, Lung(bronchus), Mammary gland, inguinal
Medulla oblongata, Muscle, quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch, ileal, Salivary gland, submandibular
Skin, inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord, thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-89(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 5103 Female 0 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Adrenal Mineralization: minimal, unilateral
with pigmented macrophage

Pancreas Focus,basophilic: minimal

Following tissues : Not remarkable

Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye, Heart
Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pituitary, Peyer's patch,ileal, Salivary gland,submandibular, Skin,inguinal
Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus, Trachea, Thyroid
Urinary bladder, Uterus, Vagina

Appendix 9-90(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 5104 Female 0 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons), Eye
Heart, Intestine, duodenum, Intestine, jejunum, Intestine, ileum, Intestine, cecum
Intestine, colon, Intestine, rectum, Kidney, Lymph node, mesenteric
Lymph node, submandibular, Liver, Lung(bronchus), Mammary gland, inguinal
Medulla oblongata, Muscle, quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch, ileal, Salivary gland, submandibular
Skin, inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord, thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-91(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 5105 Female 0 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons), Eye
Heart, Intestine, duodenum, Intestine, jejunum, Intestine, ileum, Intestine, cecum
Intestine, colon, Intestine, rectum, Kidney, Lymph node, mesenteric
Lymph node, submandibular, Liver, Lung(bronchus), Mammary gland, inguinal
Medulla oblongata, Muscle, quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch, ileal, Salivary gland, submandibular
Skin, inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord, thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-92(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 6101 Female 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-93(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 6102 Female 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons), Eye
Heart, Intestine, duodenum, Intestine, jejunum, Intestine, ileum, Intestine, cecum
Intestine, colon, Intestine, rectum, Kidney, Lymph node, mesenteric
Lymph node, submandibular, Liver, Lung(bronchus), Mammary gland, inguinal
Medulla oblongata, Muscle, quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch, ileal, Salivary gland, submandibular
Skin, inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord, thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-94(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 6103 Female 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-95(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 6104 Female 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-96(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 6105 Female 1000 mg/kg Day 43 End of dosing

Gross pathology:

All tissues Not remarkable

Histopathology:

Heart Infiltrate/Fibrosis: minimal

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-97(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1008 Male 0 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Prostate Infiltrate, inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung(bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-98(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1009 Male 0 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine,duodenum, Intestine,jejunum
Intestine,ileum, Intestine,cecum, Intestine,colon, Intestine,rectum, Kidney
Lymph node,mesenteric, Lymph node,submandibular, Liver, Lung(bronchus)
Mammary gland,inguinal, Medulla oblongata, Muscle,quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Prostate, Peyer's patch,ileal
Salivary gland,submandibular, Skin,inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord,thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-99(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1010 Male 0 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Heart Infiltrate/Fibrosis: minimal

Prostate Infiltrate,inflammatory cell: mild

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons)
Coagulating gland, Epididymis, Eye, Intestine,duodenum, Intestine,jejunum
Intestine,ileum, Intestine,cecum, Intestine,colon, Intestine,rectum, Kidney
Lymph node,mesenteric, Lymph node,submandibular, Liver, Lung(bronchus)
Mammary gland,inguinal, Medulla oblongata, Muscle,quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch,ileal
Salivary gland,submandibular, Skin,inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord,thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-100(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1011 Male 0 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Eye Retinal rosette: minimal

Prostate Infiltrate, inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung(bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-101(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 1012 Male 0 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Parathyroid No sample

Prostate Infiltrate, inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung(bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-102(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4008 Male 1000 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Heart Infiltrate/Fibrosis: minimal

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons)
Coagulating gland, Epididymis, Eye, Intestine,duodenum, Intestine,jejunum
Intestine,ileum, Intestine,cecum, Intestine,colon, Intestine,rectum, Kidney
Lymph node,mesenteric, Lymph node,submandibular, Liver, Lung(bronchus)
Mammary gland,inguinal, Medulla oblongata, Muscle,quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Prostate, Peyer's patch,ileal
Salivary gland,submandibular, Skin,inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord,thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-103(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4009 Male 1000 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Testis Atrophy,tubule: minimal, unilateral

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine,duodenum, Intestine,jejunum
Intestine,ileum, Intestine,cecum, Intestine,colon, Intestine,rectum, Kidney
Lymph node,mesenteric, Lymph node,submandibular, Liver, Lung(bronchus)
Mammary gland,inguinal, Medulla oblongata, Muscle,quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Prostate, Peyer's patch,ileal
Salivary gland,submandibular, Skin,inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord,thoracic, Thymus, Trachea, Thyroid, Urinary bladder

Appendix 9-104(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4010 Male 1000 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Heart Infiltrate/Fibrosis: minimal

Prostate Infiltrate,inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons)
Coagulating gland, Epididymis, Eye, Intestine,duodenum, Intestine,jejunum
Intestine,ileum, Intestine,cecum, Intestine,colon, Intestine,rectum, Kidney
Lymph node,mesenteric, Lymph node,submandibular, Liver, Lung(bronchus)
Mammary gland,inguinal, Medulla oblongata, Muscle,quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch,ileal
Salivary gland,submandibular, Skin,inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord,thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-105(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4011 Male 1000 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Prostate Infiltrate, inflammatory cell: minimal

Testis Atrophy, tubule: minimal, unilateral

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung(bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Thymus, Trachea, Thyroid, Urinary bladder

Appendix 9-106(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 4012 Male 1000 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Prostate Infiltrate, inflammatory cell: minimal

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons)
Coagulating gland, Epididymis, Eye, Heart, Intestine, duodenum, Intestine, jejunum
Intestine, ileum, Intestine, cecum, Intestine, colon, Intestine, rectum, Kidney
Lymph node, mesenteric, Lymph node, submandibular, Liver, Lung(bronchus)
Mammary gland, inguinal, Medulla oblongata, Muscle, quadriceps femoris
Optic nerve, Parathyroid, Pancreas, Pituitary, Peyer's patch, ileal
Salivary gland, submandibular, Skin, inguinal, Sciatic nerve, Spleen, Stomach
Seminal vesicle, Spinal cord, thoracic, Testis, Thymus, Trachea, Thyroid
Urinary bladder

Appendix 9-107(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 5106 Female 0 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons), Eye
Heart, Intestine, duodenum, Intestine, jejunum, Intestine, ileum, Intestine, cecum
Intestine, colon, Intestine, rectum, Kidney, Lymph node, mesenteric
Lymph node, submandibular, Liver, Lung(bronchus), Mammary gland, inguinal
Medulla oblongata, Muscle, quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch, ileal, Salivary gland, submandibular
Skin, inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord, thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-108(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 5107 Female 0 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Kidney Urinary cast,hyaline: minimal, bilateral
Fibrosis,interstitium: minimal, unilateral

Ovary Age-related atrophy: mild, bilateral

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Parathyroid, Pancreas
Pituitary, Peyer's patch,ileal, Salivary gland,submandibular, Skin,inguinal
Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus, Trachea, Thyroid
Urinary bladder, Uterus, Vagina

Appendix 9-109(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 5108 Female 0 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Kidney Urinary cast,hyaline: minimal, unilateral

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-110(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 5109 Female 0 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons), Eye
Heart, Intestine, duodenum, Intestine, jejunum, Intestine, ileum, Intestine, cecum
Intestine, colon, Intestine, rectum, Kidney, Lymph node, mesenteric
Lymph node, submandibular, Liver, Lung(bronchus), Mammary gland, inguinal
Medulla oblongata, Muscle, quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch, ileal, Salivary gland, submandibular
Skin, inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord, thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-111(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 5110 Female 0 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-112(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 6106 Female 1000 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-113(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 6107 Female 1000 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone, femur, Bone marrow, femur, Brain(cerebrum, cerebellum, pons), Eye
Heart, Intestine, duodenum, Intestine, jejunum, Intestine, ileum, Intestine, cecum
Intestine, colon, Intestine, rectum, Kidney, Lymph node, mesenteric
Lymph node, submandibular, Liver, Lung(bronchus), Mammary gland, inguinal
Medulla oblongata, Muscle, quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch, ileal, Salivary gland, submandibular
Skin, inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord, thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-114(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 6108 Female 1000 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-115(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 6109 Female 1000 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Eye
Heart, Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

Appendix 9-116(1/1) Dodecyloxirane: An oral combined repeated dose toxicity study with the reproduction/ developmental toxicity screening test in rats

Individual gross and histopathological findings

Animal No. 6110 Female 1000 mg/kg Day 57 End of recovery

Gross pathology:

All tissues Not remarkable

Histopathology:

Eye Retinal fold: minimal

Following tissues : Not remarkable

Adrenal, Bone,femur, Bone marrow,femur, Brain(cerebrum,cerebellum,pons), Heart
Intestine,duodenum, Intestine,jejunum, Intestine,ileum, Intestine,cecum
Intestine,colon, Intestine,rectum, Kidney, Lymph node,mesenteric
Lymph node,submandibular, Liver, Lung(bronchus), Mammary gland,inguinal
Medulla oblongata, Muscle,quadriceps femoris, Optic nerve, Ovary, Parathyroid
Pancreas, Pituitary, Peyer's patch,ileal, Salivary gland,submandibular
Skin,inguinal, Sciatic nerve, Spleen, Stomach, Spinal cord,thoracic, Thymus
Trachea, Thyroid, Urinary bladder, Uterus, Vagina

		Dose : Dodecyloxirane 0 mg/kg														Species : Rat	
		/Before mating (day)															
Animal No.		1	2	3	4	5	6	7	8	9	10	11	12	13	14		
1101		E	M	D	D	P	E	M	D	D	P	E	M	D	P		
1102		P	E	M	D	D	P	E	M	D	D	P	E	M	D		
1103		E	M	D	P	E	M	D	P	E	M	D	P	E	M		
1104		D	P	E	M	D	P	E	M	D	P	E	M	D	P		
1105		D	P	E	M	D	P	E	M	D	P	E	M	D	P		
1106		P	E	M	D	P	E	M	D	P	E	M	D	P	E		
1107		M	D	P	E	M	D	P	E	M	D	P	E	M	D		
1108		M	D	P	E	M	D	P	E	M	D	P	E	M	D		
1109		D	P	E	M	D	P	E	M	D	P	E	M	D	P		
1110		P	E	M	D	P	E	M	D	P	E	E	M	D	P		
1111		E	M	D	P	E	M	D	P	E	M	D	P	E	M		
1112		D	P	E	M	D	P	E	M	D	P	E	M	D	P		

P : Proestrus, E : Estrus, M : Metestrus, D : Diestrus, - : Smear was not sampled or missing value

Estrus cycles

Generation : F0

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Generation : F0		Dose : Dodecylxirane 0 mg/kg				Species : Rat	
Animal No.	/Before mating (day)	Length of estrous cycle			Mean	Number of estrus	Animal with abnormal estrous cycle
		(days)					
1101	E	5	5	4	4.7	4	-
1102	D	5	5		5.0	3	-
1103	D	4	4	4	4.0	4	-
1104	E	4	4	4	4.0	4	-
1105	E	4	4	4	4.0	4	-
1106	M	4	4	4	4.0	4	-
1107	P	4	4		4.0	3	-
1108	P	4	4		4.0	3	-
1109	E	4	4	4	4.0	4	-
1110	E	4	4	5	4.3	4	-
1111	D	4	4	4	4.0	4	-
1112	E	4	4	4	4.0	4	-

