

最終報告書

表　題：ペルフルオロヘキサデカン酸のラットにおける反復投与毒性・生殖発生毒性併合試験

試験番号：S R 0 7 1 2 5

株式会社 化合物安全性研究所

陳述書

表題：ペルフルオロヘキサデカン酸のラットにおける反復投与毒性・生殖発生毒性併合試験

試験番号：S R 0 7 1 2 5

1. 本試験は、「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について」(平成 15 年 11 月 21 日薬食発第 1121003 号・平成 15・11・17 製局第 3 号・環保企発第 031121004 号、最終改正 平成 20 年 7 月 4 日 厚生労働省医薬食品局長・経済産業省製造産業局長・環境省総合環境政策局長連名通知) に従い、試験方法は、OECD 試験法ガイドライン (OECD Guideline for Testing of Chemicals; Combined Repeated Dose Toxicity Study with the Reproduction/Developmental Toxicity Screening Test (422), 1996. 03. 22) に基づいて実施したものであります。
2. 本試験は、試験計画書に従って実施し、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認められませんでした。

株式会社 化合物安全性研究所

試験責任者

2012年5月31日

信頼性保証書

表題：ペルフルオロヘキサデカン酸のラットにおける反復投与毒性・生殖発生毒性併合試験

試験番号：S R 0 7 1 2 5

本試験は、株式会社 化合物安全性研究所 QAUによって、下記のとおり査察された。

査 察 段 階	査 察 日	試験責任者 への報告日	運営管理者 への報告日
試験計画書	2010年12月6日	2010年12月6日	2010年12月6日
試験計画書変更書(No. 1)	2011年6月3日	2011年6月6日	2011年6月6日
試験計画書変更書(No. 2)	2011年6月20日	2011年6月20日	2011年6月20日
試験計画書変更書(No. 3)	2011年10月3日	2011年10月3日	2011年10月3日
試験計画書変更書(No. 4)	2012年3月7日	2012年3月7日	2012年3月7日
被験物質の受入・表示・保存	2010年12月6日	2010年12月6日	2010年12月6日
投与液の調製	2010年12月15日	2010年12月15日	2010年12月15日
投与液の化学分析	2010年12月15日 2010年12月21日	2010年12月21日	2010年12月21日
動物受入・検疫・馴化	2010年12月8日	2010年12月8日	2010年12月8日
群分け	2010年12月21日	2010年12月21日	2010年12月21日
投与	2010年12月27日	2010年12月27日	2010年12月27日
一般状態観察	2010年12月27日	2010年12月27日	2010年12月27日
体重測定	2010年12月27日	2010年12月27日	2010年12月27日
摂餌量測定	2010年12月27日	2010年12月27日	2010年12月27日
性周期検査	2010年12月27日	2010年12月27日	2010年12月27日
詳細な一般状態観察	2010年12月29日	2011年1月4日	2011年1月4日
生殖能検査(交配)	2011年1月5日 2011年1月6日	2011年1月6日	2011年1月6日
機能検査	2011年1月28日	2011年1月28日	2011年1月28日
尿検査	2011年1月30日 2011年1月31日	2011年1月31日	2011年1月31日
分娩および哺育状態観察	2011年1月28日	2011年1月28日	2011年1月28日
新生児の剖検	2011年2月1日	2011年2月1日	2011年2月1日

SR07125

査 察 段 階	査 察 日	試 験 責 任 者 へ の 報 告 日	運 営 管 理 者 へ の 報 告 日
剖検・器官重量測定	2011年2月3日	2011年2月3日	2011年2月3日
血液学的検査	2011年2月3日	2011年2月3日	2011年2月3日
血液化学的検査	2011年2月3日		
	2011年2月4日	2011年2月7日	2011年2月7日
病理組織学的検査(標本作製)	2011年2月7日		
	2011年7月4日	2011年7月6日	2011年7月6日
病理組織学的検査(鏡検)	2011年7月6日		
	2011年2月8日		
病理組織学的検査(標本作製)	2011年2月14日	2011年2月15日	2011年2月15日
	2011年2月15日		
生データ	2011年3月31日	2011年3月31日	2011年3月31日
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	2011年9月7日		
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1. 本試験は、「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について」(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 厚生労働省医薬食品局長・経済産業省製造産業局長・環境省総合環境政策局長連名通知)およびOECD試験法ガイドライン(OECD Guideline for Testing of Chemicals; Combined Repeated Dose Toxicity Study with the Reproduction/Developmental Toxicity Screening Test (422), 1996.03.22)に従い実施された。
2. 本試験は、試験計画書に従って実施され、また、本報告書には当該試験に使用した方法および手順が正確に記載されており、試験成績には当該試験の実施過程において得られた生データが正確に反映していることを確認した。

株式会社 化合物安全性研究所

Q A U 責任者

2012年 5月 8日

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表題：ペルフルオロヘキサデカン酸のラットにおける反復投与毒性・生殖発生毒性併合試験

試験番号：S R 0 7 1 2 5

試験目的：この試験は、ペルフルオロヘキサデカン酸を雌雄ラットに反復経口投与してその毒性ならびに性腺機能、交尾行動、受胎および分娩等の生殖に及ぼす毒性を検討するため行われた。

試験実施基準（GLP）および試験法ガイドライン

試験実施基準（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について」
(平成 15 年 11 月 21 日薬食発第 1121003 号・平成 15・11・17 製局第 3 号・
環保企発第 031121004 号、最終改正 平成 20 年 7 月 4 日 厚生労働省医
薬食品局長・経済産業省製造産業局長・環境省総合環境政策局長連名通
知)

試験法ガイドライン：OECD 試験法ガイドライン (OECD Guideline for Testing of Chemicals;
Combined Repeated Dose Toxicity Study with the Reproduction/
Developmental Toxicity Screening Test (422), 1996. 03. 22)

動物愛護

本試験は試験施設の動物実験倫理委員会の承認を得、かつ、標準操作手順書（動物実験倫理規定）に準拠した。

法規および基準等：「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、
最終改正 平成 18 年 6 月 2 日 法律第 50 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)
「動物実験に関する指針」(昭和 62 年 5 月 22 日承認 社団法人 日本実験動
物学会)

試験委託者

名称 : 厚生労働省 医薬食品局
所在地 : 東京都千代田区霞が関 1-2-2 (〒100-8916)
連絡先 : 審査管理課 化学物質安全対策室

試験施設

名称 : 株式会社 化合物安全性研究所
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所属 : 株式会社 化合物安全性研究所 安全性研究部

試験従事者およびその業務分担

被験物質管理 : [REDACTED]
化学分析 : [REDACTED]
臨床検査 : [REDACTED]
動物管理 : [REDACTED]
検疫・馴化 : [REDACTED]
投与・観察・測定 : [REDACTED]
病理検査 : [REDACTED]

試験期間

試験開始日	: 2010年 12月 6日
被験物質受入	: 2009年 7月 1日
動物受入	: 2010年 12月 8日
実験開始(投与開始)日	: 2010年 12月 23日
交配開始	: 2011年 1月 5日
新生児剖検開始	: 2011年 2月 1日
母動物剖検開始	: 2011年 2月 3日
雄動物および回復群雌動物投与終了	: 2011年 2月 2日
雄動物投与終了時剖検	: 2011年 2月 3日
回復群雌雄動物剖検	: 2011年 2月 17日
ホルモン測定終了	: 2011年 8月 30日
病理組織学的検査終了(実験終了)	: 2012年 3月 27日
試験終了日	: 2012年 5月 31日

要 約

ペルフルオロヘキサデカン酸の0(対照、0.5%カルボキシメチルセルロースナトリウム水溶液)、4、20および100 mg/kgを1群雌雄各12匹のCr1:CD(SD)ラットに、雄に対しては交配前、交配期間および交配後を含む計42日間、雌に対しては交配前、交配および妊娠期間、ならびに分娩後5日までの期間経口投与し、雌雄動物への反復投与による影響、雌雄動物の生殖および新生児の発生に及ぼす影響について検討した。また、0(対照)および100 mg/kgについて、雄動物は各5匹を選抜し、雌動物は非交配群として各5匹を別に設け、42日間の投与終了後14日間の回復性についても併せて検討した。

1. 反復投与毒性

一般状態観察および詳細な一般状態観察では、雌雄とも被験物質投与に関連する変化はみられなかった。機能検査では、回復2週に100 mg/kg群の雌雄とも後肢の握力に低値がみられた。

体重については、100 mg/kg群の雄で、投与35および42日の体重、投与1-42日の体重増加量および体重増加率に低値がみられた。摂餌量については、100 mg/kg群の雄で回復14日に、雌で妊娠5、7、10、14日、哺育4日に低値がみられた。

尿検査および血液学的検査では、雌雄とも被験物質投与に関連する変化はみられなかった。

ホルモン測定では、投与期間終了時に雌のすべての用量群でT3に低値がみられた。

血液化学的検査では、投与期間終了時に雌雄の100 mg/kg群および雌の20 mg/kg群でクロールに高値がみられ、雌の100 mg/kg群でナトリウムおよび尿素窒素が高値であった。

投与期間終了時の剖検では被験物質投与と関連する変化はみられなかった。器官重量では100 mg/kg群で雄の肝臓の絶対および相対重量に高値がみられ、病理組織学的検査では、小葉中心性肝細胞肥大および小葉中心性脂肪化が20 mg/kg以上の投与群で認められた。雌では、小葉中心性肝細胞肥大が100 mg/kg群でみられた。回復期間終了時に100 mg/kg群では、雌のクロールの高値傾向、雄の肝臓の絶対および相対重量に高値がみられ、病理組織学的検査では小葉中心性肝細胞肥大がみられたが、変化の程度は軽く回復傾向がみられた。

以上の結果から、本試験条件下におけるペルフルオロヘキサデカン酸の無影響量(NOEL)は雄で4 mg/kg/day、雌で4 mg/kg/day未満と考えられた。

2. 生殖発生毒性

主試験群の生殖能および回復群の性周期検査では、100 mg/kg群まで被験物質投与に関連する変化はみられなかった。新生児の生後4日の体重に100 mg/kg群で低値傾向がみられた。

したがって、本試験条件下におけるペルフルオロヘキサデカン酸の親動物の生殖に対する無影

影響量 (NOEL) は 100 mg/kg/day、新生児の発生に対する無影響量 (NOEL) は 20 mg/kg/day と考えられた。

緒 言

ペルフルオロヘキサデカン酸を、0 (対照、0.5%カルボキシメチルセルロースナトリウム水溶液)、4、20 および 100 mg/kg/day の用量で、1 群雌雄各 12 匹の Crl:CD (SD) ラットに、雄に対しては交配前、交配期間および交配後を含む計 42 日間、雌に対しては交配前、交配および妊娠期間、ならびに分娩後 5 日までの期間経口投与し、雌雄動物への反復投与による影響、雌雄動物の生殖および新生児の発生に及ぼす影響について検討した。また、0 (対照) および 100 mg/kg について、雄動物は各 5 匹を選抜し、雌動物は非交配群として各 5 匹を別に設け、42 日間の投与終了後 14 日間の回復性についても併せて検討した。

材料および方法

1. 被験物質

名称 : ペルフルオロヘキサデカン酸

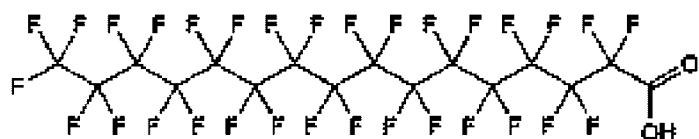
コード名 : PFHxDA

英名 : Perfluorohexadecanoic acid

CAS No. : 67905-19-5

官報公示整理番号 : 2-2658

構造式 :



分子式 : C₁₆HF₃₁O₂

分子量 : 814

物理化学的性質 : 外観 ; 白色粉末

融点 ; 125~126°C

沸点 ; 211°C/100mmHg

ロット番号 : 1262

純度 (GC) : 95.3% (Appendix 1-1)

不純物の濃度 : 不明 (データなし)

製造者 : 名称 ; Exfluor Research Corporation

所在地 ; 2350 Double Creek Drive Round Rock, Texas 78664, USA

入手量	: 978 g (関連試験と共に一括受入)
安定性および反応性	: 安定性；常温、常圧で安定 避けるべき条件；熱、粉塵 避けるべき物質；強酸化剤、強塩基、還元剤 関連試験を含め全試験操作の終了後、使用した被験物質の純度に関する分析成績を入手し、被験物質の試験期間中の安定性を確認した (Appendix 1-2)。
保存条件	: 密閉、冷所 (実測範囲 : 2~10°C) 強酸化剤、強塩基、還元剤との保管は避けた。
保存場所	: 株式会社 化合物安全性研究所の被験物質保存室
取扱い上の注意	: 吸い込んだり、眼、皮膚および衣類に触れないように、適切な保護具を着用した。
サンプリング	: 被験物質サンプルとして、約 1 g を採取し、試験施設の資料保存室に保存した。
残余被験物質の処置	: 関連試験も含めすべての試験操作終了後、焼却処分するために、産業廃棄物として回収した。
参考文献	: 日本化学物質辞書 Web ; 独立行政法人 科学技術振興機構 http://nikkajiweb.jst.go.jp/nikkaji_web/pages/top.html Material Safety Data Sheet, Exfluor Research Corporation

2. 媒体

名称	: 0.5%カルボキシメチルセルロースナトリウム水溶液 (0.5%CMC-Na)
調製方法	: 日本薬局方カルメロースナトリウム (ロット番号 0803、丸石製薬株式会社) を精秤し、日本薬局方精製水 (ロット番号 006057、008036 および 008038、ヤクハン製薬株式会社) に、所定の濃度となるように溶解した。
保存条件	: 遮光気密容器に入れ、冷蔵 (実測範囲 : 2.1~6.4°C) で保存した。
使用期限	: 調製後 15 日以内 (調製後 10 日以内に使用した)。

3. 投与液の調製および化学分析

調製方法	: 被験物質を正確に秤量し、所定の濃度となるように媒体 (0.5%CMC-Na) を加えて懸濁した。
調製頻度	: 10 日間に 1 回以上の頻度で調製した。
保存条件	: 遮光気密容器に入れ、冷蔵保存 (実測範囲 : 2.1~6.4°C) し、10 日以内に使用した。

保存場所 : 株式会社 化合物安全性研究所の被験物質保存室

調製上の注意 : 被験物質はクリーンベンチ内で取扱い、調製の際にはゴム手袋およびマスクを着用し、吸い込んだり、眼、皮膚および衣類に触れないようにした。

残余調製液の処置 : 残余の投与液は、焼却処分するために産業廃棄物として回収した。

調製液の安定性および均一性 : 投与開始前に、0.01 および 100 mg/mL の濃度の調製液について分析を実施し、被験物質の投与液中における均一性ならびに冷蔵保存 10 日間（調製日を 0 日として起算した）および室温保存 4 時間後の安定性を確認した（Appendix 2-1～2-2）。

調製液の濃度確認 : 被験物質の全濃度に関する投与液中における濃度を、初回および最終回調製時の計 2 回分析した。その結果、含有率は設定濃度の 98.5～104.5% であり、判定基準を満たしていた（Appendix 2-3 および 2-4）。

被験物質調製液の濃度分析方法 :

1) 使用機器

高速液体クロマトグラフ-質量分析計 (LC-MS/MS)

2695 (LC 部)	Waters Corporation
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Quattro Micro (MS/MS 部)	Waters Corporation
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データ処理装置	Mass Lynx V. 4.0	Waters Corporation
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電子天秤	xp205	METTLER TOLEDO
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純水製造装置	Milli-Q Labo	日本ミリポア株式会社
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2) 標準物質(気密容器に入れ、冷暗所に保存)

ペルフルオロヘキサデカン酸 (PFHxD) (被験物質)

Lot. No. 1262	Exfluor Research Corporation
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3) 試薬

¹³C-ペルフルオロオクタン酸 (¹³C-PFOA) (内標準物質)

Lot No. SCHG-007 (50 μg/mL メタノール溶液)	
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	Cambridge Isotope Laboratories, Inc.
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メタノール	高速液体クロマトグラフィー用	関東化学株式会社
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酢酸アンモニウム	試薬特級	関東化学株式会社
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4) 調製(以下の割合で調製、調製日を0日として起算)

a) 標準溶液

ペルフルオロヘキサデカン酸0.010 gを正確に量りとり、100 mLメスフラスコに入れ、メタノールを加えて100 μ g/mL溶液を調製した。さらにこの液1 mLを正確に採取し、100 mLメスフラスコに入れ、メタノールを加えて1 μ g/mL溶液を調製した(標準原液)。標準原液をメタノールで希釈し、50、100、150、200および250 ng/mL溶液を調製し、標準溶液(検量線)とした。なお、各標準溶液には、最終濃度として100 ng/mLとなるように¹³C-PFOAを内標準としてそれぞれ添加した。

b) 試料溶液

被験物質調製液の採取点数は、濃度確認試験および安定性試験については被験物質調製液の中層付近から2点とし、均一性試験については被験物質調製液の上、中、下層付近から各2点の計6点とした。なお、均一性試験の中層の濃度を安定性試験の調製時の分析結果とした。

① 被験物質調製液を採取し、¹³C-PFOAの濃度が100 ng/mLとなるように添加した後、被験物質の濃度が検量線の範囲(100 ng/mL付近)となるようにメタノールで希釈したものを試料溶液とした。

② 試料溶液の調製は、1点につき1回、HPLCへの注入は各1回とした。

c) 移動相

① 酢酸アンモニウム0.77 gを純水1 Lに溶解し、10 mmol/L酢酸アンモニウム水溶液とした。

調製後は室温で保存し、使用期限は1カ月とした。

② メタノールおよび純水はそのまま用い、溶媒の混合(グラジエント)はLC装置で行った。

d) オートサンプラ洗浄液

メタノールおよび蒸留水は等量混合したものを使用した。調製後は室温で保存し、使用期限は1カ月とした。

e) 洗浄用注入液

メタノールを使用した。

5) HPLC条件

カラム : Waters Atlantis dC18、3 μ m、2.1×150 mm

移動相 : グラジエント(下記参照)

時間(分)	純水	メタノール	10 mmol/L 酢酸アンモニウム水溶液
0~15	15	80	5

オートサンプラ洗浄液 : メタノール/蒸留水(1:1)の混合溶液

洗浄用注入液 : メタノール

カラム温度 : 50°C

カラム流量	: 0.2 mL/min
注入量	: 5 μ L
オートサンプラー温度	: 20°C
イオン化モード	: ESI-
キャピラリー電圧	: 1.0 kV
脱溶媒ガス	: 窒素
コーンガス流量	: 50 L/hr
ソース温度	: 110°C
コリジョンガス (MS/MS 部)	: ヘリウム
分析時間	: 15 分

6) システム適合性試験

測定日毎に標準溶液 (100 ng/mL) を連続して 6 回注入した。ペルフルオロヘキサデカン酸と 13C-PFOA の面積比および保持時間について変動係数を求めた。

7) 計算

Mass Lynx V. 4.0 を用いて、各標準物質および内標準物質の（子イオンの）ピーク面積を求め、それぞれのピークの面積比 (RES) と濃度から検量線を作成した。検量線から各試料溶液の被験物質濃度を求め、さらに変動係数、含有率および残存率を以下の式より算出した。

MS/MS によるモニターイオン

	親イオン	子イオン
ペルフルオロヘキサデカン酸	813.0	769.0
13C-ペルフルオロオクタン酸	421.0	376.0

$$\text{面積比(RES)} = \frac{\text{測定対象物質(または標準物質)のピーク面積}}{\text{内標準物質のピーク面積}}$$

$$\text{被験物質濃度(mg/mL)} = \frac{\text{測定濃度(ng/mL)} \times \text{希釈係数}}{1,000,000}$$

$$\text{変動係数(\%)} = \frac{\text{標準偏差}}{\text{平均値}} \times 100$$

$$\text{含有率(\%)} = \frac{\text{被験物質濃度平均値}}{\text{調製液の表示濃度}} \times 100$$

$$\text{残存率(\%)} = \frac{\text{保存後の被験物質濃度平均値}}{\text{調製時の被験物質濃度平均値}} \times 100$$

8) 数値の表示

- a) 標準溶液の濃度は秤量値より算出し、四捨五入して有効数字3桁に丸めた（計算値）。
- b) 調製液の被験物質濃度は四捨五入して有効数字3桁に丸めた。ただし、3桁以上の整数となる場合は小数点以下第1位を四捨五入し、整数で表示した。
- c) 変動係数、含有率および残存率は四捨五入して小数点以下第1位に丸めた。

9) 判定基準

- a) 濃度確認試験：含有率が85～115%の場合を適とした。
- b) 安定性試験：残存率が85～115%の場合を適とした。
- c) 均一性試験：変動係数が5%以下の場合を適とした。
- d) システム適合性試験：変動係数が2%以下の場合を適とした。

4. 試験方法

(1) 試験系

種・系統	: ラット、Cr1:CD (SD)
微生物統御	: SPF
生産業者	: 日本チャールス・リバー株式会社 厚木飼育センター
微生物モニタリング	: 動物生産業者よりデータを入手した。
動物選定理由	: ラットはこの種の試験で通常用いられている動物種であり、繁殖成績が安定していることと当研究所における背景対照データが利用できることから、この系統を選定した。
発注動物数	: 雄52匹、雌62匹
受入動物数	: 雄54匹、雌64匹
発注動物週齢	: 雌雄とも8週齢
出荷体重基準	: 雄は240～330 g、雌は160～230 g
受入時体重範囲	: 雄は248～287 g、雌は176～212 g
投与開始時週齢	: 雌雄とも10週齢
群数	: 雌雄各6群（主試験群 雌雄各4群、回復群 雌雄各2群）
各群動物数	: 主試験群 雌雄各12匹、回復群 雌雄各5匹

(2) 検疫および馴化

- 検疫方法 : 一般状態を1日1回観察し、体重を動物受入日、検疫および馴化8日ならびに検疫および馴化期間終了日（投与開始前々日）に測定した。期間中角膜の外傷が雌1例、切歯破折および不正咬合が雌1例にみられた。

他の動物には異常はみられなかった。

性周期検査	: 雌動物について、群分け日までの 9 日間に膣垢スメア塗抹法により性周期検査を行った。検査の結果、雌 1 例に性周期の異常がみられた。
期間	: 13 日間

(3) 群分け

検疫および馴化期間終了日（投与開始前々日）に、性周期の異常および一般状態に異常のみられた動物を除くすべての動物の体重に基づいて、層化無作為抽出法により各群の平均体重が均一になるように群分けを行った。これらの動物の体重範囲は、雄で 350～415 g、雌で 222～267 g であり、平均体重（雄、382.4 g；雌、242.5 g）の±20%以内であった。選択された動物の投与開始前日の一般状態の観察では、異常はみられなかった。選抜から外れた動物は試験から除外後、標準操作手順書に従って取扱った。

(4) 動物およびケージの識別

動物	: 群分け前は油性フェルトペンで尾部に印を付け、個体識別を行った。 群分け後は耳介に動物番号を入れ墨し、個体識別を行った。 新生児については、個体の識別は行わなかった。
飼育ケージ	: 群分け前は性別毎に色分けしたラベルに試験番号および受入時の動物番号を明記し、各ケージの前面に標示した。 群分け後は性別毎に色分けしたラベルに試験番号、試験群および群分け後の動物番号を明記し、各ケージの前面に標示した。

(5) 動物飼育

1) 飼育環境

飼育室番号	: 303 号室
温度・湿度	: $22 \pm 3^{\circ}\text{C}$, $50 \pm 20\%$ (実測範囲 $20 \sim 25^{\circ}\text{C}$, $41 \sim 55\%$)
換気回数	: 10～15 回／時間
照明時間	: 人工照明 12 時間 (8:00～20:00)

2) 飼育器材および飼育方法

ケージの種類	: プラケット式金属製金網床ケージ (260W×380D×180H, mm) ただし、交尾成立雌動物については妊娠 17 日から哺育 4 日まで同ケージの金網床を小型受皿に代えて実験動物用床敷 (ホワイトフレーク、日本チャールス・リバー株式会社) を使用して分娩と哺育を行わせた。
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1 ケージあたりの収容動物数 :

検疫および馴化期間中は 2 匹ずつ、群分け後は 1 匹、交配期間中は雌雄各 1 匹、分娩後は 1 腹を収容した。

ケージおよび給餌器の交換 :

群分け時に 1 回実施し、その後は 2 週に 1 回の頻度で交換した。ただし、交尾成立雌動物については妊娠 0 日および 14 日に実施した。

受皿交換 : 週 2 回洗浄滅菌済みのものと交換した。

自動給水装置の水抜き : 週 1 回実施した。

給水器の交換 : 尿検査時にのみ用いた。

小型受皿の交換 : 妊娠 20 日に実施した。

室内の清掃および清拭消毒 :

1 日 1 回実施した。清拭消毒に際しては、塩素系消毒薬およびヨウ素系消毒薬を 1 週間単位で交互に使用した。

3) 飼料

種類・名称 : 固型飼料、CRF-1

ロット番号 : 100907、101005

製造業者 : オリエンタル酵母工業株式会社

給餌方法 : 金属製給餌器を用いて自由に摂取させた。

ただし、剖検前日の夕刻からは全例を絶食させた。

汚染物質および微生物検査 :

試験に悪影響を及ぼす恐れのある汚染物質の分析あるいは微生物検査を、使用した各ロットの飼料について実施した。汚染物質の分析は Eurofins Analytics 社 (分析報告書 : AR-10-JP-001058-01、AR-10-JP-001107-01) が、微生物検査は飼料製造業者 (分析試験報告書 : No. 10G03-139、10G03-153) がそれぞれ行った。分析データを飼料製造業者からロット毎に入手した。分析項目と許容値は株式会社 化合物安全性研究所の標準操作手順書に準拠した。分析の結果、いずれの項目にも規定された許容値を超える値は認められなかった。

4) 飲料水

種類 : 札幌市水道水

給水方法 : 自動給水装置を用いて自由に摂取させた。ただし、尿検査時は給水器を用いた。

汚染物質検査 : 試験に悪影響を及ぼす恐れのある汚染物質の分析を、2010 年 10 月 1 日、2011 年 1 月 1 日および 2011 年 4 月 1 日に、当該飼育室と同系

統配管の最末端（301号室）から試料を採取して実施した（水質検査結果表：No. A223085、A224281、A230001）。分析は日本衛生株式会社において行い、分析データを入手した。分析項目と許容値は株式会社 化合物安全性研究所の標準操作手順書に準拠した。分析の結果、いずれの項目にも規定された許容値を超える値は認められなかった。

(6) 被験物質の投与

1) 投与量の設定

- 投与量 : 0 (媒体のみ)、4、20 および 100 mg/kg/day とした。
- 設定理由 : 先に実施した予備試験 (S R 0 7 1 2 5 P)¹⁾ (投与量 0、20、100、500 および 800 mg/kg/day、雌雄各 4 匹/群、2 週間反復経口投与) の結果、体重については、500 および 800 mg/kg 群の雌雄とも投与期間後期に体重の減少がみられ、投与 1-14 日の体重増加量および体重増加率は有意な低値であった。摂餌量については、800 mg/kg 群において、雌雄とも低値傾向または有意な低値がみられた。血液学的検査では、雄では 800 mg/kg 群で、雌では 500 および 800 mg/kg 群で、赤血球数およびヘモグロビン濃度に有意な高値がみられた。血液化学的検査において、20 および 100 mg/kg 群の雄で総コレステロールに有意な低値がみられた。500 および 800 mg/kg 群では、雌雄とも総蛋白および総コレステロールに有意な低値、トリグリセリドに低値傾向がみられた。剖検所見では、500 および 800 mg/kg 群において、雌雄とも肝臓の淡褐色化が全例に観察された。肝臓重量については、100、500 および 800 mg/kg 群の雄で絶対および相対重量に有意な高値または高値傾向がみられ、500 および 800 mg/kg 群の雌で相対重量に有意な高値がみられた。したがって、本試験では投与期間の延長を考慮して、100 mg/kg/day を最高用量とし、以下公比を 5 として 20 および 4 mg/kg/day を設定した。そのほかに媒体のみを同様の方法で投与する対照群を設けた。

試験群の構成 : 試験群の構成と各群の動物番号は以下の通りとした。

試験群	投与量 (mg/kg)	濃度 (mg/mL)	投与容量 (mL/kg)	動物数(動物番号)	
				雄	雌
<主試験群>					
対照群	0	0	10	12 (101～112)	12 (151～162)
低用量群	4	0.4	10	12 (201～212)	12 (251～262)
中用量群	20	2	10	12 (301～312)	12 (351～362)
高用量群	100	10	10	12 (401～412)	12 (451～462)
<回復群>					
対照群	0	0	10	5 (102, 105, 107, 108, 111)	5 (163～167)
高用量群	100	10	10	5 (405, 406, 409, 410, 411)	5 (463～467)

対照群の動物には、被験物質投与群と同様の方法で媒体のみを投与した。

回復群の雄は、主試験群から交配期間終了後に投与 28 日の体重に基づいて、全体の平均値に近似するよう集団の中央値の周辺に位置する動物を各群から 5 匹ずつ選抜した。

2) 投与

投与方法および投与経路：ディスポーザブル胃ゾンデおよびディスポーザブルシリンジを用いて強制的に胃内に経口投与した。

投与回数 : 1 日 1 回、連日投与した。

投与時刻 : 9 : 11～11 : 59

投与期間 : 雄；交配 14 日前より 42 日間

雌；交配前 14 日間および交尾成立までの交配期間、さらに交尾成立例は妊娠期間および分娩後 5 日（哺育 0 日を分娩後 0 日として起算）までの期間、妊娠 25 日まで分娩が認められない交尾成立例については妊娠 25 日までの期間

回復群については 42 日間

回復期間 : 投与期間終了後 14 日間

投与液量 : 各個体の投与液量は投与日に最も近い測定日の体重に基づいて算出した。

投与方法、投与経路、投与回数および投与期間の選定理由：
試験法ガイドラインを参考にした。

(7) 観察、測定および検査項目

I. 反復投与毒性

投与開始日を投与 1 日と起算し、さらに、主試験群の雌は交尾成立日を妊娠 0 日、分娩終了日を哺育 0 日、回復群の雌雄は投与 42 日の翌日を回復 1 日と起算した。

1) 一般状態観察

例数	: 全例
期間	: 投与 1 日から剖検日まで。
頻度	: 投与期間中は投与前および投与後の 1 日 2 回、回復期間中は午前および午後の 1 日 2 回、剖検日は午前中に 1 回観察した。
観察方法	: 個々の動物の生死、外観、行動等について観察した。異常が認められる場合は、その症状ならびに症状の持続期間を記録した。

2) 詳細な一般状態観察

例数	: 全例
時期	: 投与開始前ならびに投与 7、14、21、28、35 および 42 日。 回復群の動物については、回復 7 および 14 日。
観察方法	: あらかじめ定めたスコアリング基準を用いてスコア化した観察結果を記録した。

観察項目およびその方法 :

- ①体位・姿勢、呼吸状態、振戦・痙攣、常同行動（回転・旋回）、異常行動（自咬）をケージ外から観察した。
- ②取り出し易さ、取扱い易さ、筋収縮性、立毛、被毛の状態、皮膚、眼・眼球および粘膜の外観、瞳孔径、流涙、流涎、その他分泌物の有無について、ケージから取り出す時に観察した。
- ③歩行、運動協調性、環境刺激に対する反応、探索行動、排泄状態（排尿・排糞）、常同行動（身づくろい・くびふり）、異常行動（後ずさり・異常発声）、攻撃性について、オープンフィールド内で観察した。

3) 機能検査

例数	: 各群の体重の平均値に近似するように選抜した雄（回復群を含む）および回復群の雌の各群 5 例ならびに主試験群の雌の分娩日の早いものから順に選抜した各群 5 例。
時期	: 投与 6 週（投与 37 日）および回復 2 週（回復 12 日）。 主試験群の雌は哺育 1 日。
観察／測定方法	: あらかじめ定めたスコアリング基準を用いてスコア化した観察結果

あるいは測定機器による測定値を記録した。

観察項目およびその方法：

①刺激に対する感覚運動反応：検査台上で以下を観察した。

視覚刺激、触覚刺激、聴覚刺激、痛覚刺激、固有受容器刺激、空
中正向反射

②握力：CPU ゲージ（アイコーエンジニアリング株式会社）を用い
て前肢および後肢について各 3 回測定し、1 g 単位で記録した。

③自発運動量：自発運動量測定装置（スーパーメックスおよび
CompACT、室町機械株式会社）を用いて測定した。上記に引き続き、
10 分間隔で 1 時間測定した。

4) 体重測定

例数 : 全例

測定日 : 投与 1、3、5、7、10、14、21、28、35 および 42 日の投与前、回復
7 および 14 日ならびに剖検日。

主試験群の雌は投与 1、3、5、7、10、14 日の投与前、妊娠 0、1、3、
5、7、10、14、17 および 20 日の投与前、哺育 0、1 および 4 日の投
与前ならびに剖検日。

測定方法 : 電子式上皿天秤（GX-2000、株式会社エー・アンド・デイ）を用い
て測定し、1 g 単位で記録した。

体重増加量および体重増加率：

以下の式により算出した。

〈雄および回復群の雌〉

投与期間；

$$\text{体重増加量(g)} = \text{投与 42 日体重(g)} - \text{投与 1 日体重(g)}$$

$$\text{体重増加率(%)} = \frac{\text{体重増加量(g)}}{\text{投与 1 日体重(g)}} \times 100$$

回復期間；

$$\text{体重増加量(g)} = \text{回復 14 日体重(g)} - \text{投与 42 日体重(g)}$$

$$\text{体重増加率(%)} = \frac{\text{体重増加量(g)}}{\text{投与 42 日体重(g)}} \times 100$$

〈主試験群の雌〉

妊娠前投与期間；

$$\text{体重増加量(g)} = \text{投与 14 日体重(g)} - \text{投与 1 日体重(g)}$$

$$\text{体重増加率(%)} = \frac{\text{体重増加量(g)}}{\text{投与 1 日体重(g)}} \times 100$$

妊娠期間；

$$\text{体重増加量(g)} = \text{妊娠 20 日体重(g)} - \text{妊娠 0 日体重(g)}$$

$$\text{体重増加率(%)} = \frac{\text{体重増加量(g)}}{\text{妊娠 0 日体重(g)}} \times 100$$

哺育期間；

$$\text{体重増加量(g)} = \text{哺育 4 日体重(g)} - \text{哺育 0 日体重(g)}$$

$$\text{体重増加率(%)} = \frac{\text{体重増加量(g)}}{\text{哺育 0 日体重(g)}} \times 100$$

5) 摂餌量測定

- 例数 : 全例
- 測定日 : 雄雌とも剖検日および交配期間を除き、体重測定と同じ日に実施した。ただし、主試験群の雌については、妊娠 20 日は残量、哺育 0 日は給与量のみを測定した。
- 測定方法 : 電子式上皿天秤 (GX-2000、株式会社エー・アンド・デイ) を用いて測定し、1 g 単位で記録した。投与開始前日に適当量を測定後ケージ毎に給与し、その後は測定日に残量および給与量を測定した。ただし、最終回の測定は残量のみとした。
以下の式により、摂餌量 (g/rat/day) を算出した。

$$\text{摂餌量(g/rat/day)} = \frac{\text{給与量(g/rat)} - \text{残量(g/rat)}}{\text{測定日間の日数(day)}}$$

6) 尿検査

- 例数 : 雄は機能検査と同一の各群 5 例について、雌は回復群の各群 5 例
- 時期 : 投与 6 週 (投与 39~40 日) および回復 2 週 (回復 12~13 日)。
- 採尿方法 : 非絶食下でラット用代謝ケージ (KN-646、B-1 型、株式会社夏目製作所) を用いて採尿し、投与直後から約 3 時間の蓄尿で①~⑧を、また約 21 時間の蓄尿で⑨および⑩を実施した。採取した尿は検査終了後廃棄した。

検査項目および検査方法 :

①pH	試験紙法
②蛋白 (Protein)	試験紙法
③糖 (Glucose)	試験紙法
④ケトン体 (Ketone body)	試験紙法
⑤ウロビリノーゲン (Urobilinogen)	試験紙法
⑥ビリルビン (Bilirubin)	試験紙法
⑦潜血反応 (Occult blood)	試験紙法
⑧色調 (Color)	肉眼観察
⑨尿量 (Urine Volume)	容量測定

<u>⑩比重(Specific gravity)</u>	屈折計法
①～⑦ マルティスティックス、シーメンスヘルスケア・ダイアグノスティクス	
⑩ 尿比重屈折計ユリコン-S、アタゴ	

7) 血液学的検査

例数	: 各群 5 例 (主試験群の雄は機能検査に用いた動物以外から動物番号の若い順に選抜した。主試験群の雌は機能検査と同一例、回復群は全例。)
時期	: 剖検時 (主試験群の雄は投与 42 日の翌日、雌は分娩後 6 日、回復群は回復 14 日の翌日) に採血した。
採血方法	: 16～22 時間の絶食下でラットをエーテル麻酔し、腹部大動脈より採血した。検査項目のうち、①～⑩については EDTA・2K (ベノジェクト II 真空採血管、テルモ株式会社) で処理した血液約 1 mL を用い、⑪および⑫については 3.8% クエン酸ナトリウムで処理した血液約 1～2 mL を、3500 回転/分で 10 分間の遠心分離して得られた血漿を用いた。得られた血液および血漿は検査終了後廃棄した。なお、白血球塗抹標本 (May-Grünwald-Giemsa 染色) を作製し、保存した。白血球の分布異常はみられなかつたため、白血球塗抹標本の鏡検は行わなかつた。

検査項目および検査方法 :

①赤血球数(RBC)	電気抵抗検出法
②ヘマトクリット値(HCT)	電気抵抗検出法
③ヘモグロビン濃度(HGB)	SLS ヘモグロビン法
④平均赤血球容積(MCV)	RBC, HCT 値より算出
⑤平均赤血球ヘモグロビン量(MCH)	RBC, HGB 値より算出
⑥平均赤血球ヘモグロビン濃度(MCHC)	HCT, HGB 値より算出
⑦網赤血球数(Reticulocyte)	フローサイトメトリー法
⑧血小板数(Platelet)	電気抵抗検出法
⑨白血球数(WBC)	フローサイトメトリー法
⑩白血球分画 (Differential count of WBC)	フローサイトメトリー法
⑪プロトロンビン時間(PT)	トロンボプラスチン法
⑫活性化部分トロンボプラスチン時間(APTT)	エラジン酸法

①～⑩ 自動血球分析装置 XT-2000 iV、シスマックス
 ⑪⑫ 血液凝固自動測定装置 KC4 デルタ、トリニティ・バイオテック

8) 血液化学的検査

例数	: 血液学的検査と同じ各群 5 例
時期	: 剖検時に採血した。
採血方法	: 16～22 時間の絶食下でラットをエーテル麻酔し、腹部大動脈より採

血した。検査項目のうち、(1)および(5)については、血液 1 mL あたりヘパリンナトリウム (ヘパリンナトリウム注N「味の素」、1000 単位/mL、味の素株式会社) 約 20 単位で処理後、3500 回転/分で 10 分間の遠心分離で得られた血漿を用いて検査した。他の項目については分離剤入り試験管 (セパクリーン A、栄研器材株式会社) に血液を採取し、3500 回転/分で 10 分間の遠心分離で得られた血清を用いて検査した。また、得られた血清の一部は(20)～(22)測定用としてポリプロピレン製チューブ 4 本に約 150 μ L ずつ分注し、測定に供するまで超低温フリーザー (-80°C 設定) に保存した。これらのうち 2 本は(20)～(21)測定用として三菱化学メディエンス株式会社に送付し、測定結果を入手した。得られた血漿および血清は検査終了後 -20°C 以下で凍結保存し、試験終了日に廃棄した。

検査項目および検査方法：

(1) AST	JSCC 法
(2) ALT	JSCC 法
(3) アルカリホスファターゼ(ALP)	JSCC 法
(4) γ -GTP	L- γ -グルタミル-3-カルボキシ-4-ニトロアニリド基質法
(5) グルコース(Glucose)	ヘキソキナーゼ法
(6) 総コレステロール(T-Chol)	酵素法
(7) トリグリセリド(TG)	遊離グリセロール消去法
(8) 総ビリルビン(T-Bil)	アゾビリルビン法
(9) 尿素窒素(UN)	ウレアーゼ・GLDH 法
(10) クレアチニン(Crea)	Jaffé 法
(11) ナトリウム(Na)	イオン選択電極(ISE)法
(12) カリウム(K)	イオン選択電極(ISE)法
(13) クロール(Cl)	イオン選択電極(ISE)法
(14) カルシウム(Ca)	OCPC 法
(15) 無機リン(IP)	Fiske-Subba Row 法
(16) 総蛋白(TP)	ビウレット法
(17) 蛋白分画(Protein fraction)	セルロースアセテート膜電気泳動法
(18) A/G 比(A/G ratio)	蛋白分画より算出
(19) アルブミン(Albumin)	総蛋白と蛋白分画より算出
(20) 甲状腺刺激ホルモン(TSH)	RIA 法
(21) T3	RIA 法
(22) T4	ELISA 法

(1)～(16) 自動分析装置 7080 形、日立ハイテクノロジーズ

(17) 自動電気泳動装置 AES320、三島オリンパス

(20)～(21) 三菱化学メディエンス株式会社で測定

(22) ウエルリーダーSK601、株式会社 サイニクス

9) 割検

例数 : 全例

時期	: 主試験群の雄は投与 42 日の翌日、雌の分娩例は分娩後 6 日（哺育 4 日の翌々日）に、妊娠 25 日まで分娩が認められない交尾成立例は妊娠 26 日に、回復群は雌雄とも回復 14 日の翌日に剖検した。
検査方法	: 体外表を観察し、エーテル麻酔下で採血後、放血により安樂死させ、全身の器官・組織を肉眼的に観察した。検査終了後、以下の器官・組織を 10% 中性緩衝ホルマリンに固定・保存した。なお、眼球およびハーダー腺はデビッドソン液で固定・保存し、精巣および精巣上体はブアン液で固定、70% エタノールに保存した。肺については固定液を注入後浸漬固定した。左右のある器官については原則として左右とも固定・保存した。
器官・組織名	: 脳（大脳、小脳および脳橋）、脊髄、下垂体、胸腺、甲状腺（上皮小体を含む）、副腎、脾臓、心臓、食道、胃、肝臓、胰臓、十二指腸、空腸、回腸（ペイエル板を含む）、盲腸、結腸、直腸、気管、肺、腎臓、膀胱、精巣、精巣上体、前立腺、精囊（凝固腺含む）、卵巣、子宮（角部および頸部）、眼球およびハーダー腺、乳腺（右腹部）、大腿骨（骨髓を含む、右）、腸間膜リンパ節、頸下リンパ節、坐骨神経および肉眼的異常部位（正常組織との境界部含む）。

10) 器官重量測定

例数	: 血液学的検査と同一の各群 5 例。 精巣および精巣上体については全例。
時期	: 剖検時
測定方法	: 電子式上皿天秤（ER-180A、株式会社エー・アンド・ディ）を用いて以下の器官について重量を測定した。左右のある器官については、左右併せて測定した。
検査器官	: 脳、下垂体、甲状腺（上皮小体含む）、心臓、肝臓、腎臓、脾臓、副腎、胸腺、精巣、精巣上体、前立腺（腹葉）、精囊（凝固腺とともに分泌物含む）、卵巣
相対重量の算出	: 以下の式から相対重量を算出した。

$$\text{相対重量(%)} = \frac{\text{絶対重量(g)}}{\text{剖検日体重(g)}} \times 100$$

11) 病理組織学的検査

例数	: 剖検時に固定・保存した全例の全器官・組織について標本作製を実施し、対照群および高用量群の雌雄それぞれ 5 例（血液学的検査と同一例）について鏡検した。また、主試験群の精巣の精子形成につ
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いて精査した。さらに、肝臓については、高用量群において病理組織学的検査で異常がみられたため全投与群の雌雄全例を検査した。

前立腺については、高用量群において所見(炎症性細胞の浸潤)の増加がみられたため、全投与群の雄全例を検査した。また、剖検で異常がみられた部位の検査を実施した：4 mg/kg 群；胃境界縁白色腫瘍(動物番号 260)、20 mg/kg 群；精巣上体尾部黄白色斑(動物番号 309)、100 mg/kg 群；皮下黄色腫瘍(動物番号 460)。

検査方法 : パラフィン包埋後薄切し、ヘマトキシリン・エオジン染色標本を作製して鏡検した。なお、所見の確定のため、肝臓では PAS 染色および oil red O 染色標本を追加作製した。

II. 生殖発生毒性

1) 性周期検査

例数 : 雌の全例
 期間 : 投与開始日から主試験群は交尾成立まで、回復群は回復 14 日の翌日の剖検日までの連日。
 方法 : ギムザ染色による膣垢塗抹標本を作製し、光学顕微鏡下で性周期段階を観察した。
 判定 : 性周期の各段階(発情前期、発情期、発情後期および発情休止期)を 4 日から 6 日の間隔で繰返すものを正常と判定し、発情期間隔を算出した。発情休止期が 7 日以上継続してみられる例は連続非発情とし異常と判定した。

2) 生殖能検査

例数 : 主試験群の雌雄の全例
 時期、方法 : 投与 14 日を交配開始日、交配開始日の翌日を交配 1 日とし、交配開始日の夕刻から、同試験群内の雌雄 1 対を交尾が確認されるまで 14 日間を限度として連続同居させた。

交配組み合せ : 無作為組み合せ

交尾成立の確認方法 : 膣内または受皿上に落下した膣栓、あるいは膣垢スマア標本中の精子確認により判定した。いずれかが認められた日を妊娠 0 日とした。次式から群毎に交尾率を算出した。

$$\text{交尾率(Copulation index, \%)} = \frac{\text{交尾した雄/雌の数}}{\text{同居させた雄/雌の数}} \times 100$$

受胎能 : 妊娠の確認を分娩の有無および剖検時に子宮内の着床痕の計数によ

り行った。

次式から群毎に受胎率を算出した。

$$\text{受胎率(Fertility index, \%)} = \frac{\text{受胎動物数}}{\text{交尾した雄/雌の数}} \times 100$$

3) 分娩および哺育状態観察

例数 : 受胎した雌の全例

分娩観察 : 交尾が確認された雌動物は全例自然分娩させた。

分娩状態を妊娠 21 日から 25 日まで、毎日少なくとも 3 回 (9:00、13:00 および 17:00) 観察した。

分娩終了の確認 : 9:00 に母動物が児を巣の中に集めて腹の下に抱え込んでいるのが観察された場合に分娩終了とし、その日を哺育 0 日 (生後 0 日) とした。1 匹以上の生存児を出産したものを正常出産とした。

次式から群毎に出産率を算出した。

$$\text{出産率(Gestation index, \%)} = \frac{\text{生児出産雌数}}{\text{妊娠雌数}} \times 100$$

出産児の観察 : 生後 0 日に正常に出産した腹毎に生存児数と死亡児数を計数し、それらの合計を出産児数とした。

次式から腹毎に出生率を算出した。

$$\text{出生率(Live birth index, \%)} = \frac{\text{出産時生存児数}}{\text{出産児数}} \times 100$$

出産児の性比の算出 : 生後 0 および 4 日に個々の児動物の性を肛門と生殖突起の間の長さで判定した。死亡児も含めた生後 0 日の全出産児、死亡例を含めない生後 0 日の生存児ならびに生後 4 日の生存児を対象として以下を算出した。

$$\text{生後 0 日の全出産児の性比(Sex ratio)} = \frac{\text{雄出産児数}}{\text{雄出産児数} + \text{雌出産児数}}$$

$$\text{生後 0 日の生存児の性比(Sex ratio)} = \frac{\text{雄生存児数}}{\text{雄生存児数} + \text{雌生存児数}}$$

$$\text{生後 4 日の生存児の性比(Sex ratio)} = \frac{\text{雄生存児数}}{\text{雄生存児数} + \text{雌生存児数}}$$

妊娠期間の算出 : 妊娠 0 日から哺育 0 日までの期間の日数を計数した。

分娩率の算出 : 剖検時に各雌の子宮内の着床痕を肉眼的に計数した。

次式から腹毎に分娩率を算出した。

$$\text{分娩率(Delivery index, \%)} = \frac{\text{出産児数}}{\text{着床数}} \times 100$$

着床率の算出

: 剖検時に各雌の卵巣の黄体数を計数した。

次式から腹毎に着床率を算出した。

$$\text{着床率(Implantation index, \%)} = \frac{\text{着床数}}{\text{黄体数}} \times 100$$

哺育 4 日の哺育率の算出：次式から群毎に算出した。

$$\text{哺育率(Nursing index, \%)} = \frac{\text{哺育 4 日に生存児を持つ雌数}}{\text{生児出産雌数}} \times 100$$

4) 新生児の一般状態観察および生存率

例数 : 全例

頻度 : 1 回／日

期間 : 生後 0 日から生後 4 日（剖検日）までとした。

観察方法 : 生存または死亡を確認し、一般状態および外表について観察した。
なお、死亡例は発見後速やかに剖検し、Whole body を 10% 中性緩衝ホルマリン液で固定・保存した。

生後 4 日の新生児生存率の算出：次式により 1 腹単位で算出した。

$$\text{新生児生存率 (Viability index, \%)} = \frac{\text{生後 4 日の生存児数}}{\text{出産時生存児数}} \times 100$$

5) 新生児の体重測定

例数・時期 : 生存児全例について、生後 0、1 および 4 日に実施した。

測定方法 : 電子式上皿天秤 (GX-2000、株式会社エー・アンド・デイ) を用いて雌雄別に 1 腹まとめて測定し、0.1 g まで記録した。
雌雄毎に腹当たりの平均体重を求めた。

6) 新生児の剖検

時期・例数 : 生後 4 日に全例について実施した。

検査方法 : 体外表（口腔内を含む）を観察し、二酸化炭素吸入法により安楽死させ、全身の器官・組織を肉眼的に観察した。異常例については、Whole body を 10% 中性緩衝ホルマリン液に固定・保存した。