P : Proestrus, E : Estrus, M : Metestrus, D : Diestrus, - : Smear was not sampled or missing value

+ : Animal with abnormal estrous cycle, - : Animal with normal and regular cycle

Estrus cycles

Generation : F0

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	/Before mating (day)													
	1	2	3	4	5	6	7	8	9	10	11	12	13	14
2101	P	E	M	D	D	P	E	M	D	D	P	E	E	M
2102	P	E	M	D	P	E	M	D	P	E	M	D	P	E
2103	D	P	E	M	D	P	E	M	D	P	E	M	D	P
2104	E	M	D	P	E	M	D	P	E	M	D	P	E	M
2105	M	D	P	E	M	D	P	E	M	D	P	E	M	D
2106	D	P	E	M	D	P	E	M	D	P	E	M	D	P
2107	P	E	M	D	P	E	M	D	P	E	M	D	P	E
2108	M	D	P	E	M	D	P	E	M	D	P	E	M	D
2109	M	D	D	P	E	M	D	D	P	E	M	D	D	P
2110	D	P	E	M	D	P	E	M	D	P	E	M	D	P
2111	M	D	P	E	E	M	D	P	E	E	M	D	P	E
2112	E	M	D	P	E	M	D	P	E	M	D	P	E	M

P : Proestrus, E : Estrus, M : Metestrus, D : Diestrus, - : Smear was not sampled or missing value

		Dose : Dodecyloxirane 100 mg/kg			Species : Rat		
Animal No.	/Before mating (day)	Length of estrous cycle (days)			Mean	Number of estrus	Animal with abnormal estrous cycle
2101	D	5	5		5.0	3	+
2102	M	4	4	4	4.0	4	-
2103	E	4	4	4	4.0	4	-
2104	D	4	4	4	4.0	4	-
2105	P	4	4		4.0	3	-
2106	E	4	4	4	4.0	4	-
2107	M	4	4	4	4.0	4	-
2108	P	4	4		4.0	3	-
2109	E	5	5		5.0	3	-
2110	E	4	4	4	4.0	4	-
2111	E	5	5		5.0	3	-
2112	D	4	4	4	4.0	4	-

P : Proestrus, E : Estrus, M : Metestrus, D : Diestrus, - : Smear was not sampled or missing value
+ : Animal with abnormal estrous cycle, - : Animal with normal and regular cycle

		Dose : Dodecyloxirane 300 mg/kg														Species : Rat
		/Before mating (day)														
Animal No.		1	2	3	4	5	6	7	8	9	10	11	12	13	14	
3101		D	P	E	M	D	P	E	M	D	P	E	M	D	P	
3102		M	D	P	E	M	D	P	E	M	D	P	E	M	D	
3103		P	E	M	D	P	E	M	D	P	E	M	D	P	E	
3104		E	M	D	P	E	M	D	P	E	M	D	P	E	M	
3105		P	E	M	D	P	E	M	D	P	E	M	D	P	E	
3106		D	P	E	M	D	P	E	M	D	P	E	M	D	P	
3107		D	P	E	M	D	P	E	M	D	P	E	M	D	P	
3108		D	P	E	M	D	P	E	M	D	P	E	M	D	P	
3109		D	P	E	M	D	P	E	M	D	P	E	M	D	P	
3110		M	D	P	E	M	D	P	E	M	D	P	E	M	D	
3111		M	D	P	E	M	D	P	E	M	D	P	E	M	D	
3112		M	D	D	P	E	M	D	P	E	M	D	P	E	M	

P : Proestrus, E : Estrus, M : Metestrus, D : Diestrus, - : Smear was not sampled or missing value

Estrus cycles

Generation : F0

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

		Before mating (day)			Length of estrous cycle		Mean	Number of estrus	Animal with abnormal estrous cycle
Animal No.		15	(days)						
3101	E		4	4	4	4.0	4	-	
3102	P		4	4		4.0	3	-	
3103	M		4	4	4	4.0	4	-	
3104	D		4	4	4	4.0	4	-	
3105	E		4	4	4	4.0	4	-	
3106	E		4	4	4	4.0	4	-	
3107	E		4	4	4	4.0	4	-	
3108	E		4	4	4	4.0	4	-	
3109	E		4	4	4	4.0	4	-	
3110	P		4	4		4.0	3	-	
3111	P		4	4		4.0	3	-	
3112	D		4	4		4.0	3	-	

P : Proestrus, E : Estrus, M : Metestrus, D : Diestrus, - : Smear was not sampled or missing value

+ : Animal with abnormal estrous cycle, - : Animal with normal and regular cycle

Appendix 10 - 7

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats
Estrus cycles
Generation : F0

Study No. : R-1261

		Dose : Dodecyloxirane 1000 mg/kg														Species : Rat	
		/Before mating (day)															
Animal No.		1	2	3	4	5	6	7	8	9	10	11	12	13	14		
4101		E	M	D	P	E	M	D	P	E	M	D	P	E	M		
4102		P	E	M	D	P	E	M	D	P	E	M	D	D	P		
4103		M	D	P	E	M	D	P	E	M	D	P	E	M	D		
4104		D	P	E	E	M	D	P	E	E	M	D	P	E	E		
4105		D	P	E	M	D	P	E	M	D	P	E	M	D	P		
4106		D	P	E	M	D	D	P	E	M	D	P	E	M	D		
4107		P	E	M	D	P	E	M	D	P	E	M	D	P	E		
4108		P	E	M	D	P	E	M	D	P	E	M	D	P	E		
4109		E	M	D	P	E	M	D	P	E	M	D	P	E	M		
4110		P	E	M	D	P	E	M	D	P	E	M	D	D	P		
4111		P	E	M	D	P	E	E	M	D	P	E	M	D	P		
4112		D	P	E	M	D	P	E	M	D	P	E	M	D	P		

P : Proestrus, E : Estrus, M : Metestrus, D : Diestrus, - : Smear was not sampled or missing value

Estrus cycles

Generation : F0

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Generation : F0		Dose : Dodecyclohexane 1000 mg/kg			Species : Rat		
Animal No.	/Before mating (day)	Length of estrous cycle			Mean	Number of estrus	Animal with abnormal estrous cycle
		(days)					
4101	D	4	4	4	4.0	4	-
4102	E	4	4	5	4.3	4	-
4103	P	4	4		4.0	3	-
4104	M	5	5		5.0	3	-
4105	E	4	4	4	4.0	4	-
4106	P	5	4		4.5	3	-
4107	M	4	4	4	4.0	4	-
4108	M	4	4	4	4.0	4	-
4109	D	4	4	4	4.0	4	-
4110	E	4	4	5	4.3	4	-
4111	E	4	5	4	4.3	4	-
4112	E	4	4	4	4.0	4	-

P : Proestrus, E : Estrus, M : Metestrus, D : Diestrus, - : Smear was not sampled or missing value

+ : Animal with abnormal estrous cycle, - : Animal with normal and regular cycle

Animal No.	Elapsed day of conceiving	1st mating		
		Paired animal	Copulation a)	Fertility b)
1001	1	1101	+	+
1002	11	1102	+	+
1003	3	1103	+	+
1004	4	1104	+	+
1005	1	1105	+	+
1006	3	1106	+	+
1007	1	1107	+	+
1008	1	1108	+	+
1009	4	1109	+	+
1010	4	1110	+	+
1011	2	1111	+	+
1012	4	1112	+	+
n	12		12	12
Mean	3.3			
S.D.	2.8			
No. of positives			12	12
%			100.0 c)	100.0 d)

/// : No need to input/measure

a) + : Copulation, - : Not copulation

b) + : Pregnancy, - : Non-pregnancy

c) : (Number of copulated animals / Number of mated animals)x 100

d) : (Number of males which impregnated females / Number of copulated males)x 100

Reproductive performance

Generation : F0

Sex : Male

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	Elapsed day of conceiving	1st mating		
		Paired animal	Copulation a)	Fertility b)
2001	3	2101	+	+
2002	3	2102	+	+
2003	4	2103	+	+
2004	2	2104	+	+
2005	1	2105	+	+
2006	1	2106	+	-
2007	3	2107	+	+
2008	1	2108	+	+
2009	-	2109	-	///
2010	4	2110	+	+
2011	4	2111	+	+
2012	2	2112	+	+
n	11		12	11
Mean	2.5			
S.D.	1.2			
No. of positives			11	10
%			91.7 c)	90.9 d)

/// : No need to input/measure

a) + : Copulation, - : Not copulation

b) + : Pregnancy, - : Non-pregnancy

c) : (Number of copulated animals / Number of mated animals)x 100

d) : (Number of males which impregnated females / Number of copulated males)x 100

Reproductive performance

Generation : F0

Sex : Male

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	Elapsed day of conceiving	1st mating		
		Paired animal	Copulation a)	Fertility b)
3001	1	3101	+	+
3002	1	3102	+	+
3003	3	3103	+	+
3004	2	3104	+	+
3005	5	3105	+	+
3006	4	3106	+	+
3007	14	3107	+	+
3008	-	3108	-	///
3009	2	3109	+	-
3010	1	3110	+	+
3011	1	3111	+	+
3012	2	3112	+	+
n	11		12	11
Mean	3.3			
S.D.	3.8			
No. of positives			11	10
%			91.7 c)	90.9 d)

/// : No need to input/measure

a) + : Copulation, - : Not copulation

b) + : Pregnancy, - : Non-pregnancy

c) : (Number of copulated animals / Number of mated animals)x 100

d) : (Number of males which impregnated females / Number of copulated males)x 100

Reproductive performance

Generation : F0

Sex : Male

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Elapsed day of conceiving	1st mating		
		Paired animal	Copulation a)	Fertility b)
4001	2	4101	+	-
4002	1	4102	+	+
4003	1	4103	+	+
4004	4	4104	+	+
4005	4	4105	+	+
4006	1	4106	+	+
4007	3	4107	+	+
4008	3	4108	+	+
4009	2	4109	+	+
4010	4	4110	+	+
4011	1	4111	+	+
4012	14	4112	+	+
n	12		12	12
Mean	3.3			
S.D.	3.6			
No. of positives			12	11
%			100.0 c)	91.7 d)

/// : No need to input/measure

a) + : Copulation, - : Not copulation

b) + : Pregnancy, - : Non-pregnancy

c) : (Number of copulated animals / Number of mated animals)x 100

d) : (Number of males which impregnated females / Number of copulated males)x 100

Reproductive performance

Generation : F0

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Elapsed day of conceiving	1st mating		
		Paired animal	Copulation a)	Fertility b)
1101	1	1001	+	+
1102	11	1002	+	+
1103	3	1003	+	+
1104	4	1004	+	+
1105	1	1005	+	+
1106	3	1006	+	+
1107	1	1007	+	+
1108	1	1008	+	+
1109	4	1009	+	+
1110	4	1010	+	+
1111	2	1011	+	+
1112	4	1012	+	+
n	12		12	12
Mean	3.3			
S.D.	2.8			
No. of positives			12	12
%			100.0 c)	100.0 d)

/// : No need to input/measure

a) + : Copulation, - : Not copulation

b) + : Pregnancy, - : Non-pregnancy

c) : (Number of copulated animals / Number of mated animals)x 100

d) : (Number of pregnant females / Number of copulated females)x 100

Reproductive performance

Generation : F0

Sex : Female

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	Elapsed day of conceiving	1st mating		
		Paired animal	Copulation a)	Fertility b)
2101	3	2001	+	+
2102	3	2002	+	+
2103	4	2003	+	+
2104	2	2004	+	+
2105	1	2005	+	+
2106	1	2006	+	-
2107	3	2007	+	+
2108	1	2008	+	+
2109	-	2009	-	///
2110	4	2010	+	+
2111	4	2011	+	+
2112	2	2012	+	+
n	11		12	11
Mean	2.5			
S.D.	1.2			
No. of positives			11	10
%			91.7 c)	90.9 d)

/// : No need to input/measure

a) + : Copulation, - : Not copulation

b) + : Pregnancy, - : Non-pregnancy

c) : (Number of copulated animals / Number of mated animals)x 100

d) : (Number of pregnant females / Number of copulated females)x 100

Reproductive performance

Generation : F0

Sex : Female

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	Elapsed day of conceiving	1st mating		
		Paired animal	Copulation a)	Fertility b)
3101	1	3001	+	+
3102	1	3002	+	+
3103	3	3003	+	+
3104	2	3004	+	+
3105	5	3005	+	+
3106	4	3006	+	+
3107	14	3007	+	+
3108	-	3008	-	///
3109	2	3009	+	-
3110	1	3010	+	+
3111	1	3011	+	+
3112	2	3012	+	+
n	11		12	11
Mean	3.3			
S.D.	3.8			
No. of positives			11	10
%			91.7 c)	90.9 d)

/// : No need to input/measure

a) + : Copulation, - : Not copulation

b) + : Pregnancy, - : Non-pregnancy

c) : (Number of copulated animals / Number of mated animals)x 100

d) : (Number of pregnant females / Number of copulated females)x 100

Reproductive performance

Generation : F0

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Elapsed day of conceiving	1st mating		
		Paired animal	Copulation a)	Fertility b)
4101	2	4001	+	-
4102	1	4002	+	+
4103	1	4003	+	+
4104	4	4004	+	+
4105	4	4005	+	+
4106	1	4006	+	+
4107	3	4007	+	+
4108	3	4008	+	+
4109	2	4009	+	+
4110	4	4010	+	+
4111	1	4011	+	+
4112	14	4012	+	+
n	12		12	12
Mean	3.3			
S.D.	3.6			
No. of positives			12	11
%			100.0 c)	91.7 d)

/// : No need to input/measure

a) + : Copulation, - : Not copulation

b) + : Pregnancy, - : Non-pregnancy

c) : (Number of copulated animals / Number of mated animals)x 100

d) : (Number of pregnant females / Number of copulated females)x 100

Delivery data

Generation : F0

Dose : Dodecylloxirane 0 mg/kg

Species : Rat

Dam No.	Gestation period (day)	Number of implan- tation	Delivery index (%) a)	Live birth index (%) b)	Number of offspring	Number of live newborns			Number of dead newborns			
						M	F	Total	M	F	U	Total
1101	22.5	10(7/3)	90.0	100.0	9	3	6	9	0	0	0	0
1102	22.5	18(11/7)	27.8	100.0	5	4	1	5	0	0	0	0
1103	22.5	18(9/9)	100.0	100.0	18	13	5	18	0	0	0	0
1104	22	17(9/8)	94.1	100.0	16	7	9	16	0	0	0	0
1105	22	16(9/7)	100.0	100.0	16	9	7	16	0	0	0	0
1106	22.5	16(7/9)	56.3	88.9	9	5	3	8	0	1	0	1
1107	21.5	13(7/6)	100.0	100.0	13	9	4	13	0	0	0	0
1108	22	17(7/10)	88.2	100.0	15	8	7	15	0	0	0	0
1109	22	13(8/5)	100.0	100.0	13	5	8	13	0	0	0	0
1110	22.5	14(5/9)	85.7	100.0	12	8	4	12	0	0	0	0
1111	22	17(9/8)	82.4	100.0	14	9	5	14	0	0	0	0
1112	21.5	18(10/8)	100.0	94.4	18	11	6	17	0	1	0	1
n	12	12	12	12	12	12	12	12	12	12	12	12
Mean	22.1	15.6	85.4	98.6	13.2	7.6	5.4	13.0	0.0	0.2	0.0	0.2
S.D.	0.4	2.5	22.0	3.5	3.9	2.9	2.2	3.9	0.0	0.4	0.0	0.4

(/) : Right/Left, M : Male, F : Female, U : Unable to be sexed

a): (Number of delivered pups / Number of implantation) x 100

b): (Number of liveborns / Number of delivered pups) x 100

Delivery data

Generation : F0

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Dam No.	Gestation period (day)	Number of implantation	Delivery index (%) a)	Live birth index (%) b)	Number of offspring	Number of live newborns			Number of dead newborns			
						M	F	Total	M	F	U	Total
2101	22	16(10/6)	100.0	100.0	16	9	7	16	0	0	0	0
2102	22	14(6/8)	100.0	78.6	14	5	6	11	1	2	0	3
2103	22	16(5/11)	93.8	100.0	15	7	8	15	0	0	0	0
2104	22	17(11/6)	76.5	100.0	13	5	8	13	0	0	0	0
2105	22.5	13(7/6)	92.3	100.0	12	6	6	12	0	0	0	0
2107	21.5	14(5/9)	100.0	100.0	14	6	8	14	0	0	0	0
2108	21.5	17(9/8)	88.2	100.0	15	6	9	15	0	0	0	0
2110	22	18(7/11)	94.4	100.0	17	8	9	17	0	0	0	0
2111	23	17(12/5)	82.4	100.0	14	11	3	14	0	0	0	0
2112	22.5	15(13/2)	86.7	100.0	13	6	7	13	0	0	0	0
n	10	10	10	10	10	10	10	10	10	10	10	10
Mean	22.1	15.7	91.4	97.9	14.3	6.9	7.1	14.0	0.1	0.2	0.0	0.3
S.D.	0.5	1.6	8.0	6.8	1.5	1.9	1.8	1.8	0.3	0.6	0.0	0.9

(/) : Right/Left, M : Male, F : Female, U : Unable to be sexed

a): (Number of delivered pups / Number of implantation) x 100

b): (Number of liveborns / Number of delivered pups) x 100

Delivery data

Generation : F0

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Dam No.	Gestation period (day)	Number of implan- tation	Delivery index (%) a)	Live birth index (%) b)	Number of offspring	Number of live newborns			Number of dead newborns			
						M	F	Total	M	F	U	Total
3101	22	15(7/8)	100.0	100.0	15	9	6	15	0	0	0	0
3102	22.5	15(4/11)	100.0	100.0	15	7	8	15	0	0	0	0
3103	21.5	16(8/8)	62.5	100.0	10	5	5	10	0	0	0	0
3104	21.5	16(5/11)	100.0	100.0	16	9	7	16	0	0	0	0
3105	22	17(9/8)	94.1	100.0	16	9	7	16	0	0	0	0
3106	22	18(11/7)	88.9	100.0	16	9	7	16	0	0	0	0
3107	22	14(9/5)	92.9	100.0	13	10	3	13	0	0	0	0
3110	22	13(13/0)	92.3	100.0	12	8	4	12	0	0	0	0
3111	21.5	19(10/9)	94.7	100.0	18	12	6	18	0	0	0	0
3112	22	16(10/6)	93.8	100.0	15	7	8	15	0	0	0	0
n	10	10	10	10	10	10	10	10	10	10	10	10
Mean	21.9	15.9	91.9	100.0	14.6	8.5	6.1	14.6	0.0	0.0	0.0	0.0
S.D.	0.3	1.8	11.0	0.0	2.3	1.9	1.7	2.3	0.0	0.0	0.0	0.0

(/) : Right/Left, M : Male, F : Female, U : Unable to be sexed

a): (Number of delivered pups / Number of implantation) x 100

b): (Number of liveborns / Number of delivered pups) x 100

Delivery data

Generation : F0

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Dam No.	Gestation period (day)	Number of implan- tation	Delivery index (%) a)	Live birth index (%) b)	Number of offspring	Number of live newborns			Number of dead newborns			
						M	F	Total	M	F	U	Total
4102	22.5	14(4/10)	100.0	100.0	14	9	5	14	0	0	0	0
4103	22	18(10/8)	88.9	100.0	16	5	11	16	0	0	0	0
4104	22.5	15(6/9)	80.0	100.0	12	7	5	12	0	0	0	0
4105	22	14(8/6)	100.0	100.0	14	7	7	14	0	0	0	0
4107	22.5	17(10/7)	94.1	100.0	16	10	6	16	0	0	0	0
4108	22.5	9(9/0)	88.9	100.0	8	3	5	8	0	0	0	0
4109	22.5	16(8/8)	100.0	100.0	16	9	7	16	0	0	0	0
4110	22	16(8/8)	81.3	100.0	13	8	5	13	0	0	0	0
4111	22	14(11/3)	100.0	100.0	14	4	10	14	0	0	0	0
4112	22	16(7/9)	87.5	100.0	14	7	7	14	0	0	0	0
n	10	10	10	10	10	10	10	10	10	10	10	10
Mean	22.3	14.9	92.1	100.0	13.7	6.9	6.8	13.7	0.0	0.0	0.0	0.0
S.D.	0.3	2.5	7.9	0.0	2.4	2.3	2.1	2.4	0.0	0.0	0.0	0.0

(/) : Right/Left, M : Male, F : Female, U : Unable to be sexed

a): (Number of delivered pups / Number of implantation) x 100

b): (Number of liveborns / Number of delivered pups) x 100

Appendix 13 - 1

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

External examination of offspring

Unit : Number of anomalous offspring (incidence %)

Days after birth : 0

Generation : F0

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Dam No.	Number of offspring examined	/Findings
1101	9	No anomaly
1102	5	No anomaly
1103	18	No anomaly
1104	16	No anomaly
1105	16	No anomaly
1106	9	No anomaly
1107	13	No anomaly
1108	15	No anomaly
1109	13	No anomaly
1110	12	No anomaly
1111	14	No anomaly
1112	18	No anomaly
Total	158	
n	12	
Mean		
S.D.		

Mean : Average of incidence (%) : (Number of pups with external anomalies / Number of delivered pups) x 100

"No anomaly" was calculated as "0".

Appendix 13 - 2

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

External examination of offspring

Unit : Number of anomalous offspring (incidence %)

Days after birth : 0

Generation : F0

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Dam No.	Number of offspring examined	/Findings
2101	16	No anomaly
2102	14	No anomaly
2103	15	No anomaly
2104	13	No anomaly
2105	12	No anomaly
2107	14	No anomaly
2108	15	No anomaly
2110	17	No anomaly
2111	14	No anomaly
2112	13	No anomaly
Total	143	
n	10	
Mean		
S.D.		

Mean : Average of incidence (%) : (Number of pups with external anomalies / Number of delivered pups) x 100

"No anomaly" was calculated as "0".

Appendix 13 - 3

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

External examination of offspring

Unit : Number of anomalous offspring (incidence %)

Days after birth : 0

Generation : F0

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Dam No.	Number of offspring examined	/Findings
3101	15	No anomaly
3102	15	No anomaly
3103	10	No anomaly
3104	16	No anomaly
3105	16	No anomaly
3106	16	No anomaly
3107	13	No anomaly
3110	12	No anomaly
3111	18	No anomaly
3112	15	No anomaly
Total	146	
n	10	
Mean		
S.D.		

Mean : Average of incidence (%) : (Number of pups with external anomalies / Number of delivered pups) x 100

"No anomaly" was calculated as "0".