5. 統計学的方法

握力、自発運動量、体重、体重増加量および増加率、摂餌量、尿量、血液学的検査、血液化学的検査、器官の絶対重量および相対重量、発情期間隔、黄体数、着床数および着床率、出産児数、出産時の生存児数および死亡児数、分娩率、出生率、性比、妊娠期間、生後 4 日の生存

児数および新生児生存率の成績について平均値および標準偏差を算出し、Bartlett の検定法を行い、等分散性を解析した。等分散の場合は一元配置分散分析法で解析し、不等分散の場合は Kruskal-Wallis の検定法で解析した。一元配置分散分析の結果、有意差がみられた場合は Dunnett の検定法を用いて対照群との比較を行った。Kruskal-Wallis 法の解析の結果、有意差がみられた場合は Mann-Whitney の U-検定法を用い対照群との比較を行った。新生児の出生率、性比、新生児生存率および雌雄別体重については、1 腹を標本単位として処理した。

詳細な一般状態観察および機能検査の観察項目、尿検査の定性的項目、尿比重ならびに病理組織学的検査のうち 2 段階以上のグレードが認められた所見については、群毎の傾向を Kruskal-Wallis の検定法で解析し、有意差がみられた場合は Mann-Whitney の U-検定法を用いて対照群との比較を行った。

正常性周期の雌の出現率、交尾率、受胎率、出産率、哺育 4 日の哺育率、ならびに病理組織学的検査のうち 1 段階のグレードが認められた所見については、多試料カイ二乗検定を行い、その結果有意差が認められた場合には 2 試料カイ二乗検定で対照群との比較を行った。ただし、2 試料カイ二乗検定に不適合の場合には Fisher の直接確率検定法を用いた。

対照群との比較検定については、有意水準は 5%とした。統計学的方法に関する表示方法を Appendix 3 に示す。

成 績

I. 反復投与毒性

1. 一般状態

一般状態の成績を Table 1 および 2、 INDIVIDUAL DATA 1-1-1～1-3-2 に示す。

(1) 主試験群雄

投与期間中、対照群を含むいずれの試験群の動物にも異常はみられなかった。

(2) 主試験群雌

妊娠前期間、妊娠期間、哺育期間を通じて、対照群、4 および 20 mg/kg 群の動物には異常はみられなかった。100 mg/kg 群では、妊娠および哺育期間に皮下腫瘍が 1 例みられた。

(3) 回復群雄

回復期間中、対照群および 100 mg/kg 群の動物には異常は認められなかった。

(4) 回復群雌

回復期間中、対照群および 100 mg/kg 群の動物には異常は認められなかった。

2. 詳細な一般状態観察

詳細な一般状態観察の成績を Table 3-1～3-3 および 4-1-1～4-2-3、 INDIVIDUAL DATA 2-1-1～2-18-6、スコアリングの基準を Appendix 4 に示す。

(1) 主試験群雄

投与期間中の観察では、いずれの項目にも 4、20 および 100 mg/kg 群と対照群の間に有意な差はみられなかった。

(2) 主試験群雌

投与期間中の観察では、いずれの項目にも 4、20 および 100 mg/kg 群と対照群の間に有意な差はみられなかった。

(3) 回復群雄

100 mg/kg 群の回復期間中の観察では、いずれの項目にも対照群と比較して有意な差はみられなかった。

(4) 回復群雌

100 mg/kg 群の投与期間および回復期間中の観察では、いずれの項目にも対照群と比較して有意な差はみられなかった。

3. 機能検査

機能検査の成績を Table 5 および 6、INDIVIDUAL DATA 3-1-1～3-10-2、スコアリングの基準を Appendix 4 に示す。

(1) 主試験群雄

投与 6 週の検査では、いずれの項目にも被験物質投与群と対照群の間に有意な差はみられなかった。

(2) 主試験群雌

哺育 4 日の検査では、いずれの項目にも被験物質投与群と対照群の間に有意な差はみられなかった。

(3) 回復群雄

回復 2 週の検査では、100 mg/kg 群において、後肢の握力に対照群と比較して有意な低値がみられた。

(4) 回復群雌

投与 6 週の検査では、100 mg/kg 群において、20-30 分の間の自発運動量に対照群と比較して有意な低値がみられたが、その他の期間では有意な差はみられなかった。回復 2 週の検査では、100 mg/kg 群において、後肢の握力に対照群と比較して有意な低値がみられた。

4. 体重推移

体重推移を Figure 1～3、Table 7～11、INDIVIDUAL DATA 4-1-1～4-5-4 に示す。

(1) 主試験群雄

4 および 20 mg/kg 群では体重、体重増加量および体重増加率のいずれにも対照群と比較して有意な差はみられなかった。100 mg/kg 群では、投与 35 および 42 日の体重、投与 1-42 日の体重増加量および体重増加率に有意な低値がみられた。

(2) 主試験群雌

4、20 および 100 mg/kg のいずれの投与群においても、交配前、妊娠および哺育期間中を通して体重、体重増加量および体重増加率に対照群と比較して有意な差はみられなかった。

(3) 回復群雄

100 mg/kg 群では、回復期間中の体重、体重増加量および体重増加率は対照群と比較して有意な差はみられなかった。

(4) 回復群雌

100 mg/kg 群では、投与期間および回復期間中の体重、体重増加量および体重増加率のいずれにも対照群と比較して有意な差はみられなかった。

5. 摂餌量

摂餌量を Figure 4~6、Table 12~16、INDIVIDUAL DATA 5-1-1~5-5-4 に示す。

(1) 主試験群雄

4、20 および 100 mg/kg 群では、投与期間中に対照群と比較して有意な差はみられなかった。

(2) 主試験群雌

交配前の期間では、20 mg/kg 群における投与 5 日の有意な低値を除き、有意な差はみられなかった。妊娠期間では、100 mg/kg 群で、妊娠 5、7、10、14 日に有意な低値がみられた。妊娠 14 日では、4 および 20 mg/kg 群においても有意な低値がみられた。哺育期間中では、100 mg/kg 群で哺育 4 日に有意な低値がみられた。

(3) 回復群雄

100 mg/kg 群では、回復 14 日に対照群と比較して有意な低値がみられた。

(4) 回復群雌

100 mg/kg 群では、投与 5、10 日および回復 7 日に対照群と比較して有意な低値がみられた。

6. 尿検査

尿検査の成績を Table 17~20、INDIVIDUAL DATA 6-1-1~6-4-2 に示す。

(1) 主試験群雄

投与 6 週の検査では、いずれの項目にも被験物質投与群と対照群の間で有意な差はみられなかった。

(2) 回復群雄

100 mg/kg 群の回復 2 週の検査では、いずれの項目にも対照群と比較して有意な差はみられなかった。

(3) 回復群雌

100 mg/kg 群の投与 6 週および回復 2 週の検査では、いずれの項目にも対照群と比較して有意な差はみられなかった。

7. 血液学的検査

血液学的検査の成績を Table 21~24、INDIVIDUAL DATA 7-1-1~7-4-4 に示す。

(1) 主試験群雄

投与期間終了時には、4、20 および 100 mg/kg 群ではいずれの検査項目にも対照群と比較して有意な差はみられなかった。

(2) 主試験群雌

投与期間終了時には、4、20 および 100 mg/kg 群ではいずれの検査項目にも対照群と比較して有意な差はみられなかった。

(3) 回復群雄

回復期間終了時の 100 mg/kg 群では、網赤血球数に対照群と比較して有意な高値がみられた。

(4) 回復群雌

回復期間終了時の 100 mg/kg 群では、網赤血球数に対照群と比較して有意な高値がみられた。

8. 血液化学的検査

血液化学的検査の成績を Table 25~28、INDIVIDUAL DATA 8-1-1~8-4-4 に示す。

(1) 主試験群雄

投与期間終了時には、4 および 20 mg/kg 群ではいずれの検査項目にも対照群と比較して有意な差はみられなかった。100 mg/kg 群ではクロールに有意な高値がみられた。

(2) 主試験群雌

投与期間終了時には、4 mg/kg 群ではいずれの検査項目にも対照群と比較して有意な差はみられなかった。20 mg/kg 群ではクロールに有意な高値がみられた。100 mg/kg 群では、総ビリルビンに有意な低値、尿素窒素、ナトリウムおよびクロールに有意な高値がみられた。

(3) 回復群雄

回復期間終了時の 100 mg/kg 群では、AST および総ビリルビンに有意な低値がみられた。また、クロールは高値傾向であったが、有意な差はみられなかった。

(4) 回復群雌

回復期間終了時の 100 mg/kg 群では、総ビリルビンに有意な低値、クロールに有意な高値がみられた。

9. ホルモン測定

ホルモン測定の成績を Table 29~32、INDIVIDUAL DATA 9-1-1~9-4-2 に示す。

(1) 主試験群雄

投与期間終了時には、4、20 および 100 mg/kg 群のいずれの投与群においても、T3、T4 および TSII に対照群と比較して有意な差はみられなかった。

(2) 主試験群雌

投与期間終了時には、すべての被験物質投与群で T3 に対照群と比較して有意な低値がみられた。T4 および TSII には、いずれの被験物質投与群においても対照群と比較して有意な差はみられなかった。

(3) 回復群雄

回復期間終了時の 100 mg/kg 群では、T4 に対照群と比較して有意な低値がみられた。T3 および TSH には有意な差はみられなかった。

(4) 回復群雌

回復期間終了時の 100 mg/kg 群では、T3、T4 および TSH のいずれにも対照群と比較して有意な差はみられなかった。

10. 剖検所見

剖検所見を Table 33～35、INDIVIDUAL DATA 10-1-1～10-4-2 に示す。

(1) 主試験群雄

投与期間終了時の剖検では、対照群で肝臓 6 葉および 7 葉の萎縮および黄緑色化が 1 例、腎孟拡張が 1 例、20 mg/kg 群で精巣上体尾部の黄白色斑が 1 例に観察された。4 および 100 mg/kg 群では異常所見はみられなかった。

(2) 主試験群雌

投与期間終了時の剖検では、対照群および 20 mg/kg 群では異常所見はみられなかった。4 mg/kg 群では、胃境界縁白色腫瘍が 1 例にみられた。100 mg/kg 群では、後腹部右側皮下黄色腫瘍が 1 例にみられた。

(3) 回復群雄

回復期間終了時の剖検では、対照群で回腸の憩室が 1 例にみられた。100 mg/kg 群では異常所見はみられなかった。

(4) 回復群雌

回復期間終了時の剖検では、対照群および 100 mg/kg 群で異常所見はみられなかった。

11. 器官重量

器官の絶対重量および相対重量の成績を Table 36-1～39、INDIVIDUAL DATA 11-1-1～11-4-4 に示す。

(1) 主試験群雄

投与期間終了時に、4 mg/kg 群では、下垂体および精巣上体の絶対重量に有意な低値がみられた。20 mg/kg 群では、甲状腺の絶対および相対重量に有意な高値がみられた。100 mg/kg 群では、剖検時の体重に有意な低値がみられ、肝臓の絶対および相対重量に有意な高値がみられた。同群ではその他に、甲状腺の相対重量に有意な高値がみられた。

(2) 主試験群雌

投与期間終了時に、4、20 および 100 mg/kg のいずれの投与群においても、対照群と比較して有意な差はみられなかった。

(3) 回復群雄

回復期間終了時に、100 mg/kg 群では、肝臓の絶対および相対重量に有意な高値、副腎の絶対および相対重量に有意な低値がみられた。

(4) 回復群雌

回復期間終了時に、100 mg/kg 群では対照群と比較して有意な差はみられなかった。

12. 病理組織学的検査

病理組織学的検査の成績を Table 40~43、INDIVIDUAL DATA 12-1-1~12-8-2 に示す。

(1) 主試験群雄

全例の検査を実施した肝臓では、小葉中心性肝細胞肥大が 20 mg/kg 群で、軽度のグレードで 12 例中 5 例にみられた。100 mg/kg 群では中等度のグレードで 7 例全例にみられ、その発生頻度は有意に高かった。また、小葉中心性脂肪化が 20 mg/kg 群で、軽度のグレードで 12 例中 2 例にみられた。100 mg/kg 群では軽度のグレードで 7 例全例にみられ、その発生頻度は有意に高かった。小肉芽腫は対照群を含む各群で、3~8 例にみられた。小葉周辺性脂肪化が対照群で 1 例、塊状壊死、ヘモジデリン/ヘマトイジンの沈着、線維化および鉱質沈着が対照群で 1 例にみられた。前立腺では炎症性細胞浸潤が対照群、低用量、中用量、高用量群でそれぞれ、2/7、8/12、2/12、6/7 例にみられた。

その他に、対照群で肺胞内マクロファージ集簇、肺胞内骨化生、尿細管上皮の再生、腎孟拡張、100 mg/kg 群で肺胞内マクロファージ集簇、肺の動脈の鉱質沈着、心臓の限局性心筋変性の所見が散見された。剖検で異常がみられた部位の検査では、20 mg/kg 群において精巣上体の精子肉芽腫が 1 例にみられた。その他の器官・組織には異常はみられなかった。

(2) 主試験群雌

全例の検査を実施した肝臓では、小葉中心性肝細胞肥大が 100 mg/kg 群で、軽度のグレードで 12 例中 8 例にみられ、その発生頻度は有意に高かった。小肉芽腫は対照群を含む各群で、7~8 例にみられた。小葉周辺性脂肪化が 4、20 および 100 mg/kg 群で各 1 例にみられた。また、限局性壊死が 4 および 100 mg/kg 群で各 1 例にみられた。

その他に、対照群で肺胞内マクロファージ集簇、肺の動脈の鉱質沈着、腎孟粘膜の炎症性細胞浸潤、下垂体後葉の管状過形成、100 mg/kg 群で肺胞内マクロファージ集簇、腎臓皮髓境界部の鉱質沈着、腎臓乳頭部鉱質沈着の所見が散見された。剖検で異常がみられた部位の検査では、4 mg/kg 群において胃境界縁の扁平上皮囊胞が 1 例、100 mg/kg 群において乳腺の線維腺腫が 1 例にみられた。その他の器官・組織には異常はみられなかった。

(3) 回復群雄

100 mg/kg 群では、肝臓の小葉中心性肝細胞肥大が軽度のグレードで 5 例全例にみられ、有意に高い発生頻度であった。肺では肺胞内マクロファージ集簇が 5 例にみられ、有意に高い発生頻度であった。

その他に、対照群で肺胞内マクロファージ集簇、肺の動脈の鉱質沈着、回腸の憩室、肝臓の小葉周辺性脂肪化、小肉芽腫、前立腺の炎症性細胞浸潤、100 mg/kg 群で肝臓の小葉中心性脂

肪化、小肉芽腫、精巢上体の精子肉芽腫、前立腺の炎症性細胞浸潤、下垂体前葉の囊胞の所見が散見された。その他の器官・組織には異常はみられなかった。

(4) 回復群雌

100 mg/kg 群では、肝臓の小葉中心性肝細胞肥大が 1 例、小肉芽腫が 4 例にみられた。

その他に、対照群で肺胞内マクロファージ集簇、肺胞内骨化生、肝臓の小肉芽腫、腎盂粘膜の炎症性細胞浸潤、100 mg/kg 群で肺の動脈の鉱質沈着、肺胞内骨化生、肝臓の小肉芽腫、腎臓皮質の鉱質沈着の所見が散見された。その他の器官・組織には異常はみられなかった。

II. 生殖発生毒性

(1) 生殖能検査

生殖能検査の成績を Table 44 および 45、INDIVIDUAL DATA 13-1-1～13-2-2 および 14-1～14-4 に示す。

正常性周期を示す雌の出現率、発情期間隔、交尾率、受胎率、出産率、妊娠期間および哺育 4 日の哺育率には、被験物質投与群と対照群の間に有意な差はみられなかった。

性周期検査ではいずれの動物にも異常はみられなかった。対照群で妊娠が成立しなかった雌が 1 例にみられた。4 mg/kg 群において、哺育 4 日までに全児死亡した腹が 2 例にみられた。

(2) 妊娠、分娩、哺育状態および新生児生存率

妊娠、分娩、哺育状態および新生児生存率の成績を Table 46、INDIVIDUAL DATA 15-1～15-4 に示す。

黄体数、着床数、着床率、分娩率、出産児数、出産児の性比、出産時の生存児数、出生率、生後 4 日の生存児数および新生児生存率には、被験物質投与群と対照群の間に有意な差はみられなかった。

(3) 新生児の一般状態

新生児の一般状態の成績を Table 47、INDIVIDUAL DATA 16-1～16-4 に示す。

対照群を含む全試験群において、生後 0 日から生後 4 日までの期間の死亡または不明児（母動物に食されたと考えられる）または授乳の証拠としてのミルクバンドのみられない児が散見された。

(4) 新生児の体重推移

新生児の体重推移を Figure 7、Table 48、INDIVIDUAL DATA 17-1～17-4 に示す。

4 および 20 mg/kg 群では新生児の体重に対照群と比較して有意な差はみられなかった。100 mg/kg 群では、有意な差ではなかったが、生後 4 日の体重が雌雄とも低値傾向を示した。

(5) 新生児の剖検

新生児の剖検の成績を Table 49、INDIVIDUAL DATA 18 および 19-1～19-4 に示す。

死亡児の剖検では、いずれの投与群においても異常はみられなかった。生後 4 日の生存児の

剖検では、肝臓の黄白色斑が 100 mg/kg 群の雄で 1 例にみられたのみであった。

考 察

ペルフルオロヘキサデカン酸の 0 (対照、0.5%CMC-Na)、4、20 および 100 mg/kg を 1 群雌雄各 12 匹の Cr1:CD (SD) ラットに、雄に対しては交配前、交配期間および交配後を含む計 42 日間、雌に対しては交配前、交配および妊娠期間、ならびに分娩後 5 日までの期間経口投与し、雌雄動物への反復投与による影響、雌雄動物の生殖および新生児の発生に及ぼす影響について検討した。また、0 (対照) および 100 mg/kg について、雄動物は各 5 匹を選抜し、雌動物は非交配群として各 5 匹を別に設け、42 日間の投与終了後 14 日間の回復性についても併せて検討した。

1. 反復投与毒性

一般状態観察では、投与期間中いずれの投与群においても被験物質投与に関連する変化はみられなかった。

詳細な一般状態観察では、雌雄とも被験物質投与に関連すると考えられる変化はいずれの投与群においてもみられなかった。機能検査では、回復 2 週の検査で、100 mg/kg 群で雌雄とも後肢の握力に対照群と比較して有意な低値がみられ、被験物質投与との関連性が示唆された。回復群雌の 100 mg/kg 群で、投与 6 週の自発運動量の測定において、測定開始後 20-30 分に有意な低値が認められたが、一過性の変化であり、総測定時間の 0-60 分の値に有意な差はみられなかったことから、偶発的変動と考えられた。

体重については、100 mg/kg 群の雄で、投与 35 および 42 日の体重、投与 1-42 日の体重増加量および体重増加率に有意な低値がみられ、被験物質投与に関連する変化と考えられた。回復期間中の体重、体重増加量および体重増加率に有意な変化はみられず、回復の傾向を示した。雌では、体重にはいずれの投与群においても有意な変化はみられなかった。摂餌量については、雄では投与期間中は有意な差はみられなかつたが、100 mg/kg 群では、回復 14 日に対照群と比較して有意な低値がみられた。雌では、妊娠期間には、100 mg/kg 群で、妊娠 5、7、10、14 日に有意な低値がみられた。なお、投与 5 日に 20 mg/kg 群で有意な低値がみられたが、用量相関性のない変化であった。また、妊娠 14 日に、4 および 20 mg/kg 群においても有意な低値がみられたが、一時期のみの一過性の変化であり、偶発的変動と考えられた。哺育期間中では、100 mg/kg 群で哺育 4 日に有意な低値がみられた。

尿検査では、いずれの項目にも被験物質投与群と対照群の間に有意な差はみられなかつた。

血液学的検査では、投与期間終了時には、雌雄とも 4、20 および 100 mg/kg 群ではいずれの検査項目にも対照群と比較して有意な差はみられなかつた。回復期間終了時には 100 mg/kg 群では、

雌雄の網赤血球数に対照群と比較して有意な高値がみられた。しかし、投与期間終了時および回復期間終了時のいずれにも貧血を示唆する所見はみられていないことから、被験物質投与と関連性のない変化と考えられた。

血液化学的検査では、投与期間終了時に雌雄の 100 mg/kg 群および雌の 20 mg/kg 群で、クロールに有意な高値がみられ、雌の 100 mg/kg 群ではナトリウムも高値であったことから、被験物質投与に関連する変化と考えられた。100 mg/kg 群では尿素窒素の高値を認めたが、クレアチニン、無機リンあるいは総蛋白などの電解質以外の腎機能検査項目、あるいは腎臓の器官重量や組織学的検査には変化がみられなかった。一方、同群では摂餌量の低値が継続してみられており、妊娠動物であることを考慮すると、尿素窒素については摂餌低下などと関連した変化と推察された。回復期間終了時においてもクロールの高値傾向が観察されたが、増加の程度は弱く回復傾向がみられた。その他、投与期間終了時または回復期間終了時に総ビリルビンあるいは AST の有意な低値がみられたが、いずれも低値の場合の臨床的意義が乏しい項目であることから、毒性影響ではないと考えられた。

ホルモン測定では、投与期間終了時に、雌の 4 mg/kg 以上の投与群で T3 に対照群と比較して有意な低値がみられ、100 mg/kg 群で TSH の増加がみられることが、肝臓に病理組織学的変化が観察されていることから、被験物質投与との関連性が示唆された。回復期間終了時の 100 mg/kg 群では、雄の T4 に対照群と比較して有意な低値がみられたが、投与期間終了時にはみられなかったことから偶発的な変動と考えられた。

投与期間終了時の剖検では対照群を含む各群で幾つかの所見（腎孟拡張、精巣上体尾部の黄白色斑等）が散見されたが、いずれも 1 例のみの発生であり、被験物質投与と関係ない変化と考えられた。回復期間終了時にも投与に関連すると考えられる所見はみられなかった。器官重量の測定では 100 mg/kg 群で雄の肝臓の絶対および相対重量に有意な高値がみられた。病理組織学的検査では、雄で小葉中心性肝細胞肥大が 20 mg/kg 群で、軽度のグレードで 12 例中 5 例にみられた。100 mg/kg 群では中等度のグレードで 7 例全例にみられ、その発生頻度は有意に高かった。また、小葉中心性脂肪化が 20 mg/kg 群で、軽度のグレードで 12 例中 2 例に、100 mg/kg 群では軽度のグレードで 7 例全例にみられ、100 mg/kg 群における発生頻度は有意に高かった。雌では、小葉中心性肝細胞肥大が 100 mg/kg 群で、軽度のグレードで 12 例中 8 例にみられ、その発生頻度は有意に高かった。前立腺では、炎症性細胞の浸潤が 100 mg/kg 群で 7 例中 6 例にみられ、被験物質投与との関連性が示唆された。

病理組織学的検査では、上記以外の変化として、肺の動脈の鉱質沈着、尿細管上皮の再生、腎孟拡張、精巣上体の精子肉芽腫、前立腺の炎症性細胞浸潤等が散見されたが、発生頻度が低く、対照群との間に有意な差もないことから、被験物質投与と関連のない変化と考えられた。また、器官重量において、4 mg/kg 群で雄の下垂体および精巣上体の絶対重量に有意な低値がみられたが、用量相関性のない変化であった。20 mg/kg 群では雄の甲状腺の絶対および相対重量に有意な

高値がみられたが、関連する病理組織学的変化はみられなかった。100 mg/kg 群では雄の甲状腺の絶対重量に有意な差はなく、同群の相対重量の有意な高値は体重の低値による二次的変動と考えられた。

回復期間終了時には、100 mg/kg 群では剖検で異常は認められなかつたが、器官重量の測定では雄の肝臓の絶対および相対重量の有意な高値がみられた。また、雄の病理組織学的検査では小葉中心性肝細胞肥大が軽度のグレードで 5 例全例にみられ、有意に高い発生頻度であった。雌でも肝臓の小葉中心性肝細胞肥大が 1 例にみられた。肝臓の変化についてはグレードの低下がみられ、回復傾向にあると考えられた。なお、雄の肺では肺胞内マクロファージ集簇が 5 例にみられ、有意に高い発生頻度であったが、投与期間終了時には有意な増加は認められていないことと、自然発生する所見であるため、被験物質投与に関連しない変化と考えられた。また、器官重量では、雄の副腎の絶対および相対重量の有意な低値がみられたが、病理組織学的検査では異常はみられず、被験物質投与と関連のない変化と考えられた。

以上をまとめると、ペルフルオロヘキサデカン酸の反復投与により 4 mg/kg 群では雄で被験物質投与に関連した変化は認められなかつた。雌では T3 の低値がみられた。20 mg/kg 群では、雌のクロールの高値、T3 の低値、雄の肝臓に小葉中心性肝細胞肥大および小葉中心性脂肪化の所見が認められた。100 mg/kg 群では雄の体重の低値、雌の摂餌量の低値、雄の肝臓重量の増加、雌雄のクロールの高値、雌の T3 の低値、病理組織学的検査における雌雄の肝臓の小葉中心性肝細胞肥大および雄の小葉中心性脂肪化の所見が認められた。

したがつて、本試験条件下におけるペルフルオロヘキサデカン酸の無影響量 (NOEL) は雄で 4 mg/kg/day、雌で 4 mg/kg/day 未満と考えられた。

2. 生殖発生毒性

生殖能検査では、100 mg/kg 群まで、主試験群の正常性周期を示す雌の出現率、発情期間隔、交尾率、受胎率、着床率、出産率、分娩率、出産児数には、対照群との間に有意な差はみられなかつた。回復群の雌では、性周期に異常はみられなかつた。なお、4 mg/kg 群において、哺育 4 日までに全児死亡した腹が 2 例にみられたが、20 および 100 mg/kg 群では同様の変化はみられなかつたことから偶発的変動と考えられた。

新生児の体重では 100 mg/kg 群で生後 4 日に雌雄とも低値傾向がみられた。一般状態および剖検では、100 mg/kg 群まで、被験物質投与に関連する変化はみられなかつた。

以上をまとめると、100 mg/kg 群まで、親動物の生殖能および新生児の発生に被験物質投与に関連した変化は認められなかつた。

したがつて、本試験条件下におけるペルフルオロヘキサデカン酸の親動物の生殖に対する無影響量 (NOEL) は 100 mg/kg/day、新生児の発生に対する無影響量 (NOEL) は 20 mg/kg/day と考えられた。

参考文献

- 1) 最終報告書：ペルフルオロヘキサデカン酸のラットにおける反復投与毒性・生殖発生毒性併合試験(予備試験)(試験番号 S R 0 7 1 2 5 P), 2010 年.

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

試験成績の信頼性に影響を及ぼしたと思われる要因および試験計画書に従わなかつたことはなかった。

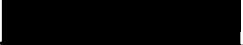
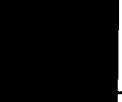
資料の保存

以下の試験関係資料を試験終了後 10 年間、株式会社 化合物安全性研究所の資料保存室に保存する。その後の保存については試験委託者との協議により決定する。

- ① 試験計画書および試験計画書変更書
- ② 生データその他の記録文書
- ③ 最終報告書
- ④ 標本：膣垢塗抹標本
血液塗抹標本
固定器官・組織
病理組織標本(パラフィン包埋ブロックおよび光顕標本)
- ⑤ 被験物質サンプル

試験責任者の記名なつ印

試験責任者

  2012年5月31日

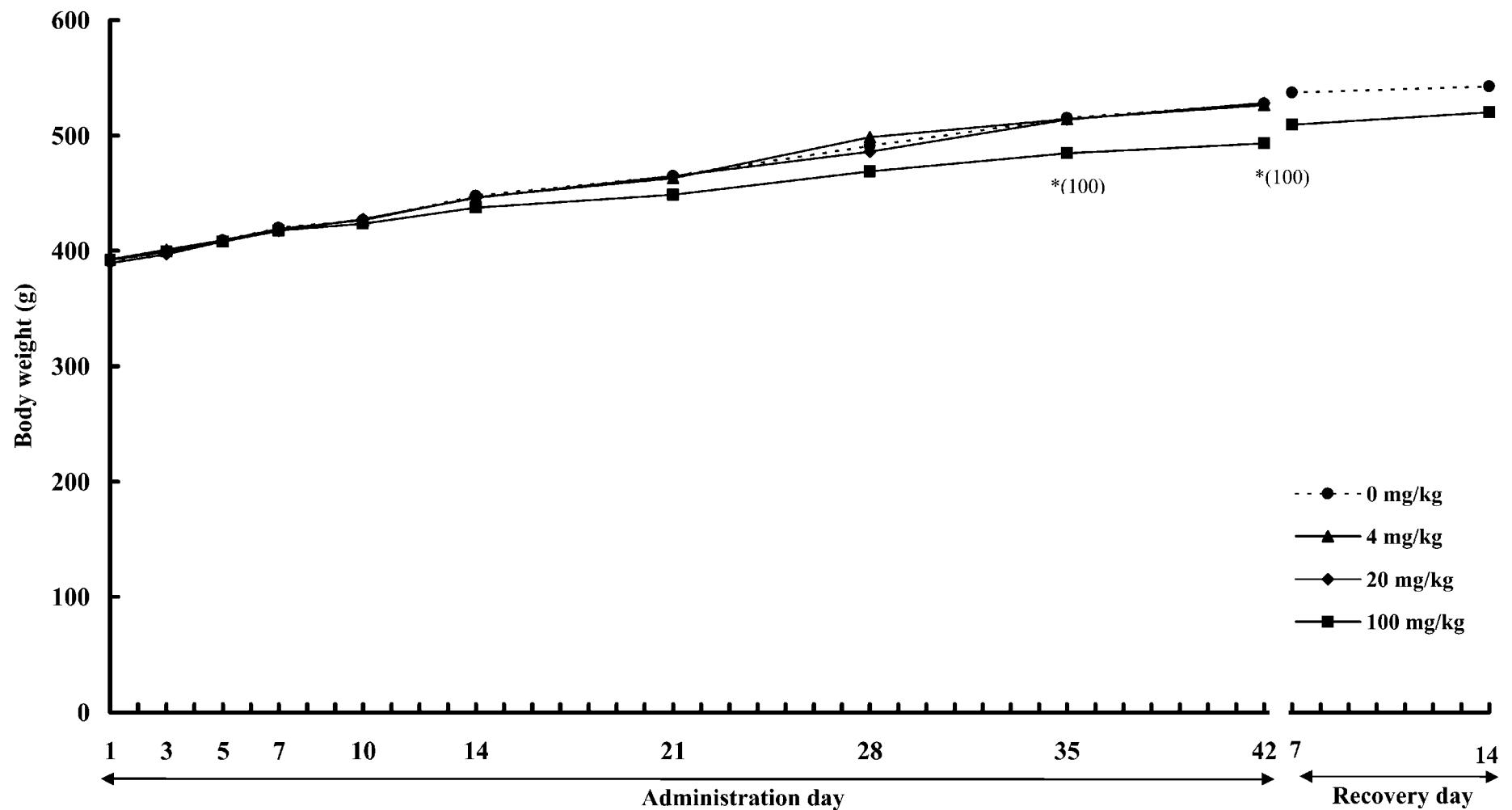


Figure 1 Body weight changes of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

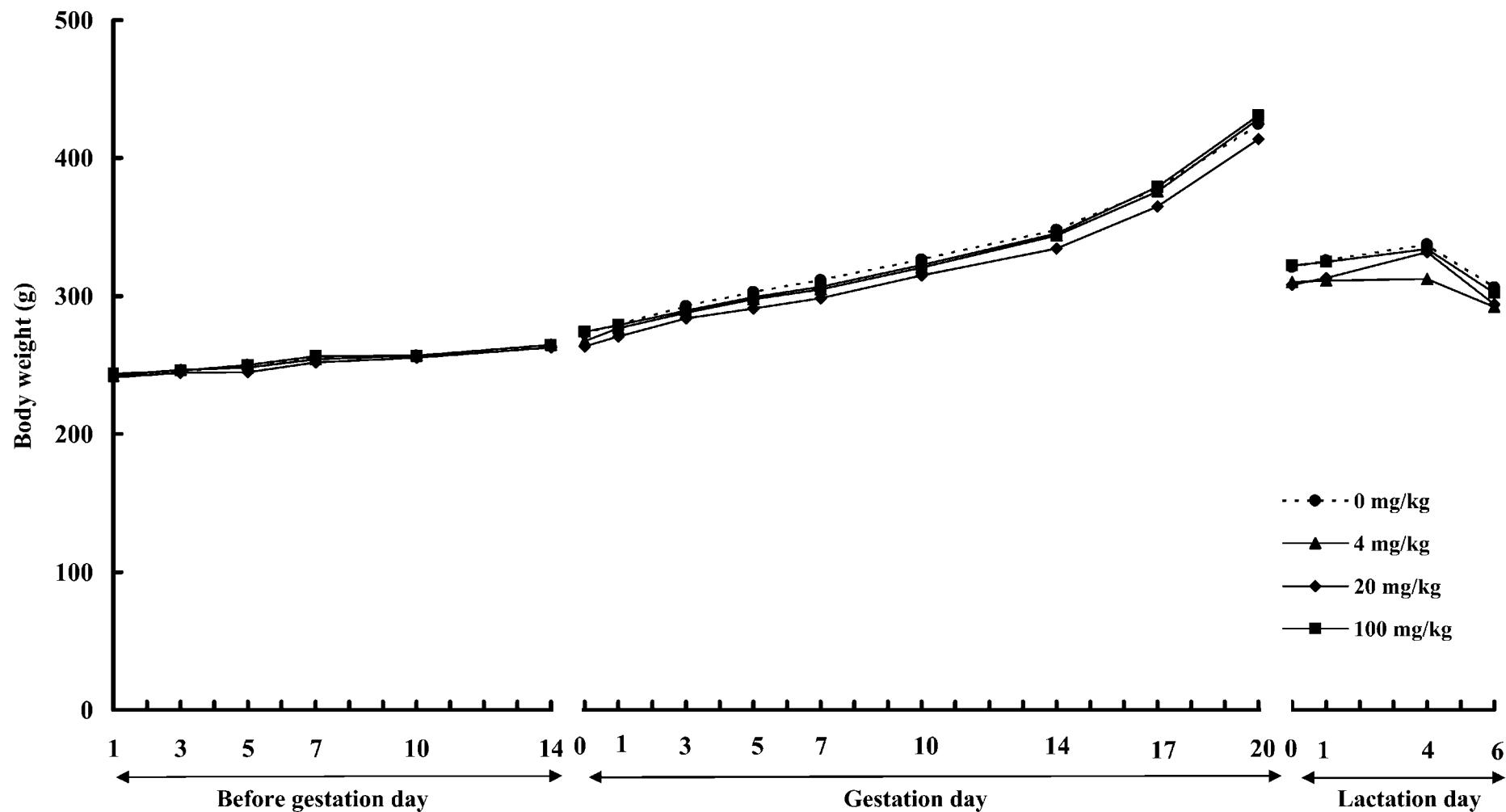


Figure 2 Body weight changes of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

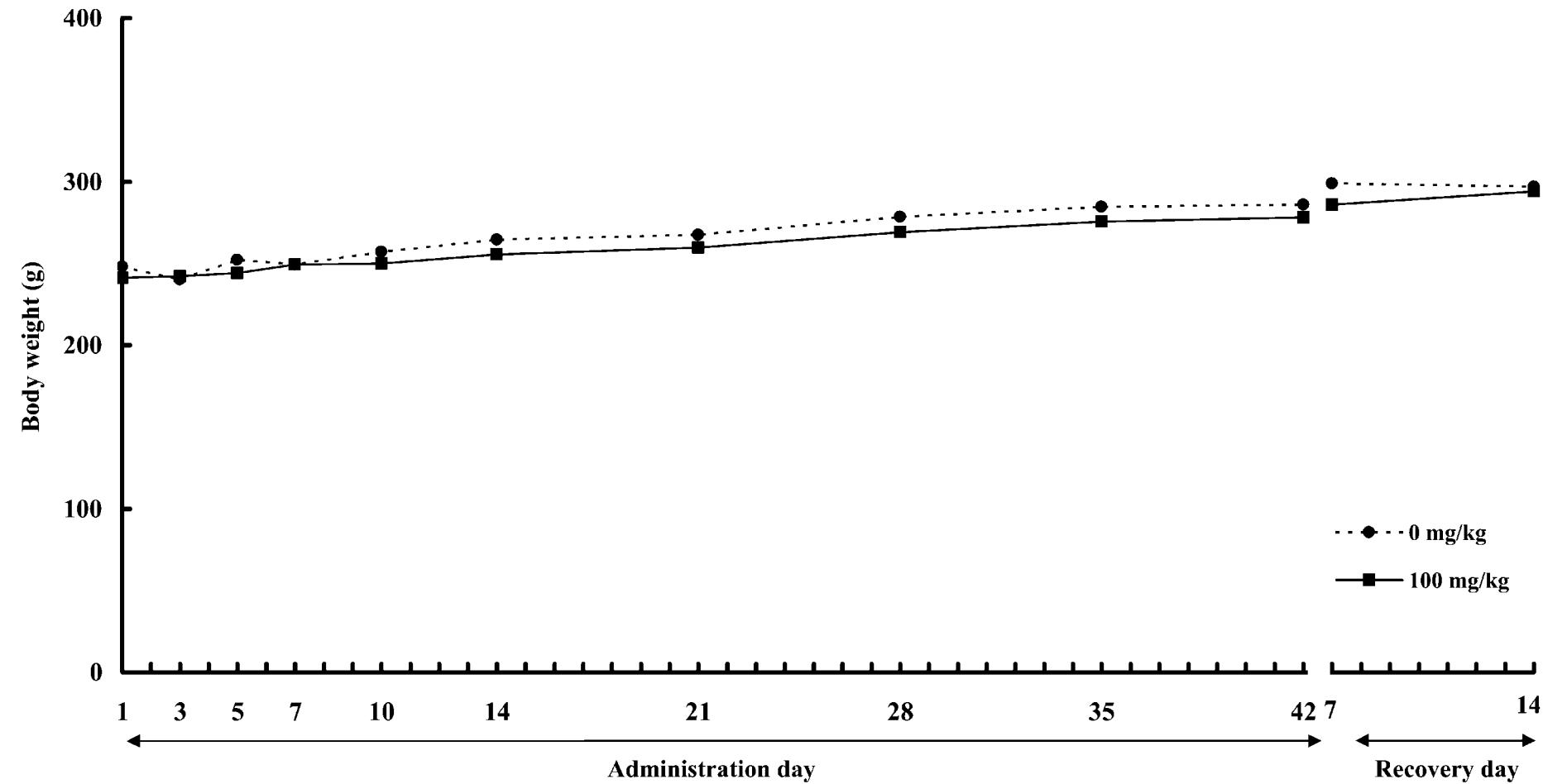


Figure 3 Body weight changes of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

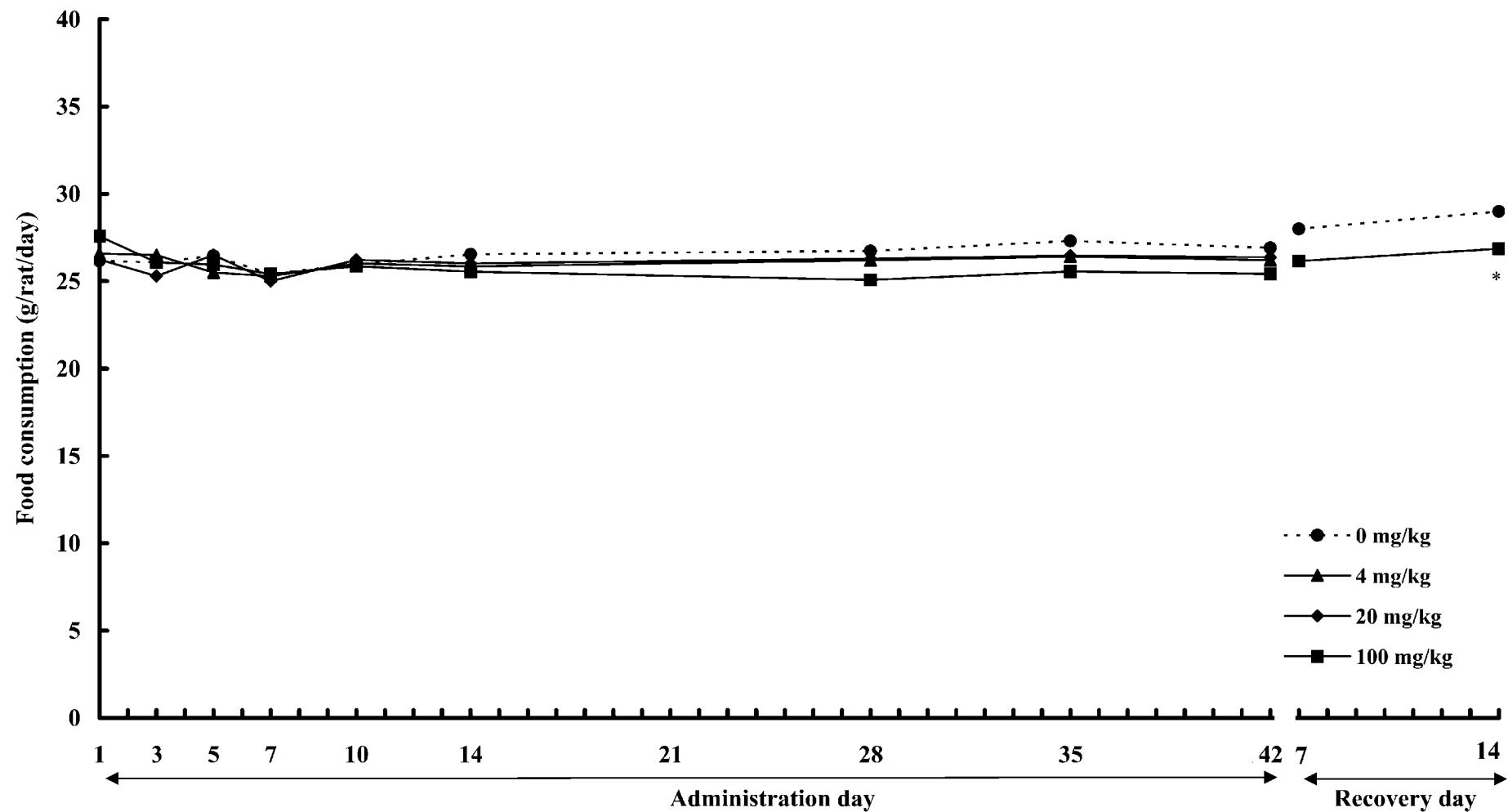


Figure 4 Food consumption of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

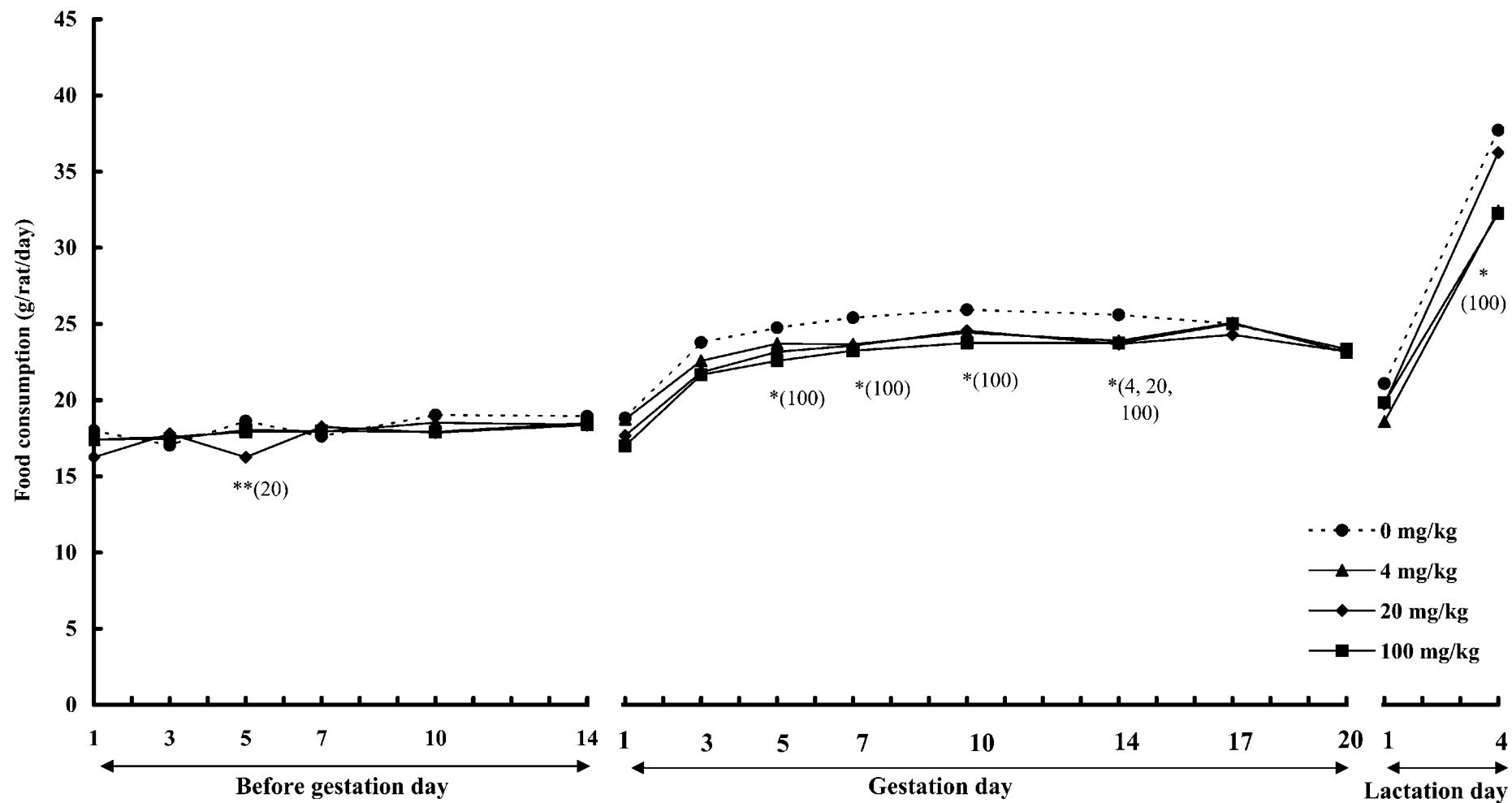


Figure 5 Food consumption of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

**: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Dunnett's test).

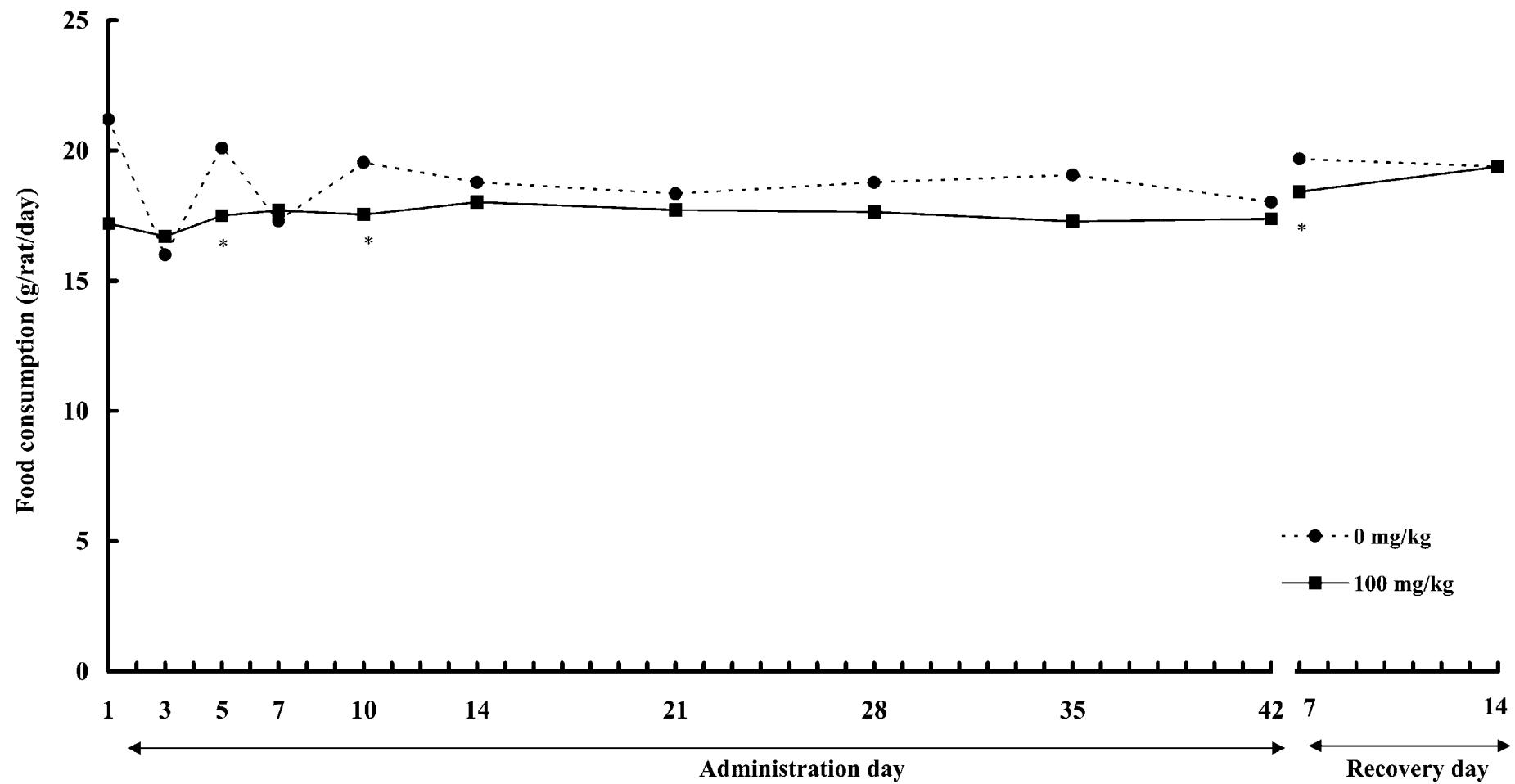


Figure 6 Food consumption of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the eproduction/developmental toxicity screening test (SR07125)

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

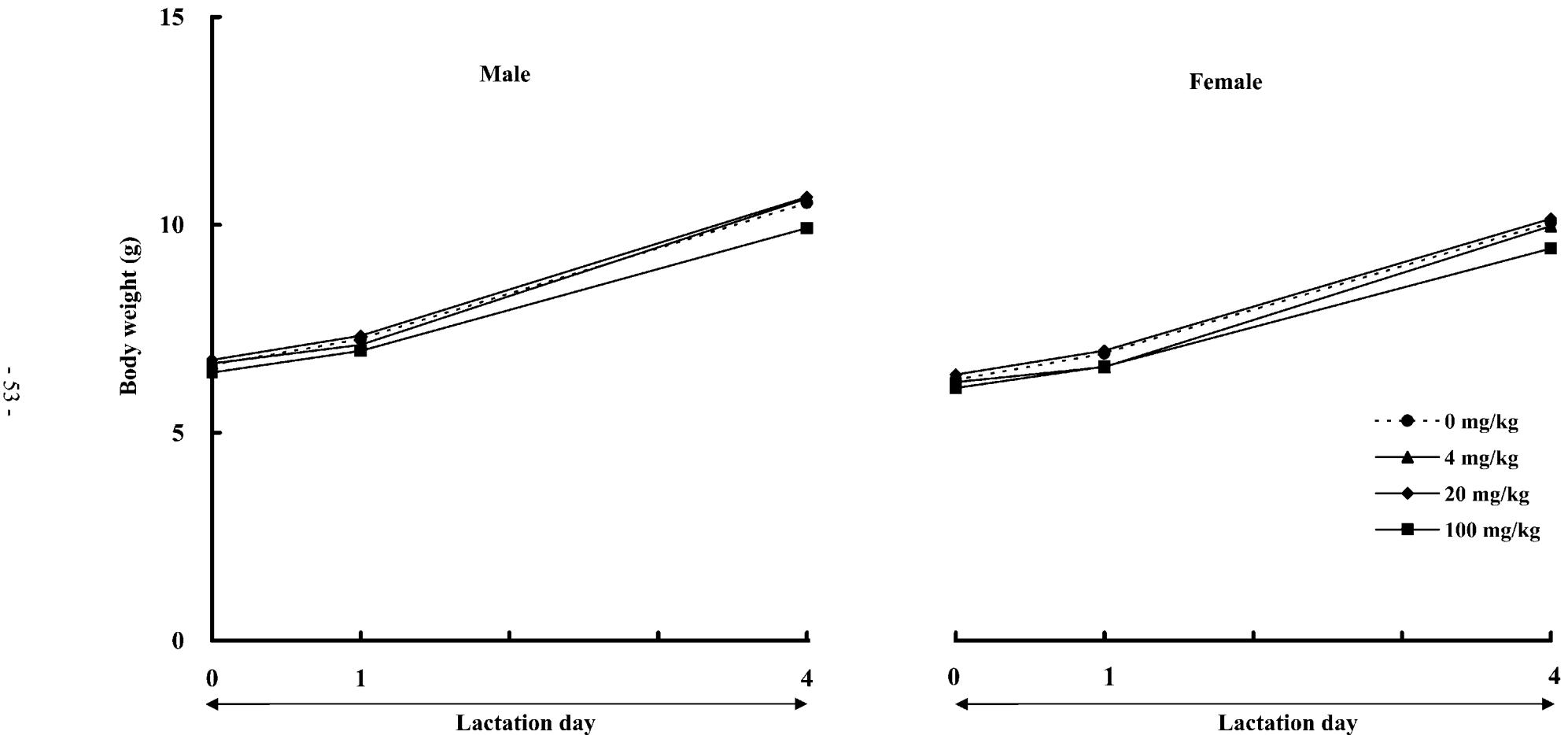


Figure 7 Body weight changes of pups in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test of perfluorohexadecanoic acid in rats (SR07125)

Table 1 General appearance of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Findings	Administration period ^a	Recovery period ^a
0 mg/kg	Number of animals examined	12	5
	No abnormal findings	12	5
4 mg/kg	Number of animals examined	12	-
	No abnormal findings	12	-
20 mg/kg	Number of animals examined	12	-
	No abnormal findings	12	-
100 mg/kg	Number of animals examined	12	5
	No abnormal findings	12	5

Values are the number of animals with findings.

a : Including autopsy day.