Appendix 13 - 4

Dodecyloxirane: An oral combined repeated dose toxicity study
with the reproduction/ developmental toxicity screening test in rats

Study No. : R-1261

External examination of offspring

Unit : Number of anomalous offspring (incidence %)

Days after birth : 0

Generation : F0

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Dam No.	Number of offspring examined	/Findings
4102	14	No anomaly
4103	16	No anomaly
4104	12	No anomaly
4105	14	No anomaly
4107	16	No anomaly
4108	8	No anomaly
4109	16	No anomaly
4110	13	No anomaly
4111	14	No anomaly
4112	14	No anomaly
Total	137	
n	10	
Mean		
S.D.		

Mean : Average of incidence (%) : (Number of pups with external anomalies / Number of delivered pups) x 100

"No anomaly" was calculated as "0".

		Generation : F0		Dose : Dodecyloxirane 0 mg/kg		Unit : mm	Species : Rat
Sex		/Days after birth					
		0					
Dam No.		n	AGD	Normalized a)			
				AGD			
1101	Male	3	2.3	1.1			
	Female	6	0.7	0.4			
1102	Male	4	2.7	1.4			
	Female	1	1.1	0.6			
1103	Male	13	2.4	1.3			
	Female	5	1.1	0.6			
1104	Male	7	2.8	1.5			
	Female	9	0.8	0.4			
1105	Male	9	2.4	1.3			
	Female	7	0.9	0.5			
1106	Male	5	2.5	1.3			
	Female	3	0.9	0.5			
1107	Male	9	2.4	1.4			
	Female	4	0.7	0.4			
1108	Male	8	2.5	1.4			
	Female	7	0.8	0.4			
1109	Male	5	2.5	1.4			
	Female	8	0.9	0.5			
1110	Male	8	2.6	1.3			
	Female	4	0.8	0.5			
1111	Male	9	2.5	1.3			
	Female	5	0.9	0.5			
1112	Male	11	2.4	1.4			
	Female	6	1.1	0.6			
n	Male	91					
	Female	65					
Mean	Male		2.5	1.3			
	Female		0.9	0.5			
S.D.	Male		0.1	0.1			
	Female		0.1	0.1			

AGD : Anogenital Distance

a): AGD (mm) / Cube root of body weight (g)

		Generation : F0		Dose : Dodecyloxirane 100 mg/kg		Unit : mm	Species : Rat
Sex		/Days after birth					
		0					
Dam No.		n	AGD	Normalized a)			
				AGD			
2101	Male	9	2.4	1.3			
	Female	7	0.8	0.5			
2102	Male	5	2.6	1.3			
	Female	6	0.9	0.5			
2103	Male	7	2.5	1.3			
	Female	8	0.8	0.4			
2104	Male	5	2.6	1.4			
	Female	8	0.9	0.5			
2105	Male	6	2.1	1.1			
	Female	6	0.7	0.4			
2107	Male	6	2.3	1.2			
	Female	8	0.9	0.5			
2108	Male	6	2.8	1.5			
	Female	9	0.8	0.4			
2110	Male	8	2.6	1.3			
	Female	9	0.8	0.4			
2111	Male	11	2.4	1.3			
	Female	3	0.9	0.5			
2112	Male	6	2.5	1.3			
	Female	7	0.9	0.5			
n	Male	69					
	Female	71					
Mean	Male		2.5	1.3			
	Female		0.8	0.5			
S.D.	Male		0.2	0.1			
	Female		0.1	0.1			

AGD : Anogenital Distance

a): AGD (mm) / Cube root of body weight (g)

		Generation : F0		Dose : Dodecyloxirane 300 mg/kg		Unit : mm	Species : Rat
Sex		/Days after birth					
		0					
Dam No.		n	AGD	Normalized a)			
				AGD			
3101	Male	9	2.4	1.3			
	Female	6	0.8	0.4			
3102	Male	7	2.2	1.2			
	Female	8	0.8	0.4			
3103	Male	5	2.5	1.3			
	Female	5	0.9	0.5			
3104	Male	9	2.4	1.4			
	Female	7	0.8	0.4			
3105	Male	9	2.5	1.3			
	Female	7	0.7	0.4			
3106	Male	9	2.7	1.5			
	Female	7	0.8	0.4			
3107	Male	10	2.4	1.3			
	Female	3	0.8	0.4			
3110	Male	8	2.6	1.4			
	Female	4	0.9	0.5			
3111	Male	12	2.6	1.4			
	Female	6	0.9	0.5			
3112	Male	7	2.5	1.3			
	Female	8	0.7	0.4			
n	Male	85					
	Female	61					
Mean	Male		2.5	1.3			
	Female		0.8	0.4			
S.D.	Male		0.1	0.1			
	Female		0.1	0.0			

AGD : Anogenital Distance

a): AGD (mm) / Cube root of body weight (g)

		Generation : F0		Dose : Dodecyloxirane 1000 mg/kg		Unit : mm	Species : Rat
Sex		/Days after birth					
		0					
Dam No.		n	AGD	Normalized a)			
				AGD			
4102	Male	9	2.3	1.2			
	Female	5	0.7	0.4			
4103	Male	5	2.5	1.4			
	Female	11	0.8	0.4			
4104	Male	7	2.4	1.3			
	Female	5	0.9	0.4			
4105	Male	7	2.6	1.4			
	Female	7	0.8	0.5			
4107	Male	10	2.5	1.3			
	Female	6	1.1	0.6			
4108	Male	3	2.7	1.4			
	Female	5	1.0	0.5			
4109	Male	9	2.7	1.4			
	Female	7	1.2	0.6			
4110	Male	8	2.6	1.4			
	Female	5	0.9	0.5			
4111	Male	4	2.4	1.3			
	Female	10	0.8	0.4			
4112	Male	7	2.3	1.2			
	Female	7	0.8	0.4			
n	Male	69					
	Female	68					
Mean	Male		2.5	1.3			
	Female		0.9	0.5			
S.D.	Male		0.1	0.1			
	Female		0.2	0.1			

AGD : Anogenital Distance

a): AGD (mm) / Cube root of body weight (g)

Generation : F0

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

		/Days after birth					
		0	4	4	4	13	13
Dam No.	Sex	Male/Female	Pre-culled Male/Female	Viability index (%) a)	Culled Male/Female	Male/Female	Viability index (%) b)
1101	Total	9 (3/6)	9 (3/6)	100.0	8 (3/5)	8 (3/5)	100.0
1102	Total	5 (4/1)	5 (4/1)	100.0	5 (4/1)	5 (4/1)	100.0
1103	Total	18 (13/5)	18 (13/5)	100.0	8 (4/4)	8 (4/4)	100.0
1104	Total	16 (7/9)	16 (7/9)	100.0	8 (4/4)	8 (4/4)	100.0
1105	Total	16 (9/7)	16 (9/7)	100.0	8 (4/4)	8 (4/4)	100.0
1106	Total	8 (5/3)	8 (5/3)	100.0	8 (5/3)	8 (5/3)	100.0
1107	Total	13 (9/4)	13 (9/4)	100.0	8 (4/4)	8 (4/4)	100.0
1108	Total	15 (8/7)	14 (7/7)	93.3	8 (4/4)	8 (4/4)	100.0
1109	Total	13 (5/8)	13 (5/8)	100.0	8 (4/4)	8 (4/4)	100.0
1110	Total	12 (8/4)	12 (8/4)	100.0	8 (4/4)	8 (4/4)	100.0
1111	Total	14 (9/5)	14 (9/5)	100.0	8 (4/4)	8 (4/4)	100.0
1112	Total	17 (11/6)	17 (11/6)	100.0	8 (4/4)	8 (4/4)	100.0
Total		156	155		93	93	
Male/Female		91/65	90/65		48/45	48/45	
n		12	12	12	12	12	12
Mean			12.9	99.4		7.8	100.0
S.D.			3.9	1.9		0.9	0.0

a): (Number of pups alive on day 4 after birth / Number of live born pups) × 100

b): (Number of pups alive on day 13 after birth / Number of pups after culling on day 4 after birth) × 100

Generation : F0

Dose : Dodecylloxirane 100 mg/kg

Species : Rat

		/Days after birth					
		0	4	4	4	13	13
Dam No.	Sex	Male/Female	Pre-culled Male/Female	Viability index (%) a)	Culled Male/Female	Male/Female	Viability index (%) b)
2101	Total	16 (9/7)	16 (9/7)	100.0	8 (4/4)	8 (4/4)	100.0
2102	Total	11 (5/6)	11 (5/6)	100.0	8 (4/4)	8 (4/4)	100.0
2103	Total	15 (7/8)	14 (6/8)	93.3	8 (4/4)	8 (4/4)	100.0
2104	Total	13 (5/8)	13 (5/8)	100.0	8 (4/4)	8 (4/4)	100.0
2105	Total	12 (6/6)	11 (6/5)	91.7	8 (4/4)	8 (4/4)	100.0
2107	Total	14 (6/8)	14 (6/8)	100.0	8 (4/4)	8 (4/4)	100.0
2108	Total	15 (6/9)	15 (6/9)	100.0	8 (4/4)	8 (4/4)	100.0
2110	Total	17 (8/9)	17 (8/9)	100.0	8 (4/4)	8 (4/4)	100.0
2111	Total	14 (11/3)	13 (11/2)	92.9	8 (6/2)	8 (6/2)	100.0
2112	Total	13 (6/7)	13 (6/7)	100.0	8 (4/4)	8 (4/4)	100.0
Total		140	137		80	80	
Male/Female		69/71	68/69		42/38	42/38	
n		10	10	10	10	10	10
Mean			13.7	97.8		8.0	100.0
S.D.			1.9	3.6		0.0	0.0

a): (Number of pups alive on day 4 after birth / Number of live born pups) × 100

b): (Number of pups alive on day 13 after birth / Number of pups after culling on day 4 after birth) × 100

Generation : F0

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

		/Days after birth					
		0	4	4	4	13	13
Dam No.	Sex	Male/Female	Pre-culled Male/Female	Viability index (%) a)	Culled Male/Female	Male/Female	Viability index (%) b)
3101	Total	15 (9/6)	15 (9/6)	100.0	8 (4/4)	8 (4/4)	100.0
3102	Total	15 (7/8)	15 (7/8)	100.0	8 (4/4)	8 (4/4)	100.0
3103	Total	10 (5/5)	10 (5/5)	100.0	8 (4/4)	8 (4/4)	100.0
3104	Total	16 (9/7)	16 (9/7)	100.0	8 (4/4)	8 (4/4)	100.0
3105	Total	16 (9/7)	16 (9/7)	100.0	8 (4/4)	8 (4/4)	100.0
3106	Total	16 (9/7)	16 (9/7)	100.0	8 (4/4)	8 (4/4)	100.0
3107	Total	13 (10/3)	13 (10/3)	100.0	8 (5/3)	8 (5/3)	100.0
3110	Total	12 (8/4)	12 (8/4)	100.0	8 (4/4)	8 (4/4)	100.0
3111	Total	18 (12/6)	17 (11/6)	94.4	8 (4/4)	8 (4/4)	100.0
3112	Total	15 (7/8)	15 (7/8)	100.0	8 (4/4)	8 (4/4)	100.0
Total		146	145		80	80	
Male/Female		85/61	84/61		41/39	41/39	
n		10	10	10	10	10	10
Mean			14.5	99.4		8.0	100.0
S.D.			2.2	1.8		0.0	0.0

a): (Number of pups alive on day 4 after birth / Number of live born pups) × 100

b): (Number of pups alive on day 13 after birth / Number of pups after culling on day 4 after birth) × 100