- : Blank.

Table 2 General appearance of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Findings	Before gestation period	Gestation period	Lactation period ^a
0 mg/kg	Number of animals examined	12	12 ^b	11
	No abnormal findings	12	12 ^b	11
4 mg/kg	Number of animals examined	12	12	12
	No abnormal findings	12	12	12
20 mg/kg	Number of animals examined	12	12	12
	No abnormal findings	12	12	12
100 mg/kg	Number of animals examined	12	12	12
	No abnormal findings	12	11	11
	Subcutaneous mass	0	1	1

Values are the number of animals with findings.

(to be continued)

a : Including autopsy day.

b : One animal that delivered no pups until gestation day 25 was euthanized on the following day.

Table 2 General appearance of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Group	Findings	Administration period	Recovery period ^a
0 mg/kg	Number of animals examined	5	5
	No abnormal findings	5	5
100 mg/kg	Number of animals examined	5	5
	No abnormal findings	5	5

Values are the number of animals with findings.

a : Including autopsy day.

Table 3-1 Detailed clinical observation of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Period	Group	In the cage								
		Number of animals	Category No.	Body position/ Posture		Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior
				1	1			1	0	
Pre	0 mg/kg	12		12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12
Day 7	0 mg/kg	12		12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12
Day 14	0 mg/kg	12		12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12
Day 21	0 mg/kg	12		12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12
Day 28	0 mg/kg	12		12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12

Category : The category number observed in each item.

(to be continued)

Pre : Pre-administration

Day 7, 14, 21 and 28 : Day 7, 14, 21 and 28 of administration.

Table 3-1 Detailed clinical observation of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	In the cage								
		Number of animals	Category No.	Body position/ Posture		Respiratory pattern		Stereotype		Bizarre behavior
				1	1	1	1	0	0	1
Day 35	0 mg/kg	12		12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12
Day 42	0 mg/kg	12		12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12
Recovery	0 mg/kg	5		5	5	5	5	5	5	5
day 7	100 mg/kg	5		5	5	5	5	5	5	5
Recovery	0 mg/kg	5		5	5	5	5	5	5	5
day 14	100 mg/kg	5		5	5	5	5	5	5	5

Category : The category number observed in each item.

Day 35 and 42 : Day 35 and 42 of administration.

Recovery day 7 and 14 : Day 7 and 14 of recovery.

Table 3-2 Detailed clinical observation of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Period	Group	Number of animals	Category No.	On the hand											
				Ease of				Mucous membranes				Pupil size		Secretions/Excretions	
				Removal	Handling	Muscle tone	Piloerection	Fur	Eyes	0	1	1	1	1	0
Pre	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
Day 7	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
Day 14	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
Day 21	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
Day 28	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12

Category : The category number observed in each item.

(to be continued)

Pre : Pre-administration

Day 7, 14, 21 and 28 : Day 7, 14, 21 and 28 of administration.

Table 3-2 Detailed clinical observation of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	Number of animals	Category No.	On the hand													
				Ease of				Muscle tone	Piloerection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacration	Salivation	Secretions/ Excretions
				Removal	Handling	1	2										
Day 35	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12	12	
Day 42	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12	12	
Recovery	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5	
day 7	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5	
Recovery	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5	
day 14	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5	

Category : The category number observed in each item.

Day 35 and 42 : Day 35 and 42 of administration.

Recovery day 7 and 14 : Day 7 and 14 of recovery.

Table 3-3 Detailed clinical observation of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Period	Group	In the open-field																	
		Number of animals	Category No.	Gait	Co-ordination of movement	Reactivity to environmental stimuli			Searching		Urination		Defecation		Stereotype		Bizarre behavior		
						1	1	1	1	0	1	0	1	0	0	Walking backward	Vocaliza- tion	Aggression	
Pre	0 mg/kg	12		12	12	12	12	12	10	2	10	2	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	12	0	10	2	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	11	1	11	1	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	10	2	11	1	12	12	12	12	12	12	
Day 7	0 mg/kg	12		12	12	12	12	12	11	1	11	1	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	10	2	11	1	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	12	0	12	0	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	11	1	9	3	12	12	12	12	12	12	
Day 14	0 mg/kg	12		12	12	12	12	12	11	1	11	1	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	12	0	12	0	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	11	1	11	1	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	12	0	12	0	12	12	12	12	12	12	
Day 21	0 mg/kg	12		12	12	12	12	12	10	2	11	1	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	11	1	11	1	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	11	1	11	1	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	11	1	10	2	12	12	12	12	12	12	
Day 28	0 mg/kg	12		12	12	12	12	12	11	1	11	1	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	10	2	12	0	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	11	1	12	0	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	11	1	12	0	12	12	12	12	12	12	

Category : The category number observed in each item.

(to be continued)

Pre : Pre-administration

Day 7, 14, 21 and 28 : Day 7, 14, 21 and 28 of administration.

Table 3-3 Detailed clinical observation of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	In the open-field															
		Number of animals	Category No.	Gait	Reactivity to environmental stimuli			Searching	Urination		Defecation		Stereotype		Bizarre behavior		
					Co-ordination of movement	1	1		1	0	1	0	1	Excessive grooming	Unusual head movement	Walking backward	Vocaliza- tion
Day 35	0 mg/kg	12		12	12	12	12	12	10	2	12	0	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	10	2	11	1	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	0	12	0	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	0	12	0	12	12	12	12	12
Day 42	0 mg/kg	12		12	12	12	12	12	12	0	12	0	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	0	11	1	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	0	12	0	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	0	12	0	12	12	12	12	12
Recovery day 7	0 mg/kg	5		5	5	5	5	5	5	0	5	0	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	0	5	0	5	5	5	5	5
Recovery day 14	0 mg/kg	5		5	5	5	5	5	3	2	5	0	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	0	5	0	5	5	5	5	5

Category : The category number observed in each item.

Day 35 and 42 : Day 35 and 42 of administration.

Recovery day 7 and 14 : Day 7 and 14 of recovery.

Table 4-1-1 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Period	Group	In the cage									
		Number of animals	Category No.	Body position/ Posture		Respiratory pattern		Tremor/ Convulsion		Stereotype	Bizarre behavior
				1	1	1	1	0	0	1	
Pre	0 mg/kg	12		12	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	12	12	
Day 7	0 mg/kg	12		12	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	12	12	
Day 14	0 mg/kg	12		12	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	12	12	
Day 21	0 mg/kg	12		12	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	12	12	
Day 28	0 mg/kg	12		12	12	12	12	12	12	12	
	4 mg/kg	12		12	12	12	12	12	12	12	
	20 mg/kg	12		12	12	12	12	12	12	12	
	100 mg/kg	12		12	12	12	12	12	12	12	

Category : The category number observed in each item.

(to be continued)

Pre : Pre-administration

Day 7, 14, 21 and 28 : Day 7, 14, 21 and 28 of administration.

Table 4-1-1 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	In the cage								
		Number of animals	Category No.	Body position/ Posture		Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior
				1	1			0	0	
Day 35	0 mg/kg	12		12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12
Day 42	0 mg/kg	11		11	11	11	11	11	11	11
	4 mg/kg	12		12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12

Category : The category number observed in each item.

Day 35 and 42 : Day 35 and 42 of administration.

Table 4-1-2 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Period	Group	Number of animals	Category No.	On the hand											
				Ease of				Mucous membranes				Pupil size		Secretions/ Excretions	
				Removal	Handling	Muscle tone	Piloerection	Fur	Eyes	0	Skin	1	1	1	0
Pre	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
Day 7	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
Day 14	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
Day 21	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
Day 28	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12

Category : The category number observed in each item.

(to be continued)

Pre : Pre-administration

Day 7, 14, 21 and 28 : Day 7, 14, 21 and 28 of administration.

Table 4-1-2 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	Number of animals	Category No.	On the hand											
				Ease of		Muscle tone	Piloerection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacration	Salivation	Secretions/ Excretions
				Removal	Handling										
Day 35	0 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
Day 42	0 mg/kg	11		11	11	11	11	11	11	11	11	11	11	11	11
	4 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	12	12	12	12	12

Category : The category number observed in each item.

Day 35 and 42 : Day 35 and 42 of administration.

Table 4-1-3 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Period	Group	Number of animals	Category No.	In the open-field																
				Gait	Co-ordination of movement	Reactivity to environmental stimuli			Searching			Urination			Defecation	Stereotype		Bizarre behavior		
						1	1	1	1	0	1	0	1	0		Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression
Pre	0 mg/kg	12		12	12	12	12	12	10	2	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	9	3	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	11	1	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	9	3	12	12	12	12	12	12	12	12	12	12
Day 7	0 mg/kg	12		12	12	12	12	12	12	0	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	10	2	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	11	1	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	11	1	12	12	12	12	12	12	12	12	12	12
Day 14	0 mg/kg	12		12	12	12	12	12	10	2	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	11	1	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	11	1	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	11	1	12	12	12	12	12	12	12	12	12	12
Day 21	0 mg/kg	12		12	12	12	12	12	12	0	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	0	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	0	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	10	2	12	12	12	12	12	12	12	12	12
Day 28	0 mg/kg	12		12	12	12	12	12	12	0	12	12	12	12	12	12	12	12	12	12
	4 mg/kg	12		12	12	12	12	12	12	0	12	12	12	12	12	12	12	12	12	12
	20 mg/kg	12		12	12	12	12	12	12	0	12	12	12	12	12	12	12	12	12	12
	100 mg/kg	12		12	12	12	12	12	12	12	0	12	12	12	12	12	12	12	12	12

Category : The category number observed in each item.

(to be continued)

Pre : Pre-administration

Day 7, 14, 21 and 28 : Day 7, 14, 21 and 28 of administration.

Table 4-1-3 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	In the open-field															
		Number of animals	Category No.	Gait	Co-ordination of movement	Reactivity to environmental stimuli			Searching	Urination	Defecation	Stereotype		Bizarre behavior			
						1	1	1				0	1	0	Walking backward	Vocaliza- tion	Aggression
Day 35	0 mg/kg	12	12	12	12	12	12	0	12	11	1	12	12	12	12	12	12
	4 mg/kg	12	12	12	12	12	11	1	12	10	2	12	12	12	12	12	12
	20 mg/kg	12	12	12	12	12	12	0	12	9	3	12	12	12	12	12	12
	100 mg/kg	12	12	12	12	12	12	0	12	12	0	12	12	12	12	12	12
Day 42	0 mg/kg	11	11	11	11	11	11	10	1	11	10	1	11	11	11	11	11
	4 mg/kg	12	12	12	12	12	12	11	1	12	12	0	12	12	12	12	12
	20 mg/kg	12	12	12	12	12	12	12	0	12	12	0	12	12	12	12	12
	100 mg/kg	12	12	12	12	12	12	12	0	12	12	0	12	12	12	12	12

Category : The category number observed in each item.

Day 35 and 42 : Day 35 and 42 of administration.

Table 4-2-1 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Period	Group	In the cage									
		Number of animals	Category No.	Body position/ Posture		Respiratory pattern		Tremor/ Convulsion		Stereotype	Bizarre behavior
				1	1	1	1	0	0	1	
Pre	0 mg/kg	5		5	5	5	5	5	5	5	
	100 mg/kg	5		5	5	5	5	5	5	5	
Day 7	0 mg/kg	5		5	5	5	5	5	5	5	
	100 mg/kg	5		5	5	5	5	5	5	5	
Day 14	0 mg/kg	5		5	5	5	5	5	5	5	
	100 mg/kg	5		5	5	5	5	5	5	5	
Day 21	0 mg/kg	5		5	5	5	5	5	5	5	
	100 mg/kg	5		5	5	5	5	5	5	5	
Day 28	0 mg/kg	5		5	5	5	5	5	5	5	
	100 mg/kg	5		5	5	5	5	5	5	5	

Category : The category number observed in each item.

(to be continued)

Pre : Pre-administration

Day 7, 14, 21 and 28 : Day 7, 14, 21 and 28 of administration.

Table 4-2-1 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	In the cage								
		Number of animals	Category No.	Body position/ Posture		Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior
				1	1			1	0	
Day 35	0 mg/kg	5		5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5
Day 42	0 mg/kg	5		5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5
Recovery day 7	0 mg/kg	5		5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5
Recovery day 14	0 mg/kg	5		5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5

Category : The category number observed in each item.

Day 35 and 42 : Day 35 and 42 of administration.

Recovery day 7 and 14 : Day 7 and 14 of recovery.

Table 4-2-2 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Period	Group	Number of animals	Category No.	On the hand												
				Ease of			Muscle tone	Piloerection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacration	Salivation	Secretions/ Excretions
				Removal	Handling	2										
Pre	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5
Day 7	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5
Day 14	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5
Day 21	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5
Day 28	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5	5

Category : The category number observed in each item.

(to be continued)

Prc : Pre-administration

Day 7, 14, 21 and 28 : Day 7, 14, 21 and 28 of administration.

Table 4-2-2 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	Number of animals	Category No.	On the hand											
				Ease of		Muscle tone	Piloerection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacration	Salivation	Secretions/ Excretions
				Removal	Handling										
Day 35	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5
Day 42	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5
Recovery day 7	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5
Recovery day 14	0 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5	5	5	5	5	5	5

Category : The category number observed in each item.

Day 35 and 42 : Day 35 and 42 of administration.

Recovery day 7 and 14 : Day 7 and 14 of recovery.

Table 4-2-3 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Period	Group	Number of animals	Category No.	In the open-field																
				Gait	Co-ordination of movement	Reactivity to environmental stimuli			Searching			Urination			Defecation	Stereotype		Bizarre behavior		
						1	1	1	1	0	1	0	1	1		Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression
Pre	0 mg/kg	5		5	5	5	5	5	5	0	5	5	5	5	5	5	5	5	5	
	100 mg/kg	5		5	5	5	5	5	5	0	5	5	5	5	5	5	5	5	5	
Day 7	0 mg/kg	5		5	5	5	5	5	5	0	5	5	5	5	5	5	5	5	5	
	100 mg/kg	5		5	5	5	5	5	5	0	5	5	5	5	5	5	5	5	5	
Day 14	0 mg/kg	5		5	5	5	5	5	5	0	5	5	5	5	5	5	5	5	5	
	100 mg/kg	5		5	5	5	5	5	5	4	1	5	5	5	5	5	5	5	5	
Day 21	0 mg/kg	5		5	5	5	5	5	5	0	5	5	5	5	5	5	5	5	5	
	100 mg/kg	5		5	5	5	5	5	5	0	5	5	5	5	5	5	5	5	5	
Day 28	0 mg/kg	5		5	5	5	5	5	5	0	5	5	5	5	5	5	5	5	5	
	100 mg/kg	5		5	5	5	5	5	5	0	5	5	5	5	5	5	5	5	5	

Category : The category number observed in each item.

(to be continued)

Pre : Pre-administration

Day 7, 14, 21 and 28 : Day 7, 14, 21 and 28 of administration.

Table 4-2-3 Detailed clinical observation of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	In the open-field															
		Number of animals	Category No.	Gait	Co-ordination of movement	Reactivity to environmental stimuli			Searching	Urination		Defecation	Stereotype		Bizarre behavior		
						1	1	1		1	0		0	1	Walking backward	Vocaliza- tion	Aggression
Day 35	0 mg/kg	5		5	5	5	5	5	0	5	5	5	0	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	0	5	4	1	5	5	5	5	5
Day 42	0 mg/kg	5		5	5	5	5	4	1	5	5	0	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	0	5	5	0	5	5	5	5	5
Recovery day 7	0 mg/kg	5		5	5	5	5	5	0	5	5	0	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	0	5	5	0	5	5	5	5	5
Recovery day 14	0 mg/kg	5		5	5	5	5	5	0	5	5	0	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	0	5	5	0	5	5	5	5	5

Category : The category number observed in each item.

Day 35 and 42 : Day 35 and 42 of administration.

Recovery day 7 and 14 : Day 7 and 14 of recovery.

Table 5 Functional test of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

(Observation on the desk)		Number of animals	Category No.	Reactivity					Righting reflex
Period	Group			Visual	Touch	Auditory	Pain	Proprioceptive	
Administration week 6	0 mg/kg	5		5	5	5	5	5	5
	4 mg/kg	5		5	5	5	5	5	5
	20 mg/kg	5		5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5
Recovery week 2	0 mg/kg	5		5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5

Category : The category number observed in each item.

(to be continued)

Values are expressed as the number of animals.

Table 5 Functional test of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	Number of animals	Grip strength (g)		Motor activity measurements							
			Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'		
Administration week 6	0 mg/kg	5	Mean	1689.42	644.80	627.6	345.2	226.2	120.6	100.8		
			S.D.	148.63	82.57	160.6	158.4	86.2	116.7	96.4		
	4 mg/kg	5	Mean	1593.34	639.00	424.8	208.8	132.8	59.0	73.8		
			S.D.	254.08	97.32	190.3	100.3	79.5	56.0	69.5		
Recovery week 2	20 mg/kg	5	Mean	1525.14	603.58	530.2	284.6	197.2	148.0	131.0		
			S.D.	241.12	75.81	315.2	236.0	215.9	150.9	124.3		
	100 mg/kg	5	Mean	1459.86	598.12	588.4	349.2	253.8	200.2	135.2		
			S.D.	290.52	64.78	196.4	63.6	82.1	59.9	66.3		
	0 mg/kg	5	Mean	1609.20	762.94	326.6	237.2	123.4	99.6	54.8		
			S.D.	130.91	31.49	129.4	109.6	60.7	59.9	63.4		
	100 mg/kg	5	Mean	1520.32	622.86*	368.8	322.0	187.8	146.4	114.6		
			S.D.	135.23	90.63	136.9	69.1	37.7	58.8	94.4		
*: Significantly different from the 0 mg/kg group at p≤0.05 (Dunnett's test).												

Table 6 Functional test of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

(Observation on the desk)		Number of animals	Category No.	Reactivity					Righting reflex
Period	Group			Visual 4	Touch 2	Auditory 1	Pain 2	Proprioceptive 1	
<Main group>									
Lactation	0 mg/kg	5		5	5	5	5	5	5
day 4	4 mg/kg	5		5	5	5	5	5	5
	20 mg/kg	5		5	5	5	5	5	5
	100 mg/kg	5		5	5	5	5	5	5
<Recovery group>									
Administration	0 mg/kg	5		5	5	5	5	5	5
week 6	100 mg/kg	5		5	5	5	5	5	5
Recovery	0 mg/kg	5		5	5	5	5	5	5
week 2	100 mg/kg	5		5	5	5	5	5	5

Category : The category number observed in each item.

(to be continued)

Values are expressed as the number of animals.

Table 6 Functional test of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125) (continued)

Period	Group	Number of animals	Grip strength (g)		Motor activity measurements					
			Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'
<Main group>										
Lactation day 4	0 mg/kg	5	Mean	1332.24	643.26	363.8	73.4	90.0	38.6	127.2
			S.D.	107.56	157.18	204.5	45.6	135.6	53.8	172.9
	4 mg/kg	5	Mean	1178.92	625.32	363.0	124.8	102.4	116.0	102.8
			S.D.	143.26	61.79	214.5	98.8	53.6	83.5	106.4
	20 mg/kg	5	Mean	1158.66	646.26	456.6	144.4	126.4	136.8	118.2
			S.D.	262.38	26.64	213.2	92.9	81.2	103.5	214.4
	100 mg/kg	5	Mean	1163.78	552.08	446.0	210.6	149.0	168.6	58.0
			S.D.	104.30	134.36	257.3	140.8	166.4	73.5	88.4
<Recovery group>										
Administration week 6	0 mg/kg	5	Mean	1106.64	499.80	761.8	694.6	550.6	344.6	217.8
			S.D.	302.58	30.55	179.3	181.4	185.2	168.6	108.0
	100 mg/kg	5	Mean	1192.28	462.54	489.4	453.6	268.8*	135.0	157.0
			S.D.	81.13	35.54	201.2	277.9	143.4	123.3	132.1
Recovery week 2	0 mg/kg	5	Mean	1205.80	652.86	570.6	341.6	229.2	191.0	162.8
			S.D.	188.38	75.20	278.9	215.0	117.5	159.2	177.6
	100 mg/kg	5	Mean	1172.74	495.46**	515.8	273.2	145.8	171.0	28.2
			S.D.	145.05	60.55	220.3	186.8	95.5	181.0	27.2

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

**: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Dunnett's test).

Table 7 Body weight changes of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Body weight (g) on administration period (day)										Body weight gain		
		1	3	5	7	10	14	21	28	35	42	Days 1-42 (g)	(%)	
0 mg/kg	12	Mean	391.4	398.9	409.2	419.8	426.6	447.9	465.0	490.8	514.9	527.5	136.1	34.728
		S.D.	20.4	22.2	24.2	25.5	26.1	28.5	31.1	37.4	40.5	42.7	31.1	7.502
4 mg/kg	12	Mean	392.5	401.3	409.2	419.2	426.8	446.3	463.2	498.3	514.3	526.5	134.0	34.263
		S.D.	16.5	14.9	14.5	16.0	16.1	18.4	17.5	23.3	18.0	20.2	19.3	5.493
20 mg/kg	12	Mean	389.3	397.0	408.1	417.2	427.3	445.9	465.0	491.2	513.8	528.1	138.8	35.720
		S.D.	18.3	18.0	18.0	18.9	19.1	22.3	22.1	25.0	28.1	31.2	23.5	6.103
100 mg/kg	12	Mean	391.9	399.1	407.8	417.8	423.4	437.5	448.6	468.7	484.4 *	493.2 *	101.3 **	25.869 **
		S.D.	16.4	16.7	16.0	17.0	16.6	17.8	19.6	25.6	27.1	26.8	20.5	5.228

Group	Number of animals	Body weight (g) on recovery period (day)		Body weight gain	
		7	14	Day 42-R14 (g)	(%)
0 mg/kg	5	Mean	537.0	542.4	24.8
		S.D.	29.8	30.4	3.8
100 mg/kg	5	Mean	509.2	520.2	25.8
		S.D.	15.5	18.7	15.3

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

**: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Dunnett's test).

R14: Recovery day 14.

Table 8 Body weight changes before gestation period of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Body weight (g) on administration period (day)						Body weight gain		
		1	3	5	7	10	14	Days 1-14 (g)	(%)	
0 mg/kg	12	Mean S.D.	243.3 14.9	245.3 12.5	249.8 16.3	255.2 14.6	256.9 15.8	264.0 17.0	20.8 9.5	8.583 3.940
4 mg/kg	12	Mean S.D.	242.2 9.0	246.6 10.4	248.0 8.2	254.0 12.0	256.7 10.7	264.5 9.1	22.3 6.0	9.262 2.577
20 mg/kg	12	Mean S.D.	240.9 11.2	244.4 12.8	244.8 9.7	251.9 12.1	255.1 10.1	262.8 11.1	21.8 5.9	9.113 2.601
100 mg/kg	12	Mean S.D.	243.6 10.7	245.8 11.8	250.1 10.4	256.3 12.2	256.9 12.0	264.6 12.0	21.0 7.3	8.658 3.061

Table 9 Body weight changes during gestation period of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Body weight (g) on gestation period (day)										Body weight gain	
		0	1	3	5	7	10	14	17	20	(g)	(%)	
0 mg/kg	11	Mean	272.8	278.8	292.5	302.7	311.9	327.4	350.8	383.6	434.6	161.8	59.515
		S.D.	17.1	15.4	15.8	16.7	16.9	15.9	19.3	21.3	24.0	16.7	7.145
4 mg/kg	12	Mean	267.3	276.7	287.8	297.8	304.7	320.7	344.0	375.8	428.2	160.9	60.259
		S.D.	9.1	8.6	13.7	11.2	11.7	13.3	14.9	14.6	14.1	9.6	3.899
20 mg/kg	12	Mean	263.3	270.8	283.9	290.8	298.6	315.0	334.4	364.8	413.7	150.3	57.145
		S.D.	13.8	12.0	13.0	13.9	15.7	17.5	19.9	23.4	26.8	19.4	6.979
100 mg/kg	12	Mean	274.1	279.0	289.6	299.0	306.8	322.6	345.3	379.2	431.1	157.0	57.279
		S.D.	10.8	11.2	12.8	13.3	13.5	16.4	18.2	19.7	22.7	15.9	5.231

Table 10 Body weight changes during lactation period of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Body weight (g) on lactation period (day)				Body weight gain		
		0	1	4	6	Days 0-4 (g)	(%)	
0 mg/kg	11	Mean	321.0	325.8	337.6	306.4	16.6	5.336
		S.D.	19.9	17.4	17.7	13.4	14.3	4.754
4 mg/kg	12	Mean	309.9	311.1	320.4	294.7	4.7	1.568
		S.D.	21.0	21.0	34.9	13.9	32.9	10.491
20 mg/kg	12	Mean	308.1	312.9	331.9	294.1	23.8	7.725
		S.D.	17.7	18.0	22.8	18.7	12.0	3.817
100 mg/kg	12	Mean	322.1	324.8	334.1	302.3	12.0	3.832
		S.D.	24.0	23.3	21.9	20.5	10.7	3.538

Values in parentheses are the number of animals examined.

Table 11 Body weight changes of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Body weight (g) on administration period (day)										Body weight gain		
		1	3	5	7	10	14	21	28	35	42	Days 1-42 (g)	(%)	
0 mg/kg	5	Mean	248.0	240.2	252.2	249.4	257.2	264.6	267.4	278.4	284.6	285.8	37.8	15.302
		S.D.	10.4	10.1	10.3	8.2	8.2	7.9	10.3	12.8	13.2	9.8	6.9	3.121
100 mg/kg	5	Mean	241.2	242.2	244.2	249.2	250.0	255.4	259.8	269.2	275.6	278.2	37.0	15.378
		S.D.	7.8	8.5	8.0	11.2	12.9	14.2	14.7	20.5	20.8	19.6	18.9	8.002

Group	Number of animals	Body weight (g) on recovery period (day)		Body weight gain		
		7	14	Day 42-R14 (g)	(%)	
0 mg/kg	5	Mean	298.8	296.8	11.0	3.864
		S.D.	10.0	9.4	3.6	1.266
100 mg/kg	5	Mean	285.8	294.0	15.8	5.796
		S.D.	15.6	16.1	5.9	2.649

R14: Recovery day 14.

Table 12 Food consumption of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Food consumption (g/rat/day) on administration period (day)									
		1	3	5	7	10	14	28	35	42	
0 mg/kg	12	Mean	26.17	26.08	26.46	25.42	25.97	26.55	26.73	27.31	26.92
		S.D.	2.17	2.27	1.70	1.58	1.19	1.61	1.99	1.95	1.57
4 mg/kg	12	Mean	26.58	26.50	25.50	25.29	26.03	25.85	26.20	26.41	26.21
		S.D.	2.50	1.73	3.15	1.51	1.27	1.52	1.40	1.26	1.57
20 mg/kg	12	Mean	26.25	25.29	26.50	25.00	26.23	26.03	26.29	26.47	26.38
		S.D.	2.86	1.50	2.17	1.98	2.02	2.31	2.25	2.40	2.29
100 mg/kg	12	Mean	27.58	26.08	25.96	25.42	25.86	25.55	25.08	25.56	25.43
		S.D.	1.98	2.11	1.42	2.01	1.55	1.85	2.04	2.17	2.04

Group	Number of animals	Food consumption (g/rat/day) on recovery period (day)		
		7	14	
0 mg/kg	5	Mean	28.00	29.00
		S.D.	1.49	1.28
100 mg/kg	5	Mean	26.16	26.86 *
		S.D.	1.09	1.13

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

Table 13 Food consumption before gestation period of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Food consumption (g/rat/day) on administration period (day)						
		1	3	5	7	10	14	
0 mg/kg	12	Mean S.D.	18.00 3.62	17.04 2.19	18.63 1.93	17.63 1.97	19.03 1.70	18.95 1.29
4 mg/kg	12	Mean S.D.	17.42 2.84	17.58 2.27	17.92 1.49	17.92 2.69	18.53 1.81	18.38 1.28
20 mg/kg	12	Mean S.D.	16.25 3.02	17.79 1.83	16.25 ** 1.44	18.25 1.89	17.87 1.73	18.37 1.37
100 mg/kg	12	Mean S.D.	17.42 3.20	17.46 2.24	18.04 1.62	17.96 2.39	17.92 1.34	18.50 1.07

**: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Dunnett's test).

Table 14 Food consumption during gestation period of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Food consumption (g/rat/day) on gestation period (day)								
		1	3	5	7	10	14	17	20	
0 mg/kg	11	Mean	18.91	24.18	25.05	25.86	26.23	26.05	25.72	24.10
		S.D.	2.88	2.08	1.72	1.50	1.35	1.90	1.53	1.40
4 mg/kg	12	Mean	18.75	22.58	23.71	23.67	24.43	23.90 *	25.08	23.13
		S.D.	1.36	3.10	2.62	2.31	2.70	2.09	1.99	1.55
20 mg/kg	12	Mean	17.67	21.83	23.17	23.58	24.57	23.68 *	24.30	23.20
		S.D.	4.01	2.73	2.04	2.84	2.52	1.94	2.05	1.84
100 mg/kg	12	Mean	17.00	21.67	22.58 *	23.25 *	23.75 *	23.75 *	25.02	23.36
		S.D.	2.89	2.00	2.05	2.13	2.39	1.85	2.34	1.81

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

Table 15 Food consumption during lactation period of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Food consumption (g/rat/day) on lactation period (day)	
		1	4
0 mg/kg	11	Mean S.D.	21.09 5.82 (10) 5.00
4 mg/kg	12	Mean S.D.	18.58 9.84 35.43 3.18
20 mg/kg	12	Mean S.D.	19.75 5.31 36.26 3.97
100 mg/kg	12	Mean S.D.	19.83 5.29 32.26 * 4.58

Values in parentheses are the number of animals examined.

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

Table 16 Food consumption of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Food consumption (g/rat/day) on administration day									
		1	3	5	7	10	14	21	28	35	42
0 mg/kg	5	Mean	21.20	16.00	20.10	17.30	19.54	18.78	18.34	18.78	19.06
		S.D.	0.84	2.89	1.24	1.60	1.00	1.32	1.39	1.20	0.52
100 mg/kg	5	Mean	17.20	16.70	17.50 *	17.70	17.54 *	18.02	17.72	17.64	17.28
		S.D.	4.44	3.03	1.27	3.67	1.66	2.29	2.46	3.03	2.20
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Group	Number of animals	Food consumption (g/rat/day) on recovery period (day)									
		7	14								
0 mg/kg	5	Mean	19.68								
		S.D.	0.85								
100 mg/kg	5	Mean	18.42 *								
		S.D.	0.73								

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

Table 17 Urinary findings of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	pH			Protein		Glucose	Ketone body	Urobilinogen 0.1 EU/dL	Bili-rubin -	Occult blood -	Color A
		7.5	8.0	8.5	±	+						
0 mg/kg	5	0	1	4	5	0	5	5	5	5	5	5
4 mg/kg	5	0	0	5	4	1	5	5	5	5	5	5
20 mg/kg	5	1	3	1	5	0	5	5	5	5	5	5
100 mg/kg	5	0	1	4	4	1	5	5	5	5	5	5

Group	Number of animals	Specific gravity				Urine volume (mL/21hr., mean±S.D.)
		1.021-	1.031-	1.041-	1.051≤	
0 mg/kg	5	0	3	2	0	16.30±4.72
4 mg/kg	5	1	3	1	0	17.20±3.47
20 mg/kg	5	1	2	1	1	17.50±9.30
100 mg/kg	5	2	1	1	1	19.00±9.31

Values are number of animals with findings.

Color: A = Pale yellow or yellow

-; Negative, ±; slight, +; moderate.

Table 18 Urinary findings of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	pH					Protein		Glucose	Ketone body	Urobilinogen 0.1 EU/dL	Bili-rubin	Occult blood	Color A
		6.5	7.0	7.5	8.0	8.5	-	±						
0 mg/kg	5	0	0	1	0	4	2	3	5	5	5	5	5	5
100 mg/kg	5	1	0	0	1	3	3	2	5	5	5	5	5	5

Group	Number of animals	Specific gravity				Urine volume (mL/21hr., mean±S.D.)
		1.031-	1.041-	1.051≤	1.040 1.050	
0 mg/kg	5	4	1	0		10.70±2.73
100 mg/kg	5	3	1	1		9.50±3.59

Values are number of animals with findings.

Color: A = Pale yellow or yellow

-; Negative, ±; slight.

Table 19 Urinary findings of male rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	pH		Protein				Glucose	Ketone body	Urobilinogen 0.1 EU/dL	Bili-rubin	Occult blood	Color A
		8.0	8.5	-	±	+	2+						
0 mg/kg	5	0	5	0	0	4	1	5	5	5	5	5	5
100 mg/kg	5	1	4	1	1	3	0	5	5	5	5	5	5

Group	Number of animals	Specific gravity			Urine volume (mL/21hr., mean±S.D.)			
		1.021-	1.031-	1.041-	1.051≤	1.030	1.040	1.050
0 mg/kg	5	0	0	3	2	12.30±4.02		
100 mg/kg	5	1	0	2	2	17.60±13.88		

Values are number of animals with findings.

Color: A = Pale yellow or yellow

-; Negative, ±; slight, +; moderate, 2+; severe.

Table 20 Urinary findings of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	pH					Protein			Glucose	Ketone body	Urobilinogen 0.1 EU/dL	Bili-rubin	Occult blood	Color A		
		6.5	7.0	7.5	8.0	8.5	-	±	+								
0 mg/kg	5	2	1	0	0	2	2	2	1	5	5	5	5	5	5		
100 mg/kg	5	0	0	0	1	4	1	1	3	5	5	5	5	5	5		
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Group	Number of animals	Specific gravity				Urine volume (mL/21hr., mean±S.D.)											
		1.021-	1.031-	1.041-	1.051≤	1.030	1.040	1.050									
0 mg/kg	5	1	1	2	1	14.00±6.92											
100 mg/kg	5	0	0	3	2	8.90±1.82											

Values are number of animals with findings.

Color: A = Pale yellow or yellow

-; Negative, ±; slight, +; moderate.

Table 21 Hematological findings of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group		Number of animals	WBC 10 ³ /µL	RBC 10 ⁶ /µL	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet 10 ³ /µL
0 mg/kg	5	Mean	96.52	926.6	16.20	45.06	48.74	17.52	35.96	125.26
		S.D.	10.43	47.3	0.66	2.00	3.60	0.93	0.86	14.39
4 mg/kg	5	Mean	113.22	917.6	15.82	43.94	47.90	17.24	36.02	123.48
		S.D.	17.58	25.7	0.48	1.07	0.80	0.17	0.41	10.74
20 mg/kg	5	Mean	110.34	939.2	16.36	45.84	48.78	17.42	35.68	116.88
		S.D.	13.25	11.2	0.44	0.91	1.15	0.54	0.29	7.21
100 mg/kg	5	Mean	110.36	897.6	15.62	43.36	48.36	17.42	36.06	109.10
		S.D.	16.87	38.4	0.28	0.64	1.68	0.53	0.19	6.48
Group		Number of animals	Reticulocyte %	PT sec	APTT sec	Differential count of WBC (10 ³ /µL)				
0 mg/kg	5	Mean	3.198	21.14	26.52	14.02	78.54	2.68	1.26	0.02
		S.D.	0.465	2.89	2.66	3.19	12.59	0.40	0.56	0.04
4 mg/kg	5	Mean	3.444	21.00	26.70	15.72	92.86	3.64	1.00	0.00
		S.D.	0.263	1.44	1.76	5.10	18.82	0.47	0.44	0.00
20 mg/kg	5	Mean	3.030	20.46	25.92	14.34	91.22	3.32	1.46	0.00
		S.D.	0.344	2.61	3.20	2.01	14.75	0.63	0.30	0.00
100 mg/kg	5	Mean	2.818	21.28	24.04	15.42	90.56	3.12	1.24	0.02
		S.D.	0.236	1.87	2.86	2.28	17.46	0.90	0.55	0.04

Table 22 Hematological findings of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group		Number of animals	WBC 10 ³ /µL	RBC 10 ⁶ /µL	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet 10 ³ /µL
0 mg/kg	5	Mean	97.30	805.2	15.20	43.56	54.16	18.90	34.88	121.04
		S.D.	22.69	35.4	0.38	0.67	2.00	0.43	0.75	11.09
4 mg/kg	5	Mean	99.90	818.0	15.32	44.02	53.84	18.72	34.78	130.02
		S.D.	20.78	20.5	0.48	1.57	1.58	0.36	0.36	6.95
20 mg/kg	5	Mean	94.72	807.6	15.62	45.48	56.38	19.36	34.34	107.28
		S.D.	9.87	47.0	0.67	1.78	1.82	0.40	0.44	15.52
100 mg/kg	5	Mean	105.86	814.6	15.34	44.44	54.58	18.84	34.54	127.82
		S.D.	19.09	50.9	0.99	3.05	2.67	0.78	0.38	6.15
Group		Number of animals	Reticulocyte %	PT sec	APTT sec	Differential count of WBC (10 ³ /µL)				
0 mg/kg	5	Mean	8.842	18.22	19.82	18.04	71.66	5.94	1.64	0.02
		S.D.	1.591	0.77	1.51	7.85	12.35	2.98	0.71	0.04
4 mg/kg	5	Mean	9.420	18.92	20.24	27.74	64.28	6.52	1.34	0.02
		S.D.	1.749	0.36	1.03	13.70	7.73	2.39	0.30	0.04
20 mg/kg	5	Mean	9.184	18.82	19.76	21.70	66.80	5.02	1.18	0.02
		S.D.	1.187	1.04	0.36	5.16	12.53	1.38	0.58	0.04
100 mg/kg	5	Mean	9.020	17.56	19.42	25.34	72.60	6.22	1.70	0.00
		S.D.	1.059	0.53	0.90	10.15	17.59	0.93	0.73	0.00

Table 23 Hematological findings of male rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group		Number of animals	WBC 10 ³ /μL	RBC 10 ⁶ /μL	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet 10 ³ /μL
0 mg/kg	5	Mean	108.66	931.2	16.46	46.64	50.10	17.68	35.30	124.96
		S.D.	24.19	22.4	0.37	1.25	1.18	0.30	0.20	13.67
100 mg/kg	5	Mean	125.28	903.4	15.94	45.06	49.88	17.66	35.38	120.14
		S.D.	14.52	43.2	0.68	1.91	0.78	0.34	0.19	13.16

Group	Number of animals	Reticulocyte %	PT sec	APTT sec	Differential count of WBC (10 ³ /μL)					
					Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil	
0 mg/kg	5	Mean	3.064	18.96	27.60	17.28	86.10	3.38	1.86	0.04
		S.D.	0.221	1.39	2.09	8.78	21.99	1.36	0.89	0.05
100 mg/kg	5	Mean	3.420 *	20.50	26.76	21.86	97.62	4.34	1.46	0.00
		S.D.	0.243	1.59	0.69	2.54	12.12	0.85	0.77	0.00

*: Significantly different from the 0 mg/kg group at p≤0.05 (Dunnett's test).

Table 24 Hematological findings of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group		Number of animals	WBC 10 ² /µL	RBC 10 ⁴ /µL	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet 10 ⁴ /µL
0 mg/kg	5	Mean	75.08	867.0	15.72	44.32	51.20	18.16	35.46	107.44
		S.D.	9.10	38.6	0.45	1.18	2.44	0.71	0.26	10.12
100 mg/kg	5	Mean	80.12	859.6	15.42	43.66	50.82	17.94	35.34	111.36
		S.D.	16.38	19.6	0.53	1.39	1.47	0.56	0.48	19.42

Group	Number of animals	Reticulocyte %	PT sec	APTT sec	Differential count of WBC (10 ² /µL)					
					Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil	
0 mg/kg	5	Mean	2.684	17.18	19.48	7.48	64.24	2.34	1.02	0.00
		S.D.	0.072	1.29	1.05	1.75	8.70	0.37	0.22	0.00
100 mg/kg	5	Mean	3.082 +	16.28	19.62	8.84	67.62	2.18	1.48	0.00
		S.D.	0.324	0.54	1.28	2.67	13.09	1.88	0.67	0.00

+: Significantly different from the 0 mg/kg group at p≤0.05 (Mann-Whitney's U-test).

Table 25 Biochemical findings of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L	
					Albumin	α_1	α_2	β	γ					
0 mg/kg	5	Mean	5.58	2.832	1.034	50.84	20.96	7.82	16.00	4.38	64.0	29.8	357.2	0.36
		S.D.	0.18	0.070	0.050	1.17	2.78	0.38	1.08	0.55	9.5	4.5	48.9	0.11
4 mg/kg	5	Mean	5.70	2.872	1.020	50.40	22.16	7.24	15.76	4.44	72.8	33.8	365.0	0.34
		S.D.	0.19	0.074	0.097	2.19	2.45	0.47	0.58	1.03	8.6	5.8	80.0	0.09
20 mg/kg	5	Mean	5.58	2.770	0.986	49.68	21.44	7.40	16.56	4.92	72.6	31.6	426.8	0.44
		S.D.	0.20	0.066	0.063	1.64	1.58	0.70	1.01	0.46	11.4	4.6	72.7	0.11
100 mg/kg	5	Mean	5.38	2.734	1.048	50.94	19.78	7.42	16.18	5.68	76.0	31.8	500.2	0.28
		S.D.	0.33	0.138	0.151	3.62	3.61	0.33	1.20	1.33	9.8	4.9	133.1	0.15

Group	Number of animals	T-Bil mg/dL	Glucose mg/dL	T-Chol mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL	
0 mg/kg	5	Mean	0.062	148.2	66.6	38.6	14.54	0.546	144.2	4.882	106.8	9.60	7.38
		S.D.	0.008	10.1	5.9	17.2	0.88	0.021	1.6	0.465	1.3	0.20	0.41
4 mg/kg	5	Mean	0.056	160.6	49.4	57.2	14.52	0.578	144.4	4.844	108.2	9.58	6.82
		S.D.	0.011	13.8	12.1	20.3	1.54	0.059	1.7	0.353	2.3	0.30	0.49
20 mg/kg	5	Mean	0.058	158.0	55.2	39.0	14.98	0.574	144.4	4.720	107.4	9.56	7.16
		S.D.	0.015	9.8	21.3	13.6	1.60	0.042	1.1	0.242	0.9	0.11	0.40
100 mg/kg	5	Mean	0.064	150.2	44.6	21.4	17.52	0.528	145.4	4.908	109.6 *	9.46	7.06
		S.D.	0.011	6.3	10.7	7.4	2.81	0.046	0.9	0.313	1.5	0.24	0.55

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

Table 26 Biochemical findings of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L	
					Albumin	Globulin	α_1	α_2	β					
0 mg/kg	5	Mean	6.50	3.368	1.078	51.86	19.48	7.08	17.00	4.58	82.4	28.8	214.0	0.84
		S.D.	0.19	0.050	0.068	1.51	2.20	0.72	1.17	1.04	17.4	6.9	31.0	0.11
4 mg/kg	5	Mean	6.50	3.244	0.996	49.84	20.52	7.64	17.44	4.56	79.2	30.0	209.8	0.78
		S.D.	0.16	0.202	0.075	1.91	0.87	0.56	1.11	1.03	6.5	3.3	15.5	0.20
20 mg/kg	5	Mean	6.32	3.236	1.052	51.12	19.60	7.90	17.26	4.12	83.8	26.2	223.2	0.80
		S.D.	0.48	0.347	0.127	3.18	1.48	1.73	1.23	0.54	11.7	3.1	64.4	0.35
100 mg/kg	5	Mean	6.44	3.336	1.078	51.84	19.94	7.48	16.58	4.16	74.8	30.6	204.8	0.88
		S.D.	0.09	0.151	0.096	2.22	1.33	1.01	1.58	0.78	4.1	1.5	62.3	0.19

Group	Number of animals	T-Bil mg/dL	Glucose mg/dL	T-Chol mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL	
0 mg/kg	5	Mean	0.080	156.2	57.6	36.0	25.78	0.648	140.8	4.882	104.0	10.94	8.72
		S.D.	0.007	24.3	10.0	18.2	2.35	0.061	0.8	0.436	0.7	0.09	0.61
4 mg/kg	5	Mean	0.076	139.0	52.8	57.0	27.82	0.672	142.2	5.044	105.0	10.86	8.74
		S.D.	0.005	11.6	13.6	19.0	2.05	0.052	0.8	0.347	0.7	0.35	0.32
20 mg/kg	5	Mean	0.072	143.4	56.8	33.6	28.22	0.676	142.6	4.738	106.2 *	10.92	8.48
		S.D.	0.013	17.2	12.7	9.5	4.41	0.086	1.5	0.524	1.3	0.49	1.31
100 mg/kg	5	Mean	0.060 ++	139.0	50.8	33.4	31.18 *	0.670	142.8 *	5.004	106.8 **	10.80	8.26
		S.D.	0.000	9.9	12.4	8.4	1.55	0.023	1.3	0.290	1.6	0.37	0.33

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

**: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Dunnett's test).

++: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Mann-Whitney's U-test).

Table 27 Biochemical findings of male rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L				
					Albumin	Globulin											
						α_1	α_2	β	γ								
0 mg/kg	5	Mean	5.84	2.858	0.958	48.88	22.46	7.32	16.20	5.14	77.6	37.6	285.6	0.52			
		S.D.	0.15	0.134	0.067	1.74	1.58	0.51	1.52	0.94	14.6	14.8	71.4	0.22			
100 mg/kg	5	Mean	5.64	2.766	0.964	49.02	20.22	7.86	17.36	5.54	63.2 +	27.8	399.0	0.70			
		S.D.	0.26	0.129	0.104	2.68	2.88	1.05	1.14	0.19	3.6	1.3	86.0	0.30			

Group	Number of animals	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL	
0 mg/kg	5	Mean	0.064	163.4	65.2	25.6	16.16	0.572	144.2	4.600	106.2	9.76	7.08
		S.D.	0.013	22.5	18.9	17.4	1.18	0.029	1.5	0.400	1.1	0.15	0.45
100 mg/kg	5	Mean	0.044 *	160.0	47.0	38.4	15.12	0.544	144.4	4.856	108.0	9.80	7.54
		S.D.	0.005	13.2	11.8	21.4	1.31	0.029	0.5	0.152	1.4	0.17	0.47

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

+: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Mann-Whitney's U-test).