Generation : F0

Dose : Dodecylloxirane 1000 mg/kg

Species : Rat

		/Days after birth					
		0	4	4	4	13	13
		Male/Female	Pre-culled Male/Female	Viability index (%) a)	Culled Male/Female	Male/Female	Viability index (%) b)
Dam No.	Sex						
4102	Total	14 (9/5)	14 (9/5)	100.0	8 (4/4)	8 (4/4)	100.0
4103	Total	16 (5/11)	15 (5/10)	93.8	8 (4/4)	8 (4/4)	100.0
4104	Total	12 (7/5)	12 (7/5)	100.0	8 (4/4)	8 (4/4)	100.0
4105	Total	14 (7/7)	13 (7/6)	92.9	8 (4/4)	8 (4/4)	100.0
4107	Total	16 (10/6)	16 (10/6)	100.0	8 (4/4)	8 (4/4)	100.0
4108	Total	8 (3/5)	8 (3/5)	100.0	8 (3/5)	8 (3/5)	100.0
4109	Total	16 (9/7)	15 (9/6)	93.8	8 (4/4)	8 (4/4)	100.0
4110	Total	13 (8/5)	13 (8/5)	100.0	8 (4/4)	8 (4/4)	100.0
4111	Total	14 (4/10)	14 (4/10)	100.0	8 (4/4)	8 (4/4)	100.0
4112	Total	14 (7/7)	14 (7/7)	100.0	8 (4/4)	8 (4/4)	100.0
Total		137	134		80	80	
Male/Female		69/68	69/65		39/41	39/41	
n		10	10	10	10	10	10
Mean			13.4	98.1		8.0	100.0
S.D.			2.2	3.1		0.0	0.0

a): (Number of pups alive on day 4 after birth / Number of live born pups) × 100

b): (Number of pups alive on day 13 after birth / Number of pups after culling on day 4 after birth) × 100

Sex ratio

Generation : F0

Dose :Dodecyloxirane 0 mg/kg

Species : Rat

Dam No.	day 4 after birth		Sex ratio (%) a)
	Number of males	Number of females	
1101	3	6	
1102	4	1	
1103	13	5	
1104	7	9	
1105	9	7	
1106	5	3	
1107	9	4	
1108	7	7	
1109	5	8	
1110	8	4	
1111	9	5	
1112	11	6	
n	12	12	
Total	90	65	90/155
(%)			58.1

a): (Number of male pups alive on day 4 after birth / Number of pups alive on day 4 after birth) × 100

Sex ratio

Generation : F0

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Dam No.	day 4 after birth		Sex ratio (%) a)
	Number of males	Number of females	
2101	9	7	
2102	5	6	
2103	6	8	
2104	5	8	
2105	6	5	
2107	6	8	
2108	6	9	
2110	8	9	
2111	11	2	
2112	6	7	
n	10	10	
Total	68	69	68/137
(%)			49.6

a): (Number of male pups alive on day 4 after birth / Number of pups alive on day 4 after birth) × 100

Sex ratio

Generation : F0

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Dam No.	day 4 after birth		Sex ratio (%) a)
	Number of males	Number of females	
3101	9	6	
3102	7	8	
3103	5	5	
3104	9	7	
3105	9	7	
3106	9	7	
3107	10	3	
3110	8	4	
3111	11	6	
3112	7	8	
n	10	10	
Mean	84	61	84/145
S.D.			57.9

a): (Number of male pups alive on day 4 after birth / Number of pups alive on day 4 after birth) × 100

Sex ratio

Generation : F0

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Dam No.	day 4 after birth		Sex ratio (%) a)
	Number of males	Number of females	
4102	9	5	
4103	5	10	
4104	7	5	
4105	7	6	
4107	10	6	
4108	3	5	
4109	9	6	
4110	8	5	
4111	4	10	
4112	7	7	
n	10	10	
Mean	69	65	69/134
S.D.			51.5

a): (Number of male pups alive on day 4 after birth / Number of pups alive on day 4 after birth) × 100

		Dose : Dodecyloxirane 0 mg/kg								Unit : g	Species : Rat
		/Days after birth									
Dam No.	Sex	0		4/Pre-culled		4/Culled		7		13	
		n	B.W.	n	B.W.	n	B.W.	n	B.W.	n	B.W.
1101	Male	3	8.0	3	13.4	3	13.4	3	20.5	3	38.9
	Female	6	7.5	6	12.7	5	12.8	5	20.0	5	37.3
1102	Male	4	7.7	4	13.1	4	13.1	4	20.1	4	37.5
	Female	1	7.3	1	12.7	1	12.7	1	19.6	1	35.6
1103	Male	13	6.5	13	8.8	4	9.7	4	16.6	4	33.1
	Female	5	6.0	5	8.3	4	9.0	4	16.0	4	31.9
1104	Male	7	6.4	7	10.6	4	11.3	4	18.4	4	37.8
	Female	9	6.1	9	10.3	4	10.4	4	16.7	4	34.0
1105	Male	9	6.9	9	11.4	4	12.1	4	19.2	4	34.8
	Female	7	6.6	7	10.6	4	10.5	4	16.4	4	32.4
1106	Male	5	8.0	5	13.8	5	13.8	5	20.4	5	35.2
	Female	3	7.6	3	13.3	3	13.3	3	19.9	3	34.1
1107	Male	9	5.9	9	10.1	4	9.9	4	16.3	4	30.1
	Female	4	5.7	4	9.8	4	9.8	4	15.8	4	30.1
1108	Male	8	6.7	7	11.8	4	12.0	4	19.2	4	36.8
	Female	7	6.2	7	10.7	4	11.3	4	18.5	4	35.3
1109	Male	5	6.7	5	11.2	4	11.6	4	18.5	4	34.2
	Female	8	6.4	8	10.8	4	11.0	4	17.4	4	32.5
1110	Male	8	7.3	8	11.6	4	11.8	4	19.2	4	34.6
	Female	4	6.5	4	10.9	4	10.9	4	18.2	4	34.6
1111	Male	9	7.0	9	11.5	4	11.7	4	18.6	4	33.7
	Female	5	6.3	5	10.8	4	11.0	4	17.0	4	31.7
1112	Male	11	5.8	11	9.5	4	10.0	4	16.7	4	32.9
	Female	6	5.5	6	8.6	4	9.1	4	15.7	4	31.6
n	Male	91		90		48		48		48	
	Female	65		65		45		45		45	
Mean	Male		6.9		11.4		11.7		18.6		35.0
	Female		6.5		10.8		11.0		17.6		33.4
S.D.	Male		0.7		1.5		1.3		1.4		2.5
	Female		0.7		1.5		1.4		1.6		2.1

		Dose : Dodecyloxirane 100 mg/kg								Unit : g	Species : Rat
		/Days after birth									
Dam No.	Sex	0		4/Pre-culled		4/Culled		7		13	
		n	B.W.	n	B.W.	n	B.W.	n	B.W.	n	B.W.
2101	Male	9	6.4	9	11.0	4	11.4	4	18.5	4	34.8
	Female	7	6.1	7	10.3	4	10.8	4	17.1	4	33.2
2102	Male	5	6.9	5	12.4	4	12.7	4	21.0	4	38.7
	Female	6	6.4	6	11.9	4	12.0	4	19.9	4	36.5
2103	Male	7	6.9	6	11.5	4	11.7	4	19.7	4	37.1
	Female	8	6.8	8	11.1	4	11.3	4	19.2	4	35.9
2104	Male	5	7.0	5	11.5	4	11.6	4	19.5	4	36.1
	Female	8	6.7	8	10.9	4	10.9	4	18.6	4	35.2
2105	Male	6	7.0	6	11.7	4	11.7	4	18.8	4	35.7
	Female	6	6.4	5	11.1	4	11.1	4	17.9	4	34.2
2107	Male	6	6.3	6	9.5	4	9.4	4	16.2	4	31.3
	Female	8	6.0	8	9.3	4	9.6	4	16.2	4	32.3
2108	Male	6	6.3	6	9.9	4	9.8	4	16.3	4	33.6
	Female	9	5.8	9	9.6	4	9.8	4	16.0	4	33.0
2110	Male	8	6.8	8	10.0	4	10.9	4	18.0	4	36.1
	Female	9	6.6	9	9.7	4	10.2	4	17.8	4	36.4
2111	Male	11	7.1	11	11.8	6	12.3	6	19.1	6	34.8
	Female	3	6.5	2	11.4	2	11.4	2	17.2	2	31.3
2112	Male	6	6.4	6	10.8	4	10.8	4	18.5	4	34.5
	Female	7	5.9	7	9.7	4	9.4	4	16.0	4	30.6
n	Male	69		68		42		42		42	
	Female	71		69		38		38		38	
Mean	Male		6.7		11.0		11.2		18.6		35.3
	Female		6.3		10.5		10.7		17.6		33.9
S.D.	Male		0.3		0.9		1.0		1.5		2.0
	Female		0.3		0.9		0.9		1.4		2.1

Body weight of offspring Generation : F0				Dose : Dodecyloxirane 300 mg/kg				Unit : g		Species : Rat	
		0 /Days after birth		4/Pre-culled		4/Culled		7		13	
Dam No.	Sex	n	B.W.	n	B.W.	n	B.W.	n	B.W.	n	B.W.
3101	Male	9	6.2	9	10.1	4	10.7	4	18.1	4	34.4
	Female	6	6.2	6	10.2	4	10.3	4	17.3	4	33.1
3102	Male	7	6.7	7	9.4	4	9.9	4	16.1	4	32.9
	Female	8	6.2	8	8.9	4	9.0	4	14.6	4	30.8
3103	Male	5	7.2	5	11.5	4	11.3	4	17.8	4	32.9
	Female	5	6.1	5	9.5	4	9.9	4	15.8	4	30.0
3104	Male	9	5.9	9	9.4	4	9.6	4	15.8	4	31.6
	Female	7	6.0	7	9.2	4	9.8	4	16.0	4	31.5
3105	Male	9	7.8	9	10.6	4	11.8	4	16.8	4	34.6
	Female	7	7.0	7	9.7	4	10.0	4	14.6	4	31.3
3106	Male	9	6.3	9	9.9	4	10.1	4	17.3	4	35.0
	Female	7	5.7	7	8.8	4	9.8	4	16.5	4	33.8
3107	Male	10	7.0	10	12.7	5	12.6	5	21.1	5	39.3
	Female	3	6.2	3	11.8	3	11.8	3	19.5	3	37.3
3110	Male	8	6.8	8	12.1	4	13.0	4	21.7	4	38.6
	Female	4	6.0	4	10.9	4	10.9	4	18.1	4	35.0
3111	Male	12	5.9	11	9.5	4	9.7	4	16.7	4	33.6
	Female	6	5.5	6	9.0	4	8.9	4	15.6	4	32.0
3112	Male	7	6.2	7	10.5	4	10.6	4	17.8	4	35.2
	Female	8	5.6	8	9.4	4	9.5	4	15.7	4	31.4
n	Male	85		84		41		41		41	
	Female	61		61		39		39		39	
Mean	Male		6.6		10.6		10.9		17.9		34.8
	Female		6.1		9.7		10.0		16.4		32.6
S.D.	Male		0.6		1.2		1.2		2.0		2.4
	Female		0.4		1.0		0.9		1.5		2.2