Table 28 Biochemical findings of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L				
					Albumin	Globulin											
						α_1	α_2	β	γ								
0 mg/kg	5	Mean	6.48	3.752	1.380	57.88	17.78	5.82	12.82	5.70	63.6	28.2	134.8	0.68			
		S.D.	0.13	0.197	0.107	1.87	1.35	0.48	0.72	1.60	4.9	5.5	25.8	0.19			
100 mg/kg	5	Mean	6.22	3.610	1.392	58.08	15.82	6.80	14.10	5.20	63.2	24.2	167.4	0.70			
		S.D.	0.25	0.177	0.134	2.35	2.33	0.89	1.55	0.79	11.1	7.6	54.0	0.22			
Group	Number of animals	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL	100				
		0 mg/kg	5	Mean	0.084	148.8	69.2	11.4	16.66	0.620	144.6	4.260	108.8	10.32	6.32		
		S.D.			0.015	5.7	10.2	1.1	1.08	0.037	1.1	0.236	0.8	0.04	0.43		
100 mg/kg	5	Mean	0.054 **	150.6	55.4	19.2	15.50	0.574	145.0	4.144	109.8 *	10.28	6.74				
		S.D.			0.011	11.7	9.3	5.7	1.09	0.055	0.7	0.611	0.4	0.28	0.49		

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

**: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Dunnett's test).

Table 29 Serum hormone levels of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals		T3 ng/mL	T4 ng/mL	TSH ng/mL
0 mg/kg	5	Mean	0.450	69.7124	3.732
		S.D.	0.070	14.9065	1.491
4 mg/kg	5	Mean	0.466	74.7306	6.586
		S.D.	0.076	10.9317	2.712
20 mg/kg	5	Mean	0.390	80.0460	7.064
		S.D.	0.060	8.6452	5.351
100 mg/kg	5	Mean	0.436	71.3830	9.682
		S.D.	0.119	3.8313	6.029

Table 30 Serum hormone levels of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals		T3 ng/mL	T4 ng/mL	TSH ng/mL
0 mg/kg	5	Mean	0.734	65.4832	4.478
		S.D.	0.023	9.3002	1.454
4 mg/kg	5	Mean	0.606 **	65.5560	5.434
		S.D.	0.036	15.8637	5.130
20 mg/kg	5	Mean	0.626 **	61.8574	4.408
		S.D.	0.068	7.5664	2.329
100 mg/kg	5	Mean	0.532 **	66.3606	8.338
		S.D.	0.040	14.8499	4.661

**: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Dunnett's test).

Table 31 Serum hormone levels of male rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals		T3 ng/mL	T4 ng/mL	TSH ng/mL
0 mg/kg	5	Mean	0.474	117.4978	13.314
		S.D.	0.123	15.0048	5.530
100 mg/kg	5	Mean	0.452	89.2546 *	13.564
		S.D.	0.061	11.8703	3.229

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

Table 32 Serum hormone levels of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals		T3 ng/mL	T4 ng/mL	TSH ng/mL
0 mg/kg	5	Mean	0.784	46.2566	3.758
		S.D.	0.143	16.6958	0.859
100 mg/kg	5	Mean	0.684	58.4804	28.772
		S.D.	0.032	7.1067	54.988

Table 33 Gross findings of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	End of administration				End of recovery	
	0 mg/kg	4 mg/kg	20 mg/kg	100 mg/kg	0 mg/kg	100 mg/kg
Number of animals examined	7	12	12	7	5	5
No abnormal findings	5	12	11	7	4	5
Organ: Findings						
Liver: Atrophy, papillary process of caudate lobe	1	0	0	0	0	0
Yellowish green discoloration, papillary process of caudate lobe	1	0	0	0	0	0
Ileum : Diverticulum	0	0	0	0	1	0
Kidney: Dilatation, renal pelvis	1	0	0	0	0	0
Epididymis: Yellowish white patch, cauda	0	0	1	0	0	0

Values are number of animals with findings.

Table 34 Gross findings of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	0 mg/kg			4 mg/kg			20 mg/kg			100 mg/kg		
	A	B	T	A	B	T	A	B	T	A	B	T
Number of animals examined	11	1	12	10	2	12	12	0	12	12	0	12
No abnormal findings	11	1	12	9	2	11	12	-	12	11	-	11
Organ: Findings												
Stomach: White mass, limiting ridge	0	0	0	1	0	1	0	-	0	0	-	0
Skin of right posterior abdominal region: Subcutaneous yellow mass	0	0	0	0	0	0	0	-	0	1	-	1

Values are the number of animals with findings.

Fate: A, animals nursing pups until lactation day 4; B, animals that were not pregnant and showed total litter loss; T, total (A+B).

-: Blank.

Table 35 Gross findings of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

	Group	0 mg/kg	100 mg/kg
Number of animals examined ^a		5	5
No abnormal findings		5	5

Values are the number of animals with findings.

a: Animals euthanized at the end of recovery.

Table 36-1 Absolute and relative organ weights of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Body weight		Liver		Kidney		Spleen		Heart		Brain		Pituitary gland	
		g	g	%	g	%	g	%	g	%	g	%	g	%	mg
0 mg/kg	5	Mean	485.4	12.152	2.502	3.248	0.672	0.806	0.166	1.478	0.308	2.200	0.456	13.90	2.884
		S.D.	44.2	1.268	0.044	0.363	0.084	0.103	0.030	0.106	0.038	0.062	0.032	1.48	0.403
4 mg/kg	5	Mean	482.6	11.808	2.446	2.946	0.612	0.748	0.156	1.352	0.280	2.124	0.440	11.80 *	2.448
		S.D.	10.1	0.545	0.095	0.190	0.038	0.094	0.021	0.140	0.025	0.086	0.014	1.35	0.330
20 mg/kg	5	Mean	486.2	12.116	2.490	2.930	0.604	0.748	0.154	1.366	0.280	2.186	0.452	13.30	2.738
		S.D.	21.6	0.849	0.151	0.144	0.021	0.124	0.022	0.065	0.016	0.107	0.029	0.82	0.153
100 mg/kg	5	Mean	444.6 +	14.504 **	3.260 **	3.058	0.688	0.750	0.170	1.320	0.298	2.138	0.480	12.22	2.752
		S.D.	12.5	0.610	0.068	0.204	0.060	0.103	0.029	0.126	0.029	0.142	0.032	1.11	0.298

Group	Number of animals	Thymus		Thyroid		Adrenal		Prostate		Seminal vesicle		
		mg	$10^{-3}\%$	mg	$10^{-3}\%$	mg	$10^{-3}\%$	mg	$10^{-3}\%$	g	%	
0 mg/kg	5	Mean	317.6	65.630	18.94	3.940	70.0	14.520	849.8	176.726	2.280	0.472
		S.D.	33.8	7.096	1.60	0.581	2.1	1.433	136.8	35.592	0.276	0.058
4 mg/kg	5	Mean	303.4	62.652	20.58	4.266	58.4	12.094	753.4	156.812	2.336	0.484
		S.D.	75.6	14.577	1.53	0.339	11.9	2.406	209.5	46.506	0.224	0.047
20 mg/kg	5	Mean	376.8	77.134	24.26 *	4.984 *	62.4	12.906	909.0	187.960	2.172	0.448
		S.D.	92.0	17.251	4.28	0.778	9.9	2.559	265.8	57.778	0.218	0.054
100 mg/kg	5	Mean	329.4	74.196	22.16	4.980 *	57.8	13.026	647.4	146.038	2.172	0.488
		S.D.	31.8	8.279	3.26	0.668	8.4	2.033	113.5	28.091	0.302	0.065

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).**: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Dunnett's test).+: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Mann-Whitney's U-test).

Table 36-2 Absolute and relative organ weights of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Body weight		Testis		Epididymis	
			g	g	%	g	%
0 mg/kg	7	Mean	500.4	3.540	0.711	1.414	0.284
		S.D.	44.8	0.283	0.086	0.109	0.037
4 mg/kg	12	Mean	492.8	3.278	0.667	1.266 *	0.256
		S.D.	19.0	0.259	0.057	0.109	0.021
20 mg/kg	12	Mean	496.0	3.380	0.683	1.326	0.269
		S.D.	26.9	0.298	0.057	0.123	0.026
100 mg/kg	7	Mean	462.9	3.314	0.719	1.356	0.294
		S.D.	34.9	0.354	0.087	0.119	0.033

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

Table 37 Absolute and relative organ weights of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Body weight		Liver		Kidney		Spleen		Heart		Brain	
		g	g	%	g	%	g	%	g	%	g	%	g
0 mg/kg	5	Mean	300.4	10.166	3.386	2.028	0.674	0.722	0.240	1.068	0.356	2.080	0.694
		S.D.	13.8	0.480	0.120	0.125	0.021	0.097	0.033	0.068	0.011	0.082	0.054
4 mg/kg	5	Mean	297.2	9.704	3.266	1.912	0.642	0.748	0.252	1.012	0.342	2.046	0.688
		S.D.	11.1	0.612	0.214	0.169	0.046	0.147	0.040	0.048	0.013	0.067	0.013
20 mg/kg	5	Mean	298.4	9.996	3.348	2.004	0.672	0.778	0.258	1.000	0.334	2.054	0.690
		S.D.	15.0	0.806	0.145	0.079	0.041	0.125	0.033	0.033	0.013	0.097	0.025
100 mg/kg	5	Mean	296.8	10.528	3.548	2.058	0.694	0.660	0.222	0.990	0.334	2.064	0.694
		S.D.	11.8	0.754	0.199	0.191	0.038	0.070	0.019	0.062	0.018	0.088	0.031

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Group	Number of animals	Pituitary gland		Thymus		Thyroid		Adrenal		Ovary		
		mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	
0 mg/kg	5	Mean	16.22	5.390	250.6	83.440	19.28	6.400	76.4	25.440	123.2	41.024
		S.D.	2.13	0.544	84.9	28.587	3.06	0.784	5.8	1.682	15.0	4.832
4 mg/kg	5	Mean	16.52	5.562	232.2	78.296	15.78	5.300	79.6	26.798	109.6	36.770
		S.D.	1.63	0.572	70.4	24.114	2.95	0.862	6.0	1.998	17.8	4.994
20 mg/kg	5	Mean	17.10	5.746	322.4	107.838	16.96	5.712	79.4	26.592	113.6	38.170
		S.D.	0.40	0.348	54.4	15.767	3.42	1.270	7.9	2.117	7.6	3.532
100 mg/kg	5	Mean	15.38	5.188	276.8	93.522	18.04	6.068	75.4	25.428	123.8	41.748
		S.D.	0.44	0.252	42.6	16.510	1.99	0.547	7.9	2.773	15.4	5.391

Table 38 Absolute and relative organ weights of male rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Body weight		Liver		Kidney		Spleen		Heart		Brain		Pituitary gland		Thymus	
		g	g	%	g	%	g	%	g	%	g	%	g	%	mg	10 ⁻³ %	mg
0 mg/kg	5	Mean	514.8	12.376	2.402	3.058	0.594	0.792	0.156	1.526	0.294	2.152	0.418	12.92	2.504	268.6	52.208
		S.D.	31.2	1.398	0.168	0.149	0.034	0.154	0.022	0.056	0.017	0.074	0.024	2.37	0.366	43.4	7.870
100 mg/kg	5	Mean	494.0	14.624 *	2.968 **	3.184	0.642	0.692	0.142	1.430	0.290	2.168	0.440	12.34	2.498	272.6	55.206
		S.D.	16.3	1.349	0.327	0.312	0.063	0.081	0.013	0.146	0.025	0.036	0.016	2.37	0.437	56.4	11.495

Group	Number of animals	Thyroid		Adrenal		Testis		Epididymis		Prostate		Seminal vesicle		
		mg	10 ⁻³ %	mg	10 ⁻³ %	g	%	g	%	mg	10 ⁻³ %	g	%	
0 mg/kg	5	Mean	21.90	4.262	70.4	13.694	3.292	0.642	1.366	0.266	773.6	150.090	2.244	0.436
		S.D.	3.98	0.819	3.8	0.717	0.411	0.083	0.180	0.034	145.5	26.611	0.328	0.061
100 mg/kg	5	Mean	22.40	4.542	55.2 **	11.170 **	3.464	0.702	1.456	0.294	788.2	159.074	2.284	0.462
		S.D.	4.10	0.838	6.1	1.144	0.258	0.057	0.150	0.035	183.2	33.800	0.253	0.048

*: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Dunnett's test).

**: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Dunnett's test).

Table 39 Absolute and relative organ weights of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of animals	Body weight		Liver		Kidney		Spleen		Heart		Brain	
		g	g	%	g	%	g	%	g	%	g	%	g
0 mg/kg	5	Mean	279.4	6.954	2.488	1.758	0.630	0.480	0.172	0.956	0.344	2.052	0.736
		S.D.	9.7	0.374	0.111	0.071	0.029	0.071	0.024	0.075	0.040	0.139	0.050
100 mg/kg	5	Mean	276.0	7.482	2.710	1.926	0.698	0.522	0.190	0.884	0.320	2.008	0.730
		S.D.	15.4	1.003	0.283	0.165	0.064	0.061	0.025	0.034	0.019	0.067	0.057

Group	Number of animals	Pituitary gland		Thymus		Thyroid		Adrenal		Ovary		
		mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	
0 mg/kg	5	Mean	15.66	5.598	276.6	99.096	18.42	6.604	67.0	23.984	102.8	36.782
		S.D.	1.55	0.451	51.4	19.098	1.97	0.793	3.5	1.109	22.8	8.034
100 mg/kg	5	Mean	15.30	5.544	315.2	114.634	16.88	6.086	74.2	26.948	97.6	35.450
		S.D.	3.01	1.052	35.1	15.823	3.46	1.007	12.4	4.704	28.5	10.513

Table 40 Histopathological findings of male rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

	Group	Control	4 mg/kg	20 mg/kg	100 mg/kg
Number of animals examined		7	12	12	7
Organ: Findings	Grade				
Lung: Aggregation, macrophage, alveolar	+	2 (5)	-	-	1 (5)
	++	0 (5)	-	-	1 (5)
Mineralization, artery	+	0 (5)	-	-	2 (5)
Metaplasia, osseous, alveoli	+	1 (5)	-	-	0 (5)
Liver: Hypertrophy, hepatocyte, centrilobular	+	0	0	5	$\begin{pmatrix} 0 \\ 7 \end{pmatrix}$ **
	++	0	0	0	
Fatty change, centrilobular	+	0	0	2	7 ^{ss}
Fatty change, periportal	+	1	0	0	0
Microgranuloma	+	5	8	8	3
Necrosis, massive	+++	1	0	0	0
Deposit, hemosiderin/ hematoidin	++	1	0	0	0
Fibrosis	++	1	0	0	0
Mineralization	++	1	0	0	0
Heart: Myocardial degeneration, focal	+	0 (5)	-	-	1 (5)
Kidney: Regeneration, tubular epithelium	+	1 (5)	-	-	0 (5)
Dilatation, renal pelvis	+	1 (5)	-	-	0 (5)
Epididymis: Granuloma, spermatic	+	0 (5)	-	1 (1)	0 (5)
Prostate: Cellular infiltration, inflammatory cell	+	2	8	2	6

Values are the number of animals with findings.

Values in parentheses are the number of animals examined.

-: Blank.

Grade; +: slight change, ++: moderate change, +++: severe change.

**: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Mann-Whitney's U-test).

ss: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Fisher's exact probability test).

Table 41 Histopathological findings of female rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

	Group	Control	4 mg/kg	20 mg/kg	100 mg/kg
Number of animals examined		12	12	12	12
Organ: Findings	Grade				
Lung: Aggregation, macrophage, alveolar	+	1 (5)	-	-	1 (5)
Mineralization, artery	+	1 (5)	-	-	0 (5)
Stomach, limiting ridge: Cyst, squamous cell	+	0 (5)	1 (1)	-	0 (5)
Liver: Hypertrophy, hepatocyte, centrilobular	+	0	0	0	8 ^{ss}
Fatty change, periportal	+	0	1	1	1
Microgranuloma	+	7	7	8	7
Necrosis, focal	+	0	1	0	1
Kidney: Cellular infiltration, inflammatory cell, renal pelvic mucosa	+	1 (5)	-	-	0 (5)
Mineralization, cortico-medullary junction	+	0 (5)	-	-	1 (5)
Mineralization, papilla	+	0 (5)	-	-	1 (5)
Pituitary gland: Tubular hyperplasia, pars nervosa	+	1 (5)	-	-	0 (5)
Mammary gland: Fibroadenoma	<+>	0 (5)	-	-	1 (6)

Values are the number of animals with findings.

Values in parentheses are the number of animals examined.

-: Blank.

Grade; +: slight change, <+>: presence in "presence or not" basis.

^{ss}: Significantly different from the 0 mg/kg group at p≤0.01 (Fisher's exact probability test).

Table 42 Histopathological findings of male rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

	Group	Control	100 mg/kg
Number of animals examined		5	5
Organ: Findings	Grade		
Lung: Aggregation, macrophage, alveolar	+	1	5 ^{\$}
Mineralization, artery	+	1	0
Ileum: Diverticulum	+	1	0
Liver: Hypertrophy, hepatocyte, centrilobular	+	0	5 ^{\$\$}
Fatty change, centrilobular	+	0	1
Fatty change, periportal	+	1	0
Microgranuloma	+	3	2
Epididymis: Granuloma, spermatic	+	0	1
Prostate: Cellular infiltration, inflammatory cell	+	1	2
Pituitary gland: Cyst, pars distalis	+	0	1

Values are the number of animals with findings.

Grade; +: slight change.

^{\$}: Significantly different from the 0 mg/kg group at $p \leq 0.05$ (Fisher's exact probability test).

^{\$\$}: Significantly different from the 0 mg/kg group at $p \leq 0.01$ (Fisher's exact probability test).

Table 43 Histopathological findings of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

	Group	Control	100 mg/kg
Number of animals examined		5	5
Organ: Findings	Grade		
Lung: Aggregation, macrophage, alveolar	+	1	0
Mineralization, artery	+	0	1
Metaplasia, osseous, alveoli	+	1	1
Liver: Hypertrophy, hepatocyte, centrilobular	+	0	1
Microgranuloma	+	1	4
Kidney: Cellular infiltration, inflammatory cell, renal pelvic mucosa	+	1	0
Mineralization, cortex	+	0	1

Values are the number of animals with findings.

Grade; +: slight change.

Table 44 Reproduction performance in parental rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Estrous cycle Normality		Copulation index		Fertility index	Gestation index	Nursing index
	Incidence	Length (days) ^a	Male Incidence	Female Incidence	Incidence	Incidence	Incidence
	(%)		(%)	(%)	(%)	(%)	(%)
0 mg/kg	12/12 (100)	4.11 0.22	12/12 (100)	12/12 (100)	11/12 (91.7)	11/11 (100)	22.3 0.5
4 mg/kg	12/12 (100)	4.18 0.32	12/12 (100)	12/12 (100)	12/12 (100)	12/12 (100)	22.3 0.5
20 mg/kg	12/12 (100)	4.03 0.09	12/12 (100)	12/12 (100)	12/12 (100)	12/12 (100)	22.3 0.5
100 mg/kg	12/12 (100)	4.00 0.00	12/12 (100)	12/12 (100)	12/12 (100)	12/12 (100)	22.2 0.4

Normal estrous cycle = (number of females with normal estrous cycle / number of females examined) x 100.

Copulation index = (number of animals with successful copulation / number of animals mated) x 100.

Fertility index = (number of pregnant females / number of pairs with successful copulation) x 100.

Gestation index = (number of females with live pups / number of pregnant females) x 100.

Nursing index = (number of females nursing live pups on lactation day 4 / number of females with live pups delivery) x 100.

a: Values are means and S.D.

Table 45 Estrus cycle of female rats in 14-day recovery test following 42-day repeated oral dose with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Administration period		Recovery period	
	Normality		Normality	
	Incidence (%)	Length (days) ^a	Incidence (%)	Length (days) ^a
0 mg/kg	5/5 (100)	4.22 0.39	5/5 (100)	4.20 0.45
100 mg/kg	5/5 (100)	4.02 0.04	5/5 (100)	4.00 0.00

Normal estrous cycle = (number of females with normal estrous cycle / number of females examined) x 100.

a: Values are means and S.D.

Table 46 Pregnancy and litter data of rats dosed orally with perfluorohexadecanoic acid in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (SR07125)

Group	Number of pregnant animals	Number of corpora lutea	Number of implantation sites	Implantation index (%)	Delivery index (%)	Lactation day 0					Lactation day 4						
						Total	Male	Female	Alive	Dead	All pups	Live pups	Sex ratio	Live birth index (%)	Number of live pups	Sex ratio	Live pups
Pup data																	
0 mg/kg	11	Mean S.D.	16.5 1.1	16.1 1.4	97.675 3.243	94.254 5.541	15.2 1.7	7.6 2.6	7.5 2.7	15.1 1.7	0.1 0.3	0.505 0.165	0.503 0.166	99.432 1.884	15.0 1.7	0.500 0.164	99.432 1.884
4 mg/kg	12	Mean S.D.	17.0 1.2	16.6 1.2	97.600 3.800	96.328 5.125	16.0 1.7	6.5 2.3	9.5 3.1	15.8 1.6	0.3 0.9	0.413 0.158	0.413 0.158	98.611 4.812	13.1 6.3	0.396 0.144	82.870 38.741
20 mg/kg	12	Mean S.D.	15.8 1.9	15.3 2.1	96.203 4.235	92.817 7.652	14.2 2.2	6.0 2.0	8.2 2.7	14.2 2.2	0.0 0.0	0.429 0.140	0.429 0.140	100.000 0.000	14.1 2.3	0.433 0.147	99.405 2.061
100 mg/kg	12	Mean S.D.	16.1 1.6	15.8 1.6	98.425 2.866	92.011 7.438	14.6 2.0	7.1 2.8	7.5 3.3	14.5 2.2	0.1 0.3	0.492 0.183	0.489 0.180	99.243 2.624	13.7 2.7	0.498 0.180	94.256 12.109

Implantation index = (number of implantation sites / number of corpora lutea) x 100.

Delivery index = (number of pups born / number of implantation sites) x 100.

Sex ratio on lactation day 0 = (number of male pups born / number of pups born) or (number of live male pups / number of live pups).

Sex ratio on lactation day 4 = number of live male pups / number of live pups.

Live birth index = (number of live pups on lactation day 0 / number of pups born) x 100.

Viability index on lactation day 4 = (number of live pups on lactation day 4 / number of live pups on lactation day 0) x 100.

Table 47 General appearance of pups in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test of perfluorohexadecanoic acid in rats (SR07125)

Group	Findings	Male					Female				
		Day of lactation					Day of lactation				
		0	1	2	3	4	0	1	2	3	4
0 mg/kg	Number of pups examined	84	83	82	82	82	83	83	83	83	83
	No abnormal findings	83	82	82	82	82	83	83	83	83	83
	Death or missing	1	1	0	0	0	0	0	0	0	0
4 mg/kg	Number of pups examined	78	77	72	64	61	114	112	103	102	96
	No abnormal findings	72	61	61	61	61	104	96	96	96	96
	Milk-band negative	5	11	3	0	0	8	7	6	0	0
	Death or missing	1	5	8	3	0	2	9	1	6	0
20 mg/kg	Number of pups examined	72	72	72	72	72	98	98	98	98	97
	No abnormal findings	72	72	72	72	72	98	98	98	97	97
	Death or missing	0	0	0	0	0	0	0	0	1	0
100 mg/kg	Number of pups examined	85	84	83	80	80	90	90	87	84	84
	No abnormal findings	83	81	80	80	80	88	86	84	84	84
	Milk-band negative	1	2	0	0	0	2	1	0	0	0
	Death or missing	1	1	3	0	0	0	3	3	0	0

Values are the number of pups with findings.

Table 48 Body weight changes of pups in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test of perfluorohexadecanoic acid in rats (SR07125)

Group	Number of litters	Male			Female			
		Body weight (g) on lactation day 0	1	4	Body weight (g) on lactation day 0	1	4	
0 mg/kg	11	Mean	6.63	7.25	10.53	6.27	6.91	10.05
		S.D.	0.58	0.56	0.85	0.51	0.51	0.79
					(10)			(10)
4 mg/kg	12	Mean	6.67	7.12	10.63	6.22	6.58	9.97
		S.D.	0.67	1.08	1.54	0.60	1.07	1.36
20 mg/kg	12	Mean	6.75	7.33	10.67	6.40	6.98	10.15
		S.D.	0.69	0.75	1.14	0.66	0.82	1.25
100 mg/kg	12	Mean	6.45	6.97	9.93	6.08	6.59	9.43
		S.D.	0.46	0.80	1.24	0.50	0.79	1.31

Values in parentheses are the number of litters examined.

Table 49 Gross findings of pups in the combined repeated dose toxicity study with the reproduction/developmental toxicity screening test of perfluorohexadecanoic acid in rats (SR07125)

Group	Male				Female			
	0 mg/kg	4 mg/kg	20 mg/kg	100 mg/kg	0 mg/kg	4 mg/kg	20 mg/kg	100 mg/kg
Findings of dead pups during lactation days 0-4								
Number of pups examined	1	12	0	4	0	13	0	3
No abnormal findings	1	12	-	4	-	13	-	3
Findings of pups euthanized on lactation day 4								
Number of pups examined	82	61	72	80	83	96	97	84
No abnormal findings	82	61	72	79	83	96	97	84
Organ: Findings								
Liver: Yellowish white patch, middle lobe	0	0	0	1	0	0	0	0

Values are the number of pups with findings.

-: Blank.

INDIVIDUAL DATA 1-1-1

STUDY NO. SR07125 TITLE : PFHxD/DA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg

- 123

N : No abnormal findings.

A : AM

· Blank

P · PM

(to be continued)

INDIVIDUAL DATA 1-1-1 (continued)

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg

Animal No.	Recovery period (day)														Autopsy day		
	1		2		3		4		5		6		7		8		
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	
101	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
102	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
103	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
104	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
105	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
106	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
107	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
108	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
109	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
110	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
111	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
112	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#

N : No abnormal findings.

A : AM.

: Blank.

P : PM.

INDIVIDUAL DATA 1-1-2

STUDY NO. SR07125 TITLE : PFHxD/DA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg

- 125

N : No abnormal findings.

A : AM.

P · PM

INDIVIDUAL DATA 1-1-3

STUDY NO. SR07125 TITLE : PFHxD/DA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg

- 126

N : No abnormal findings.

A : AM.

P · PM

INDIVIDUAL DATA 1-1-4

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg

Animal No.	Administration period (day)																																								
	1		2		3		4		5		6		7		8		9		10		11		12		13		14		15		16		17		18		19		20		21
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P					
401	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
402	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
403	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
404	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
405	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
406	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
407	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
408	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
409	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
410	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
411	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
412	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					

Animal No.	Administration period (day)																														Autopsy day											
	22		23		24		25		26		27		28		29		30		31		32		33		34		35		36		37		38		39		40		41		42	
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P						
401	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
402	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
403	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
404	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
405	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#					
406	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#					
407	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
408	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					
409	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#					
410	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#					
411	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#					
412	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N					

N : No abnormal findings.

A : AM.

: Blank.

P : PM.

(to be continued)

INDIVIDUAL DATA 1-1-4 (continued)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg

Animal No.	Recovery period (day)														Autopsy day		
	1		2		3		4		5		6		7		8		
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	
401	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
402	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
403	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
404	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
405	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
406	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
407	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
408	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#
409	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
410	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
411	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
412	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#

N : No abnormal findings.

A : AM.

: Blank.

P : PM.

INDIVIDUAL DATA 1-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

Animal No.	Pre-mating period (day)																																				
	1		2		3		4		5		6		7		8		9		10		11		12		13		14		15		16		17		18		
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P			
151	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N			
152	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#	#	#	#	#	#	#	#				
153	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N			
154	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#			
155	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#			
156	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#			
157	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#			
158	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#			
159	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#			
160	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#			
161	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#			
162	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#			

N : No abnormal findings.

A : AM.

: Blank (Copulated).

P : PM.

(to be continued)

INDIVIDUAL DATA 1-2-1 (continued-1)

STUDY NO. SR07125 **TITLE : PFHxD/DA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)**

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

N : No abnormal findings.

a : Non-pregnancy.

: Blank(Delivered).

A : AM.

P : PM.

(to be continued)

INDIVIDUAL DATA 1-2-1 (continued-2)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

Animal No.	Lactation period (day)									
	0		1		2		3	4	5	6
	A	P	A	P	A	P	A	P	A	A
151	N	N	N	N	N	N	N	N	N	N
152	N	N	N	N	N	N	N	N	N	N
153	N	N	N	N	N	N	N	N	N	N
154	N	N	N	N	N	N	N	N	N	N
155	N	N	N	N	N	N	N	N	N	N
156	N	N	N	N	N	N	N	N	N	N
157	N	N	N	N	N	N	N	N	N	N
158	N	N	N	N	N	N	N	N	N	N
159	N	N	N	N	N	N	N	N	N	N
160	N	N	N	N	N	N	N	N	N	N
161 ^a	#	#	#	#	#	#	#	#	#	#
162	N	N	N	N	N	N	N	N	N	N

N: No abnormal findings.

A : AM.

a: Non-pregnancy.

P : PM.

#: Blank.

INDIVIDUAL DATA 1-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg

Animal No.	Pre-mating period (day)																																						
	1		2		3		4		5		6		7		8		9		10		11		12		13		14		15		16		17		18		19		20
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	
251	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	
252	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	
253	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	
254	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	
255	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	
256	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	#	
257	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N		
258	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	
259	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	
260	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	
261	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	
262	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	

N : No abnormal findings.

A : AM.

(to be continued)

: Blank (Copulated).

P : PM.

INDIVIDUAL DATA 1-2-2 (continued-1)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg

Animal No.	Gestation period (day)																																												
	0		1		2		3		4		5		6		7		8		9		10		11		12		13		14		15		16		17		18		19		20		21		22
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P							
251	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#								
252	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N								
253	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#								
254	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N								
255	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#								
256	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#								
257	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N								
258	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#								
259	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#								
260	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#								
261	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N								
262	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#								

N : No abnormal findings.

: Blank(Delivered).

A : AM.

P : PM.

(to be continued)

INDIVIDUAL DATA 1-2-2 (continued-2)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg

Animal No.	Lactation period (day)									
	0		1		2		3	4	5	6
	A	P	A	P	A	P	A	P	A	A
251	N	N	N	N	N	N	N	N	N	N
252 ^a	N	N	N	N	N	N	N	N	N	N
253	N	N	N	N	N	N	N	N	N	N
254 ^a	N	N	N	N	N	N	N	N	N	N
255	N	N	N	N	N	N	N	N	N	N
256	N	N	N	N	N	N	N	N	N	N
257	N	N	N	N	N	N	N	N	N	N
258	N	N	N	N	N	N	N	N	N	N
259	N	N	N	N	N	N	N	N	N	N
260	N	N	N	N	N	N	N	N	N	N
261	N	N	N	N	N	N	N	N	N	N
262	N	N	N	N	N	N	N	N	N	N

N : No abnormal findings.

A : AM.

a : All pups died by day 3 of lactation.

P : PM.

INDIVIDUAL DATA 1-2-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg

Animal No.	Pre-mating period (day)																														
	1		2		3		4		5		6		7		8		9		10		11		12		13		14		15		16
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	
351	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#		
352	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#			
353	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#		
354	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N		
355	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#		
356	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N		
357	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N		
358	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N		
359	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N		
360	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#			
361	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#			
362	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N			

N : No abnormal findings.

A : AM.

(to be continued)

: Blank (Copulated).

P : PM.

INDIVIDUAL DATA 1-2-3 (continued-1)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg

Animal No.	Gestation period (day)																																												
	0		1		2		3		4		5		6		7		8		9		10		11		12		13		14		15		16		17		18		19		20		21		22
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P									
351	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#									
352	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#									
353	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#									
354	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#									
355	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#									
356	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N									
357	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N									
358	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N									
359	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#									
360	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#									
361	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#									
362	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#									

N : No abnormal findings.

: Blank(Delivered).

A : AM.

P : PM.

(to be continued)

INDIVIDUAL DATA 1-2-3 (continued-2)

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg

Animal No.	Lactation period (day)									
	0		1		2		3	4	5	6
	A	P	A	P	A	P	A	P	A	A
351	N	N	N	N	N	N	N	N	N	N
352	N	N	N	N	N	N	N	N	N	N
353	N	N	N	N	N	N	N	N	N	N
354	N	N	N	N	N	N	N	N	N	N
355	N	N	N	N	N	N	N	N	N	N
356	N	N	N	N	N	N	N	N	N	N
357	N	N	N	N	N	N	N	N	N	N
358	N	N	N	N	N	N	N	N	N	N
359	N	N	N	N	N	N	N	N	N	N
360	N	N	N	N	N	N	N	N	N	N
361	N	N	N	N	N	N	N	N	N	N
362	N	N	N	N	N	N	N	N	N	N

N : No abnormal findings.

A : AM.

P : PM.

INDIVIDUAL DATA 1-2-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Pre-mating period (day)																																
	1		2		3		4		5		6		7		8		9		10		11		12		13		14		15		16		17
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	
451	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#	#	#	
452	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
453	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#	#	#	
454	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
455	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	
456	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
457	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#	#	#	#	
458	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
459	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	
460	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	
461	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#
462	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#	#	#	#	#	

N : No abnormal findings.

A : AM.

(to be continued)

: Blank (Copulated).

P : PM.

INDIVIDUAL DATA 1-2-4 (continued-1)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Gestation period (day)																																																
	0		1		2		3		4		5		6		7		8		9		10		11		12		13		14		15		16		17		18		19		20		21		22				
	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P	A	P											
451	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#												
452	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#												
453	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#												
454	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N												
455	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#												
456	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N												
457	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#												
458	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#												
459	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#												
460	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	Su(#	#																													
																		35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×	35×35×		
																		20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×	20×20×		
																		(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)	(15)					
461	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#												
462	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N	#	#												

N : No abnormal findings.

A : AM.

(to be continued)

Su : Subcutaneous mass, right posterior abdominal region (size, mm).

P : PM.

: Blank(Delivered).

INDIVIDUAL DATA 1-2-4 (continued-2)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Lactation period (day)									
	0		1		2		3	4	5	6
	A	P	A	P	A	P	A	P	A	A
451	N	N	N	N	N	N	N	N	N	N
452	N	N	N	N	N	N	N	N	N	N
453	N	N	N	N	N	N	N	N	N	N
454	N	N	N	N	N	N	N	N	N	N
455	N	N	N	N	N	N	N	N	N	N
456	N	N	N	N	N	N	N	N	N	N
457	N	N	N	N	N	N	N	N	N	N
458	N	N	N	N	N	N	N	N	N	N
459	N	N	N	N	N	N	N	N	N	N
460	Su(Su(Su(Su(Su(Su(Su(Su(Su(Su(
	35× 35×	20× 20×	20× 20×	20× 20×	20× 20×	20× 20×	20× 20×	20× 20×	20×	
	20× 20×	15× 15×	15× 15×	10× 10×	10× 10×	10× 10×	10× 10×	10×		
	(15)	(15)	(5)	(5)	(5)	(5)	(5)	(5)	(5)	
461	N	N	N	N	N	N	N	N	N	N
462	N	N	N	N	N	N	N	N	N	N

N : No abnormal findings.

A : AM.

Su : Subcutaneous mass, right posterior abdominal region (size, mm).

P : PM.

INDIVIDUAL DATA 1-3-1

STUDY NO. SR07125 **TITLE : PFHxD/DA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)**

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

N : No abnormal findings.

A : AM.

P : PM.

INDIVIDUAL DATA 1-3-2

STUDY NO. SR07125 TITLE : PFHxD/DA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

General appearance ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

N : No abnormal findings.

A:AM.

P · PM

INDIVIDUAL DATA 2-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Pre-administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
101	1	1	1	0	0	1	
102	1	1	1	0	0	1	
103	1	1	1	0	0	1	
104	1	1	1	0	0	1	
105	1	1	1	0	0	1	
106	1	1	1	0	0	1	
107	1	1	1	0	0	1	
108	1	1	1	0	0	1	
109	1	1	1	0	0	1	
110	1	1	1	0	0	1	
111	1	1	1	0	0	1	
112	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Pre-administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
201	1	1	1	0	0	1	
202	1	1	1	0	0	1	
203	1	1	1	0	0	1	
204	1	1	1	0	0	1	
205	1	1	1	0	0	1	
206	1	1	1	0	0	1	
207	1	1	1	0	0	1	
208	1	1	1	0	0	1	
209	1	1	1	0	0	1	
210	1	1	1	0	0	1	
211	1	1	1	0	0	1	
212	1	1	1	0	0	1	

N

12

12

12

12

12

12

INDIVIDUAL DATA 2-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Pre-administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
301	1	1	1	0	0	1	
302	1	1	1	0	0	1	
303	1	1	1	0	0	1	
304	1	1	1	0	0	1	
305	1	1	1	0	0	1	
306	1	1	1	0	0	1	
307	1	1	1	0	0	1	
308	1	1	1	0	0	1	
309	1	1	1	0	0	1	
310	1	1	1	0	0	1	
311	1	1	1	0	0	1	
312	1	1	1	0	0	1	

N

12

12

12

12

12

12

INDIVIDUAL DATA 2-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Pre-administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
401	1	1	1	0	0	1	
402	1	1	1	0	0	1	
403	1	1	1	0	0	1	
404	1	1	1	0	0	1	
405	1	1	1	0	0	1	
406	1	1	1	0	0	1	
407	1	1	1	0	0	1	
408	1	1	1	0	0	1	
409	1	1	1	0	0	1	
410	1	1	1	0	0	1	
411	1	1	1	0	0	1	
412	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-1-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Pre-administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Removal	Handling			Fur	Eyes			Pupil size	Salivation		
101	1	1	2	1	1	1	0	1	1	1	1	0
102	1	1	2	1	1	1	0	1	1	1	1	0
103	1	1	2	1	1	1	0	1	1	1	1	0
104	1	1	2	1	1	1	0	1	1	1	1	0
105	1	1	2	1	1	1	0	1	1	1	1	0
106	1	1	2	1	1	1	0	1	1	1	1	0
107	1	1	2	1	1	1	0	1	1	1	1	0
108	1	1	2	1	1	1	0	1	1	1	1	0
109	1	1	2	1	1	1	0	1	1	1	1	0
110	1	1	2	1	1	1	0	1	1	1	1	0
111	1	1	2	1	1	1	0	1	1	1	1	0
112	1	1	2	1	1	1	0	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-1-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Pre-administration

Animal No.	On the hand												
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
201	1	1	2	1	1	1	0	1	1	1	1	1	0
202	1	1	2	1	1	1	0	1	1	1	1	1	0
203	1	1	2	1	1	1	0	1	1	1	1	1	0
204	1	1	2	1	1	1	0	1	1	1	1	1	0
205	1	1	2	1	1	1	0	1	1	1	1	1	0
206	1	1	2	1	1	1	0	1	1	1	1	1	0
207	1	1	2	1	1	1	0	1	1	1	1	1	0
208	1	1	2	1	1	1	0	1	1	1	1	1	0
209	1	1	2	1	1	1	0	1	1	1	1	1	0
210	1	1	2	1	1	1	0	1	1	1	1	1	0
211	1	1	2	1	1	1	0	1	1	1	1	1	0
212	1	1	2	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-1-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Pre-administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling		Fur	Eyes					Pupil size	Salivation		
301	1	1	2	1	1	1	0	1	1	1	1	0
302	1	1	2	1	1	0	1	1	1	1	1	0
303	1	1	2	1	1	0	1	1	1	1	1	0
304	1	1	2	1	1	0	1	1	1	1	1	0
305	1	1	2	1	1	0	1	1	1	1	1	0
306	1	1	2	1	1	0	1	1	1	1	1	0
307	1	1	2	1	1	0	1	1	1	1	1	0
308	1	1	2	1	1	0	1	1	1	1	1	0
309	1	1	2	1	1	0	1	1	1	1	1	0
310	1	1	2	1	1	0	1	1	1	1	1	0
311	1	1	2	1	1	0	1	1	1	1	1	0
312	1	1	2	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-1-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Pre-administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Removal	Handling			Fur	Eyes			Pupil size	Salivation		
401	1	1	2	1	1	1	0	1	1	1	1	0
402	1	1	2	1	1	1	0	1	1	1	1	0
403	1	1	2	1	1	1	0	1	1	1	1	0
404	1	1	2	1	1	1	0	1	1	1	1	0
405	1	1	2	1	1	1	0	1	1	1	1	0
406	1	1	2	1	1	1	0	1	1	1	1	0
407	1	1	2	1	1	1	0	1	1	1	1	0
408	1	1	2	1	1	1	0	1	1	1	1	0
409	1	1	2	1	1	1	0	1	1	1	1	0
410	1	1	2	1	1	1	0	1	1	1	1	0
411	1	1	2	1	1	1	0	1	1	1	1	0
412	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-1-9

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Pre-administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
101	1	1	1	1	0	0	0	0	1	1	1	1	
102	1	1	1	1	0	0	0	0	1	1	1	1	
103	1	1	1	1	0	0	0	0	1	1	1	1	
104	1	1	1	1	1	1	0	0	1	1	1	1	
105	1	1	1	1	0	0	0	0	1	1	1	1	
106	1	1	1	1	1	0	0	0	1	1	1	1	
107	1	1	1	1	0	1	0	0	1	1	1	1	
108	1	1	1	1	0	0	0	0	1	1	1	1	
109	1	1	1	1	0	0	0	0	1	1	1	1	
110	1	1	1	1	0	0	0	0	1	1	1	1	
111	1	1	1	1	0	0	0	0	1	1	1	1	
112	1	1	1	1	0	0	0	0	1	1	1	1	
N	12	12	12	12	12	12	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	2.4047	0.7460	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-1-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Pre-administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
201	1	1	1	1	0	1	0	0	1	1	1		
202	1	1	1	1	0	0	0	0	1	1	1		
203	1	1	1	1	0	0	0	0	1	1	1		
204	1	1	1	1	0	0	0	0	1	1	1		
205	1	1	1	1	0	0	0	0	1	1	1		
206	1	1	1	1	0	0	0	0	1	1	1		
207	1	1	1	1	0	0	0	0	1	1	1		
208	1	1	1	1	0	0	0	0	1	1	1		
209	1	1	1	1	0	0	0	0	1	1	1		
210	1	1	1	1	0	1	0	0	1	1	1		
211	1	1	1	1	0	0	0	0	1	1	1		
212	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12

INDIVIDUAL DATA 2-1-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Pre-administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
301	1	1	1	1	0	0	0	0	1	1	1	
302	1	1	1	1	0	0	0	0	1	1	1	
303	1	1	1	1	0	0	0	0	1	1	1	
304	1	1	1	1	0	0	0	0	1	1	1	
305	1	1	1	1	0	0	0	0	1	1	1	
306	1	1	1	1	1	1	0	0	1	1	1	
307	1	1	1	1	0	0	0	0	1	1	1	
308	1	1	1	1	0	0	0	0	1	1	1	
309	1	1	1	1	0	0	0	0	1	1	1	
310	1	1	1	1	0	0	0	0	1	1	1	
311	1	1	1	1	0	0	0	0	1	1	1	
312	1	1	1	1	0	0	0	0	1	1	1	

INDIVIDUAL DATA 2-1-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Pre-administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
401	1	1	1	1	1	0	0	0	1	1	1		
402	1	1	1	1	0	0	0	0	1	1	1		
403	1	1	1	1	0	0	0	0	1	1	1		
404	1	1	1	1	0	0	0	0	1	1	1		
405	1	1	1	1	0	0	0	0	1	1	1		
406	1	1	1	1	0	0	0	0	1	1	1		
407	1	1	1	1	0	0	0	0	1	1	1		
408	1	1	1	1	1	1	0	0	1	1	1		
409	1	1	1	1	0	0	0	0	1	1	1		
410	1	1	1	1	0	0	0	0	1	1	1		
411	1	1	1	1	0	0	0	0	1	1	1		
412	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12

INDIVIDUAL DATA 2-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Pre-administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
151	1	1	1	0	0	1	
152	1	1	1	0	0	1	
153	1	1	1	0	0	1	
154	1	1	1	0	0	1	
155	1	1	1	0	0	1	
156	1	1	1	0	0	1	
157	1	1	1	0	0	1	
158	1	1	1	0	0	1	
159	1	1	1	0	0	1	
160	1	1	1	0	0	1	
161	1	1	1	0	0	1	
162	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Pre-administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
251	1	1	1	0	0	1	
252	1	1	1	0	0	1	
253	1	1	1	0	0	1	
254	1	1	1	0	0	1	
255	1	1	1	0	0	1	
256	1	1	1	0	0	1	
257	1	1	1	0	0	1	
258	1	1	1	0	0	1	
259	1	1	1	0	0	1	
260	1	1	1	0	0	1	
261	1	1	1	0	0	1	
262	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-2-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Pre-administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
351	1	1	1	0	0	1	
352	1	1	1	0	0	1	
353	1	1	1	0	0	1	
354	1	1	1	0	0	1	
355	1	1	1	0	0	1	
356	1	1	1	0	0	1	
357	1	1	1	0	0	1	
358	1	1	1	0	0	1	
359	1	1	1	0	0	1	
360	1	1	1	0	0	1	
361	1	1	1	0	0	1	
362	1	1	1	0	0	1	

N

12

12

12

12

12

12

INDIVIDUAL DATA 2-2-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Pre-administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
451	1	1	1	0	0	1	
452	1	1	1	0	0	1	
453	1	1	1	0	0	1	
454	1	1	1	0	0	1	
455	1	1	1	0	0	1	
456	1	1	1	0	0	1	
457	1	1	1	0	0	1	
458	1	1	1	0	0	1	
459	1	1	1	0	0	1	
460	1	1	1	0	0	1	
461	1	1	1	0	0	1	
462	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-2-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Pre-administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
151	1	1	2	1	1	1	1	0	1	1	1	1	1	0
152	1	1	2	1	1	1	1	0	1	1	1	1	1	0
153	1	1	2	1	1	1	1	0	1	1	1	1	1	0
154	1	1	2	1	1	1	1	0	1	1	1	1	1	0
155	1	1	2	1	1	1	1	0	1	1	1	1	1	0
156	1	1	2	1	1	1	1	0	1	1	1	1	1	0
157	1	1	2	1	1	1	1	0	1	1	1	1	1	0
158	1	1	2	1	1	1	1	0	1	1	1	1	1	0
159	1	1	2	1	1	1	1	0	1	1	1	1	1	0
160	1	1	2	1	1	1	1	0	1	1	1	1	1	0
161	1	1	2	1	1	1	1	0	1	1	1	1	1	0
162	1	1	2	1	1	1	1	0	1	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-2-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Pre-administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
251	1	1	2	1	1	1	1	0	1	1	1	1	1	0
252	1	1	2	1	1	1	1	0	1	1	1	1	1	0
253	1	1	2	1	1	1	1	0	1	1	1	1	1	0
254	1	1	2	1	1	1	1	0	1	1	1	1	1	0
255	1	1	2	1	1	1	1	0	1	1	1	1	1	0
256	1	1	2	1	1	1	1	0	1	1	1	1	1	0
257	1	1	2	1	1	1	1	0	1	1	1	1	1	0
258	1	1	2	1	1	1	1	0	1	1	1	1	1	0
259	1	1	2	1	1	1	1	0	1	1	1	1	1	0
260	1	1	2	1	1	1	1	0	1	1	1	1	1	0
261	1	1	2	1	1	1	1	0	1	1	1	1	1	0
262	1	1	2	1	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-2-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Pre-administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
351	1	1	2	1	1	1	1	0	1	1	1	1	1	0
352	1	1	2	1	1	1	1	0	1	1	1	1	1	0
353	1	1	2	1	1	1	1	0	1	1	1	1	1	0
354	1	1	2	1	1	1	1	0	1	1	1	1	1	0
355	1	1	2	1	1	1	1	0	1	1	1	1	1	0
356	1	1	2	1	1	1	1	0	1	1	1	1	1	0
357	1	1	2	1	1	1	1	0	1	1	1	1	1	0
358	1	1	2	1	1	1	1	0	1	1	1	1	1	0
359	1	1	2	1	1	1	1	0	1	1	1	1	1	0
360	1	1	2	1	1	1	1	0	1	1	1	1	1	0
361	1	1	2	1	1	1	1	0	1	1	1	1	1	0
362	1	1	2	1	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-2-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Pre-administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling				Fur	Eyes				Pupil size	Salivation	
451	1	1	2	1	1	1	0	1	1	1	1	0
452	1	1	2	1	1	1	0	1	1	1	1	0
453	1	1	2	1	1	1	0	1	1	1	1	0
454	1	1	2	1	1	1	0	1	1	1	1	0
455	1	1	2	1	1	1	0	1	1	1	1	0
456	1	1	2	1	1	1	0	1	1	1	1	0
457	1	1	2	1	1	1	0	1	1	1	1	0
458	1	1	2	1	1	1	0	1	1	1	1	0
459	1	1	2	1	1	1	0	1	1	1	1	0
460	1	1	2	1	1	1	0	1	1	1	1	0
461	1	1	2	1	1	1	0	1	1	1	1	0
462	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-2-9

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Pre-administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
151	1	1	1	1	0	0	0	0	1	1	1	1	
152	1	1	1	1	0	0	0	0	1	1	1	1	
153	1	1	1	1	0	0	0	0	1	1	1	1	
154	1	1	1	1	1	0	0	0	1	1	1	1	
155	1	1	1	1	0	0	0	0	1	1	1	1	
156	1	1	1	1	0	0	0	0	1	1	1	1	
157	1	1	1	1	0	0	0	0	1	1	1	1	
158	1	1	1	1	0	0	0	0	1	1	1	1	
159	1	1	1	1	0	0	0	0	1	1	1	1	
160	1	1	1	1	0	0	0	0	1	1	1	1	
161	1	1	1	1	1	0	0	0	1	1	1	1	
162	1	1	1	1	0	0	0	0	1	1	1	1	
N	12	12	12	12	12	12	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	1.4729	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-2-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Pre-administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
251	1	1	1	1	0	0	0	0	1	1	1	
252	1	1	1	1	1	0	0	0	1	1	1	
253	1	1	1	1	0	0	0	0	1	1	1	
254	1	1	1	1	0	0	0	0	1	1	1	
255	1	1	1	1	0	0	0	0	1	1	1	
256	1	1	1	1	1	0	0	0	1	1	1	
257	1	1	1	1	0	0	0	0	1	1	1	
258	1	1	1	1	0	0	0	0	1	1	1	
259	1	1	1	1	0	0	0	0	1	1	1	
260	1	1	1	1	0	0	0	0	1	1	1	
261	1	1	1	1	1	0	0	0	1	1	1	
262	1	1	1	1	0	0	0	0	1	1	1	

N	12	12	12	12	12	12	12	12	12	12	12	12
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INDIVIDUAL DATA 2-2-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Pre-administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
351	1	1	1	1	1	0	0	0	1	1	1		
352	1	1	1	1	0	0	0	0	1	1	1		
353	1	1	1	1	0	0	0	0	1	1	1		
354	1	1	1	1	0	0	0	0	1	1	1		
355	1	1	1	1	0	0	0	0	1	1	1		
356	1	1	1	1	0	0	0	0	1	1	1		
357	1	1	1	1	0	0	0	0	1	1	1		
358	1	1	1	1	0	0	0	0	1	1	1		
359	1	1	1	1	0	0	0	0	1	1	1		
360	1	1	1	1	0	0	0	0	1	1	1		
361	1	1	1	1	0	0	0	0	1	1	1		
362	1	1	1	1	0	0	0	0	1	1	1		

INDIVIDUAL DATA 2-2-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Pre-administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
451	1	1	1	1	0	0	0	0	1	1	1	
452	1	1	1	1	0	0	0	0	1	1	1	
453	1	1	1	1	0	0	0	0	1	1	1	
454	1	1	1	1	1	0	0	0	1	1	1	
455	1	1	1	1	0	0	0	0	1	1	1	
456	1	1	1	1	0	0	0	0	1	1	1	
457	1	1	1	1	1	0	0	0	1	1	1	
458	1	1	1	1	0	0	0	0	1	1	1	
459	1	1	1	1	0	0	0	0	1	1	1	
460	1	1	1	1	1	0	0	0	1	1	1	
461	1	1	1	1	0	0	0	0	1	1	1	
462	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-2-13

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Pre-administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
163	1	1	1	0	0	1
164	1	1	1	0	0	1
165	1	1	1	0	0	1
166	1	1	1	0	0	1
167	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-2-14

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Pre-administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
463	1	1	1	0	0	1
464	1	1	1	0	0	1
465	1	1	1	0	0	1
466	1	1	1	0	0	1
467	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-2-15

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Pre-administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
163	1	1	2	1	1	1	0	1	1	1	1
164	1	1	2	1	1	1	0	1	1	1	1
165	1	1	2	1	1	1	0	1	1	1	1
166	1	1	2	1	1	1	0	1	1	1	1
167	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-2-16

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Pre-administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
463	1	1	2	1	1	1	0	1	1	1	1
464	1	1	2	1	1	1	0	1	1	1	1
465	1	1	2	1	1	1	0	1	1	1	1
466	1	1	2	1	1	1	0	1	1	1	1
467	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-2-17

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Pre-administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
163	1	1	1	1	0	0	0	0	1	1	1		
164	1	1	1	1	0	0	0	0	1	1	1		
165	1	1	1	1	0	0	0	0	1	1	1		
166	1	1	1	1	0	0	0	0	1	1	1		
167	1	1	1	1	0	0	0	0	1	1	1		
N	5	5	5	5	5	5	5	5	5	5	5		
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000		

INDIVIDUAL DATA 2-2-18

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Pre-administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
463	1	1	1	1	0	0	0	0	1	1	1	
464	1	1	1	1	0	0	0	0	1	1	1	
465	1	1	1	1	0	0	0	0	1	1	1	
466	1	1	1	1	0	0	0	0	1	1	1	
467	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	

INDIVIDUAL DATA 2-3-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 7 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
101	1	1	1	0	0	1	
102	1	1	1	0	0	1	
103	1	1	1	0	0	1	
104	1	1	1	0	0	1	
105	1	1	1	0	0	1	
106	1	1	1	0	0	1	
107	1	1	1	0	0	1	
108	1	1	1	0	0	1	
109	1	1	1	0	0	1	
110	1	1	1	0	0	1	
111	1	1	1	0	0	1	
112	1	1	1	0	0	1	
<hr/>							
N	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 7 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
201	1	1	1	0	0	1	
202	1	1	1	0	0	1	
203	1	1	1	0	0	1	
204	1	1	1	0	0	1	
205	1	1	1	0	0	1	
206	1	1	1	0	0	1	
207	1	1	1	0	0	1	
208	1	1	1	0	0	1	
209	1	1	1	0	0	1	
210	1	1	1	0	0	1	
211	1	1	1	0	0	1	
212	1	1	1	0	0	1	

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INDIVIDUAL DATA 2-3-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 7 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
301	1	1	1	0	0	1	
302	1	1	1	0	0	1	
303	1	1	1	0	0	1	
304	1	1	1	0	0	1	
305	1	1	1	0	0	1	
306	1	1	1	0	0	1	
307	1	1	1	0	0	1	
308	1	1	1	0	0	1	
309	1	1	1	0	0	1	
310	1	1	1	0	0	1	
311	1	1	1	0	0	1	
312	1	1	1	0	0	1	

N

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INDIVIDUAL DATA 2-3-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 7 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
401	1	1	1	0	0	1	
402	1	1	1	0	0	1	
403	1	1	1	0	0	1	
404	1	1	1	0	0	1	
405	1	1	1	0	0	1	
406	1	1	1	0	0	1	
407	1	1	1	0	0	1	
408	1	1	1	0	0	1	
409	1	1	1	0	0	1	
410	1	1	1	0	0	1	
411	1	1	1	0	0	1	
412	1	1	1	0	0	1	

N

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INDIVIDUAL DATA 2-3-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 7 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
101	1	1	2	1	1	1	1	0	1	1	1	1	1	0
102	1	1	2	1	1	1	1	0	1	1	1	1	1	0
103	1	1	2	1	1	1	1	0	1	1	1	1	1	0
104	1	1	2	1	1	1	1	0	1	1	1	1	1	0
105	1	1	2	1	1	1	1	0	1	1	1	1	1	0
106	1	1	2	1	1	1	1	0	1	1	1	1	1	0
107	1	1	2	1	1	1	1	0	1	1	1	1	1	0
108	1	1	2	1	1	1	1	0	1	1	1	1	1	0
109	1	1	2	1	1	1	1	0	1	1	1	1	1	0
110	1	1	2	1	1	1	1	0	1	1	1	1	1	0
111	1	1	2	1	1	1	1	0	1	1	1	1	1	0
112	1	1	2	1	1	1	1	0	1	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-3-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 7 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
201	1	1	2	1	1	1	1	1	0	1	1	1	1	0
202	1	1	2	1	1	1	1	0	1	1	1	1	1	0
203	1	1	2	1	1	1	1	0	1	1	1	1	1	0
204	1	1	2	1	1	1	1	0	1	1	1	1	1	0
205	1	1	2	1	1	1	1	0	1	1	1	1	1	0
206	1	1	2	1	1	1	1	0	1	1	1	1	1	0
207	1	1	2	1	1	1	1	0	1	1	1	1	1	0
208	1	1	2	1	1	1	1	0	1	1	1	1	1	0
209	1	1	2	1	1	1	1	0	1	1	1	1	1	0
210	1	1	2	1	1	1	1	0	1	1	1	1	1	0
211	1	1	2	1	1	1	1	0	1	1	1	1	1	0
212	1	1	2	1	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-3-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 7 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
301	1	1	2	1	1	1	1	1	0	1	1	1	1	0
302	1	1	2	1	1	1	1	0	1	1	1	1	1	0
303	1	1	2	1	1	1	1	0	1	1	1	1	1	0
304	1	1	2	1	1	1	1	0	1	1	1	1	1	0
305	1	1	2	1	1	1	1	0	1	1	1	1	1	0
306	1	1	2	1	1	1	1	0	1	1	1	1	1	0
307	1	1	2	1	1	1	1	0	1	1	1	1	1	0
308	1	1	2	1	1	1	1	0	1	1	1	1	1	0
309	1	1	2	1	1	1	1	0	1	1	1	1	1	0
310	1	1	2	1	1	1	1	0	1	1	1	1	1	0
311	1	1	2	1	1	1	1	0	1	1	1	1	1	0
312	1	1	2	1	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-3-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 7 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Removal	Handling			Fur	Eyes			Pupil size	Salivation		
401	1	1	2	1	1	1	0	1	1	1	1	0
402	1	1	2	1	1	1	0	1	1	1	1	0
403	1	1	2	1	1	1	0	1	1	1	1	0
404	1	1	2	1	1	1	0	1	1	1	1	0
405	1	1	2	1	1	1	0	1	1	1	1	0
406	1	1	2	1	1	1	0	1	1	1	1	0
407	1	1	2	1	1	1	0	1	1	1	1	0
408	1	1	2	1	1	1	0	1	1	1	1	0
409	1	1	2	1	1	1	0	1	1	1	1	0
410	1	1	2	1	1	1	0	1	1	1	1	0
411	1	1	2	1	1	1	0	1	1	1	1	0
412	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-3-9

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 7 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli				Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
				Searching	Urination	Defecation							
101	1	1	1	1	0	0	0	0	1	1	1	1	
102	1	1	1	1	0	0	0	0	1	1	1	1	
103	1	1	1	1	0	0	0	0	1	1	1	1	
104	1	1	1	1	0	0	0	0	1	1	1	1	
105	1	1	1	1	0	0	0	0	1	1	1	1	
106	1	1	1	1	0	0	0	0	1	1	1	1	
107	1	1	1	1	1	1	0	0	1	1	1	1	
108	1	1	1	1	0	0	0	0	1	1	1	1	
109	1	1	1	1	0	0	0	0	1	1	1	1	
110	1	1	1	1	0	0	0	0	1	1	1	1	
111	1	1	1	1	0	0	0	0	1	1	1	1	
112	1	1	1	1	0	0	0	0	1	1	1	1	
N	12	12	12	12	12	12	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	2.1364	4.1535	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-3-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 7 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
201	1	1	1	1	0	0	0	0	1	1	1		
202	1	1	1	1	0	0	0	0	1	1	1		
203	1	1	1	1	0	0	0	0	1	1	1		
204	1	1	1	1	1	0	0	0	1	1	1		
205	1	1	1	1	0	0	0	0	1	1	1		
206	1	1	1	1	0	0	0	0	1	1	1		
207	1	1	1	1	1	1	0	0	1	1	1		
208	1	1	1	1	0	0	0	0	1	1	1		
209	1	1	1	1	0	0	0	0	1	1	1		
210	1	1	1	1	0	0	0	0	1	1	1		
211	1	1	1	1	0	0	0	0	1	1	1		
212	1	1	1	1	0	0	0	0	1	1	1		

N	12	12	12	12	12	12	12	12	12	12	12	12	12
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INDIVIDUAL DATA 2-3-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 7 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
301	1	1	1	1	0	0	0	0	1	1	1		
302	1	1	1	1	0	0	0	0	1	1	1		
303	1	1	1	1	0	0	0	0	1	1	1		
304	1	1	1	1	0	0	0	0	1	1	1		
305	1	1	1	1	0	0	0	0	1	1	1		
306	1	1	1	1	0	0	0	0	1	1	1		
307	1	1	1	1	0	0	0	0	1	1	1		
308	1	1	1	1	0	0	0	0	1	1	1		
309	1	1	1	1	0	0	0	0	1	1	1		
310	1	1	1	1	0	0	0	0	1	1	1		
311	1	1	1	1	0	0	0	0	1	1	1		
312	1	1	1	1	0	0	0	0	1	1	1		

N 12 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-3-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 7 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
401	1	1	1	1	0	0	0	0	1	1	1		
402	1	1	1	1	0	1	0	0	1	1	1		
403	1	1	1	1	0	0	0	0	1	1	1		
404	1	1	1	1	0	0	0	0	1	1	1		
405	1	1	1	1	0	0	0	0	1	1	1		
406	1	1	1	1	0	0	0	0	1	1	1		
407	1	1	1	1	0	0	0	0	1	1	1		
408	1	1	1	1	1	1	0	0	1	1	1		
409	1	1	1	1	0	0	0	0	1	1	1		
410	1	1	1	1	0	0	0	0	1	1	1		
411	1	1	1	1	0	0	0	0	1	1	1		
412	1	1	1	1	0	1	0	0	1	1	1		

N 12 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-4-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 7 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
151	1	1	1	0	0	1	
152	1	1	1	0	0	1	
153	1	1	1	0	0	1	
154	1	1	1	0	0	1	
155	1	1	1	0	0	1	
156	1	1	1	0	0	1	
157	1	1	1	0	0	1	
158	1	1	1	0	0	1	
159	1	1	1	0	0	1	
160	1	1	1	0	0	1	
161	1	1	1	0	0	1	
162	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 7 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
251	1	1	1	0	0	1	
252	1	1	1	0	0	1	
253	1	1	1	0	0	1	
254	1	1	1	0	0	1	
255	1	1	1	0	0	1	
256	1	1	1	0	0	1	
257	1	1	1	0	0	1	
258	1	1	1	0	0	1	
259	1	1	1	0	0	1	
260	1	1	1	0	0	1	
261	1	1	1	0	0	1	
262	1	1	1	0	0	1	

N

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INDIVIDUAL DATA 2-4-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 7 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
351	1	1	1	0	0	1	
352	1	1	1	0	0	1	
353	1	1	1	0	0	1	
354	1	1	1	0	0	1	
355	1	1	1	0	0	1	
356	1	1	1	0	0	1	
357	1	1	1	0	0	1	
358	1	1	1	0	0	1	
359	1	1	1	0	0	1	
360	1	1	1	0	0	1	
361	1	1	1	0	0	1	
362	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-4-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 7 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
451	1	1	1	0	0	1	
452	1	1	1	0	0	1	
453	1	1	1	0	0	1	
454	1	1	1	0	0	1	
455	1	1	1	0	0	1	
456	1	1	1	0	0	1	
457	1	1	1	0	0	1	
458	1	1	1	0	0	1	
459	1	1	1	0	0	1	
460	1	1	1	0	0	1	
461	1	1	1	0	0	1	
462	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-4-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 7 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling		Fur	Eyes					Pupil size	Salivation		
151	1	1	2	1	1	1	0	1	1	1	1	0
152	1	1	2	1	1	1	0	1	1	1	1	0
153	1	1	2	1	1	1	0	1	1	1	1	0
154	1	1	2	1	1	1	0	1	1	1	1	0
155	1	1	2	1	1	1	0	1	1	1	1	0
156	1	1	2	1	1	1	0	1	1	1	1	0
157	1	1	2	1	1	1	0	1	1	1	1	0
158	1	1	2	1	1	1	0	1	1	1	1	0
159	1	1	2	1	1	1	0	1	1	1	1	0
160	1	1	2	1	1	1	0	1	1	1	1	0
161	1	1	2	1	1	1	0	1	1	1	1	0
162	1	1	2	1	1	1	0	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-4-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 7 of administration

Animal No.	On the hand												
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
251	1	1	2	1	1	1	0	1	1	1	1	1	0
252	1	1	2	1	1	1	0	1	1	1	1	1	0
253	1	1	2	1	1	1	0	1	1	1	1	1	0
254	1	1	2	1	1	1	0	1	1	1	1	1	0
255	1	1	2	1	1	1	0	1	1	1	1	1	0
256	1	1	2	1	1	1	0	1	1	1	1	1	0
257	1	1	2	1	1	1	0	1	1	1	1	1	0
258	1	1	2	1	1	1	0	1	1	1	1	1	0
259	1	1	2	1	1	1	0	1	1	1	1	1	0
260	1	1	2	1	1	1	0	1	1	1	1	1	0
261	1	1	2	1	1	1	0	1	1	1	1	1	0
262	1	1	2	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-4-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 7 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
351	1	1	2	1	1	1	0	1	1	1	1	0
352	1	1	2	1	1	1	0	1	1	1	1	0
353	1	1	2	1	1	1	0	1	1	1	1	0
354	1	1	2	1	1	1	0	1	1	1	1	0
355	1	1	2	1	1	1	0	1	1	1	1	0
356	1	1	2	1	1	1	0	1	1	1	1	0
357	1	1	2	1	1	1	0	1	1	1	1	0
358	1	1	2	1	1	1	0	1	1	1	1	0
359	1	1	2	1	1	1	0	1	1	1	1	0
360	1	1	2	1	1	1	0	1	1	1	1	0
361	1	1	2	1	1	1	0	1	1	1	1	0
362	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-4-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 7 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
451	1	1	2	1	1	1	0	1	1	1	1
452	1	1	2	1	1	1	0	1	1	1	1
453	1	1	2	1	1	1	0	1	1	1	1
454	1	1	2	1	1	1	0	1	1	1	1
455	1	1	2	1	1	1	0	1	1	1	1
456	1	1	2	1	1	1	0	1	1	1	1
457	1	1	2	1	1	1	0	1	1	1	1
458	1	1	2	1	1	1	0	1	1	1	1
459	1	1	2	1	1	1	0	1	1	1	1
460	1	1	2	1	1	1	0	1	1	1	1
461	1	1	2	1	1	1	0	1	1	1	1
462	1	1	2	1	1	1	0	1	1	1	1

INDIVIDUAL DATA 2-4-9

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 7 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
151	1	1	1	1	0	0	0	0	1	1	1		
152	1	1	1	1	0	0	0	0	1	1	1		
153	1	1	1	1	0	0	0	0	1	1	1		
154	1	1	1	1	0	0	0	0	1	1	1		
155	1	1	1	1	0	0	0	0	1	1	1		
156	1	1	1	1	0	0	0	0	1	1	1		
157	1	1	1	1	0	0	0	0	1	1	1		
158	1	1	1	1	0	0	0	0	1	1	1		
159	1	1	1	1	0	0	0	0	1	1	1		
160	1	1	1	1	0	0	0	0	1	1	1		
161	1	1	1	1	0	0	0	0	1	1	1		
162	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	2.1364	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-4-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 7 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
251	1	1	1	1	0	0	0	0	1	1	1	
252	1	1	1	1	0	0	0	0	1	1	1	
253	1	1	1	1	1	0	0	0	1	1	1	
254	1	1	1	1	1	0	0	0	1	1	1	
255	1	1	1	1	0	0	0	0	1	1	1	
256	1	1	1	1	0	0	0	0	1	1	1	
257	1	1	1	1	0	0	0	0	1	1	1	
258	1	1	1	1	0	0	0	0	1	1	1	
259	1	1	1	1	0	0	0	0	1	1	1	
260	1	1	1	1	0	0	0	0	1	1	1	
261	1	1	1	1	0	0	0	0	1	1	1	
262	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-4-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 7 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
351	1	1	1	1	0	0	0	0	1	1	1	
352	1	1	1	1	0	0	0	0	1	1	1	
353	1	1	1	1	0	0	0	0	1	1	1	
354	1	1	1	1	0	0	0	0	1	1	1	
355	1	1	1	1	0	0	0	0	1	1	1	
356	1	1	1	1	0	0	0	0	1	1	1	
357	1	1	1	1	0	0	0	0	1	1	1	
358	1	1	1	1	0	0	0	0	1	1	1	
359	1	1	1	1	1	0	0	0	1	1	1	
360	1	1	1	1	0	0	0	0	1	1	1	
361	1	1	1	1	0	0	0	0	1	1	1	
362	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-4-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 7 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
451	1	1	1	1	0	0	0	0	1	1	1	
452	1	1	1	1	0	0	0	0	1	1	1	
453	1	1	1	1	0	0	0	0	1	1	1	
454	1	1	1	1	1	0	0	0	1	1	1	
455	1	1	1	1	0	0	0	0	1	1	1	
456	1	1	1	1	0	0	0	0	1	1	1	
457	1	1	1	1	0	0	0	0	1	1	1	
458	1	1	1	1	0	0	0	0	1	1	1	
459	1	1	1	1	0	0	0	0	1	1	1	
460	1	1	1	1	0	0	0	0	1	1	1	
461	1	1	1	1	0	0	0	0	1	1	1	
462	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-4-13

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 7 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
163	1	1	1	0	0	1
164	1	1	1	0	0	1
165	1	1	1	0	0	1
166	1	1	1	0	0	1
167	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-4-14

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 7 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
463	1	1	1	0	0	1
464	1	1	1	0	0	1
465	1	1	1	0	0	1
466	1	1	1	0	0	1
467	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-4-15

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 7 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
163	1	1	2	1	1	1	0	1	1	1	1
164	1	1	2	1	1	1	0	1	1	1	1
165	1	1	2	1	1	1	0	1	1	1	1
166	1	1	2	1	1	1	0	1	1	1	1
167	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-4-16

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 7 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
463	1	1	2	1	1	1	0	1	1	1	1
464	1	1	2	1	1	1	0	1	1	1	1
465	1	1	2	1	1	1	0	1	1	1	1
466	1	1	2	1	1	1	0	1	1	1	1
467	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-4-17

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 7 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
163	1	1	1	1	0	0	0	0	1	1	1		
164	1	1	1	1	0	0	0	0	1	1	1		
165	1	1	1	1	0	0	0	0	1	1	1		
166	1	1	1	1	0	0	0	0	1	1	1		
167	1	1	1	1	0	0	0	0	1	1	1		
N	5	5	5	5	5	5	5	5	5	5	5		
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000		

INDIVIDUAL DATA 2-4-18

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 7 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
463	1	1	1	1	0	0	0	0	1	1	1	
464	1	1	1	1	0	0	0	0	1	1	1	
465	1	1	1	1	0	0	0	0	1	1	1	
466	1	1	1	1	0	0	0	0	1	1	1	
467	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	

INDIVIDUAL DATA 2-5-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 14 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
101	1	1	1	0	0	1	
102	1	1	1	0	0	1	
103	1	1	1	0	0	1	
104	1	1	1	0	0	1	
105	1	1	1	0	0	1	
106	1	1	1	0	0	1	
107	1	1	1	0	0	1	
108	1	1	1	0	0	1	
109	1	1	1	0	0	1	
110	1	1	1	0	0	1	
111	1	1	1	0	0	1	
112	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-5-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 14 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
201	1	1	1	0	0	1	
202	1	1	1	0	0	1	
203	1	1	1	0	0	1	
204	1	1	1	0	0	1	
205	1	1	1	0	0	1	
206	1	1	1	0	0	1	
207	1	1	1	0	0	1	
208	1	1	1	0	0	1	
209	1	1	1	0	0	1	
210	1	1	1	0	0	1	
211	1	1	1	0	0	1	
212	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-5-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 14 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
301	1	1	1	0	0	1	
302	1	1	1	0	0	1	
303	1	1	1	0	0	1	
304	1	1	1	0	0	1	
305	1	1	1	0	0	1	
306	1	1	1	0	0	1	
307	1	1	1	0	0	1	
308	1	1	1	0	0	1	
309	1	1	1	0	0	1	
310	1	1	1	0	0	1	
311	1	1	1	0	0	1	
312	1	1	1	0	0	1	

N

12

12

12

12

12

12

INDIVIDUAL DATA 2-5-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 14 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
401	1	1	1	0	0	1	
402	1	1	1	0	0	1	
403	1	1	1	0	0	1	
404	1	1	1	0	0	1	
405	1	1	1	0	0	1	
406	1	1	1	0	0	1	
407	1	1	1	0	0	1	
408	1	1	1	0	0	1	
409	1	1	1	0	0	1	
410	1	1	1	0	0	1	
411	1	1	1	0	0	1	
412	1	1	1	0	0	1	

N

12

12

12

12

12

12

INDIVIDUAL DATA 2-5-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 14 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling		Fur	Eyes					Pupil size	Salivation		
101	1	1	2	1	1	1	0	1	1	1	1	0
102	1	1	2	1	1	1	0	1	1	1	1	0
103	1	1	2	1	1	1	0	1	1	1	1	0
104	1	1	2	1	1	1	0	1	1	1	1	0
105	1	1	2	1	1	1	0	1	1	1	1	0
106	1	1	2	1	1	1	0	1	1	1	1	0
107	1	1	2	1	1	1	0	1	1	1	1	0
108	1	1	2	1	1	1	0	1	1	1	1	0
109	1	1	2	1	1	1	0	1	1	1	1	0
110	1	1	2	1	1	1	0	1	1	1	1	0
111	1	1	2	1	1	1	0	1	1	1	1	0
112	1	1	2	1	1	1	0	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-5-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 14 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling				Fur	Eyes			Pupil size		Salivation	
201	1	1	2	1	1	1	0	1	1	1	1	0
202	1	1	2	1	1	1	0	1	1	1	1	0
203	1	1	2	1	1	1	0	1	1	1	1	0
204	1	1	2	1	1	1	0	1	1	1	1	0
205	1	1	2	1	1	1	0	1	1	1	1	0
206	1	1	2	1	1	1	0	1	1	1	1	0
207	1	1	2	1	1	1	0	1	1	1	1	0
208	1	1	2	1	1	1	0	1	1	1	1	0
209	1	1	2	1	1	1	0	1	1	1	1	0
210	1	1	2	1	1	1	0	1	1	1	1	0
211	1	1	2	1	1	1	0	1	1	1	1	0
212	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-5-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 14 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
301	1	1	2	1	1	1	1	1	0	1	1	1	1	0
302	1	1	2	1	1	1	1	0	1	1	1	1	1	0
303	1	1	2	1	1	1	1	0	1	1	1	1	1	0
304	1	1	2	1	1	1	1	0	1	1	1	1	1	0
305	1	1	2	1	1	1	1	0	1	1	1	1	1	0
306	1	1	2	1	1	1	1	0	1	1	1	1	1	0
307	1	1	2	1	1	1	1	0	1	1	1	1	1	0
308	1	1	2	1	1	1	1	0	1	1	1	1	1	0
309	1	1	2	1	1	1	1	0	1	1	1	1	1	0
310	1	1	2	1	1	1	1	0	1	1	1	1	1	0
311	1	1	2	1	1	1	1	0	1	1	1	1	1	0
312	1	1	2	1	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-5-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 14 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Removal	Handling			Fur	Eyes			Pupil size	Salivation		
401	1	1	2	1	1	1	0	1	1	1	1	0
402	1	1	2	1	1	1	0	1	1	1	1	0
403	1	1	2	1	1	1	0	1	1	1	1	0
404	1	1	2	1	1	1	0	1	1	1	1	0
405	1	1	2	1	1	1	0	1	1	1	1	0
406	1	1	2	1	1	1	0	1	1	1	1	0
407	1	1	2	1	1	1	0	1	1	1	1	0
408	1	1	2	1	1	1	0	1	1	1	1	0
409	1	1	2	1	1	1	0	1	1	1	1	0
410	1	1	2	1	1	1	0	1	1	1	1	0
411	1	1	2	1	1	1	0	1	1	1	1	0
412	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-5-9

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 14 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
101	1	1	1	1	0	0	0	0	1	1	1		
102	1	1	1	1	0	0	0	0	1	1	1		
103	1	1	1	1	0	0	0	0	1	1	1		
104	1	1	1	1	0	0	0	0	1	1	1		
105	1	1	1	1	0	0	0	0	1	1	1		
106	1	1	1	1	0	0	0	0	1	1	1		
107	1	1	1	1	1	1	0	0	1	1	1		
108	1	1	1	1	0	0	0	0	1	1	1		
109	1	1	1	1	0	0	0	0	1	1	1		
110	1	1	1	1	0	0	0	0	1	1	1		
111	1	1	1	1	0	0	0	0	1	1	1		
112	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	2.0435	2.0435	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-5-10

STUDY NO. SR07125 TITLE : PFHxD_A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 14 of administration

Animal No.	In the open-field							Stereotype			Bizarre behavior		
	Co-ordination of movement		Reactivity to environmental stimuli		Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
	Gait												
201	1	1	1	1	0	0	0	0	0	1	1	1	
202	1	1	1	1	0	0	0	0	0	1	1	1	
203	1	1	1	1	0	0	0	0	0	1	1	1	
204	1	1	1	1	0	0	0	0	0	1	1	1	
205	1	1	1	1	0	0	0	0	0	1	1	1	
206	1	1	1	1	0	0	0	0	0	1	1	1	
207	1	1	1	1	0	0	0	0	0	1	1	1	
208	1	1	1	1	0	0	0	0	0	1	1	1	
209	1	1	1	1	0	0	0	0	0	1	1	1	
210	1	1	1	1	0	0	0	0	0	1	1	1	
211	1	1	1	1	0	0	0	0	0	1	1	1	
212	1	1	1	1	0	0	0	0	0	1	1	1	

INDIVIDUAL DATA 2-5-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 14 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
301	1	1	1	1	0	0	0	0	1	1	1	1	
302	1	1	1	1	0	0	0	0	1	1	1	1	
303	1	1	1	1	0	0	0	0	1	1	1	1	
304	1	1	1	1	0	1	0	0	1	1	1	1	
305	1	1	1	1	0	0	0	0	1	1	1	1	
306	1	1	1	1	1	0	0	0	1	1	1	1	
307	1	1	1	1	0	0	0	0	1	1	1	1	
308	1	1	1	1	0	0	0	0	1	1	1	1	
309	1	1	1	1	0	0	0	0	1	1	1	1	
310	1	1	1	1	0	0	0	0	1	1	1	1	
311	1	1	1	1	0	0	0	0	1	1	1	1	
312	1	1	1	1	0	0	0	0	1	1	1	1	
N	12	12	12	12	12	12	12	12	12	12	12	12	

INDIVIDUAL DATA 2-5-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 14 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
401	1	1	1	1	0	0	0	0	1	1	1	1	
402	1	1	1	1	0	0	0	0	1	1	1	1	
403	1	1	1	1	0	0	0	0	1	1	1	1	
404	1	1	1	1	0	0	0	0	1	1	1	1	
405	1	1	1	1	0	0	0	0	1	1	1	1	
406	1	1	1	1	0	0	0	0	1	1	1	1	
407	1	1	1	1	0	0	0	0	1	1	1	1	
408	1	1	1	1	0	0	0	0	1	1	1	1	
409	1	1	1	1	0	0	0	0	1	1	1	1	
410	1	1	1	1	0	0	0	0	1	1	1	1	
411	1	1	1	1	0	0	0	0	1	1	1	1	
412	1	1	1	1	0	0	0	0	1	1	1	1	
N	12	12	12	12	12	12	12	12	12	12	12	12	

INDIVIDUAL DATA 2-6-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 14 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
151	1	1	1	0	0	1	
152	1	1	1	0	0	1	
153	1	1	1	0	0	1	
154	1	1	1	0	0	1	
155	1	1	1	0	0	1	
156	1	1	1	0	0	1	
157	1	1	1	0	0	1	
158	1	1	1	0	0	1	
159	1	1	1	0	0	1	
160	1	1	1	0	0	1	
161	1	1	1	0	0	1	
162	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-6-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 14 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
251	1	1	1	0	0	1	
252	1	1	1	0	0	1	
253	1	1	1	0	0	1	
254	1	1	1	0	0	1	
255	1	1	1	0	0	1	
256	1	1	1	0	0	1	
257	1	1	1	0	0	1	
258	1	1	1	0	0	1	
259	1	1	1	0	0	1	
260	1	1	1	0	0	1	
261	1	1	1	0	0	1	
262	1	1	1	0	0	1	

N

12

12

12

12

12

12

INDIVIDUAL DATA 2-6-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 14 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
351	1	1	1	0	0	1	
352	1	1	1	0	0	1	
353	1	1	1	0	0	1	
354	1	1	1	0	0	1	
355	1	1	1	0	0	1	
356	1	1	1	0	0	1	
357	1	1	1	0	0	1	
358	1	1	1	0	0	1	
359	1	1	1	0	0	1	
360	1	1	1	0	0	1	
361	1	1	1	0	0	1	
362	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-6-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 14 of administration

Animal No.	In the cage					
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
451	1	1	1	0	0	1
452	1	1	1	0	0	1
453	1	1	1	0	0	1
454	1	1	1	0	0	1
455	1	1	1	0	0	1
456	1	1	1	0	0	1
457	1	1	1	0	0	1
458	1	1	1	0	0	1
459	1	1	1	0	0	1
460	1	1	1	0	0	1
461	1	1	1	0	0	1
462	1	1	1	0	0	1

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-6-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 14 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
151	1	1	2	1	1	1	1	0	1	1	1	1	1	0
152	1	1	2	1	1	1	1	0	1	1	1	1	1	0
153	1	1	2	1	1	1	1	0	1	1	1	1	1	0
154	1	1	2	1	1	1	1	0	1	1	1	1	1	0
155	1	1	2	1	1	1	1	0	1	1	1	1	1	0
156	1	1	2	1	1	1	1	0	1	1	1	1	1	0
157	1	1	2	1	1	1	1	0	1	1	1	1	1	0
158	1	1	2	1	1	1	1	0	1	1	1	1	1	0
159	1	1	2	1	1	1	1	0	1	1	1	1	1	0
160	1	1	2	1	1	1	1	0	1	1	1	1	1	0
161	1	1	2	1	1	1	1	0	1	1	1	1	1	0
162	1	1	2	1	1	1	1	0	1	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-6-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 14 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
251	1	1	2	1	1	1	0	1	1	1	1	0
252	1	1	2	1	1	1	0	1	1	1	1	0
253	1	1	2	1	1	1	0	1	1	1	1	0
254	1	1	2	1	1	1	0	1	1	1	1	0
255	1	1	2	1	1	1	0	1	1	1	1	0
256	1	1	2	1	1	1	0	1	1	1	1	0
257	1	1	2	1	1	1	0	1	1	1	1	0
258	1	1	2	1	1	1	0	1	1	1	1	0
259	1	1	2	1	1	1	0	1	1	1	1	0
260	1	1	2	1	1	1	0	1	1	1	1	0
261	1	1	2	1	1	1	0	1	1	1	1	0
262	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-6-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 14 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
351	1	1	2	1	1	1	0	1	1	1	1	0
352	1	1	2	1	1	1	0	1	1	1	1	0
353	1	1	2	1	1	1	0	1	1	1	1	0
354	1	1	2	1	1	1	0	1	1	1	1	0
355	1	1	2	1	1	1	0	1	1	1	1	0
356	1	1	2	1	1	1	0	1	1	1	1	0
357	1	1	2	1	1	1	0	1	1	1	1	0
358	1	1	2	1	1	1	0	1	1	1	1	0
359	1	1	2	1	1	1	0	1	1	1	1	0
360	1	1	2	1	1	1	0	1	1	1	1	0
361	1	1	2	1	1	1	0	1	1	1	1	0
362	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-6-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 14 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
451	1	1	2	1	1	1	0	1	1	1	1
452	1	1	2	1	1	1	0	1	1	1	1
453	1	1	2	1	1	1	0	1	1	1	1
454	1	1	2	1	1	1	0	1	1	1	1
455	1	1	2	1	1	1	0	1	1	1	1
456	1	1	2	1	1	1	0	1	1	1	1
457	1	1	2	1	1	1	0	1	1	1	1
458	1	1	2	1	1	1	0	1	1	1	1
459	1	1	2	1	1	1	0	1	1	1	1
460	1	1	2	1	1	1	0	1	1	1	1
461	1	1	2	1	1	1	0	1	1	1	1
462	1	1	2	1	1	1	0	1	1	1	1

N 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-6-9

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 14 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
151	1	1	1	1	1	0	0	0	1	1	1	1	
152	1	1	1	1	0	0	0	0	1	1	1	1	
153	1	1	1	1	1	0	0	0	1	1	1	1	
154	1	1	1	1	0	0	0	0	1	1	1	1	
155	1	1	1	1	0	0	0	0	1	1	1	1	
156	1	1	1	1	0	0	0	0	1	1	1	1	
157	1	1	1	1	0	0	0	0	1	1	1	1	
158	1	1	1	1	0	0	0	0	1	1	1	1	
159	1	1	1	1	0	0	0	0	1	1	1	1	
160	1	1	1	1	0	0	0	0	1	1	1	1	
161	1	1	1	1	0	0	0	0	1	1	1	1	
162	1	1	1	1	0	0	0	0	1	1	1	1	
N	12	12	12	12	12	12	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.6558	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-6-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 14 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
251	1	1	1	1	1	0	0	0	1	1	1	
252	1	1	1	1	0	0	0	0	1	1	1	
253	1	1	1	1	0	0	0	0	1	1	1	
254	1	1	1	1	0	0	0	0	1	1	1	
255	1	1	1	1	0	0	0	0	1	1	1	
256	1	1	1	1	0	0	0	0	1	1	1	
257	1	1	1	1	0	0	0	0	1	1	1	
258	1	1	1	1	0	0	0	0	1	1	1	
259	1	1	1	1	0	0	0	0	1	1	1	
260	1	1	1	1	0	0	0	0	1	1	1	
261	1	1	1	1	0	0	0	0	1	1	1	
262	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-6-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 14 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
351	1	1	1	1	0	0	0	0	1	1	1	
352	1	1	1	1	0	0	0	0	1	1	1	
353	1	1	1	1	1	0	0	0	1	1	1	
354	1	1	1	1	0	0	0	0	1	1	1	
355	1	1	1	1	0	0	0	0	1	1	1	
356	1	1	1	1	0	0	0	0	1	1	1	
357	1	1	1	1	0	0	0	0	1	1	1	
358	1	1	1	1	0	0	0	0	1	1	1	
359	1	1	1	1	0	0	0	0	1	1	1	
360	1	1	1	1	0	0	0	0	1	1	1	
361	1	1	1	1	0	0	0	0	1	1	1	
362	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-6-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 14 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
451	1	1	1	1	0	0	0	0	1	1	1		
452	1	1	1	1	0	0	0	0	1	1	1		
453	1	1	1	1	0	0	0	0	1	1	1		
454	1	1	1	1	1	0	0	0	1	1	1		
455	1	1	1	1	0	0	0	0	1	1	1		
456	1	1	1	1	0	0	0	0	1	1	1		
457	1	1	1	1	0	0	0	0	1	1	1		
458	1	1	1	1	0	0	0	0	1	1	1		
459	1	1	1	1	0	0	0	0	1	1	1		
460	1	1	1	1	0	0	0	0	1	1	1		
461	1	1	1	1	0	0	0	0	1	1	1		
462	1	1	1	1	0	0	0	0	1	1	1		

N 12 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-6-13

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 14 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
163	1	1	1	0	0	1
164	1	1	1	0	0	1
165	1	1	1	0	0	1
166	1	1	1	0	0	1
167	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-6-14

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 14 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
463	1	1	1	0	0	1
464	1	1	1	0	0	1
465	1	1	1	0	0	1
466	1	1	1	0	0	1
467	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-6-15

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 14 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
163	1	1	2	1	1	1	0	1	1	1	1
164	1	1	2	1	1	1	0	1	1	1	1
165	1	1	2	1	1	1	0	1	1	1	1
166	1	1	2	1	1	1	0	1	1	1	1
167	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-6-16

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 14 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
463	1	1	2	1	1	1	0	1	1	1	1
464	1	1	2	1	1	1	0	1	1	1	1
465	1	1	2	1	1	1	0	1	1	1	1
466	1	1	2	1	1	1	0	1	1	1	1
467	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-6-17

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 14 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
163	1	1	1	1	0	0	0	0	1	1	1	
164	1	1	1	1	0	0	0	0	1	1	1	
165	1	1	1	1	0	0	0	0	1	1	1	
166	1	1	1	1	0	0	0	0	1	1	1	
167	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	
H	0.0000	0.0000	0.0000	0.0000	1.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-6-18

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 14 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
463	1	1	1	1	0	0	0	0	1	1	1	
464	1	1	1	1	1	0	0	0	1	1	1	
465	1	1	1	1	0	0	0	0	1	1	1	
466	1	1	1	1	0	0	0	0	1	1	1	
467	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	

INDIVIDUAL DATA 2-7-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 21 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
101	1	1	1	0	0	1	
102	1	1	1	0	0	1	
103	1	1	1	0	0	1	
104	1	1	1	0	0	1	
105	1	1	1	0	0	1	
106	1	1	1	0	0	1	
107	1	1	1	0	0	1	
108	1	1	1	0	0	1	
109	1	1	1	0	0	1	
110	1	1	1	0	0	1	
111	1	1	1	0	0	1	
112	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-7-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 21 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
201	1	1	1	0	0	1	
202	1	1	1	0	0	1	
203	1	1	1	0	0	1	
204	1	1	1	0	0	1	
205	1	1	1	0	0	1	
206	1	1	1	0	0	1	
207	1	1	1	0	0	1	
208	1	1	1	0	0	1	
209	1	1	1	0	0	1	
210	1	1	1	0	0	1	
211	1	1	1	0	0	1	
212	1	1	1	0	0	1	

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INDIVIDUAL DATA 2-7-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 21 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
301	1	1	1	0	0	1	
302	1	1	1	0	0	1	
303	1	1	1	0	0	1	
304	1	1	1	0	0	1	
305	1	1	1	0	0	1	
306	1	1	1	0	0	1	
307	1	1	1	0	0	1	
308	1	1	1	0	0	1	
309	1	1	1	0	0	1	
310	1	1	1	0	0	1	
311	1	1	1	0	0	1	
312	1	1	1	0	0	1	

N

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INDIVIDUAL DATA 2-7-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 21 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
401	1	1	1	0	0	1	
402	1	1	1	0	0	1	
403	1	1	1	0	0	1	
404	1	1	1	0	0	1	
405	1	1	1	0	0	1	
406	1	1	1	0	0	1	
407	1	1	1	0	0	1	
408	1	1	1	0	0	1	
409	1	1	1	0	0	1	
410	1	1	1	0	0	1	
411	1	1	1	0	0	1	
412	1	1	1	0	0	1	

N

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INDIVIDUAL DATA 2-7-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 21 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
101	1	1	2	1	1	1	1	0	1	1	1	1	1	0
102	1	1	2	1	1	1	1	0	1	1	1	1	1	0
103	1	1	2	1	1	1	1	0	1	1	1	1	1	0
104	1	1	2	1	1	1	1	0	1	1	1	1	1	0
105	1	1	2	1	1	1	1	0	1	1	1	1	1	0
106	1	1	2	1	1	1	1	0	1	1	1	1	1	0
107	1	1	2	1	1	1	1	0	1	1	1	1	1	0
108	1	1	2	1	1	1	1	0	1	1	1	1	1	0
109	1	1	2	1	1	1	1	0	1	1	1	1	1	0
110	1	1	2	1	1	1	1	0	1	1	1	1	1	0
111	1	1	2	1	1	1	1	0	1	1	1	1	1	0
112	1	1	2	1	1	1	1	0	1	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-7-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 21 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling		Fur	Eyes					Pupil size	Salivation		
201	1	1	2	1	1	1	0	1	1	1	1	0
202	1	1	2	1	1	1	0	1	1	1	1	0
203	1	1	2	1	1	1	0	1	1	1	1	0
204	1	1	2	1	1	1	0	1	1	1	1	0
205	1	1	2	1	1	1	0	1	1	1	1	0
206	1	1	2	1	1	1	0	1	1	1	1	0
207	1	1	2	1	1	1	0	1	1	1	1	0
208	1	1	2	1	1	1	0	1	1	1	1	0
209	1	1	2	1	1	1	0	1	1	1	1	0
210	1	1	2	1	1	1	0	1	1	1	1	0
211	1	1	2	1	1	1	0	1	1	1	1	0
212	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-7-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 21 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
301	1	1	2	1	1	1	1	1	0	1	1	1	1	0
302	1	1	2	1	1	1	1	0	1	1	1	1	1	0
303	1	1	2	1	1	1	1	0	1	1	1	1	1	0
304	1	1	2	1	1	1	1	0	1	1	1	1	1	0
305	1	1	2	1	1	1	1	0	1	1	1	1	1	0
306	1	1	2	1	1	1	1	0	1	1	1	1	1	0
307	1	1	2	1	1	1	1	0	1	1	1	1	1	0
308	1	1	2	1	1	1	1	0	1	1	1	1	1	0
309	1	1	2	1	1	1	1	0	1	1	1	1	1	0
310	1	1	2	1	1	1	1	0	1	1	1	1	1	0
311	1	1	2	1	1	1	1	0	1	1	1	1	1	0
312	1	1	2	1	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-7-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 21 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Removal	Handling			Fur	Eyes			Pupil size	Salivation		
401	1	1	2	1	1	1	0	1	1	1	1	0
402	1	1	2	1	1	1	0	1	1	1	1	0
403	1	1	2	1	1	1	0	1	1	1	1	0
404	1	1	2	1	1	1	0	1	1	1	1	0
405	1	1	2	1	1	1	0	1	1	1	1	0
406	1	1	2	1	1	1	0	1	1	1	1	0
407	1	1	2	1	1	1	0	1	1	1	1	0
408	1	1	2	1	1	1	0	1	1	1	1	0
409	1	1	2	1	1	1	0	1	1	1	1	0
410	1	1	2	1	1	1	0	1	1	1	1	0
411	1	1	2	1	1	1	0	1	1	1	1	0
412	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-7-9

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 21 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
101	1	1	1	1	0	0	0	0	1	1	1		
102	1	1	1	1	0	0	0	0	1	1	1		
103	1	1	1	1	0	0	0	0	1	1	1		
104	1	1	1	1	0	0	0	0	1	1	1		
105	1	1	1	1	0	0	0	0	1	1	1		
106	1	1	1	1	0	0	0	0	1	1	1		
107	1	1	1	1	1	1	0	0	1	1	1		
108	1	1	1	1	0	0	0	0	1	1	1		
109	1	1	1	1	1	0	0	0	1	1	1		
110	1	1	1	1	0	0	0	0	1	1	1		
111	1	1	1	1	0	0	0	0	1	1	1		
112	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.6558	0.6558	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-7-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 21 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
201	1	1	1	1	0	0	0	0	1	1	1		
202	1	1	1	1	0	0	0	0	1	1	1		
203	1	1	1	1	0	0	0	0	1	1	1		
204	1	1	1	1	1	1	0	0	1	1	1		
205	1	1	1	1	0	0	0	0	1	1	1		
206	1	1	1	1	0	0	0	0	1	1	1		
207	1	1	1	1	0	0	0	0	1	1	1		
208	1	1	1	1	0	0	0	0	1	1	1		
209	1	1	1	1	0	0	0	0	1	1	1		
210	1	1	1	1	0	0	0	0	1	1	1		
211	1	1	1	1	0	0	0	0	1	1	1		
212	1	1	1	1	0	0	0	0	1	1	1		

N	12	12	12	12	12	12	12	12	12	12	12	12	12
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INDIVIDUAL DATA 2-7-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 21 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
301	1	1	1	1	0	0	0	0	1	1	1		
302	1	1	1	1	0	0	0	0	1	1	1		
303	1	1	1	1	0	0	0	0	1	1	1		
304	1	1	1	1	1	1	0	0	1	1	1		
305	1	1	1	1	0	0	0	0	1	1	1		
306	1	1	1	1	0	0	0	0	1	1	1		
307	1	1	1	1	0	0	0	0	1	1	1		
308	1	1	1	1	0	0	0	0	1	1	1		
309	1	1	1	1	0	0	0	0	1	1	1		
310	1	1	1	1	0	0	0	0	1	1	1		
311	1	1	1	1	0	0	0	0	1	1	1		
312	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	

INDIVIDUAL DATA 2-7-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 21 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
401	1	1	1	1	0	1	0	0	1	1	1		
402	1	1	1	1	0	0	0	0	1	1	1		
403	1	1	1	1	1	1	0	0	1	1	1		
404	1	1	1	1	0	0	0	0	1	1	1		
405	1	1	1	1	0	0	0	0	1	1	1		
406	1	1	1	1	0	0	0	0	1	1	1		
407	1	1	1	1	0	0	0	0	1	1	1		
408	1	1	1	1	0	0	0	0	1	1	1		
409	1	1	1	1	0	0	0	0	1	1	1		
410	1	1	1	1	0	0	0	0	1	1	1		
411	1	1	1	1	0	0	0	0	1	1	1		
412	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12

INDIVIDUAL DATA 2-8-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 21 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
151	1	1	1	0	0	1	
152	1	1	1	0	0	1	
153	1	1	1	0	0	1	
154	1	1	1	0	0	1	
155	1	1	1	0	0	1	
156	1	1	1	0	0	1	
157	1	1	1	0	0	1	
158	1	1	1	0	0	1	
159	1	1	1	0	0	1	
160	1	1	1	0	0	1	
161	1	1	1	0	0	1	
162	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-8-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 21 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
251	1	1	1	0	0	1	
252	1	1	1	0	0	1	
253	1	1	1	0	0	1	
254	1	1	1	0	0	1	
255	1	1	1	0	0	1	
256	1	1	1	0	0	1	
257	1	1	1	0	0	1	
258	1	1	1	0	0	1	
259	1	1	1	0	0	1	
260	1	1	1	0	0	1	
261	1	1	1	0	0	1	
262	1	1	1	0	0	1	

N

12

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12

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INDIVIDUAL DATA 2-8-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 21 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
351	1	1	1	0	0	1	
352	1	1	1	0	0	1	
353	1	1	1	0	0	1	
354	1	1	1	0	0	1	
355	1	1	1	0	0	1	
356	1	1	1	0	0	1	
357	1	1	1	0	0	1	
358	1	1	1	0	0	1	
359	1	1	1	0	0	1	
360	1	1	1	0	0	1	
361	1	1	1	0	0	1	
362	1	1	1	0	0	1	