		Dose : Dodecyloxirane 1000 mg/kg								Unit : g	Species : Rat
		/Days after birth									
Dam No.	Sex	0		4/Pre-culled		4/Culled		7		13	
		n	B.W.	n	B.W.	n	B.W.	n	B.W.	n	B.W.
4102	Male	9	6.8	9	11.2	4	11.2	4	18.0	4	33.7
	Female	5	6.2	5	10.1	4	9.9	4	16.0	4	29.8
4103	Male	5	6.0	5	9.0	4	9.5	4	15.5	4	28.2
	Female	11	5.9	10	9.7	4	10.1	4	16.6	4	29.2
4104	Male	7	6.6	7	11.2	4	11.3	4	18.1	4	33.6
	Female	5	6.5	5	10.9	4	11.0	4	17.2	4	32.1
4105	Male	7	6.1	7	9.9	4	10.2	4	17.2	4	30.3
	Female	7	5.6	6	9.4	4	9.6	4	15.9	4	27.7
4107	Male	10	6.6	10	9.7	4	10.0	4	16.5	4	29.3
	Female	6	6.4	6	9.3	4	9.4	4	15.2	4	28.8
4108	Male	3	7.7	3	12.7	3	12.7	3	18.0	3	29.4
	Female	5	7.0	5	11.9	5	11.9	5	16.5	5	27.7
4109	Male	9	7.5	9	11.9	4	12.1	4	21.0	4	38.8
	Female	7	7.2	6	12.0	4	11.8	4	20.3	4	37.6
4110	Male	8	6.5	8	10.4	4	10.8	4	17.6	4	33.3
	Female	5	6.4	5	10.6	4	10.7	4	17.6	4	33.1
4111	Male	4	6.4	4	9.7	4	9.7	4	14.9	4	27.9
	Female	10	6.0	10	9.5	4	11.1	4	17.5	4	31.7
4112	Male	7	7.1	7	11.5	4	11.5	4	18.7	4	32.8
	Female	7	6.5	7	10.9	4	10.7	4	17.9	4	31.1
n	Male	69		69		39		39		39	
	Female	68		65		41		41		41	
Mean	Male		6.7		10.7		10.9		17.6		31.7
	Female		6.4		10.4		10.6		17.1		30.9
S.D.	Male		0.6		1.2		1.1		1.7		3.4
	Female		0.5		1.0		0.9		1.4		3.0

Dose : Dodecyloxirane 0 mg/kg

Base : Day 4 after birth

Unit : g

Species : Rat

Dam No.	Sex	/Days after birth /Before culling		/After culling			
		n	0-4	n	4-7	n	4-13
1101	Male	3	5.4	3	7.1	3	25.5
	Female	6	5.2	5	7.2	5	24.5
1102	Male	4	5.4	4	7.1	4	24.4
	Female	1	5.4	1	6.9	1	22.9
1103	Male	13	2.3	4	6.9	4	23.4
	Female	5	2.3	4	7.0	4	22.9
1104	Male	7	4.2	4	7.1	4	26.5
	Female	9	4.2	4	6.4	4	23.6
1105	Male	9	4.5	4	7.1	4	22.6
	Female	7	4.0	4	6.0	4	22.0
1106	Male	5	5.8	5	6.5	5	21.4
	Female	3	5.7	3	6.6	3	20.8
1107	Male	9	4.2	4	6.5	4	20.3
	Female	4	4.1	4	6.0	4	20.3
1108	Male	7	5.1	4	7.2	4	24.9
	Female	7	4.5	4	7.3	4	24.0
1109	Male	5	4.5	4	7.0	4	22.7
	Female	8	4.4	4	6.4	4	21.5
1110	Male	8	4.3	4	7.3	4	22.8
	Female	4	4.4	4	7.3	4	23.7
1111	Male	9	4.5	4	6.9	4	22.0
	Female	5	4.5	4	6.1	4	20.7
1112	Male	11	3.7	4	6.8	4	22.9
	Female	6	3.1	4	6.6	4	22.5
n	Male	90		48		48	
	Female	65		45		45	
Mean	Male		4.5		7.0		23.3
	Female		4.3		6.7		22.5
S.D.	Male		0.9		0.3		1.8
	Female		0.9		0.5		1.4

Dose : Dodecyloxirane 100 mg/kg

Base : Day 4 after birth

Unit : g

Species : Rat

Dam No.	Sex	/Days after birth /Before culling		/After culling			
		n	0-4	n	4-7	n	4-13
2101	Male	9	4.6	4	7.1	4	23.4
	Female	7	4.2	4	6.3	4	22.4
2102	Male	5	5.5	4	8.3	4	26.0
	Female	6	5.5	4	8.0	4	24.5
2103	Male	6	4.6	4	8.0	4	25.4
	Female	8	4.3	4	8.0	4	24.6
2104	Male	5	4.5	4	7.9	4	24.5
	Female	8	4.2	4	7.7	4	24.3
2105	Male	6	4.7	4	7.1	4	24.0
	Female	5	4.7	4	6.8	4	23.1
2107	Male	6	3.2	4	6.7	4	21.9
	Female	8	3.3	4	6.6	4	22.6
2108	Male	6	3.6	4	6.6	4	23.9
	Female	9	3.8	4	6.2	4	23.2
2110	Male	8	3.2	4	7.2	4	25.2
	Female	9	3.1	4	7.6	4	26.2
2111	Male	11	4.7	6	6.8	6	22.5
	Female	2	4.9	2	5.8	2	20.0
2112	Male	6	4.4	4	7.7	4	23.7
	Female	7	3.8	4	6.6	4	21.2
n	Male	68		42		42	
	Female	69		38		38	
Mean	Male		4.3		7.3		24.1
	Female		4.2		7.0		23.2
S.D.	Male		0.7		0.6		1.3
	Female		0.7		0.8		1.8

Dose : Dodecyloxirane 300 mg/kg

Base : Day 4 after birth

Unit : g

Species : Rat

Dam No.	Sex	/Days after birth /Before culling		/After culling			
		n	0-4	n	4-7	n	4-13
3101	Male	9	3.9	4	7.4	4	23.7
	Female	6	4.0	4	7.0	4	22.8
3102	Male	7	2.7	4	6.2	4	23.0
	Female	8	2.7	4	5.6	4	21.8
3103	Male	5	4.3	4	6.6	4	21.7
	Female	5	3.4	4	5.9	4	20.1
3104	Male	9	3.5	4	6.2	4	22.0
	Female	7	3.2	4	6.2	4	21.7
3105	Male	9	2.8	4	5.1	4	22.8
	Female	7	2.7	4	4.6	4	21.3
3106	Male	9	3.6	4	7.2	4	24.9
	Female	7	3.1	4	6.7	4	24.0
3107	Male	10	5.7	5	8.5	5	26.7
	Female	3	5.6	3	7.7	3	25.5
3110	Male	8	5.3	4	8.7	4	25.7
	Female	4	4.9	4	7.2	4	24.0
3111	Male	11	3.6	4	7.1	4	24.0
	Female	6	3.5	4	6.7	4	23.1
3112	Male	7	4.3	4	7.2	4	24.7
	Female	8	3.8	4	6.2	4	22.0
n	Male	84		41		41	
	Female	61		39		39	
Mean	Male		4.0		7.0		23.9
	Female		3.7		6.4		22.6
S.D.	Male		1.0		1.1		1.6
	Female		0.9		0.9		1.6

Dose : Dodecyloxirane 1000 mg/kg

Base : Day 4 after birth

Unit : g

Species : Rat

Dam No.	Sex	/Days after birth /Before culling		/After culling			
		n	0-4	n	4-7	n	4-13
4102	Male	9	4.4	4	6.8	4	22.5
	Female	5	3.9	4	6.1	4	19.9
4103	Male	5	3.0	4	6.0	4	18.7
	Female	10	3.8	4	6.5	4	19.2
4104	Male	7	4.6	4	6.8	4	22.3
	Female	5	4.4	4	6.2	4	21.1
4105	Male	7	3.8	4	7.0	4	20.1
	Female	6	3.8	4	6.3	4	18.1
4107	Male	10	3.1	4	6.6	4	19.4
	Female	6	2.9	4	5.8	4	19.5
4108	Male	3	5.0	3	5.3	3	16.7
	Female	5	4.9	5	4.5	5	15.8
4109	Male	9	4.4	4	9.0	4	26.7
	Female	6	4.8	4	8.4	4	25.8
4110	Male	8	3.9	4	6.8	4	22.5
	Female	5	4.2	4	6.9	4	22.4
4111	Male	4	3.3	4	5.3	4	18.3
	Female	10	3.5	4	6.4	4	20.6
4112	Male	7	4.4	4	7.2	4	21.3
	Female	7	4.4	4	7.2	4	20.4
n	Male	69		39		39	
	Female	65		41		41	
Mean	Male		4.0		6.7		20.9
	Female		4.1		6.4		20.3
S.D.	Male		0.7		1.1		2.8
	Female		0.6		1.0		2.6

Appendix 18 - 1

DodecDodecyloxirane: An oral combined repeated dose toxicity study
with tr with the reproduction/ developmental toxicity screening test in rats
Externally for gross abnormalities in dead pups

Study No. : R-1261

Dose : Dodecyloxirane 0 mg/kg			Species : Rat
Animal No.	Number of pups examined Total (Male + Female)	Findings	
1101	0		
1102	0		
1103	0		
1104	0		
1105	0		
1106	0		
1107	0		
1108	0		
1109	0		
1110	0		
1111	0		
1112	0		
NAF : No abnormal findings			

		Dose : Dodecyloxirane 100 mg/kg		Species : Rat
Animal No.	Number of pups examined Total (Male + Female)	Findings		
2101	0			
2102	0			
2103	0			
2104	0			
2105	1 (0+1)	NAF		
2107	0			
2108	0			
2110	0			
2111	0			
2112	0			

NAF : No abnormal findings

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	Number of pups examined Total (Male + Female)		Findings
3101	0		
3102	0		
3103	0		
3104	0		
3105	0		
3106	0		
3107	0		
3110	0		
3111	1	(1+0)	NAF
3112	0		

NAF : No abnormal findings

Dose : Dodecyloxirane 1000 mg/kg			Species : Rat
Animal No.	Number of pups examined Total (Male + Female)	Findings	
4102	0		
4103	0		
4104	0		
4105	0		
4107	0		
4108	0		
4109	1 (0+1)	NAF	
4110	0		
4111	0		
4112	0		

NAF : No abnormal findings

Nipple retention of male pups						Days after birth : 13			
Generation : F0		Dose : Dodecyloxirane 0 mg/kg				Unit : No.	Species : Rat		
Dam No.	Sex	Offspring No. :	Nipple retention				Mean	S.D.	n
1101	Male	1101M01 : 0	1101M02 : 0	1101M03 : 0			0.0	0.0	3
1102	Male	1102M01 : 0	1102M02 : 0	1102M03 : 0	1102M04 : 0		0.0	0.0	4
1103	Male	1103M01 : 0	1103M02 : 0	1103M03 : 0	1103M04 : 0		0.0	0.0	4
1104	Male	1104M01 : 0	1104M02 : 0	1104M03 : 0	1104M04 : 0		0.0	0.0	4
1105	Male	1105M01 : 0	1105M02 : 0	1105M03 : 0	1105M04 : 0		0.0	0.0	4
1106	Male	1106M01 : 0	1106M02 : 0	1106M03 : 0	1106M04 : 0				
		1106M05 : 0					0.0	0.0	5
1107	Male	1107M01 : 0	1107M02 : 0	1107M03 : 0	1107M04 : 0		0.0	0.0	4
1108	Male	1108M01 : 0	1108M02 : 0	1108M03 : 0	1108M04 : 0		0.0	0.0	4
1109	Male	1109M01 : 0	1109M02 : 0	1109M03 : 0	1109M04 : 0		0.0	0.0	4
1110	Male	1110M01 : 0	1110M02 : 0	1110M03 : 0	1110M04 : 0		0.0	0.0	4
1111	Male	1111M01 : 0	1111M02 : 0	1111M03 : 0	1111M04 : 0		0.0	0.0	4
1112	Male	1112M01 : 0	1112M02 : 0	1112M03 : 0	1112M04 : 0		0.0	0.0	4