N

12

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12

12

INDIVIDUAL DATA 2-8-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 21 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
451	1	1	1	0	0	1	
452	1	1	1	0	0	1	
453	1	1	1	0	0	1	
454	1	1	1	0	0	1	
455	1	1	1	0	0	1	
456	1	1	1	0	0	1	
457	1	1	1	0	0	1	
458	1	1	1	0	0	1	
459	1	1	1	0	0	1	
460	1	1	1	0	0	1	
461	1	1	1	0	0	1	
462	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-8-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 21 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
151	1	1	2	1	1	1	1	0	1	1	1	1	1	0
152	1	1	2	1	1	1	1	0	1	1	1	1	1	0
153	1	1	2	1	1	1	1	0	1	1	1	1	1	0
154	1	1	2	1	1	1	1	0	1	1	1	1	1	0
155	1	1	2	1	1	1	1	0	1	1	1	1	1	0
156	1	1	2	1	1	1	1	0	1	1	1	1	1	0
157	1	1	2	1	1	1	1	0	1	1	1	1	1	0
158	1	1	2	1	1	1	1	0	1	1	1	1	1	0
159	1	1	2	1	1	1	1	0	1	1	1	1	1	0
160	1	1	2	1	1	1	1	0	1	1	1	1	1	0
161	1	1	2	1	1	1	1	0	1	1	1	1	1	0
162	1	1	2	1	1	1	1	0	1	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-8-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 21 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling				Fur	Eyes			Pupil size		Salivation	
251	1	1	2	1	1	1	0	1	1	1	1	0
252	1	1	2	1	1	1	0	1	1	1	1	0
253	1	1	2	1	1	1	0	1	1	1	1	0
254	1	1	2	1	1	1	0	1	1	1	1	0
255	1	1	2	1	1	1	0	1	1	1	1	0
256	1	1	2	1	1	1	0	1	1	1	1	0
257	1	1	2	1	1	1	0	1	1	1	1	0
258	1	1	2	1	1	1	0	1	1	1	1	0
259	1	1	2	1	1	1	0	1	1	1	1	0
260	1	1	2	1	1	1	0	1	1	1	1	0
261	1	1	2	1	1	1	0	1	1	1	1	0
262	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-8-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 21 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
351	1	1	2	1	1	1	0	1	1	1	1	0
352	1	1	2	1	1	1	0	1	1	1	1	0
353	1	1	2	1	1	1	0	1	1	1	1	0
354	1	1	2	1	1	1	0	1	1	1	1	0
355	1	1	2	1	1	1	0	1	1	1	1	0
356	1	1	2	1	1	1	0	1	1	1	1	0
357	1	1	2	1	1	1	0	1	1	1	1	0
358	1	1	2	1	1	1	0	1	1	1	1	0
359	1	1	2	1	1	1	0	1	1	1	1	0
360	1	1	2	1	1	1	0	1	1	1	1	0
361	1	1	2	1	1	1	0	1	1	1	1	0
362	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-8-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 21 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
451	1	1	2	1	1	1	0	1	1	1	1
452	1	1	2	1	1	0	1	1	1	1	0
453	1	1	2	1	1	0	1	1	1	1	0
454	1	1	2	1	1	0	1	1	1	1	0
455	1	1	2	1	1	0	1	1	1	1	0
456	1	1	2	1	1	0	1	1	1	1	0
457	1	1	2	1	1	0	1	1	1	1	0
458	1	1	2	1	1	0	1	1	1	1	0
459	1	1	2	1	1	0	1	1	1	1	0
460	1	1	2	1	1	0	1	1	1	1	0
461	1	1	2	1	1	0	1	1	1	1	0
462	1	1	2	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-8-9

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 21 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
151	1	1	1	1	0	0	0	0	1	1	1		
152	1	1	1	1	0	0	0	0	1	1	1		
153	1	1	1	1	0	0	0	0	1	1	1		
154	1	1	1	1	0	0	0	0	1	1	1		
155	1	1	1	1	0	0	0	0	1	1	1		
156	1	1	1	1	0	0	0	0	1	1	1		
157	1	1	1	1	0	0	0	0	1	1	1		
158	1	1	1	1	0	0	0	0	1	1	1		
159	1	1	1	1	0	0	0	0	1	1	1		
160	1	1	1	1	0	0	0	0	1	1	1		
161	1	1	1	1	0	0	0	0	1	1	1		
162	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	6.1304	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-8-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 21 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
251	1	1	1	1	0	0	0	0	1	1	1	
252	1	1	1	1	0	0	0	0	1	1	1	
253	1	1	1	1	0	0	0	0	1	1	1	
254	1	1	1	1	0	0	0	0	1	1	1	
255	1	1	1	1	0	0	0	0	1	1	1	
256	1	1	1	1	0	0	0	0	1	1	1	
257	1	1	1	1	0	0	0	0	1	1	1	
258	1	1	1	1	0	0	0	0	1	1	1	
259	1	1	1	1	0	0	0	0	1	1	1	
260	1	1	1	1	0	0	0	0	1	1	1	
261	1	1	1	1	0	0	0	0	1	1	1	
262	1	1	1	1	0	0	0	0	1	1	1	

N	12	12	12	12	12	12	12	12	12	12	12	12
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INDIVIDUAL DATA 2-8-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 21 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
351	1	1	1	1	0	0	0	0	1	1	1	
352	1	1	1	1	0	0	0	0	1	1	1	
353	1	1	1	1	0	0	0	0	1	1	1	
354	1	1	1	1	0	0	0	0	1	1	1	
355	1	1	1	1	0	0	0	0	1	1	1	
356	1	1	1	1	0	0	0	0	1	1	1	
357	1	1	1	1	0	0	0	0	1	1	1	
358	1	1	1	1	0	0	0	0	1	1	1	
359	1	1	1	1	0	0	0	0	1	1	1	
360	1	1	1	1	0	0	0	0	1	1	1	
361	1	1	1	1	0	0	0	0	1	1	1	
362	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-8-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 21 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
451	1	1	1	1	1	0	0	0	1	1	1	
452	1	1	1	1	0	0	0	0	1	1	1	
453	1	1	1	1	0	0	0	0	1	1	1	
454	1	1	1	1	0	0	0	0	1	1	1	
455	1	1	1	1	0	0	0	0	1	1	1	
456	1	1	1	1	0	0	0	0	1	1	1	
457	1	1	1	1	0	0	0	0	1	1	1	
458	1	1	1	1	0	0	0	0	1	1	1	
459	1	1	1	1	1	0	0	0	1	1	1	
460	1	1	1	1	0	0	0	0	1	1	1	
461	1	1	1	1	0	0	0	0	1	1	1	
462	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-8-13

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 21 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
163	1	1	1	0	0	1
164	1	1	1	0	0	1
165	1	1	1	0	0	1
166	1	1	1	0	0	1
167	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-8-14

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 21 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
463	1	1	1	0	0	1
464	1	1	1	0	0	1
465	1	1	1	0	0	1
466	1	1	1	0	0	1
467	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-8-15

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 21 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
163	1	1	2	1	1	1	0	1	1	1	1
164	1	1	2	1	1	1	0	1	1	1	1
165	1	1	2	1	1	1	0	1	1	1	1
166	1	1	2	1	1	1	0	1	1	1	1
167	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-8-16

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 21 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
463	1	1	2	1	1	1	0	1	1	1	1
464	1	1	2	1	1	1	0	1	1	1	1
465	1	1	2	1	1	1	0	1	1	1	1
466	1	1	2	1	1	1	0	1	1	1	1
467	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-8-17

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 21 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
163	1	1	1	1	0	0	0	0	1	1	1	
164	1	1	1	1	0	0	0	0	1	1	1	
165	1	1	1	1	0	0	0	0	1	1	1	
166	1	1	1	1	0	0	0	0	1	1	1	
167	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-8-18

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 21 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
463	1	1	1	1	0	0	0	0	1	1	1	
464	1	1	1	1	0	0	0	0	1	1	1	
465	1	1	1	1	0	0	0	0	1	1	1	
466	1	1	1	1	0	0	0	0	1	1	1	
467	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	

INDIVIDUAL DATA 2-9-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 28 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
101	1	1	1	0	0	1	
102	1	1	1	0	0	1	
103	1	1	1	0	0	1	
104	1	1	1	0	0	1	
105	1	1	1	0	0	1	
106	1	1	1	0	0	1	
107	1	1	1	0	0	1	
108	1	1	1	0	0	1	
109	1	1	1	0	0	1	
110	1	1	1	0	0	1	
111	1	1	1	0	0	1	
112	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-9-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 28 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
201	1	1	1	0	0	1	
202	1	1	1	0	0	1	
203	1	1	1	0	0	1	
204	1	1	1	0	0	1	
205	1	1	1	0	0	1	
206	1	1	1	0	0	1	
207	1	1	1	0	0	1	
208	1	1	1	0	0	1	
209	1	1	1	0	0	1	
210	1	1	1	0	0	1	
211	1	1	1	0	0	1	
212	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-9-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 28 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
301	1	1	1	0	0	1	
302	1	1	1	0	0	1	
303	1	1	1	0	0	1	
304	1	1	1	0	0	1	
305	1	1	1	0	0	1	
306	1	1	1	0	0	1	
307	1	1	1	0	0	1	
308	1	1	1	0	0	1	
309	1	1	1	0	0	1	
310	1	1	1	0	0	1	
311	1	1	1	0	0	1	
312	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-9-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 28 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
401	1	1	1	0	0	1	
402	1	1	1	0	0	1	
403	1	1	1	0	0	1	
404	1	1	1	0	0	1	
405	1	1	1	0	0	1	
406	1	1	1	0	0	1	
407	1	1	1	0	0	1	
408	1	1	1	0	0	1	
409	1	1	1	0	0	1	
410	1	1	1	0	0	1	
411	1	1	1	0	0	1	
412	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-9-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 28 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling		Fur	Eyes					Pupil size	Salivation		
101	1	1	2	1	1	1	0	1	1	1	1	0
102	1	1	2	1	1	1	0	1	1	1	1	0
103	1	1	2	1	1	1	0	1	1	1	1	0
104	1	1	2	1	1	1	0	1	1	1	1	0
105	1	1	2	1	1	1	0	1	1	1	1	0
106	1	1	2	1	1	1	0	1	1	1	1	0
107	1	1	2	1	1	1	0	1	1	1	1	0
108	1	1	2	1	1	1	0	1	1	1	1	0
109	1	1	2	1	1	1	0	1	1	1	1	0
110	1	1	2	1	1	1	0	1	1	1	1	0
111	1	1	2	1	1	1	0	1	1	1	1	0
112	1	1	2	1	1	1	0	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-9-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 28 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
201	1	1	2	1	1	1	0	1	1	1	1	0
202	1	1	2	1	1	1	0	1	1	1	1	0
203	1	1	2	1	1	1	0	1	1	1	1	0
204	1	1	2	1	1	1	0	1	1	1	1	0
205	1	1	2	1	1	1	0	1	1	1	1	0
206	1	1	2	1	1	1	0	1	1	1	1	0
207	1	1	2	1	1	1	0	1	1	1	1	0
208	1	1	2	1	1	1	0	1	1	1	1	0
209	1	1	2	1	1	1	0	1	1	1	1	0
210	1	1	2	1	1	1	0	1	1	1	1	0
211	1	1	2	1	1	1	0	1	1	1	1	0
212	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-9-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 28 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
301	1	1	2	1	1	1	1	1	0	1	1	1	1	0
302	1	1	2	1	1	1	1	0	1	1	1	1	1	0
303	1	1	2	1	1	1	1	0	1	1	1	1	1	0
304	1	1	2	1	1	1	1	0	1	1	1	1	1	0
305	1	1	2	1	1	1	1	0	1	1	1	1	1	0
306	1	1	2	1	1	1	1	0	1	1	1	1	1	0
307	1	1	2	1	1	1	1	0	1	1	1	1	1	0
308	1	1	2	1	1	1	1	0	1	1	1	1	1	0
309	1	1	2	1	1	1	1	0	1	1	1	1	1	0
310	1	1	2	1	1	1	1	0	1	1	1	1	1	0
311	1	1	2	1	1	1	1	0	1	1	1	1	1	0
312	1	1	2	1	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-9-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 28 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Removal	Handling			Fur	Eyes			Pupil size	Salivation		
401	1	1	2	1	1	1	0	1	1	1	1	0
402	1	1	2	1	1	1	0	1	1	1	1	0
403	1	1	2	1	1	1	0	1	1	1	1	0
404	1	1	2	1	1	1	0	1	1	1	1	0
405	1	1	2	1	1	1	0	1	1	1	1	0
406	1	1	2	1	1	1	0	1	1	1	1	0
407	1	1	2	1	1	1	0	1	1	1	1	0
408	1	1	2	1	1	1	0	1	1	1	1	0
409	1	1	2	1	1	1	0	1	1	1	1	0
410	1	1	2	1	1	1	0	1	1	1	1	0
411	1	1	2	1	1	1	0	1	1	1	1	0
412	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-9-9

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 28 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
101	1	1	1	1	0	0	0	0	1	1	1		
102	1	1	1	1	0	0	0	0	1	1	1		
103	1	1	1	1	0	0	0	0	1	1	1		
104	1	1	1	1	0	0	0	0	1	1	1		
105	1	1	1	1	0	0	0	0	1	1	1		
106	1	1	1	1	0	0	0	0	1	1	1		
107	1	1	1	1	1	1	0	0	1	1	1		
108	1	1	1	1	0	0	0	0	1	1	1		
109	1	1	1	1	0	0	0	0	1	1	1		
110	1	1	1	1	0	0	0	0	1	1	1		
111	1	1	1	1	0	0	0	0	1	1	1		
112	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.6558	3.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-9-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 28 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
201	1	1	1	1	1	0	0	0	1	1	1		
202	1	1	1	1	0	0	0	0	1	1	1		
203	1	1	1	1	0	0	0	0	1	1	1		
204	1	1	1	1	1	0	0	0	1	1	1		
205	1	1	1	1	0	0	0	0	1	1	1		
206	1	1	1	1	0	0	0	0	1	1	1		
207	1	1	1	1	0	0	0	0	1	1	1		
208	1	1	1	1	0	0	0	0	1	1	1		
209	1	1	1	1	0	0	0	0	1	1	1		
210	1	1	1	1	0	0	0	0	1	1	1		
211	1	1	1	1	0	0	0	0	1	1	1		
212	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12

INDIVIDUAL DATA 2-9-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 28 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
301	1	1	1	1	1	0	0	0	1	1	1		
302	1	1	1	1	0	0	0	0	1	1	1		
303	1	1	1	1	0	0	0	0	1	1	1		
304	1	1	1	1	0	0	0	0	1	1	1		
305	1	1	1	1	0	0	0	0	1	1	1		
306	1	1	1	1	0	0	0	0	1	1	1		
307	1	1	1	1	0	0	0	0	1	1	1		
308	1	1	1	1	0	0	0	0	1	1	1		
309	1	1	1	1	0	0	0	0	1	1	1		
310	1	1	1	1	0	0	0	0	1	1	1		
311	1	1	1	1	0	0	0	0	1	1	1		
312	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	

INDIVIDUAL DATA 2-9-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 28 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
401	1	1	1	1	0	0	0	0	1	1	1		
402	1	1	1	1	0	0	0	0	1	1	1		
403	1	1	1	1	0	0	0	0	1	1	1		
404	1	1	1	1	0	0	0	0	1	1	1		
405	1	1	1	1	0	0	0	0	1	1	1		
406	1	1	1	1	0	0	0	0	1	1	1		
407	1	1	1	1	0	0	0	0	1	1	1		
408	1	1	1	1	1	0	0	0	1	1	1		
409	1	1	1	1	0	0	0	0	1	1	1		
410	1	1	1	1	0	0	0	0	1	1	1		
411	1	1	1	1	0	0	0	0	1	1	1		
412	1	1	1	1	0	0	0	0	1	1	1		

N	12	12	12	12	12	12	12	12	12	12	12	12	12
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INDIVIDUAL DATA 2-10-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 28 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
151	1	1	1	0	0	1	
152	1	1	1	0	0	1	
153	1	1	1	0	0	1	
154	1	1	1	0	0	1	
155	1	1	1	0	0	1	
156	1	1	1	0	0	1	
157	1	1	1	0	0	1	
158	1	1	1	0	0	1	
159	1	1	1	0	0	1	
160	1	1	1	0	0	1	
161	1	1	1	0	0	1	
162	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-10-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 28 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
251	1	1	1	0	0	1	
252	1	1	1	0	0	1	
253	1	1	1	0	0	1	
254	1	1	1	0	0	1	
255	1	1	1	0	0	1	
256	1	1	1	0	0	1	
257	1	1	1	0	0	1	
258	1	1	1	0	0	1	
259	1	1	1	0	0	1	
260	1	1	1	0	0	1	
261	1	1	1	0	0	1	
262	1	1	1	0	0	1	

N

12

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INDIVIDUAL DATA 2-10-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 28 of administration

Animal No.	In the cage						
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
351	1	1	1	0	0	1	
352	1	1	1	0	0	1	
353	1	1	1	0	0	1	
354	1	1	1	0	0	1	
355	1	1	1	0	0	1	
356	1	1	1	0	0	1	
357	1	1	1	0	0	1	
358	1	1	1	0	0	1	
359	1	1	1	0	0	1	
360	1	1	1	0	0	1	
361	1	1	1	0	0	1	
362	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-10-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 28 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
451	1	1	1	0	0	1
452	1	1	1	0	0	1
453	1	1	1	0	0	1
454	1	1	1	0	0	1
455	1	1	1	0	0	1
456	1	1	1	0	0	1
457	1	1	1	0	0	1
458	1	1	1	0	0	1
459	1	1	1	0	0	1
460	1	1	1	0	0	1
461	1	1	1	0	0	1
462	1	1	1	0	0	1

N

12

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INDIVIDUAL DATA 2-10-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 28 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
151	1	1	2	1	1	1	1	0	1	1	1	1	1	0
152	1	1	2	1	1	1	1	0	1	1	1	1	1	0
153	1	1	2	1	1	1	1	0	1	1	1	1	1	0
154	1	1	2	1	1	1	1	0	1	1	1	1	1	0
155	1	1	2	1	1	1	1	0	1	1	1	1	1	0
156	1	1	2	1	1	1	1	0	1	1	1	1	1	0
157	1	1	2	1	1	1	1	0	1	1	1	1	1	0
158	1	1	2	1	1	1	1	0	1	1	1	1	1	0
159	1	1	2	1	1	1	1	0	1	1	1	1	1	0
160	1	1	2	1	1	1	1	0	1	1	1	1	1	0
161	1	1	2	1	1	1	1	0	1	1	1	1	1	0
162	1	1	2	1	1	1	1	0	1	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-10-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 28 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
251	1	1	2	1	1	1	0	1	1	1	1	0
252	1	1	2	1	1	1	0	1	1	1	1	0
253	1	1	2	1	1	1	0	1	1	1	1	0
254	1	1	2	1	1	1	0	1	1	1	1	0
255	1	1	2	1	1	1	0	1	1	1	1	0
256	1	1	2	1	1	1	0	1	1	1	1	0
257	1	1	2	1	1	1	0	1	1	1	1	0
258	1	1	2	1	1	1	0	1	1	1	1	0
259	1	1	2	1	1	1	0	1	1	1	1	0
260	1	1	2	1	1	1	0	1	1	1	1	0
261	1	1	2	1	1	1	0	1	1	1	1	0
262	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-10-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 28 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling				Fur	Eyes			Pupil size		Salivation	
351	1	1	2	1	1	1	0	1	1	1	1	0
352	1	1	2	1	1	1	0	1	1	1	1	0
353	1	1	2	1	1	1	0	1	1	1	1	0
354	1	1	2	1	1	1	0	1	1	1	1	0
355	1	1	2	1	1	1	0	1	1	1	1	0
356	1	1	2	1	1	1	0	1	1	1	1	0
357	1	1	2	1	1	1	0	1	1	1	1	0
358	1	1	2	1	1	1	0	1	1	1	1	0
359	1	1	2	1	1	1	0	1	1	1	1	0
360	1	1	2	1	1	1	0	1	1	1	1	0
361	1	1	2	1	1	1	0	1	1	1	1	0
362	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-10-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 28 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
451	1	1	2	1	1	1	0	1	1	1	1
452	1	1	2	1	1	1	0	1	1	1	1
453	1	1	2	1	1	1	0	1	1	1	1
454	1	1	2	1	1	1	0	1	1	1	1
455	1	1	2	1	1	1	0	1	1	1	1
456	1	1	2	1	1	1	0	1	1	1	1
457	1	1	2	1	1	1	0	1	1	1	1
458	1	1	2	1	1	1	0	1	1	1	1
459	1	1	2	1	1	1	0	1	1	1	1
460	1	1	2	1	1	1	0	1	1	1	1
461	1	1	2	1	1	1	0	1	1	1	1
462	1	1	2	1	1	1	0	1	1	1	1

INDIVIDUAL DATA 2-10-9

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 28 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
151	1	1	1	1	0	0	0	0	1	1	1		
152	1	1	1	1	0	0	0	0	1	1	1		
153	1	1	1	1	0	0	0	0	1	1	1		
154	1	1	1	1	0	0	0	0	1	1	1		
155	1	1	1	1	0	0	0	0	1	1	1		
156	1	1	1	1	0	0	0	0	1	1	1		
157	1	1	1	1	0	0	0	0	1	1	1		
158	1	1	1	1	0	0	0	0	1	1	1		
159	1	1	1	1	0	0	0	0	1	1	1		
160	1	1	1	1	0	0	0	0	1	1	1		
161	1	1	1	1	0	0	0	0	1	1	1		
162	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-10-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 28 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
251	1	1	1	1	0	0	0	0	1	1	1	
252	1	1	1	1	0	0	0	0	1	1	1	
253	1	1	1	1	0	0	0	0	1	1	1	
254	1	1	1	1	0	0	0	0	1	1	1	
255	1	1	1	1	0	0	0	0	1	1	1	
256	1	1	1	1	0	0	0	0	1	1	1	
257	1	1	1	1	0	0	0	0	1	1	1	
258	1	1	1	1	0	0	0	0	1	1	1	
259	1	1	1	1	0	0	0	0	1	1	1	
260	1	1	1	1	0	0	0	0	1	1	1	
261	1	1	1	1	0	0	0	0	1	1	1	
262	1	1	1	1	0	0	0	0	1	1	1	
N	12	12	12	12	12	12	12	12	12	12	12	

INDIVIDUAL DATA 2-10-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 28 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
351	1	1	1	1	0	0	0	0	1	1	1	
352	1	1	1	1	0	0	0	0	1	1	1	
353	1	1	1	1	0	0	0	0	1	1	1	
354	1	1	1	1	0	0	0	0	1	1	1	
355	1	1	1	1	0	0	0	0	1	1	1	
356	1	1	1	1	0	0	0	0	1	1	1	
357	1	1	1	1	0	0	0	0	1	1	1	
358	1	1	1	1	0	0	0	0	1	1	1	
359	1	1	1	1	0	0	0	0	1	1	1	
360	1	1	1	1	0	0	0	0	1	1	1	
361	1	1	1	1	0	0	0	0	1	1	1	
362	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-10-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 28 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
451	1	1	1	1	0	0	0	0	1	1	1	
452	1	1	1	1	0	0	0	0	1	1	1	
453	1	1	1	1	0	0	0	0	1	1	1	
454	1	1	1	1	0	0	0	0	1	1	1	
455	1	1	1	1	0	0	0	0	1	1	1	
456	1	1	1	1	0	0	0	0	1	1	1	
457	1	1	1	1	0	0	0	0	1	1	1	
458	1	1	1	1	0	0	0	0	1	1	1	
459	1	1	1	1	0	0	0	0	1	1	1	
460	1	1	1	1	0	0	0	0	1	1	1	
461	1	1	1	1	0	0	0	0	1	1	1	
462	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-10-13

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 28 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
163	1	1	1	0	0	1
164	1	1	1	0	0	1
165	1	1	1	0	0	1
166	1	1	1	0	0	1
167	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-10-14

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 28 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
463	1	1	1	0	0	1
464	1	1	1	0	0	1
465	1	1	1	0	0	1
466	1	1	1	0	0	1
467	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-10-15

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 28 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
163	1	1	2	1	1	1	0	1	1	1	1
164	1	1	2	1	1	1	0	1	1	1	1
165	1	1	2	1	1	1	0	1	1	1	1
166	1	1	2	1	1	1	0	1	1	1	1
167	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-10-16

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 28 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
463	1	1	2	1	1	1	0	1	1	1	0
464	1	1	2	1	1	1	0	1	1	1	0
465	1	1	2	1	1	1	0	1	1	1	0
466	1	1	2	1	1	1	0	1	1	1	0
467	1	1	2	1	1	1	0	1	1	1	0
N	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-10-17

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 28 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
163	1	1	1	1	0	0	0	0	1	1	1		
164	1	1	1	1	0	0	0	0	1	1	1		
165	1	1	1	1	0	0	0	0	1	1	1		
166	1	1	1	1	0	0	0	0	1	1	1		
167	1	1	1	1	0	0	0	0	1	1	1		
N	5	5	5	5	5	5	5	5	5	5	5		
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000		

INDIVIDUAL DATA 2-10-18

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 28 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
463	1	1	1	1	0	0	0	0	1	1	1		
464	1	1	1	1	0	0	0	0	1	1	1		
465	1	1	1	1	0	0	0	0	1	1	1		
466	1	1	1	1	0	0	0	0	1	1	1		
467	1	1	1	1	0	0	0	0	1	1	1		
N	5	5	5	5	5	5	5	5	5	5	5		

INDIVIDUAL DATA 2-11-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 35 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
101	1	1	1	0	0	1	
102	1	1	1	0	0	1	
103	1	1	1	0	0	1	
104	1	1	1	0	0	1	
105	1	1	1	0	0	1	
106	1	1	1	0	0	1	
107	1	1	1	0	0	1	
108	1	1	1	0	0	1	
109	1	1	1	0	0	1	
110	1	1	1	0	0	1	
111	1	1	1	0	0	1	
112	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-11-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 35 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
201	1	1	1	0	0	1	
202	1	1	1	0	0	1	
203	1	1	1	0	0	1	
204	1	1	1	0	0	1	
205	1	1	1	0	0	1	
206	1	1	1	0	0	1	
207	1	1	1	0	0	1	
208	1	1	1	0	0	1	
209	1	1	1	0	0	1	
210	1	1	1	0	0	1	
211	1	1	1	0	0	1	
212	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-11-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 35 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
301	1	1	1	0	0	1	
302	1	1	1	0	0	1	
303	1	1	1	0	0	1	
304	1	1	1	0	0	1	
305	1	1	1	0	0	1	
306	1	1	1	0	0	1	
307	1	1	1	0	0	1	
308	1	1	1	0	0	1	
309	1	1	1	0	0	1	
310	1	1	1	0	0	1	
311	1	1	1	0	0	1	
312	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-11-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 35 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
401	1	1	1	0	0	1	
402	1	1	1	0	0	1	
403	1	1	1	0	0	1	
404	1	1	1	0	0	1	
405	1	1	1	0	0	1	
406	1	1	1	0	0	1	
407	1	1	1	0	0	1	
408	1	1	1	0	0	1	
409	1	1	1	0	0	1	
410	1	1	1	0	0	1	
411	1	1	1	0	0	1	
412	1	1	1	0	0	1	

N

12

12

12

12

12

12

INDIVIDUAL DATA 2-11-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 35 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
101	1	1	2	1	1	1	1	0	1	1	1	1	1	0
102	1	1	2	1	1	1	1	0	1	1	1	1	1	0
103	1	1	2	1	1	1	1	0	1	1	1	1	1	0
104	1	1	2	1	1	1	1	0	1	1	1	1	1	0
105	1	1	2	1	1	1	1	0	1	1	1	1	1	0
106	1	1	2	1	1	1	1	0	1	1	1	1	1	0
107	1	1	2	1	1	1	1	0	1	1	1	1	1	0
108	1	1	2	1	1	1	1	0	1	1	1	1	1	0
109	1	1	2	1	1	1	1	0	1	1	1	1	1	0
110	1	1	2	1	1	1	1	0	1	1	1	1	1	0
111	1	1	2	1	1	1	1	0	1	1	1	1	1	0
112	1	1	2	1	1	1	1	0	1	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-11-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 35 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling		Fur	Eyes					Pupil size	Salivation		
201	1	1	2	1	1	1	0	1	1	1	1	0
202	1	1	2	1	1	1	0	1	1	1	1	0
203	1	1	2	1	1	1	0	1	1	1	1	0
204	1	1	2	1	1	1	0	1	1	1	1	0
205	1	1	2	1	1	1	0	1	1	1	1	0
206	1	1	2	1	1	1	0	1	1	1	1	0
207	1	1	2	1	1	1	0	1	1	1	1	0
208	1	1	2	1	1	1	0	1	1	1	1	0
209	1	1	2	1	1	1	0	1	1	1	1	0
210	1	1	2	1	1	1	0	1	1	1	1	0
211	1	1	2	1	1	1	0	1	1	1	1	0
212	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-11-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 35 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Removal	Handling			Fur	Eyes			Pupil size	Salivation		
301	1	1	2	1	1	1	0	1	1	1	1	0
302	1	1	2	1	1	1	0	1	1	1	1	0
303	1	1	2	1	1	1	0	1	1	1	1	0
304	1	1	2	1	1	1	0	1	1	1	1	0
305	1	1	2	1	1	1	0	1	1	1	1	0
306	1	1	2	1	1	1	0	1	1	1	1	0
307	1	1	2	1	1	1	0	1	1	1	1	0
308	1	1	2	1	1	1	0	1	1	1	1	0
309	1	1	2	1	1	1	0	1	1	1	1	0
310	1	1	2	1	1	1	0	1	1	1	1	0
311	1	1	2	1	1	1	0	1	1	1	1	0
312	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-11-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 35 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling				Fur	Eyes			Pupil size		Salivation	
401	1	1	2	1	1	1	0	1	1	1	1	0
402	1	1	2	1	1	1	0	1	1	1	1	0
403	1	1	2	1	1	1	0	1	1	1	1	0
404	1	1	2	1	1	1	0	1	1	1	1	0
405	1	1	2	1	1	1	0	1	1	1	1	0
406	1	1	2	1	1	1	0	1	1	1	1	0
407	1	1	2	1	1	1	0	1	1	1	1	0
408	1	1	2	1	1	1	0	1	1	1	1	0
409	1	1	2	1	1	1	0	1	1	1	1	0
410	1	1	2	1	1	1	0	1	1	1	1	0
411	1	1	2	1	1	1	0	1	1	1	1	0
412	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-11-9

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 35 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
101	1	1	1	1	0	0	0	0	1	1	1		
102	1	1	1	1	1	0	0	0	1	1	1		
103	1	1	1	1	0	0	0	0	1	1	1		
104	1	1	1	1	0	0	0	0	1	1	1		
105	1	1	1	1	0	0	0	0	1	1	1		
106	1	1	1	1	0	0	0	0	1	1	1		
107	1	1	1	1	0	0	0	0	1	1	1		
108	1	1	1	1	0	0	0	0	1	1	1		
109	1	1	1	1	1	0	0	0	1	1	1		
110	1	1	1	1	0	0	0	0	1	1	1		
111	1	1	1	1	0	0	0	0	1	1	1		
112	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	4.2727	3.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-11-10

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 35 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
201	1	1	1	1	1	0	0	0	1	1	1		
202	1	1	1	1	0	0	0	0	1	1	1		
203	1	1	1	1	0	0	0	0	1	1	1		
204	1	1	1	1	1	1	0	0	1	1	1		
205	1	1	1	1	0	0	0	0	1	1	1		
206	1	1	1	1	0	0	0	0	1	1	1		
207	1	1	1	1	0	0	0	0	1	1	1		
208	1	1	1	1	0	0	0	0	1	1	1		
209	1	1	1	1	0	0	0	0	1	1	1		
210	1	1	1	1	0	0	0	0	1	1	1		
211	1	1	1	1	0	0	0	0	1	1	1		
212	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	

INDIVIDUAL DATA 2-11-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 35 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
301	1	1	1	1	0	0	0	0	1	1	1		
302	1	1	1	1	0	0	0	0	1	1	1		
303	1	1	1	1	0	0	0	0	1	1	1		
304	1	1	1	1	0	0	0	0	1	1	1		
305	1	1	1	1	0	0	0	0	1	1	1		
306	1	1	1	1	0	0	0	0	1	1	1		
307	1	1	1	1	0	0	0	0	1	1	1		
308	1	1	1	1	0	0	0	0	1	1	1		
309	1	1	1	1	0	0	0	0	1	1	1		
310	1	1	1	1	0	0	0	0	1	1	1		
311	1	1	1	1	0	0	0	0	1	1	1		
312	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	

INDIVIDUAL DATA 2-11-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 35 of administration

Animal No.	In the open-field						Stereotype					Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Stereotype		Unusual head movement	Walking backward	Vocalization	Aggression		
							Excessive grooming							
401	1	1	1	1	0	0	0	0	0	1	1	1		
402	1	1	1	1	0	0	0	0	0	1	1	1		
403	1	1	1	1	0	0	0	0	0	1	1	1		
404	1	1	1	1	0	0	0	0	0	1	1	1		
405	1	1	1	1	0	0	0	0	0	1	1	1		
406	1	1	1	1	0	0	0	0	0	1	1	1		
407	1	1	1	1	0	0	0	0	0	1	1	1		
408	1	1	1	1	0	0	0	0	0	1	1	1		
409	1	1	1	1	0	0	0	0	0	1	1	1		
410	1	1	1	1	0	0	0	0	0	1	1	1		
411	1	1	1	1	0	0	0	0	0	1	1	1		
412	1	1	1	1	0	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12		

INDIVIDUAL DATA 2-12-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 35 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
151	1	1	1	0	0	1	
152	1	1	1	0	0	1	
153	1	1	1	0	0	1	
154	1	1	1	0	0	1	
155	1	1	1	0	0	1	
156	1	1	1	0	0	1	
157	1	1	1	0	0	1	
158	1	1	1	0	0	1	
159	1	1	1	0	0	1	
160	1	1	1	0	0	1	
161	1	1	1	0	0	1	
162	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-12-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 35 of administration

Animal No.	In the cage					
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
251	1	1	1	0	0	1
252	1	1	1	0	0	1
253	1	1	1	0	0	1
254	1	1	1	0	0	1
255	1	1	1	0	0	1
256	1	1	1	0	0	1
257	1	1	1	0	0	1
258	1	1	1	0	0	1
259	1	1	1	0	0	1
260	1	1	1	0	0	1
261	1	1	1	0	0	1
262	1	1	1	0	0	1

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-12-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 35 of administration

Animal No.	In the cage						
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
351	1	1	1	0	0	1	
352	1	1	1	0	0	1	
353	1	1	1	0	0	1	
354	1	1	1	0	0	1	
355	1	1	1	0	0	1	
356	1	1	1	0	0	1	
357	1	1	1	0	0	1	
358	1	1	1	0	0	1	
359	1	1	1	0	0	1	
360	1	1	1	0	0	1	
361	1	1	1	0	0	1	
362	1	1	1	0	0	1	

N	12	12	12	12	12	12
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INDIVIDUAL DATA 2-12-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 35 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
451	1	1	1	0	0	1	
452	1	1	1	0	0	1	
453	1	1	1	0	0	1	
454	1	1	1	0	0	1	
455	1	1	1	0	0	1	
456	1	1	1	0	0	1	
457	1	1	1	0	0	1	
458	1	1	1	0	0	1	
459	1	1	1	0	0	1	
460	1	1	1	0	0	1	
461	1	1	1	0	0	1	
462	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-12-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 35 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
151	1	1	2	1	1	1	1	0	1	1	1	1	1	0
152	1	1	2	1	1	1	1	0	1	1	1	1	1	0
153	1	1	2	1	1	1	1	0	1	1	1	1	1	0
154	1	1	2	1	1	1	1	0	1	1	1	1	1	0
155	1	1	2	1	1	1	1	0	1	1	1	1	1	0
156	1	1	2	1	1	1	1	0	1	1	1	1	1	0
157	1	1	2	1	1	1	1	0	1	1	1	1	1	0
158	1	1	2	1	1	1	1	0	1	1	1	1	1	0
159	1	1	2	1	1	1	1	0	1	1	1	1	1	0
160	1	1	2	1	1	1	1	0	1	1	1	1	1	0
161	1	1	2	1	1	1	1	0	1	1	1	1	1	0
162	1	1	2	1	1	1	1	0	1	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-12-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 35 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
251	1	1	2	1	1	1	0	1	1	1	1	0
252	1	1	2	1	1	1	0	1	1	1	1	0
253	1	1	2	1	1	1	0	1	1	1	1	0
254	1	1	2	1	1	1	0	1	1	1	1	0
255	1	1	2	1	1	1	0	1	1	1	1	0
256	1	1	2	1	1	1	0	1	1	1	1	0
257	1	1	2	1	1	1	0	1	1	1	1	0
258	1	1	2	1	1	1	0	1	1	1	1	0
259	1	1	2	1	1	1	0	1	1	1	1	0
260	1	1	2	1	1	1	0	1	1	1	1	0
261	1	1	2	1	1	1	0	1	1	1	1	0
262	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-12-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 35 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
351	1	1	2	1	1	1	1	1	0	1	1	1	1	0
352	1	1	2	1	1	1	1	0	1	1	1	1	1	0
353	1	1	2	1	1	1	1	0	1	1	1	1	1	0
354	1	1	2	1	1	1	1	0	1	1	1	1	1	0
355	1	1	2	1	1	1	1	0	1	1	1	1	1	0
356	1	1	2	1	1	1	1	0	1	1	1	1	1	0
357	1	1	2	1	1	1	1	0	1	1	1	1	1	0
358	1	1	2	1	1	1	1	0	1	1	1	1	1	0
359	1	1	2	1	1	1	1	0	1	1	1	1	1	0
360	1	1	2	1	1	1	1	0	1	1	1	1	1	0
361	1	1	2	1	1	1	1	0	1	1	1	1	1	0
362	1	1	2	1	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-12-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 35 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
451	1	1	2	1	1	1	1	1	0	1	1	1	1	0
452	1	1	2	1	1	1	1	0	1	1	1	1	1	0
453	1	1	2	1	1	1	1	0	1	1	1	1	1	0
454	1	1	2	1	1	1	1	0	1	1	1	1	1	0
455	1	1	2	1	1	1	1	0	1	1	1	1	1	0
456	1	1	2	1	1	1	1	0	1	1	1	1	1	0
457	1	1	2	1	1	1	1	0	1	1	1	1	1	0
458	1	1	2	1	1	1	1	0	1	1	1	1	1	0
459	1	1	2	1	1	1	1	0	1	1	1	1	1	0
460	1	1	2	1	1	1	1	0	1	1	1	1	1	0
461	1	1	2	1	1	1	1	0	1	1	1	1	1	0
462	1	1	2	1	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-12-9

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 35 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli				Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
				Searching	Urination	Defecation							
151	1	1	1	1	0	0	1	0	1	1	1	1	
152	1	1	1	1	0	0	0	0	1	1	1	1	
153	1	1	1	1	0	0	0	0	1	1	1	1	
154	1	1	1	1	0	0	0	0	1	1	1	1	
155	1	1	1	1	0	0	0	0	1	1	1	1	
156	1	1	1	1	0	0	0	0	1	1	1	1	
157	1	1	1	1	0	0	0	0	1	1	1	1	
158	1	1	1	1	0	0	0	0	1	1	1	1	
159	1	1	1	1	0	0	0	0	1	1	1	1	
160	1	1	1	1	0	0	0	0	1	1	1	1	
161	1	1	1	1	0	0	0	0	1	1	1	1	
162	1	1	1	1	0	0	0	0	1	1	1	1	
N	12	12	12	12	12	12	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	3.0000	0.0000	3.7302	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-12-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 35 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
251	1	1	1	1	0	0	0	0	1	1	1		
252	1	1	1	1	0	0	0	0	1	1	1		
253	1	1	1	1	0	0	0	0	1	1	1		
254	1	1	1	1	0	0	0	0	1	1	1		
255	1	1	1	1	0	0	1	0	1	1	1		
256	1	1	1	1	0	0	1	0	1	1	1		
257	1	1	1	1	0	0	0	0	1	1	1		
258	1	1	1	1	0	0	0	0	1	1	1		
259	1	1	1	1	0	0	0	0	1	1	1		
260	1	1	1	1	0	0	0	0	1	1	1		
261	1	1	1	1	0	0	0	0	1	1	1		
262	1	1	1	1	1	0	0	0	1	1	1		

N 12 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-12-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 35 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
351	1	1	1	1	0	0	1	0	1	1	1	
352	1	1	1	1	0	0	0	0	1	1	1	
353	1	1	1	1	0	0	1	0	1	1	1	
354	1	1	1	1	0	0	0	0	1	1	1	
355	1	1	1	1	0	0	0	0	1	1	1	
356	1	1	1	1	0	0	0	0	1	1	1	
357	1	1	1	1	0	0	0	0	1	1	1	
358	1	1	1	1	0	0	0	0	1	1	1	
359	1	1	1	1	0	0	1	0	1	1	1	
360	1	1	1	1	0	0	0	0	1	1	1	
361	1	1	1	1	0	0	0	0	1	1	1	
362	1	1	1	1	0	0	0	0	1	1	1	

N	12	12	12	12	12	12	12	12	12	12	12	12
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INDIVIDUAL DATA 2-12-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 35 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
451	1	1	1	1	0	0	0	0	1	1	1	
452	1	1	1	1	0	0	0	0	1	1	1	
453	1	1	1	1	0	0	0	0	1	1	1	
454	1	1	1	1	0	0	0	0	1	1	1	
455	1	1	1	1	0	0	0	0	1	1	1	
456	1	1	1	1	0	0	0	0	1	1	1	
457	1	1	1	1	0	0	0	0	1	1	1	
458	1	1	1	1	0	0	0	0	1	1	1	
459	1	1	1	1	0	0	0	0	1	1	1	
460	1	1	1	1	0	0	0	0	1	1	1	
461	1	1	1	1	0	0	0	0	1	1	1	
462	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-12-13

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 35 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
163	1	1	1	0	0	1
164	1	1	1	0	0	1
165	1	1	1	0	0	1
166	1	1	1	0	0	1
167	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-12-14

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 35 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
463	1	1	1	0	0	1
464	1	1	1	0	0	1
465	1	1	1	0	0	1
466	1	1	1	0	0	1
467	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-12-15

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 35 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
163	1	1	2	1	1	1	0	1	1	1	1
164	1	1	2	1	1	1	0	1	1	1	1
165	1	1	2	1	1	1	0	1	1	1	1
166	1	1	2	1	1	1	0	1	1	1	1
167	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-12-16

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 35 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
463	1	1	2	1	1	1	0	1	1	1	0
464	1	1	2	1	1	1	0	1	1	1	0
465	1	1	2	1	1	1	0	1	1	1	0
466	1	1	2	1	1	1	0	1	1	1	0
467	1	1	2	1	1	1	0	1	1	1	0
N	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-12-17

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 35 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
163	1	1	1	1	0	0	0	0	1	1	1	
164	1	1	1	1	0	0	0	0	1	1	1	
165	1	1	1	1	0	0	0	0	1	1	1	
166	1	1	1	1	0	0	0	0	1	1	1	
167	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	1.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-12-18

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 35 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
463	1	1	1	1	0	0	0	0	1	1	1	
464	1	1	1	1	0	0	0	0	1	1	1	
465	1	1	1	1	0	0	0	0	1	1	1	
466	1	1	1	1	0	0	1	0	1	1	1	
467	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	

INDIVIDUAL DATA 2-13-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 42 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
101	1	1	1	0	0	1	
102	1	1	1	0	0	1	
103	1	1	1	0	0	1	
104	1	1	1	0	0	1	
105	1	1	1	0	0	1	
106	1	1	1	0	0	1	
107	1	1	1	0	0	1	
108	1	1	1	0	0	1	
109	1	1	1	0	0	1	
110	1	1	1	0	0	1	
111	1	1	1	0	0	1	
112	1	1	1	0	0	1	
N	12	12	12	12	12	12	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-13-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 42 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
201	1	1	1	0	0	1	
202	1	1	1	0	0	1	
203	1	1	1	0	0	1	
204	1	1	1	0	0	1	
205	1	1	1	0	0	1	
206	1	1	1	0	0	1	
207	1	1	1	0	0	1	
208	1	1	1	0	0	1	
209	1	1	1	0	0	1	
210	1	1	1	0	0	1	
211	1	1	1	0	0	1	
212	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-13-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 42 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
301	1	1	1	0	0	1	
302	1	1	1	0	0	1	
303	1	1	1	0	0	1	
304	1	1	1	0	0	1	
305	1	1	1	0	0	1	
306	1	1	1	0	0	1	
307	1	1	1	0	0	1	
308	1	1	1	0	0	1	
309	1	1	1	0	0	1	
310	1	1	1	0	0	1	
311	1	1	1	0	0	1	
312	1	1	1	0	0	1	

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INDIVIDUAL DATA 2-13-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 42 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
401	1	1	1	0	0	1	
402	1	1	1	0	0	1	
403	1	1	1	0	0	1	
404	1	1	1	0	0	1	
405	1	1	1	0	0	1	
406	1	1	1	0	0	1	
407	1	1	1	0	0	1	
408	1	1	1	0	0	1	
409	1	1	1	0	0	1	
410	1	1	1	0	0	1	
411	1	1	1	0	0	1	
412	1	1	1	0	0	1	

N

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INDIVIDUAL DATA 2-13-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 42 of administration

Animal No.	On the hand													
	Ease of Removal		Handling		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
101	1	1	2	1	1	1	1	0	1	1	1	1	1	0
102	1	1	2	1	1	1	1	0	1	1	1	1	1	0
103	1	1	2	1	1	1	1	0	1	1	1	1	1	0
104	1	1	2	1	1	1	1	0	1	1	1	1	1	0
105	1	1	2	1	1	1	1	0	1	1	1	1	1	0
106	1	1	2	1	1	1	1	0	1	1	1	1	1	0
107	1	1	2	1	1	1	1	0	1	1	1	1	1	0
108	1	1	2	1	1	1	1	0	1	1	1	1	1	0
109	1	1	2	1	1	1	1	0	1	1	1	1	1	0
110	1	1	2	1	1	1	1	0	1	1	1	1	1	0
111	1	1	2	1	1	1	1	0	1	1	1	1	1	0
112	1	1	2	1	1	1	1	0	1	1	1	1	1	0
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-13-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 42 of administration

Animal No.	On the hand												
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation	Secretions/ Excretions
201	1	1	2	1	1	1	0	1	1	1	1	1	0
202	1	1	2	1	1	1	0	1	1	1	1	1	0
203	1	1	2	1	1	1	0	1	1	1	1	1	0
204	1	1	2	1	1	1	0	1	1	1	1	1	0
205	1	1	2	1	1	1	0	1	1	1	1	1	0
206	1	1	2	1	1	1	0	1	1	1	1	1	0
207	1	1	2	1	1	1	0	1	1	1	1	1	0
208	1	1	2	1	1	1	0	1	1	1	1	1	0
209	1	1	2	1	1	1	0	1	1	1	1	1	0
210	1	1	2	1	1	1	0	1	1	1	1	1	0
211	1	1	2	1	1	1	0	1	1	1	1	1	0
212	1	1	2	1	1	1	0	1	1	1	1	1	0

INDIVIDUAL DATA 2-13-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 42 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Handling				Fur	Eyes			Pupil size		Salivation	
301	1	1	2	1	1	1	0	1	1	1	1	0
302	1	1	2	1	1	1	0	1	1	1	1	0
303	1	1	2	1	1	1	0	1	1	1	1	0
304	1	1	2	1	1	1	0	1	1	1	1	0
305	1	1	2	1	1	1	0	1	1	1	1	0
306	1	1	2	1	1	1	0	1	1	1	1	0
307	1	1	2	1	1	1	0	1	1	1	1	0
308	1	1	2	1	1	1	0	1	1	1	1	0
309	1	1	2	1	1	1	0	1	1	1	1	0
310	1	1	2	1	1	1	0	1	1	1	1	0
311	1	1	2	1	1	1	0	1	1	1	1	0
312	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-13-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 42 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone		Pilo-erection		Mucous membranes		Skin		Lacrimation	Secretions/ Excretions
	Removal	Handling			Fur	Eyes			Pupil size	Salivation		
401	1	1	2	1	1	1	0	1	1	1	1	0
402	1	1	2	1	1	1	0	1	1	1	1	0
403	1	1	2	1	1	1	0	1	1	1	1	0
404	1	1	2	1	1	1	0	1	1	1	1	0
405	1	1	2	1	1	1	0	1	1	1	1	0
406	1	1	2	1	1	1	0	1	1	1	1	0
407	1	1	2	1	1	1	0	1	1	1	1	0
408	1	1	2	1	1	1	0	1	1	1	1	0
409	1	1	2	1	1	1	0	1	1	1	1	0
410	1	1	2	1	1	1	0	1	1	1	1	0
411	1	1	2	1	1	1	0	1	1	1	1	0
412	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-13-9

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 42 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
101	1	1	1	1	0	0	0	0	1	1	1		
102	1	1	1	1	0	0	0	0	1	1	1		
103	1	1	1	1	0	0	0	0	1	1	1		
104	1	1	1	1	0	0	0	0	1	1	1		
105	1	1	1	1	0	0	0	0	1	1	1		
106	1	1	1	1	0	0	0	0	1	1	1		
107	1	1	1	1	0	0	0	0	1	1	1		
108	1	1	1	1	0	0	0	0	1	1	1		
109	1	1	1	1	0	0	0	0	1	1	1		
110	1	1	1	1	0	0	0	0	1	1	1		
111	1	1	1	1	0	0	0	0	1	1	1		
112	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12
H	0.0000	0.0000	0.0000	0.0000	0.0000	3.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-13-10

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Day 42 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
201	1	1	1	1	0	0	0	0	1	1	1		
202	1	1	1	1	0	0	0	0	1	1	1		
203	1	1	1	1	0	0	0	0	1	1	1		
204	1	1	1	1	0	1	0	0	1	1	1		
205	1	1	1	1	0	0	0	0	1	1	1		
206	1	1	1	1	0	0	0	0	1	1	1		
207	1	1	1	1	0	0	0	0	1	1	1		
208	1	1	1	1	0	0	0	0	1	1	1		
209	1	1	1	1	0	0	0	0	1	1	1		
210	1	1	1	1	0	0	0	0	1	1	1		
211	1	1	1	1	0	0	0	0	1	1	1		
212	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	

INDIVIDUAL DATA 2-13-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Day 42 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
301	1	1	1	1	0	0	0	0	1	1	1		
302	1	1	1	1	0	0	0	0	1	1	1		
303	1	1	1	1	0	0	0	0	1	1	1		
304	1	1	1	1	0	0	0	0	1	1	1		
305	1	1	1	1	0	0	0	0	1	1	1		
306	1	1	1	1	0	0	0	0	1	1	1		
307	1	1	1	1	0	0	0	0	1	1	1		
308	1	1	1	1	0	0	0	0	1	1	1		
309	1	1	1	1	0	0	0	0	1	1	1		
310	1	1	1	1	0	0	0	0	1	1	1		
311	1	1	1	1	0	0	0	0	1	1	1		
312	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	

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INDIVIDUAL DATA 2-13-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 42 of administration

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
401	1	1	1	1	0	0	0	0	1	1	1		
402	1	1	1	1	0	0	0	0	1	1	1		
403	1	1	1	1	0	0	0	0	1	1	1		
404	1	1	1	1	0	0	0	0	1	1	1		
405	1	1	1	1	0	0	0	0	1	1	1		
406	1	1	1	1	0	0	0	0	1	1	1		
407	1	1	1	1	0	0	0	0	1	1	1		
408	1	1	1	1	0	0	0	0	1	1	1		
409	1	1	1	1	0	0	0	0	1	1	1		
410	1	1	1	1	0	0	0	0	1	1	1		
411	1	1	1	1	0	0	0	0	1	1	1		
412	1	1	1	1	0	0	0	0	1	1	1		
N	12	12	12	12	12	12	12	12	12	12	12	12	12

INDIVIDUAL DATA 2-14-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 42 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
151	1	1	1	0	0	1	
152	1	1	1	0	0	1	
153	1	1	1	0	0	1	
154	1	1	1	0	0	1	
155	1	1	1	0	0	1	
156	1	1	1	0	0	1	
157	1	1	1	0	0	1	
158	1	1	1	0	0	1	
159	1	1	1	0	0	1	
160	1	1	1	0	0	1	
161 ^a	#	#	#	#	#	#	
162	1	1	1	0	0	1	
N	11	11	11	11	11	11	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

a: Non-pregnancy.

#: Blank.

INDIVIDUAL DATA 2-14-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 42 of administration

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
251	1	1	1	0	0	1	
252	1	1	1	0	0	1	
253	1	1	1	0	0	1	
254	1	1	1	0	0	1	
255	1	1	1	0	0	1	
256	1	1	1	0	0	1	
257	1	1	1	0	0	1	
258	1	1	1	0	0	1	
259	1	1	1	0	0	1	
260	1	1	1	0	0	1	
261	1	1	1	0	0	1	
262	1	1	1	0	0	1	

N

12

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INDIVIDUAL DATA 2-14-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg

Animal No.	In the cage						
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation	
				Rolling	Repetitive circling		
351	1	1	1	0	0	1	
352	1	1	1	0	0	1	
353	1	1	1	0	0	1	
354	1	1	1	0	0	1	
355	1	1	1	0	0	1	
356	1	1	1	0	0	1	
357	1	1	1	0	0	1	
358	1	1	1	0	0	1	
359	1	1	1	0	0	1	
360	1	1	1	0	0	1	
361	1	1	1	0	0	1	
362	1	1	1	0	0	1	

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-14-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 42 of administration

Animal No.	In the cage					
	Body position/ Posture	Respira- tory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
451	1	1	1	0	0	1
452	1	1	1	0	0	1
453	1	1	1	0	0	1
454	1	1	1	0	0	1
455	1	1	1	0	0	1
456	1	1	1	0	0	1
457	1	1	1	0	0	1
458	1	1	1	0	0	1
459	1	1	1	0	0	1
460	1	1	1	0	0	1
461	1	1	1	0	0	1
462	1	1	1	0	0	1

N 12 12 12 12 12 12

INDIVIDUAL DATA 2-14-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 42 of administration

Animal No.	On the hand										Secretions/Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
151	1	1	2	1	1	1	0	1	1	1	1
152	1	1	2	1	1	0	1	1	1	1	0
153	1	1	2	1	1	0	1	1	1	1	0
154	1	1	2	1	1	0	1	1	1	1	0
155	1	1	2	1	1	0	1	1	1	1	0
156	1	1	2	1	1	0	1	1	1	1	0
157	1	1	2	1	1	0	1	1	1	1	0
158	1	1	2	1	1	0	1	1	1	1	0
159	1	1	2	1	1	0	1	1	1	1	0
160	1	1	2	1	1	0	1	1	1	1	0
161 ^a	#	#	#	#	#	#	#	#	#	#	#
162	1	1	2	1	1	1	0	1	1	1	0
N	11	11	11	11	11	11	11	11	11	11	11
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

a: Non-pregnancy.

#: Blank.

INDIVIDUAL DATA 2-14-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 42 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
251	1	1	2	1	1	1	0	1	1	1	1	0
252	1	1	2	1	1	1	0	1	1	1	1	0
253	1	1	2	1	1	1	0	1	1	1	1	0
254	1	1	2	1	1	1	0	1	1	1	1	0
255	1	1	2	1	1	1	0	1	1	1	1	0
256	1	1	2	1	1	1	0	1	1	1	1	0
257	1	1	2	1	1	1	0	1	1	1	1	0
258	1	1	2	1	1	1	0	1	1	1	1	0
259	1	1	2	1	1	1	0	1	1	1	1	0
260	1	1	2	1	1	1	0	1	1	1	1	0
261	1	1	2	1	1	1	0	1	1	1	1	0
262	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-14-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 42 of administration

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
351	1	1	2	1	1	1	0	1	1	1	1	0
352	1	1	2	1	1	1	0	1	1	1	1	0
353	1	1	2	1	1	1	0	1	1	1	1	0
354	1	1	2	1	1	1	0	1	1	1	1	0
355	1	1	2	1	1	1	0	1	1	1	1	0
356	1	1	2	1	1	1	0	1	1	1	1	0
357	1	1	2	1	1	1	0	1	1	1	1	0
358	1	1	2	1	1	1	0	1	1	1	1	0
359	1	1	2	1	1	1	0	1	1	1	1	0
360	1	1	2	1	1	1	0	1	1	1	1	0
361	1	1	2	1	1	1	0	1	1	1	1	0
362	1	1	2	1	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-14-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 42 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
451	1	1	2	1	1	1	0	1	1	1	1
452	1	1	2	1	1	0	1	1	1	1	0
453	1	1	2	1	1	0	1	1	1	1	0
454	1	1	2	1	1	0	1	1	1	1	0
455	1	1	2	1	1	0	1	1	1	1	0
456	1	1	2	1	1	0	1	1	1	1	0
457	1	1	2	1	1	0	1	1	1	1	0
458	1	1	2	1	1	0	1	1	1	1	0
459	1	1	2	1	1	0	1	1	1	1	0
460	1	1	2	1	1	0	1	1	1	1	0
461	1	1	2	1	1	0	1	1	1	1	0
462	1	1	2	1	1	0	1	1	1	1	0

INDIVIDUAL DATA 2-14-9

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 42 of administration

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Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
151	1	1	1	1	1	0	0	0	1	1	1	
152	1	1	1	1	0	0	0	0	1	1	1	
153	1	1	1	1	0	0	1	0	1	1	1	
154	1	1	1	1	0	0	0	0	1	1	1	
155	1	1	1	1	0	0	0	0	1	1	1	
156	1	1	1	1	0	0	0	0	1	1	1	
157	1	1	1	1	0	0	0	0	1	1	1	
158	1	1	1	1	0	0	0	0	1	1	1	
159	1	1	1	1	0	0	0	0	1	1	1	
160	1	1	1	1	0	0	0	0	1	1	1	
161 ^a	#	#	#	#	#	#	#	#	#	#	#	
162	1	1	1	1	0	0	0	0	1	1	1	
N	11	11	11	11	11	11	11	11	11	11	11	
H	0.0000	0.0000	0.0000	0.0000	2.1412	0.0000	3.2727	0.0000	0.0000	0.0000	0.0000	

a: Non-pregnancy.

#: Blank.

INDIVIDUAL DATA 2-14-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 42 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
251	1	1	1	1	0	0	0	0	1	1	1	
252	1	1	1	1	0	0	0	0	1	1	1	
253	1	1	1	1	0	0	0	0	1	1	1	
254	1	1	1	1	0	0	0	0	1	1	1	
255	1	1	1	1	0	0	0	0	1	1	1	
256	1	1	1	1	1	0	0	0	1	1	1	
257	1	1	1	1	0	0	0	0	1	1	1	
258	1	1	1	1	0	0	0	0	1	1	1	
259	1	1	1	1	0	0	0	0	1	1	1	
260	1	1	1	1	0	0	0	0	1	1	1	
261	1	1	1	1	0	0	0	0	1	1	1	
262	1	1	1	1	0	0	0	0	1	1	1	
N	12	12	12	12	12	12	12	12	12	12	12	

INDIVIDUAL DATA 2-14-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 42 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
351	1	1	1	1	0	0	0	0	1	1	1	
352	1	1	1	1	0	0	0	0	1	1	1	
353	1	1	1	1	0	0	0	0	1	1	1	
354	1	1	1	1	0	0	0	0	1	1	1	
355	1	1	1	1	0	0	0	0	1	1	1	
356	1	1	1	1	0	0	0	0	1	1	1	
357	1	1	1	1	0	0	0	0	1	1	1	
358	1	1	1	1	0	0	0	0	1	1	1	
359	1	1	1	1	0	0	0	0	1	1	1	
360	1	1	1	1	0	0	0	0	1	1	1	
361	1	1	1	1	0	0	0	0	1	1	1	
362	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-14-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 42 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
451	1	1	1	1	0	0	0	0	1	1	1	
452	1	1	1	1	0	0	0	0	1	1	1	
453	1	1	1	1	0	0	0	0	1	1	1	
454	1	1	1	1	0	0	0	0	1	1	1	
455	1	1	1	1	0	0	0	0	1	1	1	
456	1	1	1	1	0	0	0	0	1	1	1	
457	1	1	1	1	0	0	0	0	1	1	1	
458	1	1	1	1	0	0	0	0	1	1	1	
459	1	1	1	1	0	0	0	0	1	1	1	
460	1	1	1	1	0	0	0	0	1	1	1	
461	1	1	1	1	0	0	0	0	1	1	1	
462	1	1	1	1	0	0	0	0	1	1	1	

N 12 12 12 12 12 12 12 12 12 12 12 12 12

INDIVIDUAL DATA 2-14-13

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 42 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
163	1	1	1	0	0	1
164	1	1	1	0	0	1
165	1	1	1	0	0	1
166	1	1	1	0	0	1
167	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-14-14

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 42 of administration

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
463	1	1	1	0	0	1
464	1	1	1	0	0	1
465	1	1	1	0	0	1
466	1	1	1	0	0	1
467	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-14-15

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 42 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
163	1	1	2	1	1	1	0	1	1	1	1
164	1	1	2	1	1	1	0	1	1	1	1
165	1	1	2	1	1	1	0	1	1	1	1
166	1	1	2	1	1	1	0	1	1	1	1
167	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-14-16

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 42 of administration

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
463	1	1	2	1	1	1	0	1	1	1	1
464	1	1	2	1	1	1	0	1	1	1	1
465	1	1	2	1	1	1	0	1	1	1	1
466	1	1	2	1	1	1	0	1	1	1	1
467	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-14-17

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 42 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
163	1	1	1	1	0	0	0	0	1	1	1	
164	1	1	1	1	1	0	0	0	1	1	1	
165	1	1	1	1	0	0	0	0	1	1	1	
166	1	1	1	1	0	0	0	0	1	1	1	
167	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	
H	0.0000	0.0000	0.0000	0.0000	1.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-14-18

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 42 of administration

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
463	1	1	1	1	0	0	0	0	1	1	1	
464	1	1	1	1	0	0	0	0	1	1	1	
465	1	1	1	1	0	0	0	0	1	1	1	
466	1	1	1	1	0	0	0	0	1	1	1	
467	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	

INDIVIDUAL DATA 2-15-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 7 of recovery

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
102	1	1	1	0	0	1
105	1	1	1	0	0	1
107	1	1	1	0	0	1
108	1	1	1	0	0	1
111	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-15-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 7 of recovery

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
405	1	1	1	0	0	1
406	1	1	1	0	0	1
409	1	1	1	0	0	1
410	1	1	1	0	0	1
411	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-15-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 7 of recovery

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
102	1	1	2	1	1	1	0	1	1	1	1
105	1	1	2	1	1	1	0	1	1	1	1
107	1	1	2	1	1	1	0	1	1	1	1
108	1	1	2	1	1	1	0	1	1	1	1
111	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-15-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 7 of recovery

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
405	1	1	2	1	1	1	0	1	1	1	1	0
406	1	1	2	1	1	1	0	1	1	1	1	0
409	1	1	2	1	1	1	0	1	1	1	1	0
410	1	1	2	1	1	1	0	1	1	1	1	0
411	1	1	2	1	1	1	0	1	1	1	1	0
N	5	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-15-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 7 of recovery

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
102	1	1	1	1	0	0	0	0	1	1	1		
105	1	1	1	1	0	0	0	0	1	1	1		
107	1	1	1	1	0	0	0	0	1	1	1		
108	1	1	1	1	0	0	0	0	1	1	1		
111	1	1	1	1	0	0	0	0	1	1	1		
N	5	5	5	5	5	5	5	5	5	5	5		
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000		

INDIVIDUAL DATA 2-15-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 7 of recovery

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
405	1	1	1	1	0	0	0	0	1	1	1	
406	1	1	1	1	0	0	0	0	1	1	1	
409	1	1	1	1	0	0	0	0	1	1	1	
410	1	1	1	1	0	0	0	0	1	1	1	
411	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	

INDIVIDUAL DATA 2-16-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 7 of recovery

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
163	1	1	1	0	0	1
164	1	1	1	0	0	1
165	1	1	1	0	0	1
166	1	1	1	0	0	1
167	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-16-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 7 of recovery

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
463	1	1	1	0	0	1
464	1	1	1	0	0	1
465	1	1	1	0	0	1
466	1	1	1	0	0	1
467	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-16-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 7 of recovery

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
163	1	1	2	1	1	1	0	1	1	1	1
164	1	1	2	1	1	1	0	1	1	1	1
165	1	1	2	1	1	1	0	1	1	1	1
166	1	1	2	1	1	1	0	1	1	1	1
167	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-16-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 7 of recovery

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
463	1	1	2	1	1	1	0	1	1	1	1
464	1	1	2	1	1	1	0	1	1	1	1
465	1	1	2	1	1	1	0	1	1	1	1
466	1	1	2	1	1	1	0	1	1	1	1
467	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-16-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 7 of recovery

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
163	1	1	1	1	0	0	0	0	1	1	1	
164	1	1	1	1	0	0	0	0	1	1	1	
165	1	1	1	1	0	0	0	0	1	1	1	
166	1	1	1	1	0	0	0	0	1	1	1	
167	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-16-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 7 of recovery

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
463	1	1	1	1	0	0	0	0	1	1	1	
464	1	1	1	1	0	0	0	0	1	1	1	
465	1	1	1	1	0	0	0	0	1	1	1	
466	1	1	1	1	0	0	0	0	1	1	1	
467	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	

INDIVIDUAL DATA 2-17-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 14 of recovery

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
102	1	1	1	0	0	1
105	1	1	1	0	0	1
107	1	1	1	0	0	1
108	1	1	1	0	0	1
111	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-17-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 14 of recovery

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
405	1	1	1	0	0	1
406	1	1	1	0	0	1
409	1	1	1	0	0	1
410	1	1	1	0	0	1
411	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-17-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 14 of recovery

Animal No.	On the hand											
	Ease of Removal		Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes		Skin	Pupil size	Lacrimation	Salivation
102	1	1	2	1	1	1	0	1	1	1	1	0
105	1	1	2	1	1	1	0	1	1	1	1	0
107	1	1	2	1	1	1	0	1	1	1	1	0
108	1	1	2	1	1	1	0	1	1	1	1	0
111	1	1	2	1	1	1	0	1	1	1	1	0
N	5	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-17-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 14 of recovery

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
405	1	1	2	1	1	1	0	1	1	1	1
406	1	1	2	1	1	1	0	1	1	1	1
409	1	1	2	1	1	1	0	1	1	1	1
410	1	1	2	1	1	1	0	1	1	1	1
411	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-17-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Day 14 of recovery

Animal No.	In the open-field						Stereotype				Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression		
102	1	1	1	1	1	0	0	0	1	1	1		
105	1	1	1	1	0	0	0	0	1	1	1		
107	1	1	1	1	1	0	0	0	1	1	1		
108	1	1	1	1	0	0	0	0	1	1	1		
111	1	1	1	1	0	0	0	0	1	1	1		
N	5	5	5	5	5	5	5	5	5	5	5		
H	0.0000	0.0000	0.0000	0.0000	2.2500	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000		

INDIVIDUAL DATA 2-17-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Day 14 of recovery

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
405	1	1	1	1	0	0	0	0	1	1	1	
406	1	1	1	1	0	0	0	0	1	1	1	
409	1	1	1	1	0	0	0	0	1	1	1	
410	1	1	1	1	0	0	0	0	1	1	1	
411	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	

INDIVIDUAL DATA 2-18-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 14 of recovery

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
163	1	1	1	0	0	1
164	1	1	1	0	0	1
165	1	1	1	0	0	1
166	1	1	1	0	0	1
167	1	1	1	0	0	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-18-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 14 of recovery

Animal No.	In the cage					
	Body position/ Posture	Respiratory pattern	Tremor/ Convulsion	Stereotype		Bizarre behavior Biting/Selfmutilation
				Rolling	Repetitive circling	
463	1	1	1	0	0	1
464	1	1	1	0	0	1
465	1	1	1	0	0	1
466	1	1	1	0	0	1
467	1	1	1	0	0	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 2-18-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 14 of recovery

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
163	1	1	2	1	1	1	0	1	1	1	1
164	1	1	2	1	1	1	0	1	1	1	1
165	1	1	2	1	1	1	0	1	1	1	1
166	1	1	2	1	1	1	0	1	1	1	1
167	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 2-18-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 14 of recovery

Animal No.	On the hand										Secretions/ Excretions
	Ease of Removal	Handling	Muscle tone	Pilo-erection	Fur	Eyes	Mucous membranes	Skin	Pupil size	Lacrimation	
463	1	1	2	1	1	1	0	1	1	1	1
464	1	1	2	1	1	1	0	1	1	1	1
465	1	1	2	1	1	1	0	1	1	1	1
466	1	1	2	1	1	1	0	1	1	1	1
467	1	1	2	1	1	1	0	1	1	1	1
N	5	5	5	5	5	5	5	5	5	5	5

INDIVIDUAL DATA 2-18-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 14 of recovery

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
163	1	1	1	1	0	0	0	0	1	1	1	
164	1	1	1	1	0	0	0	0	1	1	1	
165	1	1	1	1	0	0	0	0	1	1	1	
166	1	1	1	1	0	0	0	0	1	1	1	
167	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	

INDIVIDUAL DATA 2-18-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Detailed clinical observation, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 14 of recovery

Animal No.	In the open-field						Stereotype			Bizarre behavior		
	Gait	Co-ordination of movement	Reactivity to environmental stimuli	Searching	Urination	Defecation	Excessive grooming	Unusual head movement	Walking backward	Vocalization	Aggression	
463	1	1	1	1	0	0	0	0	1	1	1	
464	1	1	1	1	0	0	0	0	1	1	1	
465	1	1	1	1	0	0	0	0	1	1	1	
466	1	1	1	1	0	0	0	0	1	1	1	
467	1	1	1	1	0	0	0	0	1	1	1	
N	5	5	5	5	5	5	5	5	5	5	5	

INDIVIDUAL DATA 3-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Week 6 of administration

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
102	4	2	1	2	1	1
105	4	2	1	2	1	1
107	4	2	1	2	1	1
108	4	2	1	2	1	1
111	4	2	1	2	1	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 3-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Week 6 of administration

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
202	4	2	1	2	1	1
207	4	2	1	2	1	1
209	4	2	1	2	1	1
210	4	2	1	2	1	1
212	4	2	1	2	1	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 3-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Week 6 of administration

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
301	4	2	1	2	1	1
304	4	2	1	2	1	1
307	4	2	1	2	1	1
309	4	2	1	2	1	1
310	4	2	1	2	1	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 3-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Week 6 of administration

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
405	4	2	1	2	1	1
406	4	2	1	2	1	1
409	4	2	1	2	1	1
410	4	2	1	2	1	1
411	4	2	1	2	1	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 3-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Week 6 of administration

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
163	4	2	1	2	1	1
164	4	2	1	2	1	1
165	4	2	1	2	1	1
166	4	2	1	2	1	1
167	4	2	1	2	1	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 3-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Week 6 of administration

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
463	4	2	1	2	1	1
464	4	2	1	2	1	1
465	4	2	1	2	1	1
466	4	2	1	2	1	1
467	4	2	1	2	1	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 3-3-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 4 of lactation

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
152	4	2	1	2	1	1
153	4	2	1	2	1	1
155	4	2	1	2	1	1
156	4	2	1	2	1	1
162	4	2	1	2	1	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 3-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 4 of lactation

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
251	4	2	1	2	1	1
253	4	2	1	2	1	1
255	4	2	1	2	1	1
256	4	2	1	2	1	1
262	4	2	1	2	1	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 3-3-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 4 of lactation

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
351	4	2	1	2	1	1
352	4	2	1	2	1	1
353	4	2	1	2	1	1
355	4	2	1	2	1	1
361	4	2	1	2	1	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 3-3-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 4 of lactation

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
451	4	2	1	2	1	1
453	4	2	1	2	1	1
455	4	2	1	2	1	1
457	4	2	1	2	1	1
462	4	2	1	2	1	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 3-4-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Week 2 of recovery

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
102	4	2	1	2	1	1
105	4	2	1	2	1	1
107	4	2	1	2	1	1
108	4	2	1	2	1	1
111	4	2	1	2	1	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 3-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Week 2 of recovery

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
405	4	2	1	2	1	1
406	4	2	1	2	1	1
409	4	2	1	2	1	1
410	4	2	1	2	1	1
411	4	2	1	2	1	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 3-5-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Week 2 of recovery

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
163	4	2	1	2	1	1
164	4	2	1	2	1	1
165	4	2	1	2	1	1
166	4	2	1	2	1	1
167	4	2	1	2	1	1
N	5	5	5	5	5	5
H	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000

INDIVIDUAL DATA 3-5-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Week 2 of recovery

Animal No.	On the desk					
	Reactivity					
	Visual	Touch	Auditory	Pain	Proprio- ceptive	Righting reflex
463	4	2	1	2	1	1
464	4	2	1	2	1	1
465	4	2	1	2	1	1
466	4	2	1	2	1	1
467	4	2	1	2	1	1
N	5	5	5	5	5	5

INDIVIDUAL DATA 3-6-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Week 6 of administration

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
102	1461.7	539.0	579	155	263	57	72	116	1242
105	1773.0	723.0	829	568	340	311	254	216	2518
107	1753.7	672.3	441	258	153	134	54	61	1101
108	1624.0	577.0	533	425	247	96	0	39	1340
111	1834.7	712.7	756	320	128	5	124	234	1567
N	5	5	5	5	5	5	5	5	5
MEAN	1689.42	644.80	627.6	345.2	226.2	120.6	100.8	133.2	1553.6
S.D.	148.63	82.57	160.6	158.4	86.2	116.7	96.4	88.6	565.2
S.E.	66.47	36.93	71.8	70.9	38.6	52.2	43.1	39.6	252.8
M/C	1.5646	0.6234	2.0135	6.2137	6.0905	4.9043	1.9359	4.3092	11.2104 *
F	0.8410	0.4363	0.7789	0.9119	0.8024	1.6034	0.4850	0.7630	
H									8.1086 *

INDIVIDUAL DATA 3-6-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Week 6 of administration

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
202	1717.7	640.3	398	322	248	117	98	66	1249
207	1628.7	601.7	337	121	33	3	0	3	497
209	1799.0	765.7	494	274	104	66	146	85	1169
210	1668.0	503.3	703	239	163	0	0	4	1109
212	1153.3	684.0	192	88	116	109	125	142	772
N	5	5	5	5	5	5	5	5	5
MEAN	1593.34	639.00	424.8	208.8	132.8	59.0	73.8	60.0	959.2
S.D.	254.08	97.32	190.3	100.3	79.5	56.0	69.5	58.7	315.9
S.E.	113.63	43.52	85.1	44.9	35.5	25.0	31.1	26.2	141.3
U								4.0000	

INDIVIDUAL DATA 3-6-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Week 6 of administration

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
301	1457.0	557.3	500	223	114	149	114	57	1157
304	1447.0	537.3	287	78	91	39	13	21	529
307	1486.7	634.7	321	198	109	77	70	39	814
309	1298.7	566.3	1069	692	583	408	340	378	3470
310	1936.3	722.3	474	232	89	67	118	80	1060
N	5	5	5	5	5	5	5	5	5
MEAN	1525.14	603.58	530.2	284.6	197.2	148.0	131.0	115.0	1406.0
S.D.	241.12	75.81	315.2	236.0	215.9	150.9	124.3	148.6	1179.1
S.E.	107.83	33.90	140.9	105.5	96.6	67.5	55.6	66.5	527.3
U								6.0000	

INDIVIDUAL DATA 3-6-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Week 6 of administration

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
405	1254.0	631.0	791	446	113	101	159	134	1744
406	1633.0	544.3	617	292	320	216	64	155	1664
409	1180.3	647.7	417	358	270	235	65	50	1395
410	1353.0	513.3	760	359	303	194	201	168	1985
411	1879.0	654.3	357	291	263	255	187	219	1572
N	5	5	5	5	5	5	5	5	5
MEAN	1459.86	598.12	588.4	349.2	253.8	200.2	135.2	145.2	1672.0
S.D.	290.52	64.78	196.4	63.6	82.1	59.9	66.3	61.7	217.9
S.E.	129.92	28.97	87.8	28.5	36.7	26.8	29.6	27.6	97.5

U 6.0000

INDIVIDUAL DATA 3-7-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Week 6 of administration

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
163	911.0	466.3	603	396	334	128	111	73	1645
164	1158.3	482.7	835	872	704	486	399	209	3505
165	1532.3	524.0	1042	794	649	351	188	98	3122
166	737.3	486.7	648	684	365	230	212	259	2398
167	1194.3	539.3	681	727	701	528	179	102	2918
N	5	5	5	5	5	5	5	5	5
MEAN	1106.64	499.80	761.8	694.6	550.6	344.6	217.8	148.2	2717.6
S.D.	302.58	30.55	179.3	181.4	185.2	168.6	108.0	81.1	720.5
S.E.	135.32	13.66	80.2	81.1	82.8	75.4	48.3	36.3	322.2
M/C	4.9252 *	0.0812	0.0472	0.6277	0.2297	0.3419	0.1436	0.0343	0.2156
F		3.1609	5.1090 †	2.6364	7.2358 *	5.0330 †	0.6345	0.5769	4.4672 †
H		0.5345							

INDIVIDUAL DATA 3-7-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Week 6 of administration

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
463	1209.0	446.3	311	192	156	78	114	45	896
464	1076.7	457.7	733	888	478	333	307	253	2992
465	1155.7	508.0	401	312	137	0	0	63	913
466	1293.0	415.7	678	563	349	123	281	133	2127
467	1227.0	485.0	324	313	224	141	83	42	1127
N	5	5	5	5	5	5	5	5	5
MEAN	1192.28	462.54	489.4	453.6	268.8	135.0	157.0	107.2	1611.0
S.D.	81.13	35.54	201.2	277.9	143.4	123.3	132.1	89.4	922.8
S.E.	36.28	15.89	90.0	124.3	64.1	55.2	59.1	40.0	412.7
t'			2.2603		2.6899 *	2.2434		2.1136	

INDIVIDUAL DATA 3-8-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 4 of lactation

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
152	1251.3	787.0	612	75	317	133	278	224	1639
153	1354.3	462.0	277	18	0	11	5	0	311
155	1265.3	822.7	197	118	1	0	0	0	316
156	1511.0	611.3	554	118	115	23	351	170	1331
162	1279.3	533.3	179	38	17	26	2	7	269
N	5	5	5	5	5	5	5	5	5
MEAN	1332.24	643.26	363.8	73.4	90.0	38.6	127.2	80.2	773.2
S.D.	107.56	157.18	204.5	45.6	135.6	53.8	172.9	108.4	659.1
S.E.	48.10	70.29	91.5	20.4	60.6	24.0	77.3	48.5	294.8
M/C	4.4691	10.0794 *	0.2396	3.9447	4.8840	1.5258	3.4815	2.4841	0.5889
F	1.2324		0.2601	1.5983	0.2466	2.3582	0.1988	0.5215	0.3891
H		2.1771							

INDIVIDUAL DATA 3-8-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 4 of lactation

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
251	1172.3	537.0	306	86	31	14	0	0	437
253	1195.3	610.3	440	83	62	58	0	12	655
255	1132.3	620.0	121	86	146	114	108	11	586
256	1396.0	654.0	688	301	153	219	248	197	1806
262	998.7	705.3	260	68	120	175	158	69	850
N	5	5	5	5	5	5	5	5	5
MEAN	1178.92	625.32	363.0	124.8	102.4	116.0	102.8	57.8	866.8
S.D.	143.26	61.79	214.5	98.8	53.6	83.5	106.4	82.4	545.6
S.E.	64.07	27.63	95.9	44.2	24.0	37.3	47.6	36.8	244.0

INDIVIDUAL DATA 3-8-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 4 of lactation

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
351	1048.0	647.3	625	32	121	43	0	0	821
352	1451.3	681.0	267	108	47	132	31	9	594
353	767.0	609.0	632	287	262	312	499	424	2416
355	1296.0	636.0	573	138	115	112	61	150	1149
361	1231.0	658.0	186	157	87	85	0	0	515
N	5	5	5	5	5	5	5	5	5
MEAN	1158.66	646.26	456.6	144.4	126.4	136.8	118.2	116.6	1099.0
S.D.	262.38	26.64	213.2	92.9	81.2	103.5	214.4	183.3	776.3
S.E.	117.34	11.92	95.4	41.5	36.3	46.3	95.9	82.0	347.2

INDIVIDUAL DATA 3-8-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 4 of lactation

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
451	1329.3	677.7	554	275	72	146	5	165	1217
453	1198.3	602.3	278	79	0	120	2	6	485
455	1119.0	332.7	80	40	17	86	58	39	320
457	1107.0	623.0	649	345	352	253	14	366	1979
462	1065.3	524.7	669	314	304	238	211	211	1947
N	5	5	5	5	5	5	5	5	5
MEAN	1163.78	552.08	446.0	210.6	149.0	168.6	58.0	157.4	1189.6
S.D.	104.30	134.36	257.3	140.8	166.4	73.5	88.4	144.4	782.7
S.E.	46.65	60.09	115.1	63.0	74.4	32.9	39.6	64.6	350.0

INDIVIDUAL DATA 3-9-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Week 2 of recovery

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
102	1641.3	726.0	350	257	129	135	143	56	1070
105	1603.0	790.3	509	381	219	147	101	181	1538
107	1467.3	788.0	182	77	52	0	0	0	311
108	1524.7	731.7	224	261	106	88	16	5	700
111	1809.7	778.7	368	210	111	128	14	44	875
N	5	5	5	5	5	5	5	5	5
MEAN	1609.20	762.94	326.6	237.2	123.4	99.6	54.8	57.2	898.8
S.D.	130.91	31.49	129.4	109.6	60.7	59.9	63.4	73.3	453.7
S.E.	58.54	14.08	57.9	49.0	27.1	26.8	28.4	32.8	202.9
M/C	0.0038	3.3995	0.0113	0.7316	0.7749	0.0013	0.5495	2.3662	0.8931
F	1.1150	10.6582 *	0.2510	2.1424	4.0629 †	1.5547	1.3818	0.9125	1.9694

INDIVIDUAL DATA 3-9-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Functional test ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Week 2 of recovery

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
405	1630.3	746.3	546	424	165	157	19	73	1384
406	1585.0	546.0	403	288	202	151	48	125	1217
409	1289.0	655.7	250	351	227	103	153	106	1190
410	1578.3	522.0	213	242	134	85	96	47	817
411	1519.0	644.3	432	305	211	236	257	105	1546
N	5	5	5	5	5	5	5	5	5
MEAN	1520.32	622.86	368.8	322.0	187.8	146.4	114.6	91.2	1230.8
S.D.	135.23	90.63	136.9	69.1	37.7	58.8	94.4	31.0	272.0
S.E.	60.48	40.53	61.2	30.9	16.9	26.3	42.2	13.9	121.6
t'	3.2647 *			2.0157					

INDIVIDUAL DATA 3-10-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Week 2 of recovery

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
163	1178.3	700.3	829	476	353	166	192	72	2088
164	1175.3	620.7	336	342	311	143	88	119	1339
165	1520.7	760.3	342	185	146	110	78	71	932
166	1013.7	580.0	914	619	265	468	456	467	3189
167	1141.0	603.0	432	86	71	68	0	0	657
N	5	5	5	5	5	5	5	5	5
MEAN	1205.80	652.86	570.6	341.6	229.2	191.0	162.8	145.8	1641.0
S.D.	188.38	75.20	278.9	215.0	117.5	159.2	177.6	184.5	1019.6
S.E.	84.25	33.63	124.7	96.1	52.6	71.2	79.4	82.5	456.0
M/C	0.2402	0.1658	0.1964	0.0700	0.1517	0.0587	8.5868 **	0.4614	0.2464
F	0.0967	13.2889 **	0.1189	0.2884	1.5162	0.0344		0.1649	0.4892
H							2.8098 †		

INDIVIDUAL DATA 3-10-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Functional test, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Week 2 of recovery

Animal No.	Grip strength (g)		Motor activity measurements						
	Forelimb	Hindlimb	0'-10'	10'-20'	20'-30'	30'-40'	40'-50'	50'-60'	0'-60'
463	1278.0	506.3	215	88	8	0	1	0	312
464	1294.7	440.0	558	381	231	202	61	187	1620
465	969.7	562.7	554	155	156	130	46	42	1083
466	1252.0	426.3	821	545	235	465	0	290	2356
467	1069.3	542.0	431	197	99	58	33	6	824
N	5	5	5	5	5	5	5	5	5
MEAN	1172.74	495.46	515.8	273.2	145.8	171.0	28.2	105.0	1239.0
S.D.	145.05	60.55	220.3	186.8	95.5	181.0	27.2	128.2	782.4
S.E.	64.87	27.08	98.5	83.5	42.7	81.0	12.1	57.3	349.9

t' 3.6454 **

4.5000

INDIVIDUAL DATA 4-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Body weight ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg

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Animal No.	Body weight (g) on administration period (day)										Body weight gain	
	1	3	5	7	10	14	21	28	35	42	Days 1-42	(g)
101	359	365	374	383	391	417	436	461	482	500	141	39.28
102	369	374	383	395	403	422	439	466	490	495	126	34.15
103	375	379	388	399	407	433	441	462	492	512	137	36.53
104	379	384	389	399	406	423	441	464	483	502	123	32.45
105	383	387	399	409	413	433	447	467	487	497	114	29.77
106	387	392	399	404	409	420	439	453	465	473	86	22.22
107	392	401	410	418	420	439	447	465	493	493	101	25.77
108	399	409	419	428	433	455	471	498	534	546	147	36.84
109	399	418	432	448	459	496	518	552	590	609	210	52.63
110	416	425	440	449	461	477	500	528	547	562	146	35.10
111	409	418	430	445	456	473	496	527	545	557	148	36.19
112	430	435	447	460	461	487	505	547	571	584	154	35.81
N	12	12	12	12	12	12	12	12	12	12	12	12
MEAN	391.4	398.9	409.2	419.8	426.6	447.9	465.0	490.8	514.9	527.5	136.1	34.728
S.D.	20.4	22.2	24.2	25.5	26.1	28.5	31.1	37.4	40.5	42.7	31.1	7.502
S.E.	5.9	6.4	7.0	7.4	7.5	8.2	9.0	10.8	11.7	12.3	9.0	2.166
M/C	0.6969	1.8337	3.2891	2.9844	3.3762	3.1302	4.2177	3.1955	6.6715	6.0692	3.0873	1.7167
F	0.0747	0.1101	0.0188	0.0435	0.0947	0.5387	1.4132	2.4677 †	3.0735 *	3.5799 *	6.4554 **	6.6025 **

INDIVIDUAL DATA 4-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 4 mg/kg

Animal No.	Body weight (g) on administration period (day)										Body weight gain	
	1	3	5	7	10	14	21	28	35	42	Days 1-42	(g)
201	365	375	386	392	405	422	447	475	494	502	137	37.53
202	379	387	398	409	422	443	469	500	519	537	158	41.69
203	373	389	391	401	408	421	448	554	495	507	134	35.92
204	384	396	400	408	410	425	435	477	509	532	148	38.54
205	382	393	401	411	418	442	462	485	512	529	147	38.48
206	394	394	403	409	414	429	452	480	498	508	114	28.93
207	394	399	416	424	430	456	466	494	514	514	120	30.46
208	398	410	422	434	435	456	466	482	504	521	123	30.90
209	408	417	425	438	444	466	477	510	538	532	124	30.39
210	400	414	424	431	441	464	485	513	536	552	152	38.00
211	415	418	414	436	453	471	497	523	549	571	156	37.59
212	418	423	430	437	441	461	454	487	503	513	95	22.73
N	12	12	12	12	12	12	12	12	12	12	12	12
MEAN	392.5	401.3	409.2	419.2	426.8	446.3	463.2	498.3	514.3	526.5	134.0	34.263
S.D.	16.5	14.9	14.5	16.0	16.1	18.4	17.5	23.3	18.0	20.2	19.3	5.493
S.E.	4.8	4.3	4.2	4.6	4.6	5.3	5.0	6.7	5.2	5.8	5.6	1.586
t'								0.6477	0.0553	0.0782	0.2122	0.1854

INDIVIDUAL DATA 4-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg

Animal No.	Body weight (g) on administration period (day)										Body weight gain	
	1	3	5	7	10	14	21	28	35	42	Days 1-42	(g)
301	356	364	380	393	406	429	461	490	511	518	162	45.51
302	367	375	382	387	394	406	421	444	464	480	113	30.79
303	373	385	392	402	412	430	454	479	502	523	150	40.21
304	381	385	404	410	423	440	463	493	518	539	158	41.47
305	383	391	404	407	421	436	458	479	499	508	125	32.64
306	388	395	405	416	426	440	461	479	502	520	132	34.02
307	393	402	408	414	425	438	453	485	502	516	123	31.30
308	399	408	417	433	444	466	477	506	527	543	144	36.09
309	398	404	411	421	425	450	461	483	502	504	106	26.63
310	403	410	424	434	443	456	468	495	522	526	123	30.52
311	413	420	433	442	446	469	490	511	533	554	141	34.14
312	417	425	437	447	463	491	513	550	583	606	189	45.32
N	12	12	12	12	12	12	12	12	12	12	12	12
MEAN	389.3	397.0	408.1	417.2	427.3	445.9	465.0	491.2	513.8	528.1	138.8	35.720
S.D.	18.3	18.0	18.0	18.9	19.1	22.3	22.1	25.0	28.1	31.2	23.5	6.103
S.E.	5.3	5.2	5.2	5.5	5.5	6.4	6.4	7.2	8.1	9.0	6.8	1.762
t'								0.0288	0.0967	0.0456	0.2801	0.3953

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INDIVIDUAL DATA 4-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg

Animal No.	Body weight (g) on administration period (day)										Body weight gain	
	1	3	5	7	10	14	21	28	35	42	Days 1-42	(g)
401	369	379	388	398	405	420	435	448	458	464	95	25.75
402	372	378	388	394	401	417	425	450	478	492	120	32.26
403	381	386	400	412	423	426	441	453	456	464	83	21.78
404	381	388	396	407	414	427	446	455	466	484	103	27.03
405	378	386	395	405	413	434	443	466	493	492	114	30.16
406	386	394	404	417	421	440	438	456	474	482	96	24.87
407	398	406	408	418	415	421	425	439	454	464	66	16.58
408	396	401	409	417	424	444	462	492	508	523	127	32.07
409	405	408	416	424	429	438	449	470	480	485	80	19.75
410	405	411	421	427	430	445	455	477	495	510	105	25.93
411	413	423	429	446	446	461	470	486	503	503	90	21.79
412	419	429	439	449	460	477	494	532	548	555	136	32.46
N	12	12	12	12	12	12	12	12	12	12	12	12
MEAN	391.9	399.1	407.8	417.8	423.4	437.5	448.6	468.7	484.4	493.2	101.3	25.869
S.D.	16.4	16.7	16.0	17.0	16.6	17.8	19.6	25.6	27.1	26.8	20.5	5.228
S.E.	4.7	4.8	4.6	4.9	4.8	5.1	5.7	7.4	7.8	7.7	5.9	1.509
t'								1.9142	2.5287 *	2.6847 *	3.5478 **	3.5315 **

408

INDIVIDUAL DATA 4-1-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 0 mg/kg

Animal No.	Body weight (g) on recovery period (day)		Body weight gain	
	7	14	Day 42-R14 (g)	(%)
102	512	517	22	4.44
105	514	520	23	4.63
107	521	524	31	6.29
108	562	572	26	4.76
111	576	579	22	3.95
N	5	5	5	5
MEAN	537.0	542.4	24.8	4.814
S.D.	29.8	30.4	3.8	0.881
S.E.	13.3	13.6	1.7	0.394
M/C	1.4331	0.8173	5.3468 *	4.6443 *
F	3.4251	1.9350		
H		0.1006	0.0982	

INDIVIDUAL DATA 4-1-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 100 mg/kg

Animal No.	Body weight (g) on recovery period (day)		Body weight gain	
	7	14	Day 42-R14 (g)	(%)
405	524	540	48	9.76
406	488	490	8	1.66
409	498	518	33	6.8
410	516	528	18	3.53
411	520	525	22	4.37
N	5	5	5	5
MEAN	509.2	520.2	25.8	5.224
S.D.	15.5	18.7	15.3	3.137
S.E.	6.9	8.3	6.8	1.403

INDIVIDUAL DATA 4-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Before gestation

Animal No.	Body weight (g) on administration period (day)						Body weight gain	
	1	3	5	7	10	14	Days 1-14 (g)	(%)
151	235	229	239	237	251	255	20	8.51
152	234	228	238	237	248	248	14	5.98
153	220	229	221	233	234	240	20	9.09
154	239	241	246	248	246	256	17	7.11
155	225	239	234	252	249	257	32	14.22
156	236	247	243	255	252	262	26	11.02
157	247	245	252	255	245	254	7	2.83
158	239	250	251	262	266	271	32	13.39
159	257	253	260	261	255	261	4	1.56
160	263	256	270	274	277	286	23	8.75
161	258	260	269	273	270	278	20	7.75
162	266	266	275	275	290	300	34	12.78
N	12	12	12	12	12	12	12	12
MEAN	243.3	245.3	249.8	255.2	256.9	264.0	20.8	8.583
S.D.	14.9	12.5	16.3	14.6	15.8	17.0	9.5	3.940
S.E.	4.3	3.6	4.7	4.2	4.6	4.9	2.7	1.137
M/C	2.9392	0.5405	5.9796	0.6305	2.6662	4.5668	3.2855	2.6682
F	0.1282	0.0709	0.5266	0.2612	0.0612	0.0540	0.1205	0.1402

INDIVIDUAL DATA 4-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Before gestation

Animal No.	Body weight (g) on administration period (day)						Body weight gain	
	1	3	5	7	10	14	Days 1-14 (g)	(%)
251	235	233	248	243	254	262	27	11.49
252	226	234	232	243	245	253	27	11.95
253	251	238	254	244	246	260	9	3.59
254	243	242	248	244	255	262	19	7.82
255	241	242	249	238	252	258	17	7.05
256	238	246	243	254	247	256	18	7.56
257	234	245	241	257	252	262	28	11.97
258	242	248	246	255	257	265	23	9.50
259	239	247	244	261	255	263	24	10.04
260	246	255	249	263	263	270	24	9.76
261	252	267	258	276	276	283	31	12.30
262	259	262	264	270	278	280	21	8.11
N	12	12	12	12	12	12	12	12
MEAN	242.2	246.6	248.0	254.0	256.7	264.5	22.3	9.262
S.D.	9.0	10.4	8.2	12.0	10.7	9.1	6.0	2.577
S.E.	2.6	3.0	2.4	3.5	3.1	2.6	1.7	0.744

INDIVIDUAL DATA 4-2-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Before gestation

Animal No.	Body weight (g) on administration period (day)						Body weight gain	
	1	3	5	7	10	14	Days 1-14 (g)	(%)
351	229	221	237	232	244	251	22	9.61
352	226	227	231	236	237	243	17	7.52
353	245	235	250	242	258	264	19	7.76
354	222	233	227	240	242	251	29	13.06
355	249	242	252	250	255	262	13	5.22
356	244	253	244	257	255	260	16	6.56
357	242	251	246	259	260	268	26	10.74
358	236	248	240	251	253	259	23	9.75
359	237	250	245	260	258	266	29	12.24
360	249	255	255	263	265	274	25	10.04
361	258	255	259	263	261	272	14	5.43
362	254	263	252	270	273	283	29	11.42
N	12	12	12	12	12	12	12	12
MEAN	240.9	244.4	244.8	251.9	255.1	262.8	21.8	9.113
S.D.	11.2	12.8	9.7	12.1	10.1	11.1	5.9	2.601
S.E.	3.2	3.7	2.8	3.5	2.9	3.2	1.7	0.751

INDIVIDUAL DATA 4-2-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Before gestation

Animal No.	Body weight (g) on administration period (day)						Body weight gain	
	1	3	5	7	10	14	Days 1-14 (g)	(%)
451	236	226	242	238	249	261	25	10.59
452	228	230	236	239	236	245	17	7.46
453	230	235	236	243	245	256	26	11.30
454	245	243	248	252	250	253	8	3.27
455	229	238	236	248	249	253	24	10.48
456	246	244	252	256	248	257	11	4.47
457	258	248	265	262	269	276	18	6.98
458	252	254	257	264	263	269	17	6.75
459	242	254	252	264	263	269	27	11.16
460	250	260	258	272	270	281	31	12.40
461	249	261	258	271	272	279	30	12.05
462	258	257	261	267	269	276	18	6.98
N	12	12	12	12	12	12	12	12
MEAN	243.6	245.8	250.1	256.3	256.9	264.6	21.0	8.658
S.D.	10.7	11.8	10.4	12.2	12.0	12.0	7.3	3.061
S.E.	3.1	3.4	3.0	3.5	3.5	3.5	2.1	0.884

INDIVIDUAL DATA 4-3-1

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Gestation

Animal No.	Body weight (g) on gestation period (day)									Body weight gain	
	0	1	3	5	7	10	14	17	20	Days 0-20	(g)
151	269	280	291	305	313	326	353	384	445	176	65.43
152	258	265	273	284	287	304	322	353	405	147	56.98
153	246	253	272	281	292	313	331	367	419	173	70.33
154	276	286	303	309	329	346	368	405	457	181	65.58
155	262	267	281	292	304	316	339	372	424	162	61.83
156	269	274	293	300	311	328	350	381	432	163	60.59
157	267	274	285	292	301	315	332	359	390	123	46.07
158	273	279	289	297	301	317	343	372	431	158	57.88
159	277	282	294	308	321	339	368	404	453	176	63.54
160	296	299	314	328	337	351	381	414	465	169	57.09
161 ^a	(274)	(284)	(295)	(304)	(307)	(317)	(316)	(318)	(312)	#	#
162	308	308	323	334	335	346	372	409	460	152	49.35
- 415 -	N	11	11	11	11	11	11	11	11	11	11
	MEAN	272.8	278.8	292.5	302.7	311.9	327.4	350.8	383.6	434.6	161.8
	S.D.	17.1	15.4	15.8	16.7	16.9	15.9	19.3	21.3	24.0	16.7
	S.E.	5.2	4.6	4.8	5.0	5.1	4.8	5.8	6.4	7.2	5.0
	M/C	4.6201	3.4525	0.5722	1.6232	1.5895	0.8148	1.0018	2.3565	4.2878	4.8972
	F	1.7618	1.2083	0.7845	1.5196	1.6566	1.2019	1.6432	1.8987	1.9884	1.2889

a: Non-pregnancy (values in parentheses are excluded from calculation).

#: Blank.