Nipple retention: Number of nipple / areolae

Experimental toxicity screening test in rats						Days after birth : 13		
Nipple retention of male pups		Dose : Dodecyloxirane 100 mg/kg				Species : Rat		
Generation : F0		Unit : No.				Species : Rat		
Dam No.	Sex	Offspring No. :	Nipple retention			Mean	S.D.	n
2101	Male	2101M01 : 0	2101M02 : 0	2101M03 : 0	2101M04 : 0	0.0	0.0	4
2102	Male	2102M01 : 0	2102M02 : 0	2102M03 : 0	2102M04 : 0	0.0	0.0	4
2103	Male	2103M01 : 0	2103M02 : 0	2103M03 : 0	2103M04 : 0	0.0	0.0	4
2104	Male	2104M01 : 0	2104M02 : 0	2104M03 : 0	2104M04 : 0	0.0	0.0	4
2105	Male	2105M01 : 0	2105M02 : 0	2105M03 : 0	2105M04 : 0	0.0	0.0	4
2107	Male	2107M01 : 0	2107M02 : 0	2107M03 : 0	2107M04 : 0	0.0	0.0	4
2108	Male	2108M01 : 0	2108M02 : 0	2108M03 : 0	2108M04 : 0	0.0	0.0	4
2110	Male	2110M01 : 0	2110M02 : 0	2110M03 : 0	2110M04 : 0	0.0	0.0	4
2111	Male	2111M01 : 0	2111M02 : 0	2111M03 : 0	2111M04 : 0			
		2111M05 : 0	2111M06 : 0			0.0	0.0	6
2112	Male	2112M01 : 0	2112M02 : 0	2112M03 : 0	2112M04 : 0	0.0	0.0	4
Nipple retention: Number of nipple / areolae								

Nipple retention: Number of nipple / areolae

Nipple retention of male pups							Days after birth : 13		
Generation : F0		Dose : Dodecyloxirane 300 mg/kg				Unit : No.	Species : Rat		
Dam No.	Sex	Offspring No. :	Nipple retention				Mean	S.D.	n
3101	Male	3101M01 : 0	3101M02 : 0	3101M03 : 0	3101M04 : 0	0.0	0.0	4	
3102	Male	3102M01 : 0	3102M02 : 0	3102M03 : 0	3102M04 : 0	0.0	0.0	4	
3103	Male	3103M01 : 0	3103M02 : 0	3103M03 : 0	3103M04 : 0	0.0	0.0	4	
3104	Male	3104M01 : 0	3104M02 : 0	3104M03 : 0	3104M04 : 0	0.0	0.0	4	
3105	Male	3105M01 : 0	3105M02 : 0	3105M03 : 0	3105M04 : 0	0.0	0.0	4	
3106	Male	3106M01 : 0	3106M02 : 0	3106M03 : 0	3106M04 : 0	0.0	0.0	4	
3107	Male	3107M01 : 0	3107M02 : 0	3107M03 : 0	3107M04 : 0				
	Male	3107M05 : 0				0.0	0.0	5	
3110	Male	3110M01 : 0	3110M02 : 0	3110M03 : 0	3110M04 : 0	0.0	0.0	4	
3111	Male	3111M01 : 0	3111M02 : 0	3111M03 : 0	3111M04 : 0	0.0	0.0	4	
3112	Male	3112M01 : 0	3112M02 : 0	3112M03 : 0	3112M04 : 0	0.0	0.0	4	

Nipple retention: Number of nipple / areolae

		Dose : Dodecyloxirane 1000 mg/kg				Days after birth : 13		
		Unit : No.				Species : Rat		
		Generation : F0						
Dam No.	Sex	Offspring No. : Nipple retention				Mean	S.D.	n
4102	Male	4102M01 : 0	4102M02 : 0	4102M03 : 0	4102M04 : 0	0.0	0.0	4
4103	Male	4103M01 : 0	4103M02 : 0	4103M03 : 0	4103M04 : 0	0.0	0.0	4
4104	Male	4104M01 : 0	4104M02 : 0	4104M03 : 0	4104M04 : 0	0.0	0.0	4
4105	Male	4105M01 : 0	4105M02 : 0	4105M03 : 0	4105M04 : 0	0.0	0.0	4
4107	Male	4107M01 : 0	4107M02 : 0	4107M03 : 0	4107M04 : 0	0.0	0.0	4
4108	Male	4108M01 : 0	4108M02 : 0	4108M03 : 0		0.0	0.0	3
4109	Male	4109M01 : 0	4109M02 : 0	4109M03 : 0	4109M04 : 0	0.0	0.0	4
4110	Male	4110M01 : 0	4110M02 : 0	4110M03 : 0	4110M04 : 0	0.0	0.0	4
4111	Male	4111M01 : 0	4111M02 : 0	4111M03 : 0	4111M04 : 0	0.0	0.0	4
4112	Male	4112M01 : 0	4112M02 : 0	4112M03 : 0	4112M04 : 0	0.0	0.0	4

Nipple retention: Number of nipple / areolae

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Number of pups examined		Findings
	Total	(Male + Female)	
1101	8	(3+5)	NAF
1102	5	(4+1)	NAF
1103	8	(4+4)	NAF
1104	8	(4+4)	NAF
1105	8	(4+4)	NAF
1106	8	(5+3)	NAF
1107	8	(4+4)	NAF
1108	8	(4+4)	NAF
1109	8	(4+4)	NAF
1110	8	(4+4)	NAF
1111	8	(4+4)	NAF
1112	8	(4+4)	NAF

NAF : No abnormal findings

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	Number of pups examined		Findings
	Total	(Male + Female)	
2101	8	(4+4)	NAF
2102	8	(4+4)	NAF
2103	8	(4+4)	NAF
2104	8	(4+4)	NAF
2105	8	(4+4)	NAF
2107	8	(4+4)	NAF
2108	8	(4+4)	NAF
2110	8	(4+4)	NAF
2111	8	(6+2)	NAF
2112	8	(4+4)	NAF

NAF : No abnormal findings

			Dose : Dodecyloxirane 300 mg/kg	Species : Rat
Animal No.	Number of pups examined			
	Total	(Male + Female)		
3101	8	(4+4)	NAF	
3102	8	(4+4)	NAF	
3103	8	(4+4)	NAF	
3104	8	(4+4)	NAF	
3105	8	(4+4)	NAF	
3106	8	(4+4)	NAF	
3107	8	(5+3)	NAF	
3110	8	(4+4)	NAF	
3111	8	(4+4)	NAF	
3112	8	(4+4)	NAF	

NAF : No abnormal findings

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Number of pups examined		Findings
	Total	(Male + Female)	
4102	8	(4+4)	NAF
4103	8	(4+4)	NAF
4104	8	(4+4)	NAF
4105	8	(4+4)	NAF
4107	8	(4+4)	NAF
4108	8	(3+5)	NAF
4109	8	(4+4)	NAF
4110	8	(4+4)	NAF
4111	8	(4+4)	NAF
4112	8	(4+4)	NAF

NAF : No abnormal findings

Sex : Male

Stage : After culled (F1 males)
Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Body weight	Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
1101M01	38.4	1.3	3.4	1.4	3.6	2.7	7.0
1102M01	39.4	3.7	9.4	2.8	7.1	6.5	16.5
1103M01	33.5	2.2	6.6	2.4	7.2	4.6	13.7
1104M01	37.8	1.5	4.0	2.0	5.3	3.5	9.3
1105M01	37.9	2.3	6.1	1.6	4.2	3.9	10.3
1106M01	36.0	2.6	7.2	2.3	6.4	4.9	13.6
1107M01	30.9	1.5	4.9	1.7	5.5	3.2	10.4
1108M01	37.1	1.3	3.5	1.5	4.0	2.8	7.5
1109M01	37.1	2.2	5.9	2.8	7.5	5.0	13.5
1110M01	35.6	1.7	4.8	2.1	5.9	3.8	10.7
1111M01	36.4	2.9	8.0	2.4	6.6	5.3	14.6
1112M01	32.1	1.9	5.9	2.1	6.5	4.0	12.5
n	12	12	12	12	12	12	12
Mean	36.0	2.1	5.8	2.1	5.8	4.2	11.6
S.D.	2.6	0.7	1.8	0.5	1.3	1.1	2.9

AB : Absolute weight, RE : Relative weight by body weight

Animal No.	Body weight	Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
2101M01	38.0	1.4	3.7	2.3	6.1	3.7	9.7
2102M01	39.0	2.7	6.9	2.0	5.1	4.7	12.1
2103M01	37.7	3.0	8.0	1.9	5.0	4.9	13.0
2104M01	35.1	3.1	8.8	2.3	6.6	5.4	15.4
2105M01	35.6	2.2	6.2	1.7	4.8	3.9	11.0
2107M01	29.3	2.1	7.2	2.0	6.8	4.1	14.0
2108M01	34.8	2.4	6.9	2.7	7.8	5.1	14.7
2110M01	42.4	2.3	5.4	2.1	5.0	4.4	10.4
2111M01	37.5	2.2	5.9	2.8	7.5	5.0	13.3
2112M01	36.6	2.1	5.7	1.5	4.1	3.6	9.8
n	10	10	10	10	10	10	10
Mean	36.6	2.4	6.5	2.1	5.9	4.5	12.3
S.D.	3.4	0.5	1.4	0.4	1.3	0.6	2.1

AB : Absolute weight, RE : Relative weight by body weight

Animal No.	Body weight	Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
3101M01	35.4	2.2	6.2	2.6	7.3	4.8	13.6
3102M01	33.7	1.6	4.7	1.8	5.3	3.4	10.1
3103M01	33.9	2.6	7.7	2.9	8.6	5.5	16.2
3104M01	31.7	1.5	4.7	1.5	4.7	3.0	9.5
3105M01	33.6	2.0	6.0	2.0	6.0	4.0	11.9
3106M01	34.5	1.6	4.6	1.7	4.9	3.3	9.6
3107M01	39.9	3.1	7.8	2.8	7.0	5.9	14.8
3110M01	39.3	2.8	7.1	2.3	5.9	5.1	13.0
3111M01	34.6	3.8	11.0	3.3	9.5	7.1	20.5
3112M01	35.5	2.5	7.0	2.2	6.2	4.7	13.2
n	10	10	10	10	10	10	10
Mean	35.2	2.4	6.7	2.3	6.5	4.7	13.2
S.D.	2.6	0.7	1.9	0.6	1.6	1.3	3.4

AB : Absolute weight, RE : Relative weight by body weight

Animal No.	Body weight	Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
4102M01	33.3	2.7	8.1	1.5	4.5	4.2	12.6
4103M01	29.1	2.7	9.3	2.7	9.3	5.4	18.6
4104M01	34.1	2.0	5.9	2.4	7.0	4.4	12.9
4105M01	28.4	1.8	6.3	2.1	7.4	3.9	13.7
4107M01	29.8	2.9	9.7	2.9	9.7	5.8	19.5
4108M01	30.9	4.7	15.2	2.6	8.4	7.3	23.6
4109M01	38.5	2.9	7.5	2.1	5.5	5.0	13.0
4110M01	33.8	2.3	6.8	2.9	8.6	5.2	15.4
4111M01	29.6	2.5	8.4	2.4	8.1	4.9	16.6
4112M01	34.5	2.3	6.7	2.7	7.8	5.0	14.5
n	10	10	10	10	10	10	10
Mean	32.2	2.7	8.4	2.4	7.6	5.1	16.0
S.D.	3.2	0.8	2.7	0.4	1.6	1.0	3.6

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : After culled (F1 females)

Sex : Female

Dose : Dodecyloxirane 0 mg/kg

Species : Rat

Animal No.	Body	Thyroid-R		Thyroid-L		Thyroid-RL	
	weight	AB	RE	AB	RE	AB	RE
	g	mg	mg/100g	mg	mg/100g	mg	mg/100g
1101F01	37.8	1.6	4.2	2.2	5.8	3.8	10.1
1102F01	35.6	0.9	2.5	1.0	2.8	1.9	5.3
1103F01	31.2	2.7	8.7	2.1	6.7	4.8	15.4
1104F01	35.0	2.5	7.1	2.5	7.1	5.0	14.3
1105F01	32.8	2.0	6.1	2.4	7.3	4.4	13.4
1106F01	34.5	2.4	7.0	2.4	7.0	4.8	13.9
1107F01	30.6	3.3	10.8	2.8	9.2	6.1	19.9
1108F01	35.8	2.4	6.7	2.5	7.0	4.9	13.7
1109F01	30.3	2.7	8.9	1.5	5.0	4.2	13.9
1110F01	34.6	1.7	4.9	1.8	5.2	3.5	10.1
1111F01	30.8	3.6	11.7	2.3	7.5	5.9	19.2
1112F01	33.7	1.6	4.7	2.2	6.5	3.8	11.3
n	12	12	12	12	12	12	12
Mean	33.6	2.3	6.9	2.1	6.4	4.4	13.4
S.D.	2.4	0.8	2.7	0.5	1.6	1.1	4.0

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : After culled (F1 females)

Sex : Female

Dose : Dodecyloxirane 100 mg/kg

Species : Rat

Animal No.	Body weight	Thyroid-R		Thyroid-L		Thyroid-RL	
	g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
2101F01	35.9	1.7	4.7	2.5	7.0	4.2	11.7
2102F01	36.8	3.6	9.8	2.0	5.4	5.6	15.2
2103F01	33.1	2.9	8.8	3.2	9.7	6.1	18.4
2104F01	36.3	2.7	7.4	2.3	6.3	5.0	13.8
2105F01	34.5	3.3	9.6	2.6	7.5	5.9	17.1
2107F01	33.0	1.5	4.5	1.4	4.2	2.9	8.8
2108F01	34.0	2.1	6.2	2.7	7.9	4.8	14.1
2110F01	36.7	2.1	5.7	2.2	6.0	4.3	11.7
2111F01	30.8	3.3	10.7	2.1	6.8	5.4	17.5
2112F01	31.5	2.4	7.6	2.2	7.0	4.6	14.6
n	10	10	10	10	10	10	10
Mean	34.3	2.6	7.5	2.3	6.8	4.9	14.3
S.D.	2.2	0.7	2.2	0.5	1.5	1.0	3.0

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : After culled (F1 females)

Sex : Female

Dose : Dodecyloxirane 300 mg/kg

Species : Rat

Animal No.	Body	Thyroid-R		Thyroid-L		Thyroid-RL	
	weight	AB	RE	AB	RE	AB	RE
	g	mg	mg/100g	mg	mg/100g	mg	mg/100g
3101F01	33.1	1.7	5.1	3.2	9.7	4.9	14.8
3102F01	30.0	2.3	7.7	2.8	9.3	5.1	17.0
3103F01	30.2	2.6	8.6	2.4	7.9	5.0	16.6
3104F01	31.5	2.6	8.3	2.9	9.2	5.5	17.5
3105F01	31.7	2.5	7.9	1.7	5.4	4.2	13.2
3106F01	36.4	2.7	7.4	2.8	7.7	5.5	15.1
3107F01	35.4	3.1	8.8	3.5	9.9	6.6	18.6
3110F01	37.1	2.6	7.0	2.8	7.5	5.4	14.6
3111F01	33.3	3.6	10.8	3.1	9.3	6.7	20.1
3112F01	32.5	2.9	8.9	2.2	6.8	5.1	15.7
n	10	10	10	10	10	10	10
Mean	33.1	2.7	8.1	2.7	8.3	5.4	16.3
S.D.	2.5	0.5	1.5	0.5	1.5	0.8	2.1

AB : Absolute weight, RE : Relative weight by body weight

Organ weight

Stage : After culled (F1 females)

Sex : Female

Dose : Dodecyloxirane 1000 mg/kg

Species : Rat

Animal No.	Body	Thyroid-R		Thyroid-L		Thyroid-RL	
	weight						
	g	AB mg	RE mg/100g	AB mg	RE mg/100g	AB mg	RE mg/100g
4102F01	32.3	2.4	7.4	2.1	6.5	4.5	13.9
4103F01	30.5	1.5	4.9	2.3	7.5	3.8	12.5
4104F01	33.0	2.7	8.2	2.4	7.3	5.1	15.5
4105F01	28.4	2.0	7.0	1.3	4.6	3.3	11.6
4107F01	31.5	1.4	4.4	1.1	3.5	2.5	7.9
4108F01	28.7	2.1	7.3	1.8	6.3	3.9	13.6
4109F01	36.6	2.8	7.7	2.2	6.0	5.0	13.7
4110F01	33.4	3.0	9.0	2.9	8.7	5.9	17.7
4111F01	33.8	2.0	5.9	1.8	5.3	3.8	11.2
4112F01	32.8	2.5	7.6	2.6	7.9	5.1	15.5
n	10	10	10	10	10	10	10
Mean	32.1	2.2	6.9	2.1	6.4	4.3	13.3
S.D.	2.5	0.5	1.4	0.6	1.6	1.0	2.7

AB : Absolute weight, RE : Relative weight by body weight

信頼性保証書（1/4）

試験番号 : R-1261

試験表題 : ドデシルオキシラン：ラットを用いた経口投与による反復投与毒性・生殖発生毒性併合試験

本試験は以下に示す基準に従って実施されたことを保証致します。

なお、一部委託部分（血清中ホルモン濃度測定）については試験場所信頼性保証部門により保証されていることを確認致しております。

- ・ 「新規化学物質等に係る試験を実施する試験施設に関する基準」
（平成 23 年 3 月 31 日：薬食発 0331 第 8 号、平成 23・03・29 製局第 6 号、環保企発第 110331010 号）

なお、調査は下記の通り実施し、報告致しました。

試験の調査

項 目	担当者	調 査 日	試験責任者及び 運営管理者への 報告日
試験計画書		2020 年 10 月 2 日	2020 年 10 月 2 日
作業予定表・ コンピュータプロトコール		2020 年 10 月 9 日	2020 年 10 月 9 日
改善確認		2020 年 10 月 13 日	2020 年 10 月 13 日
被験物質の安定性確認 （投与期間開始前）		2020 年 10 月 16 日	2020 年 10 月 16 日
試験計画書変更書（1）		2020 年 10 月 16 日	2020 年 10 月 16 日
試験計画書変更書（2）		2020 年 10 月 16 日	2020 年 10 月 16 日
調製・保存（被験物質）		2020 年 10 月 22 日	2020 年 10 月 22 日
被験液の分析		2020 年 10 月 22 日	2020 年 10 月 22 日
群分け		2020 年 10 月 26 日	2020 年 10 月 26 日
腔垢検査		2020 年 10 月 27 日	2020 年 10 月 27 日

信頼性保証書 (2/4)

項 目	担当者	調 査 日	試験責任者及び 運営管理者への 報告日
投与・一般状態の観察		2020 年 10 月 27 日	2020 年 10 月 27 日
詳細な一般状態の観察		2020 年 10 月 28 日	2020 年 10 月 28 日
交配		2020 年 11 月 11 日	2020 年 11 月 11 日
機能検査・握力測定・ 自発運動量測定		2020 年 12 月 2 日	2020 年 12 月 2 日
分娩・哺育観察		2020 年 12 月 2 日	2020 年 12 月 2 日
尿検査（尿量・色調・定性・沈渣）		2020 年 12 月 3 日	2020 年 12 月 3 日
摂水量測定		2020 年 12 月 4 日	2020 年 12 月 4 日
尿検査（浸透圧）		2020 年 12 月 4 日	2020 年 12 月 4 日
哺育児数調整		2020 年 12 月 7 日	2020 年 12 月 7 日
採血（生後 4 日出生児）		2020 年 12 月 7 日	2020 年 12 月 7 日
採血・剖検（雄）		2020 年 12 月 8 日	2020 年 12 月 8 日
病理組織学検査（切り出し）		2020 年 12 月 11 日	2020 年 12 月 11 日
乳頭及び乳輪数の観察		2020 年 12 月 15 日	2020 年 12 月 15 日
剖検（生後 13 日出生児）		2020 年 12 月 15 日	2020 年 12 月 15 日
剖検（母動物）		2020 年 12 月 16 日	2020 年 12 月 16 日
被験物質の安定性確認 （投与期間終了後）		2021 年 1 月 12 日	2021 年 1 月 12 日
最終報告書草案・図・表・付表		2021 年 3 月 5 日	
		2021 年 3 月 8 日	
		2021 年 3 月 9 日	
		2021 年 3 月 10 日	
		2021 年 3 月 11 日	
		2021 年 3 月 12 日	2021 年 3 月 13 日

信頼性保証書 (3/4)

項 目	担当者	調 査 日	試験責任者及び 運営管理者への 報告日
生データ		2021 年 3 月 5 日	
		2021 年 3 月 8 日	
		2021 年 3 月 9 日	
		2021 年 3 月 10 日	
		2021 年 3 月 11 日	
		2021 年 3 月 15 日	
		2021 年 3 月 16 日	
		2021 年 3 月 17 日	
		2021 年 3 月 18 日	2021 年 3 月 19 日
最終報告書		2021 年 3 月 23 日	2021 年 3 月 23 日

信頼性保証書 (4/4)

プロセス調査

項 目	試験番号	担当者	調 査 日	試験責任者及び 運営管理者への 報告日
動物入荷	M-1596		2020 年 8 月 19 日	2020 年 8 月 20 日
検疫・馴化	M-1596		2020 年 8 月 19 日	2020 年 8 月 20 日
	M-1596		2020 年 8 月 25 日	2020 年 8 月 25 日
体重・摂餌量測定	R-1257		2020 年 9 月 23 日	2020 年 9 月 23 日
生体試料の発送	B-8767		2020 年 11 月 4 日	2020 年 11 月 4 日
尿検査（電解質）	B-8705		2020 年 10 月 7 日	2020 年 10 月 7 日
血液学検査・ 血液化学検査	B-8705		2020 年 10 月 13 日	2020 年 10 月 13 日
病理組織学検査	B-8664		2020 年 10 月 14 日	2020 年 10 月 14 日
（包埋・薄切・染色・ 鏡検）	B-8664		2020 年 10 月 15 日	2020 年 10 月 15 日
	B-8664		2020 年 10 月 26 日	2020 年 10 月 26 日
	B-8649		2020 年 10 月 28 日	2020 年 10 月 28 日

2021 年 3 月 23 日

株式会社ボゾリサーチセンター

主信頼性保証部門