INDIVIDUAL DATA 4-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Gestation

Animal No.	Body weight (g) on gestation period (day)									Body weight gain	
	0	1	3	5	7	10	14	17	20	Days 0-20	(g)
251	261	276	287	298	297	315	340	369	420	159	60.92
252	266	273	277	286	294	305	332	363	429	163	61.28
253	256	273	286	289	299	315	331	371	423	167	65.23
254	266	275	292	299	307	317	345	374	424	158	59.40
255	260	267	256	290	293	307	331	358	413	153	58.85
256	260	271	283	291	299	316	337	370	423	163	62.69
257	265	273	291	302	312	335	354	390	443	178	67.17
258	270	273	286	286	298	309	336	368	418	148	54.81
259	265	271	285	292	294	317	330	363	415	150	56.60
260	272	282	294	304	312	325	350	380	423	151	55.51
261	290	297	311	316	326	342	367	402	456	166	57.24
262	276	289	305	320	325	345	375	401	451	175	63.41
N	12	12	12	12	12	12	12	12	12	12	12
MEAN	267.3	276.7	287.8	297.8	304.7	320.7	344.0	375.8	428.2	160.9	60.259
S.D.	9.1	8.6	13.7	11.2	11.7	13.3	14.9	14.6	14.1	9.6	3.899
S.E.	2.6	2.5	4.0	3.2	3.4	3.8	4.3	4.2	4.1	2.8	1.126

INDIVIDUAL DATA 4-3-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Gestation

Animal No.	Body weight (g) on gestation period (day)									Body weight gain	
	0	1	3	5	7	10	14	17	20	Days 0-20	(g)
351	246	256	270	282	287	302	318	340	383	137	55.69
352	246	249	265	272	279	302	326	356	414	168	68.29
353	266	280	283	286	292	305	323	349	407	141	53.01
354	243	259	274	278	290	297	318	352	397	154	63.37
355	264	272	280	288	297	314	336	366	417	153	57.95
356	264	273	289	297	303	315	335	368	407	143	54.17
357	255	264	274	279	288	308	326	351	402	147	57.65
358	264	270	290	289	300	319	332	370	401	137	51.89
359	274	278	285	292	297	313	330	356	402	128	46.72
360	272	274	288	296	298	312	326	353	405	133	48.90
361	278	286	295	307	311	330	353	389	443	165	59.35
362	288	289	314	323	341	363	390	427	486	198	68.75
N	12	12	12	12	12	12	12	12	12	12	12
MEAN	263.3	270.8	283.9	290.8	298.6	315.0	334.4	364.8	413.7	150.3	57.145
S.D.	13.8	12.0	13.0	13.9	15.7	17.5	19.9	23.4	26.8	19.4	6.979
S.E.	4.0	3.5	3.8	4.0	4.5	5.1	5.7	6.7	7.7	5.6	2.015

INDIVIDUAL DATA 4-3-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Gestation

Animal No.	Body weight (g) on gestation period (day)								Body weight gain	
	0	1	3	5	7	10	14	17	Days 0-20	(g)
451	260	272	279	295	291	310	328	357	404	144
452	261	268	276	280	291	305	327	359	404	143
453	261	266	274	285	294	303	325	366	425	164
454	274	280	289	301	312	327	354	389	442	168
455	262	261	275	285	292	307	334	366	422	160
456	276	279	288	295	304	315	336	364	413	137
457	277	287	292	304	312	327	349	387	441	164
458	279	281	288	293	305	317	335	366	410	131
459	280	288	304	315	322	338	359	390	430	150
460	280	280	301	307	316	337	356	389	449	169
461	293	302	316	327	334	358	388	425	478	185
462	286	284	293	301	308	327	353	392	455	169
N	12	12	12	12	12	12	12	12	12	12
MEAN	274.1	279.0	289.6	299.0	306.8	322.6	345.3	379.2	431.1	157.0
S.D.	10.8	11.2	12.8	13.3	13.5	16.4	18.2	19.7	22.7	15.9
S.E.	3.1	3.2	3.7	3.9	3.9	4.7	5.3	5.7	6.6	4.6
										5.231
										1.510

INDIVIDUAL DATA 4-4-1

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Lactation

Animal No.	Body weight (g) on lactation period (day)				Body weight gain	
	0	1	4	6	Days 0-4 (g)	(%)
151	329	334	345	309	16	4.86
152	288	300	328	292	40	13.89
153	304	308	320	295	16	5.26
154	343	347	365	330	22	6.41
155	303	313	331	290	28	9.24
156	317	320	329	301	12	3.79
157	321	324	332	300	11	3.43
158	304	308	332	298	28	9.21
159	346	336	361	314	15	4.34
160	328	344	311	317	-17	-5.18
161 ^a	#	#	#	#	#	#
162	348	350	360	324	12	3.45
N	11	11	11	11	11	11
MEAN	321.0	325.8	337.6	306.4	16.6	5.336
S.D.	19.9	17.4	17.7	13.4	14.3	4.754
S.E.	6.0	5.3	5.3	4.0	4.3	1.433
M/C	1.0372	1.1300	4.7693	2.5887	17.4007 **	16.5484 **
F	1.4512	1.7308	0.9395	1.3591		
H					4.6890	5.2262

a: Non-pregnancy.

#: Blank.

INDIVIDUAL DATA 4-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Lactation

Animal No.	Body weight (g) on lactation period (day)				Body weight gain	
	0	1	4	6	Days 0-4 (g)	(%)
251	294	303	328	290	34	11.56
252 ^a	271	279	(289)	(271)	#	#
253	294	308	324	284	30	10.20
254 ^a	291	278	(253)	(288)	#	#
255	322	310	317	298	-5	-1.55
256	307	316	336	301	29	9.45
257	329	327	347	304	18	5.47
258	293	307	265	273	-28	-9.56
259	324	298	251	278	-73	-22.53
260	325	320	334	292	9	2.77
261	337	345	352	314	15	4.45
262	332	342	350	313	18	5.42
N	12	12	10	10	10	10
MEAN	309.9	311.1	320.4	294.7	4.7	1.568
S.D.	21.0	21.0	34.9	13.9	32.9	10.491
S.E.	6.0	6.1	11.0	4.4	10.4	3.318

a : All pups died by day 3 of lactation (values in parentheses are excluded from calculation).

#: Blank.

INDIVIDUAL DATA 4-4-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Lactation

Animal No.	Body weight (g) on lactation period (day)				Body weight gain	
	0	1	4	6	Days 0-4 (g)	(%)
351	312	313	316	282	4	1.28
352	302	312	339	292	37	12.25
353	314	320	335	294	21	6.69
354	287	292	304	268	17	5.92
355	303	313	332	302	29	9.57
356	298	302	313	288	15	5.03
357	301	306	309	282	8	2.66
358	304	304	332	288	28	9.21
359	303	309	333	287	30	9.90
360	296	293	322	287	26	8.78
361	320	333	367	322	47	14.69
362	357	358	381	337	24	6.72
N	12	12	12	12	12	12
MEAN	308.1	312.9	331.9	294.1	23.8	7.725
S.D.	17.7	18.0	22.8	18.7	12.0	3.817
S.E.	5.1	5.2	6.6	5.4	3.5	1.102

INDIVIDUAL DATA 4-4-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Lactation

Animal No.	Body weight (g) on lactation period (day)				Body weight gain	
	0	1	4	6	Days 0-4 (g)	(%)
451	311	313	331	295	20	6.43
452	311	315	327	286	16	5.14
453	295	303	325	290	30	10.17
454	322	324	341	312	19	5.90
455	320	319	316	282	-4	-1.25
456	299	301	306	281	7	2.34
457	323	334	340	311	17	5.26
458	315	314	311	285	-4	-1.27
459	348	352	354	325	6	1.72
460	326	328	335	304	9	2.76
461	385	384	389	350	4	1.04
462	310	311	334	306	24	7.74
N	12	12	12	12	12	12
MEAN	322.1	324.8	334.1	302.3	12.0	3.832
S.D.	24.0	23.3	21.9	20.5	10.7	3.538
S.E.	6.9	6.7	6.3	5.9	3.1	1.021

INDIVIDUAL DATA 4-5-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

Animal No.	Body weight (g) on administration period (day)										Body weight gain Days 1-42	
	1	3	5	7	10	14	21	28	35	42	(g)	(%)
163	239	232	245	244	252	261	262	282	284	283	44	18.41
164	244	232	247	242	249	258	265	274	275	274	30	12.30
165	246	236	247	246	260	265	269	284	301	292	46	18.70
166	245	246	252	253	255	261	257	259	269	281	36	14.69
167	266	255	270	262	270	278	284	293	294	299	33	12.41
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	248.0	240.2	252.2	249.4	257.2	264.6	267.4	278.4	284.6	285.8	37.8	15.302
S.D.	10.4	10.1	10.3	8.2	8.2	7.9	10.3	12.8	13.2	9.8	6.9	3.121
S.E.	4.7	4.5	4.6	3.7	3.7	3.5	4.6	5.7	5.9	4.4	3.1	1.396
M/C	0.3041	0.0968	0.2256	0.3538	0.7058	1.1631	0.4561	0.7605	0.7217	1.5974	3.0987	2.7740
F	1.3705	0.1150	1.8890	0.0010	1.1020	1.6024	0.8955	0.7271	0.6677	0.6012	0.0079	0.0004

INDIVIDUAL DATA 4-5-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Body weight (g) on administration period (day)										Body weight gain	
	1	3	5	7	10	14	21	28	35	42	Days 1-42	(g)
463	249	235	252	242	254	258	260	259	265	279	30	12.05
464	230	242	233	245	250	254	252	268	276	280	50	21.74
465	247	245	250	259	257	265	269	282	290	281	34	13.77
466	237	234	239	237	228	232	240	242	247	248	11	4.64
467	243	255	247	263	261	268	278	295	300	303	60	24.69
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	241.2	242.2	244.2	249.2	250.0	255.4	259.8	269.2	275.6	278.2	37.0	15.378
S.D.	7.8	8.5	8.0	11.2	12.9	14.2	14.7	20.5	20.8	19.6	18.9	8.002
S.E.	3.5	3.8	3.6	5.0	5.8	6.4	6.6	9.2	9.3	8.8	8.5	3.579

INDIVIDUAL DATA 4-5-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

Animal No.	Body weight (g) on recovery period (day)		Body weight gain	
	7	14	Day 42-R14 (g)	(%)
163	287	294	11	3.89
164	299	286	12	4.38
165	311	308	16	5.48
166	291	291	10	3.56
167	306	305	6	2.01
N	5	5	5	5
MEAN	298.8	296.8	11.0	3.864
S.D.	10.0	9.4	3.6	1.266
S.E.	4.5	4.2	1.6	0.566
M/C	0.6802	0.9985	0.8477	1.7853
F	2.4571	0.1129	2.3900	2.1651

R14: Recovery day 14.

INDIVIDUAL DATA 4-5-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Body weight (g) on recovery period (day)		Body weight gain	
	7	14	Day 42-R14 (g)	(%)
463	291	291	12	4.30
464	281	292	12	4.29
465	286	294	13	4.63
466	264	274	26	10.48
467	307	319	16	5.28
N	5	5	5	5
MEAN	285.8	294.0	15.8	5.796
S.D.	15.6	16.1	5.9	2.649
S.E.	7.0	7.2	2.7	1.185

R14: Recovery day 14.

INDIVIDUAL DATA 5-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg

Animal No.	Food consumption (g/rat/day) on administration period (day)								
	1	3	5	7	10	14	28	35	42
101	27.0	24.0	25.0	22.0	25.7	26.5	28.0	27.1	26.4
102	24.0	25.5	25.5	24.5	25.7	26.8	26.9	27.0	24.7
103	24.0	23.5	24.0	24.0	24.3	25.8	22.9	26.7	27.4
104	25.0	24.5	25.0	25.5	25.3	26.3	27.0	27.0	27.4
105	25.0	24.0	26.5	24.5	25.3	25.3	25.6	25.3	25.8
106	25.0	25.5	25.5	24.5	24.0	23.5	25.3	24.4	24.3
107	26.0	26.5	26.5	26.5	26.3	25.5	24.4	26.6	27.3
108	31.0	29.5	28.0	27.0	27.7	29.0	28.1	30.7	29.8
109	24.0	27.5	27.0	26.0	27.3	28.3	28.4	29.4	27.6
110	27.0	25.5	27.0	27.0	26.0	25.8	26.7	26.3	25.9
111	27.0	26.0	27.0	26.0	26.3	26.8	27.1	26.6	28.0
112	29.0	31.0	30.5	27.5	27.7	29.0	30.4	30.6	28.4
N	12	12	12	12	12	12	12	12	12
MEAN	26.17	26.08	26.46	25.42	25.97	26.55	26.73	27.31	26.92
S.D.	2.17	2.27	1.70	1.58	1.19	1.61	1.99	1.95	1.57
S.E.	0.63	0.66	0.49	0.46	0.34	0.46	0.57	0.56	0.45
M/C	1.7028	2.1982	7.7938	1.3694	3.7948	2.3098	2.4264	4.3658	2.2948
F	0.8802	0.8234	0.5473	0.1456	0.1192	0.6196	1.5874	1.5508	1.2727

Food consumption of animal nos. 102, 105, 107, 108 and 111 on administration day 42 is the mean value of 6 days because of urinary examination.

INDIVIDUAL DATA 5-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg

Animal No.	Food consumption (g/rat/day) on administration period (day)								
	1	3	5	7	10	14	28	35	42
201	22.0	26.0	25.5	23.0	25.0	25.5	24.3	24.7	23.4
202	27.0	26.0	27.0	27.0	27.7	26.8	28.7	27.0	28.2
203	25.0	27.0	24.5	25.5	23.7	25.0	26.6	26.9	26.1
204	27.0	27.0	26.0	25.0	26.0	24.3	28.3	29.3	28.0
205	24.0	25.5	24.5	25.5	25.3	26.0	25.7	26.3	25.9
206	28.0	23.0	25.0	23.0	24.7	23.5	23.9	24.7	24.0
207	27.0	27.5	28.5	25.5	27.3	28.3	26.4	26.0	25.3
208	24.0	28.5	27.5	25.0	26.3	27.5	25.1	26.0	26.7
209	30.0	30.0	28.5	27.0	27.0	26.0	26.1	26.3	25.7
210	26.0	25.5	26.0	24.5	26.3	26.0	26.6	27.4	28.5
211	30.0	26.0	16.5	28.0	27.7	27.5	26.4	26.9	26.9
212	29.0	26.0	26.5	24.5	25.3	23.8	26.3	25.4	25.8
N	12	12	12	12	12	12	12	12	12
MEAN	26.58	26.50	25.50	25.29	26.03	25.85	26.20	26.41	26.21
S.D.	2.50	1.73	3.15	1.51	1.27	1.52	1.40	1.26	1.57
S.E.	0.72	0.50	0.91	0.44	0.37	0.44	0.40	0.36	0.45

Food consumption of animal nos. 202, 207, 209, 210 and 212 on administration day 42 is the mean value of 6 days because of urinary examination.

INDIVIDUAL DATA 5-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg

Animal No.	Food consumption (g/rat/day) on administration period (day)								
	1	3	5	7	10	14	28	35	42
301	23.0	23.5	25.0	26.0	26.7	26.5	26.0	25.9	27.0
302	22.0	23.0	24.0	21.5	23.7	23.0	23.9	24.0	24.1
303	22.0	24.5	24.0	23.5	24.3	24.8	24.7	25.0	24.7
304	28.0	25.0	28.0	25.5	27.3	27.3	27.1	27.3	28.8
305	27.0	24.0	26.0	23.5	25.3	23.8	24.7	24.9	24.3
306	27.0	25.0	26.5	24.0	24.7	23.0	23.7	23.9	23.9
307	24.0	25.5	24.5	23.0	25.7	25.0	26.4	25.1	25.8
308	30.0	27.0	27.5	28.0	27.7	28.3	28.0	27.9	26.4
309	26.0	26.0	25.0	25.0	25.0	26.8	25.0	25.4	24.7
310	28.0	25.0	27.5	26.0	27.0	26.3	27.0	27.4	27.0
311	28.0	27.0	29.0	26.0	26.0	26.5	27.1	28.4	28.4
312	30.0	28.0	31.0	28.0	31.3	31.0	31.9	32.4	31.4
N	12	12	12	12	12	12	12	12	12
MEAN	26.25	25.29	26.50	25.00	26.23	26.03	26.29	26.47	26.38
S.D.	2.86	1.50	2.17	1.98	2.02	2.31	2.25	2.40	2.29
S.E.	0.83	0.43	0.63	0.57	0.58	0.67	0.65	0.69	0.66

Food consumption of animal nos. 301, 304, 307, 309 and 310 on administration day 42 is the mean value of 6 days because of urinary examination.

INDIVIDUAL DATA 5-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg

Animal No.	Food consumption (g/rat/day) on administration period (day)								
	1	3	5	7	10	14	28	35	42
401	27.0	25.0	24.5	24.5	25.0	25.0	25.7	25.3	24.6
402	25.0	24.0	24.5	23.5	25.0	23.3	24.1	24.9	25.1
403	28.0	26.0	27.0	25.0	27.7	23.8	23.4	22.7	22.3
404	25.0	22.5	23.0	22.5	24.7	24.3	22.3	21.9	23.6
405	29.0	25.0	25.5	24.5	25.0	25.5	25.0	26.3	23.3
406	27.0	26.0	27.0	27.0	26.3	28.3	27.0	28.0	28.0
407	31.0	28.5	26.0	26.5	25.3	23.8	23.1	24.4	24.3
408	27.0	26.0	27.0	25.0	26.0	27.3	28.1	27.7	28.6
409	29.0	27.5	27.0	26.5	26.0	25.0	23.6	23.6	24.2
410	25.0	24.5	25.5	23.0	23.3	24.5	24.7	26.1	26.7
411	30.0	28.0	26.5	28.0	26.7	27.0	25.0	26.9	27.0
412	28.0	30.0	28.0	29.0	29.3	28.8	28.9	28.9	27.4
N	12	12	12	12	12	12	12	12	12
MEAN	27.58	26.08	25.96	25.42	25.86	25.55	25.08	25.56	25.43
S.D.	1.98	2.11	1.42	2.01	1.55	1.85	2.04	2.17	2.04
S.E.	0.57	0.61	0.41	0.58	0.45	0.53	0.59	0.63	0.59

Food consumption of animal nos. 405, 406, 409, 410 and 411 on administration day 42 is the mean value of 6 days because of urinary examination.

INDIVIDUAL DATA 5-1-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg

Animal No.	Food consumption (g/rat/day) on recovery period (day)	
	7	14
102	27.1	29.0
105	26.0	26.8
107	28.6	29.7
108	29.9	30.0
111	28.4	29.5
N	5	5
MEAN	28.00	29.00
S.D.	1.49	1.28
S.E.	0.67	0.57
M/C	0.3539	0.0601
F	4.9526 †	7.8606 *

Food consumption on recovery day 14 is the mean value of 6 days because of urinary examination.

INDIVIDUAL DATA 5-1-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg

Animal No.	Food consumption (g/rat/day) on recovery period (day)	
	7	14
405	26.7	28.5
406	25.4	25.5
409	24.7	26.3
410	26.6	27.3
411	27.4	26.7
N	5	5
MEAN	26.16	26.86
S.D.	1.09	1.13
S.E.	0.49	0.50
t'	2.2254	2.8037 *

Food consumption on recovery day 14 is the mean value of 6 days because of urinary examination.

INDIVIDUAL DATA 5-2-1

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Before gestation

Animal No.	Food consumption (g/rat/day) on administration period (day)					
	1	3	5	7	10	14
151	22.0	16.0	18.5	17.0	21.3	19.3
152	19.0	14.5	18.0	16.0	19.3	17.3
153	13.0	17.0	15.0	17.0	17.0	17.0
154	16.0	13.5	17.0	14.5	17.0	18.8
155	11.0	20.5	16.0	21.0	18.0	18.3
156	17.0	19.0	18.0	18.0	17.7	18.8
157	21.0	16.0	20.5	17.5	18.3	19.5
158	15.0	20.5	19.0	19.0	19.7	19.8
159	20.0	15.5	19.5	15.5	17.7	17.5
160	22.0	16.5	21.0	20.5	20.0	20.5
161	19.0	17.5	20.0	16.5	20.0	19.3
162	21.0	18.0	21.0	19.0	22.3	21.3
N	12	12	12	12	12	12
MEAN	18.00	17.04	18.63	17.63	19.03	18.95
S.D.	3.62	2.19	1.93	1.97	1.70	1.29
S.E.	1.04	0.63	0.56	0.57	0.49	0.37
M/C	0.6891	0.6171	1.1579	1.7615	1.0558	0.7107
F	0.6377	0.2618	4.6973 **	0.1542	1.3220	0.5669

INDIVIDUAL DATA 5-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Before gestation

Animal No.	Food consumption (g/rat/day) on administration period (day)					
	1	3	5	7	10	14
251	19.0	16.5	19.5	16.0	20.0	17.5
252	16.0	16.5	17.0	17.0	16.0	17.0
253	22.0	13.5	20.5	14.5	18.3	18.3
254	21.0	17.0	18.5	15.0	19.7	17.5
255	17.0	15.5	20.0	14.0	20.0	16.0
256	14.0	15.5	15.5	17.5	15.0	17.8
257	14.0	19.0	17.5	20.5	17.7	19.8
258	18.0	17.5	16.5	18.0	18.3	18.8
259	16.0	19.0	18.0	19.5	17.3	18.8
260	14.0	20.0	18.0	20.5	19.0	19.3
261	17.0	21.5	17.0	22.0	19.7	20.5
262	21.0	19.5	17.0	20.5	21.3	19.3
N	12	12	12	12	12	12
MEAN	17.42	17.58	17.92	17.92	18.53	18.38
S.D.	2.84	2.27	1.49	2.69	1.81	1.28
S.E.	0.82	0.66	0.43	0.78	0.52	0.37

t' 1.0642

INDIVIDUAL DATA 5-2-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Before gestation

Animal No.	Food consumption (g/rat/day) on administration period (day)					
	1	3	5	7	10	14
351	18.0	16.0	17.0	17.0	19.7	18.3
352	16.0	16.5	15.0	18.0	16.0	16.3
353	21.0	18.0	19.5	18.0	22.0	20.0
354	11.0	17.5	14.5	18.0	16.3	18.0
355	17.0	15.0	16.5	16.0	17.7	17.0
356	16.0	17.5	14.0	17.0	15.7	17.0
357	17.0	20.0	17.0	20.0	18.3	18.5
358	15.0	19.0	16.0	18.5	17.7	19.0
359	13.0	18.5	16.5	19.5	17.0	19.0
360	21.0	21.0	17.0	21.5	18.7	19.5
361	17.0	15.5	16.5	15.0	17.3	17.0
362	13.0	19.0	15.5	20.5	18.0	20.8
N	12	12	12	12	12	12
MEAN	16.25	17.79	16.25	18.25	17.87	18.37
S.D.	3.02	1.83	1.44	1.89	1.73	1.37
S.E.	0.87	0.53	0.42	0.55	0.50	0.40

t' 3.5682 **

INDIVIDUAL DATA 5-2-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Before gestation

Animal No.	Food consumption (g/rat/day) on administration period (day)					
	1	3	5	7	10	14
451	18.0	13.5	18.0	15.0	18.7	17.5
452	17.0	14.5	19.0	14.0	16.7	18.0
453	15.0	18.0	14.5	18.0	17.0	17.8
454	18.0	17.0	18.5	16.0	16.3	16.8
455	12.0	16.5	15.5	17.0	16.3	17.5
456	21.0	16.5	19.5	16.5	17.0	18.5
457	23.0	16.0	20.0	17.0	20.3	19.0
458	20.0	17.5	19.5	20.0	18.7	19.8
459	15.0	20.5	18.0	21.0	18.3	18.3
460	16.0	20.0	18.0	20.5	20.0	20.5
461	14.0	20.0	18.5	20.0	18.0	19.3
462	20.0	19.5	17.5	20.5	17.7	19.0
N	12	12	12	12	12	12
MEAN	17.42	17.46	18.04	17.96	17.92	18.50
S.D.	3.20	2.24	1.62	2.39	1.34	1.07
S.E.	0.92	0.65	0.47	0.69	0.39	0.31

t' 0.8764

INDIVIDUAL DATA 5-3-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Gestation

Animal No.	Food consumption (g/rat/day) on gestation period (day)							
	1	3	5	7	10	14	17	20
151	22.0	25.0	27.5	27.0	28.3	28.8	25.3	24.3
152	22.0	20.0	24.0	24.0	24.7	23.0	22.7	23.0
153	16.0	24.0	24.0	25.0	25.7	26.0	25.3	24.7
154	22.0	25.0	25.0	28.0	26.3	25.5	26.3	23.7
155	18.0	26.0	25.5	27.0	24.7	25.5	27.3	25.0
156	14.0	23.5	22.0	24.5	25.0	24.5	25.0	22.7
157	21.0	23.0	24.0	25.0	25.7	24.3	24.0	21.3
158	19.0	22.0	24.0	23.5	25.7	25.5	25.3	26.0
159	16.0	24.0	26.0	27.0	28.7	29.3	27.7	25.7
160	21.0	26.0	28.0	27.0	26.7	27.3	27.3	25.0
161 ^a	(18.0)	(19.5)	(21.5)	(20.5)	(22.7)	(20.8)	(17.3)	(14.0)
162	17.0	27.5	25.5	26.5	27.0	26.8	26.7	23.7
N	11	11	11	11	11	11	11	11
MEAN	18.91	24.18	25.05	25.86	26.23	26.05	25.72	24.10
S.D.	2.88	2.08	1.72	1.50	1.35	1.90	1.53	1.40
S.E.	0.87	0.63	0.52	0.45	0.41	0.57	0.46	0.42
M/C	10.5824 *	2.7757	1.8785	3.8538	4.7567	0.1816	1.7640	1.0019
F		2.3368 †	2.7412 †	3.1517 *	2.3343 †	3.8370 *	0.9608	0.8046
H	4.0155							

a: Non-pregnancy (values in parentheses are excluded from calculation).

INDIVIDUAL DATA 5-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Gestation

Animal No.	Food consumption (g/rat/day) on gestation period (day)							
	1	3	5	7	10	14	17	20
251	20.0	20.5	24.0	21.0	23.7	23.5	24.7	24.0
252	19.0	18.5	20.5	20.5	21.0	22.0	23.3	21.0
253	21.0	23.0	22.0	23.5	23.7	21.5	25.0	21.3
254	19.0	26.5	25.0	27.0	25.7	25.3	25.3	24.0
255	20.0	18.0	25.5	24.5	24.0	23.8	23.0	24.7
256	20.0	19.0	21.0	22.5	22.0	21.0	22.0	22.0
257	18.0	26.5	26.5	26.5	29.3	26.0	28.7	23.3
258	16.0	22.0	19.5	23.0	21.7	22.8	24.3	22.3
259	18.0	22.0	23.5	21.5	22.7	22.8	28.0	26.0
260	18.0	26.0	25.0	23.0	25.7	25.0	26.3	22.0
261	18.0	24.5	23.5	23.5	24.3	24.8	26.3	24.7
262	18.0	24.5	28.5	27.5	29.3	28.3	24.0	22.3
N	12	12	12	12	12	12	12	12
MEAN	18.75	22.58	23.71	23.67	24.43	23.90	25.08	23.13
S.D.	1.36	3.10	2.62	2.31	2.70	2.09	1.99	1.55
S.E.	0.39	0.89	0.76	0.67	0.78	0.60	0.57	0.45
t'		1.5146	1.4945	2.3302	1.8632	2.6402 *		

INDIVIDUAL DATA 5-3-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Gestation

Animal No.	Food consumption (g/rat/day) on gestation period (day)							
	1	3	5	7	10	14	17	20
351	22.0	21.0	24.0	24.0	25.3	23.8	25.0	23.0
352	16.0	20.0	21.5	20.5	22.7	23.3	23.0	23.7
353	26.0	22.5	25.0	25.5	24.7	25.3	27.0	26.0
354	17.0	23.5	24.5	25.0	23.7	22.5	22.7	19.7
355	17.0	18.5	20.5	22.5	22.0	22.8	22.0	22.7
356	15.0	23.0	23.5	23.5	24.7	23.5	23.0	22.0
357	13.0	19.0	22.5	24.0	25.7	22.8	26.3	26.0
358	15.0	25.0	22.5	23.0	24.3	23.0	24.0	21.3
359	20.0	21.0	21.0	21.0	23.7	22.8	25.0	23.0
360	13.0	22.5	22.5	24.0	24.0	22.3	21.3	22.3
361	22.0	18.5	22.5	19.5	22.3	22.8	24.3	24.0
362	16.0	27.5	28.0	30.5	31.7	29.3	28.0	24.7
N	12	12	12	12	12	12	12	12
MEAN	17.67	21.83	23.17	23.58	24.57	23.68	24.30	23.20
S.D.	4.01	2.73	2.04	2.84	2.52	1.94	2.05	1.84
S.E.	1.16	0.79	0.59	0.82	0.73	0.56	0.59	0.53
t'	2.2252	2.0999	2.4185	1.7168	2.9069 *			

INDIVIDUAL DATA 5-3-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Gestation

Animal No.	Food consumption (g/rat/day) on gestation period (day)							
	1	3	5	7	10	14	17	20
451	21.0	19.5	21.5	20.0	21.3	22.8	22.3	22.7
452	19.0	20.0	21.0	22.5	22.7	22.8	23.7	22.0
453	16.0	21.5	22.0	23.5	22.0	23.3	24.7	25.3
454	18.0	20.0	20.5	22.5	21.7	23.3	24.0	21.7
455	12.0	21.0	21.5	20.0	22.0	21.8	25.3	23.0
456	16.0	22.0	22.0	22.5	22.0	23.3	21.0	21.3
457	21.0	20.5	24.0	25.5	24.0	23.5	25.3	22.3
458	17.0	21.0	21.0	25.0	23.7	24.3	25.3	22.3
459	18.0	24.5	26.5	25.0	26.7	23.3	26.3	22.3
460	16.0	23.5	23.0	23.5	25.3	23.8	27.0	25.0
461	18.0	26.0	26.5	27.0	29.3	29.3	30.3	26.7
462	12.0	20.5	21.5	22.0	24.3	23.5	25.0	25.7
N	12	12	12	12	12	12	12	12
MEAN	17.00	21.67	22.58	23.25	23.75	23.75	25.02	23.36
S.D.	2.89	2.00	2.05	2.13	2.39	1.85	2.34	1.81
S.E.	0.83	0.58	0.59	0.61	0.69	0.53	0.67	0.52
t'	2.3831	2.7519 *	2.7721 *	2.5611 *	2.8248 *			

INDIVIDUAL DATA 5-4-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Lactation

Animal No.	Food consumption (g/rat/day) on lactation period (day)	
	1	4
151	24.0	36.3
152	18.0	39.0
153	19.0	36.0
154	19.0	37.7
155	23.0	47.0
156	17.0	33.3
157	21.0	31.3
158	29.0	42.7
159	9.0	35.0
160	30.0	44.0
161 ^a	#	#
162	23.0	32.7
N	11	11
MEAN	21.09	37.73
S.D.	5.82	5.00
S.E.	1.76	1.51
M/C	6.4146	2.0034
F	0.2561	3.4244 *

a: Non-pregnancy.

#: Blank.

INDIVIDUAL DATA 5-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Lactation

Animal No.	Food consumption (g/rat/day) on lactation period (day)	
	1	4
251	31.0	34.7
252 ^a	7.0	(17.7)
253	30.0	34.3
254 ^a	1.0	(17.0)
255	13.0	35.7
256	19.0	33.7
257	23.0	38.0
258	23.0	36.3
259	8.0	35.3
260	18.0	34.0
261	31.0	42.3
262	19.0	30.0
N	12	10
MEAN	18.58	35.43
S.D.	9.84	3.18
S.E.	2.84	1.01
t'		1.2329

a : All pups died by day 3 of lactation (values in parentheses are excluded from calculation).

INDIVIDUAL DATA 5-4-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Lactation

Animal No.	Food consumption (g/rat/day) on lactation period (day)	
	1	4
351	19.0	29.0
352	23.0	39.0
353	27.0	40.0
354	19.0	32.0
355	18.0	39.7
356	14.0	36.0
357	20.0	35.7
358	21.0	40.0
359	21.0	38.7
360	8.0	30.7
361	28.0	40.0
362	19.0	34.3
N	12	12
MEAN	19.75	36.26
S.D.	5.31	3.97
S.E.	1.53	1.15
t'		0.8252

INDIVIDUAL DATA 5-4-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Lactation

Animal No.	Food consumption (g/rat/day) on lactation period (day)	
	1	4
451	20.0	33.7
452	18.0	30.0
453	16.0	35.7
454	24.0	37.7
455	20.0	29.0
456	16.0	30.0
457	25.0	34.7
458	10.0	21.0
459	20.0	29.7
460	21.0	34.3
461	31.0	36.0
462	17.0	35.3
N	12	12
MEAN	19.83	32.26
S.D.	5.29	4.58
S.E.	1.53	1.32

t' 3.0723 *

INDIVIDUAL DATA 5-5-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Food consumption, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

Animal No.	Food consumption (g/rat/day) on administration period (day)									
	1	3	5	7	10	14	21	28	35	42
163	22.0	17.5	22.0	19.0	19.3	19.3	19.4	20.6	19.1	17.5
164	21.0	13.5	20.0	16.5	19.7	20.0	19.4	18.4	19.3	18.2
165	20.0	13.0	19.0	16.5	20.0	17.0	17.9	18.0	19.7	16.3
166	21.0	20.0	19.0	19.0	18.0	17.8	16.1	17.6	18.3	18.8
167	22.0	16.0	20.5	15.5	20.7	19.8	18.9	19.3	18.9	19.3
N	5	5	5	5	5	5	5	5	5	5
MEAN	21.20	16.00	20.10	17.30	19.54	18.78	18.34	18.78	19.06	18.02
S.D.	0.84	2.89	1.24	1.60	1.00	1.32	1.39	1.20	0.52	1.17
S.E.	0.37	1.29	0.56	0.72	0.45	0.59	0.62	0.54	0.23	0.52
M/C	7.1852 **	0.0078	0.0020	2.1944	0.8701	1.0352	1.0845	2.7016	5.7376 *	1.0704
F		0.1394	10.6457 *	0.0499	5.3248 *	0.4147	0.2410	0.6136		1.1228
H		2.4847							1.0976	

Food consumption on administration day 42 is the mean value of 6 days because of urinary examination.

INDIVIDUAL DATA 5-5-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Food consumption (g/rat/day) on administration period (day)									
	1	3	5	7	10	14	21	28	35	42
463	19.0	14.0	19.0	14.0	18.0	16.3	15.1	14.3	15.3	16.7
464	14.0	18.0	16.0	18.0	19.0	17.5	18.0	18.3	17.7	18.2
465	24.0	18.5	18.5	22.0	18.3	20.5	20.1	19.9	19.4	17.5
466	16.0	13.0	17.5	14.0	14.7	15.5	15.3	14.7	14.7	16.7
467	13.0	20.0	16.5	20.5	17.7	20.3	20.1	21.0	19.3	17.8
N	5	5	5	5	5	5	5	5	5	5
MEAN	17.20	16.70	17.50	17.70	17.54	18.02	17.72	17.64	17.28	17.38
S.D.	4.44	3.03	1.27	3.67	1.66	2.29	2.46	3.03	2.20	0.67
S.E.	1.98	1.36	0.57	1.64	0.74	1.02	1.10	1.35	0.98	0.30
t'			3.2628 *		2.3076 *					

Food consumption on administration day 42 is the mean value of 6 days because of urinary examination.

INDIVIDUAL DATA 5-5-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Food consumption, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

Animal No.	Food consumption (g/rat/day) on recovery period (day)	
	7	14
163	19.3	19.2
164	20.4	18.7
165	19.9	19.3
166	20.4	20.0
167	18.4	19.7
N	5	5
MEAN	19.68	19.38
S.D.	0.85	0.50
S.E.	0.38	0.22
M/C	0.0839	4.1384 *
F	6.3810 *	
H		0.0439

Food consumption on recovery day 14 is the mean value of 6 days because of urinary examination.

INDIVIDUAL DATA 5-5-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Food consumption, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Food consumption (g/rat/day) on recovery period (day)	
	7	14
463	17.3	17.2
464	18.6	19.7
465	18.6	21.2
466	18.3	18.3
467	19.3	20.5
N	5	5
MEAN	18.42	19.38
S.D.	0.73	1.63
S.E.	0.32	0.73

t' 2.5261 *

Food consumption on recovery day 14 is the mean value of 6 days because of urinary examination.

INDIVIDUAL DATA 6-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Urinary findings ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Week 6 of administration

Animal No.	pH	Protein	Glucose	Ketone body	Urobilinogen EU/dL	Bilirubin	Occult blood	Color	Specific gravity	Urine volume mL/21hr
102	8.5	±	—	—	0.1	—	—	A	1.045	13.0
105	8.5	±	—	—	0.1	—	—	A	1.034	18.0
107	8.5	±	—	—	0.1	—	—	A	1.035	11.5
108	8.5	±	—	—	0.1	—	—	A	1.043	15.5
111	8.0	±	—	—	0.1	—	—	A	1.032	23.5
N	5	5	5	5	5	5	5	5	5	5
MEAN										16.30
S.D.										4.72
S.E.										2.11
M/C										4.6270
F										0.1214
H	8.5467 *	2.1111	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.1523	

Color: A = Pale yellow or yellow.

-; Negative, ±; slight.

INDIVIDUAL DATA 6-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Urinary findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : Week 6 of administration

Animal No.	pH	Protein	Glucose	Ketone body	Urobilinogen EU/dL	Bilirubin	Occult blood	Color	Specific gravity	Urine volume mL/21hr
202	8.5	±	—	—	0.1	—	—	A	1.036	20.0
207	8.5	+	—	—	0.1	—	—	A	1.040	13.0
209	8.5	±	—	—	0.1	—	—	A	1.039	16.0
210	8.5	±	—	—	0.1	—	—	A	1.044	15.5
212	8.5	±	—	—	0.1	—	—	A	1.030	21.5
N	5	5	5	5	5	5	5	5	5	5
MEAN										17.20
S.D.										3.47
S.E.										1.55

U 10.0000

Color: A = Pale yellow or yellow.

-; Negative, ±; slight, +; moderate.

INDIVIDUAL DATA 6-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Urinary findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : Week 6 of administration

Animal No.	pH	Protein	Glucose	Ketone body	Urobilinogen EU/dL	Bilirubin	Occult blood	Color	Specific gravity	Urine volume mL/21hr
301	8.0	±	—	—	0.1	—	—	A	1.036	14.0
304	8.0	±	—	—	0.1	—	—	A	1.028	33.0
307	8.0	±	—	—	0.1	—	—	A	1.038	18.0
309	8.5	±	—	—	0.1	—	—	A	1.044	14.0
310	7.5	±	—	—	0.1	—	—	A	1.051 ^a	8.5
N	5	5	5	5	5	5	5	5	5	5
MEAN										17.50
S.D.										9.30
S.E.										4.16
U	4.5000									

Color: A = Pale yellow or yellow.

a: Over the maximum value of scale.

-; Negative, ±; slight.

INDIVIDUAL DATA 6-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Urinary findings ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Week 6 of administration

Animal No.	pH	Protein	Glucose	Ketone body	Urobilinogen EU/dL	Bilirubin	Occult blood	Color	Specific gravity	Urine volume mL/21hr
405	8.5	+	—	—	0.1	—	—	A	1.051 ^a	8.0
406	8.5	±	—	—	0.1	—	—	A	1.026	31.0
409	8.0	±	—	—	0.1	—	—	A	1.030	26.0
410	8.5	±	—	—	0.1	—	—	A	1.044	15.5
411	8.5	±	—	—	0.1	—	—	A	1.038	14.5
N	5	5	5	5	5	5	5	5	5	5
MEAN										19.00
S.D.										9.31
S.E.										4.16
U		12.5000								

Color: A = Pale yellow or yellow.

a: Over the maximum value of scale.

-; Negative, ±; slight, +; moderate.

INDIVIDUAL DATA 6-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Urinary findings, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Week 6 of administration

Animal No.	pH	Protein	Glucose	Ketone body	Urobilinogen EU/dL			Occult blood	Color	Specific gravity	Urine volume mL/21hr
					Bilirubin						
163	7.5	—	—	—	0.1	—	—	A	1.037	14.0	
164	8.5	±	—	—	0.1	—	—	A	1.036	7.5	
165	8.5	±	—	—	0.1	—	—	A	1.038	13.0	
166	8.5	—	—	—	0.1	—	—	A	1.031	10.0	
167	8.5	±	—	—	0.1	—	—	A	1.041	9.0	
N	5	5	5	5	5	5	5	5	5	5	5
MEAN											10.70
S.D.											2.73
S.E.											1.22
M/C											0.2628
F											0.3542
H	0.4128	0.3600	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.3975		

Color: A = Pale yellow or yellow.

-; Negative, ±; slight.

INDIVIDUAL DATA 6-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Urinary findings, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Week 6 of administration

Animal No.	pH	Protein	Glucose	Ketone body	Urobilinogen EU/dL	Bilirubin	Occult blood	Color	Specific gravity	Urine volume mL/21hr
463	6.5	—	—	—	0.1	—	—	A	1.051 ^a	4.5
464	8.5	—	—	—	0.1	—	—	A	1.032	10.0
465	8.5	±	—	—	0.1	—	—	A	1.032	10.0
466	8.5	±	—	—	0.1	—	—	A	1.048	8.5
467	8.0	—	—	—	0.1	—	—	A	1.038	14.5
N	5	5	5	5	5	5	5	5	5	5
MEAN										9.50
S.D.										3.59
S.E.										1.60

Color: A = Pale yellow or yellow.

a: Over the maximum value of scale.

-, Negative, ±; slight.

INDIVIDUAL DATA 6-3-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Urinary findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : Week 2 of recovery

Animal No.	pH	Protein	Glucose	Ketone body	Urobilinogen EU/dL	Bilirubin	Occult blood	Color	Specific gravity	Urine volume mL/21hr
102	8.5	2+	—	—	0.1	—	—	A	1.051 ^a	9.0
105	8.5	+	—	—	0.1	—	—	A	1.043	13.0
107	8.5	+	—	—	0.1	—	—	A	1.042	10.0
108	8.5	+	—	—	0.1	—	—	A	1.051 ^a	10.5
111	8.5	+	—	—	0.1	—	—	A	1.041	19.0
N	5	5	5	5	5	5	5	5	5	5
MEAN										12.30
S.D.										4.02
S.E.										1.80
M/C										4.4486 *
H	1.0000	2.7908 †	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	0.1756

Color: A = Pale yellow or yellow.

a: Over the maximum value of scale.

-; Negative, +; moderate, 2+; severe.

INDIVIDUAL DATA 6-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Urinary findings ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : Week 2 of recovery

Animal No.	pH	Protein	Glucose	Ketone body	Urobilinogen EU/dL	Bilirubin	Occult blood	Color	Specific gravity	Urine volume mL/21hr
405	8.5	+	—	—	0.1	—	—	A	1.051 ^a	8.0
406	8.5	—	—	—	0.1	—	—	A	1.025	42.0
409	8.5	±	—	—	0.1	—	—	A	1.049	12.5
410	8.5	+	—	—	0.1	—	—	A	1.051 ^a	10.5
411	8.0	+	—	—	0.1	—	—	A	1.042	15.0
N	5	5	5	5	5	5	5	5	5	5
MEAN										17.60
S.D.										13.88
S.E.										6.21
U			6.0000							

Color: A = Pale yellow or yellow.

a: Over the maximum value of scale.

—; Negative, ±; slight, +; moderate.

INDIVIDUAL DATA 6-4-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Urinary findings, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Week 2 of recovery

Animal No.	pH	Protein	Glucose	Ketone body	Urobilinogen EU/dL			Occult blood	Color	Specific gravity	Urine volume mL/21hr
					Bilirubin						
163	6.5	±	—	—	0.1	—	—	A	1.037	16.5	
164	7.0	+	—	—	0.1	—	—	A	1.044	8.5	
165	8.5	±	—	—	0.1	—	—	A	1.046	11.0	
166	8.5	—	—	—	0.1	—	—	A	1.027	25.0	
167	6.5	—	—	—	0.1	—	—	A	1.051 ^a	9.0	
N	5	5	5	5	5	5	5	5	5	5	5
MEAN										14.00	
S.D.										6.92	
S.E.										3.09	
M/C										5.0549 *	
II	2.3581	1.2245	0.0000	0.0000	0.0000	0.0000	0.0000	0.0000	2.8800 †	2.4695	

Color: A = Pale yellow or yellow.

a: Over the maximum value of scale.

-; Negative, ±; slight, +; moderate.

INDIVIDUAL DATA 6-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Urinary findings, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Week 2 of recovery

Animal No.	pH	Protein	Glucose	Ketone body	Urobilinogen EU/dL	Bilirubin	Occult blood	Color	Specific gravity	Urine volume mL/21hr
463	8.5	—	—	—	0.1	—	—	A	1.051 ^a	7.0
464	8.0	+	—	—	0.1	—	—	A	1.049	8.0
465	8.5	+	—	—	0.1	—	—	A	1.046	11.5
466	8.5	+	—	—	0.1	—	—	A	1.051 ^a	8.0
467	8.5	±	—	—	0.1	—	—	A	1.047	10.0
N	5	5	5	5	5	5	5	5	5	5
MEAN										8.90
S.D.										1.82
S.E.										0.81
U								4.5000		

Color: A = Pale yellow or yellow.

a: Over the maximum value of scale.

—; Negative, ±; slight, +; moderate.

INDIVIDUAL DATA 7-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
101	94.9	950	15.9	43.3	45.6	16.7	36.7	135.9
103	100.7	873	15.6	44.2	50.6	17.9	35.3	111.2
104	80.1	878	16.3	46.9	53.4	18.6	34.8	108.5
106	98.5	971	15.9	43.4	44.7	16.4	36.6	131.3
109	108.4	961	17.3	47.5	49.4	18.0	36.4	139.4
N	5	5	5	5	5	5	5	5
MEAN	96.52	926.6	16.20	45.06	48.74	17.52	35.96	125.26
S.D.	10.43	47.3	0.66	2.00	3.60	0.93	0.86	14.39
S.E.	4.66	21.1	0.30	0.89	1.61	0.42	0.38	6.44
M/C	1.1746	6.4159	2.5469	5.1545	9.1739 *	8.0129 *	8.5711 *	2.9401
F	1.2866	1.3653	2.4515	3.8871 *				2.5838 †
II					1.8571	0.8518	2.6476	

INDIVIDUAL DATA 7-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
201	95.8	931	16.1	44.0	47.3	17.3	36.6	125.9
203	122.4	924	16.2	45.3	49.0	17.5	35.8	115.2
204	94.7	931	15.9	44.0	47.3	17.1	36.1	109.7
205	135.2	872	15.0	42.3	48.5	17.2	35.5	134.7
206	118.0	930	15.9	44.1	47.4	17.1	36.1	131.9
N	5	5	5	5	5	5	5	5
MEAN	113.22	917.6	15.82	43.94	47.90	17.24	36.02	123.48
S.D.	17.58	25.7	0.48	1.07	0.80	0.17	0.41	10.74
S.E.	7.86	11.5	0.21	0.48	0.36	0.07	0.18	4.80
t'				1.4036			0.2759	

INDIVIDUAL DATA 7-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
302	94.4	942	16.7	46.2	49.0	17.7	36.1	114.6
303	127.9	937	16.6	46.5	49.6	17.7	35.7	109.7
305	117.7	930	15.7	44.4	47.7	16.9	35.4	111.8
306	110.6	957	16.1	45.5	47.5	16.8	35.4	121.0
308	101.1	930	16.7	46.6	50.1	18.0	35.8	127.3
N	5	5	5	5	5	5	5	5
MEAN	110.34	939.2	16.36	45.84	48.78	17.42	35.68	116.88
S.D.	13.25	11.2	0.44	0.91	1.15	0.54	0.29	7.21
S.E.	5.92	5.0	0.20	0.41	0.51	0.24	0.13	3.23
t'				0.9775				1.2988

INDIVIDUAL DATA 7-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of administration

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
401	101.7	925	15.9	44.2	47.8	17.2	36.0	119.9
402	95.7	852	15.2	42.5	49.9	17.8	35.8	110.1
403	127.2	881	15.5	43.1	48.9	17.6	36.0	104.8
404	130.1	948	15.7	43.3	45.7	16.6	36.3	104.0
407	97.1	882	15.8	43.7	49.5	17.9	36.2	106.7
N	5	5	5	5	5	5	5	5
MEAN	110.36	897.6	15.62	43.36	48.36	17.42	36.06	109.10
S.D.	16.87	38.4	0.28	0.64	1.68	0.53	0.19	6.48
S.E.	7.55	17.2	0.12	0.29	0.75	0.24	0.09	2.90
t'				2.1305			2.5045	

INDIVIDUAL DATA 7-1-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

Animal No.	Reticu- locyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
101	3.18	24.9	27.1	14.7	76.9	2.6	0.7	0.0
103	3.66	23.2	26.9	8.9	88.1	2.7	1.0	0.0
104	3.09	19.3	29.0	17.6	58.3	3.3	0.9	0.0
106	2.49	20.5	27.6	15.1	79.5	2.2	1.7	0.0
109	3.57	17.8	22.0	13.8	89.9	2.6	2.0	0.1
N	5	5	5	5	5	5	5	5
MEAN	3.198	21.14	26.52	14.02	78.54	2.68	1.26	0.02
S.D.	0.465	2.89	2.66	3.19	12.59	0.40	0.56	0.04
S.E.	0.208	1.29	1.19	1.43	5.63	0.18	0.25	0.02
M/C	2.0657	2.0401	1.2737	3.9812	0.6743	2.9035	1.5092	∞ **
F	3.0552 †	0.1242	1.0351	0.2975	0.8351	2.0387	0.7847	
H							2.1111	

INDIVIDUAL DATA 7-1-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	Reticulocyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
201	3.77	21.6	27.9	15.2	75.6	3.4	1.6	0.0
203	3.07	20.5	28.4	13.3	105.4	3.1	0.6	0.0
204	3.59	21.5	23.9	14.2	76.1	3.5	0.9	0.0
205	3.44	22.6	27.0	11.4	118.6	3.9	1.3	0.0
206	3.35	18.8	26.3	24.5	88.6	4.3	0.6	0.0
N	5	5	5	5	5	5	5	5
MEAN	3.444	21.00	26.70	15.72	92.86	3.64	1.00	0.00
S.D.	0.263	1.44	1.76	5.10	18.82	0.47	0.44	0.00
S.E.	0.118	0.64	0.79	2.28	8.42	0.21	0.20	0.00
t'	1.1483							

INDIVIDUAL DATA 7-1-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	Reticulocyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
302	3.41	18.5	27.2	14.2	74.7	4.0	1.5	0.0
303	3.35	18.6	20.6	11.9	110.8	4.0	1.2	0.0
305	2.94	21.9	25.8	14.2	99.0	2.7	1.8	0.0
306	2.85	18.9	26.9	13.9	92.7	2.9	1.1	0.0
308	2.60	24.4	29.1	17.5	78.9	3.0	1.7	0.0
N	5	5	5	5	5	5	5	5
MEAN	3.030	20.46	25.92	14.34	91.22	3.32	1.46	0.00
S.D.	0.344	2.61	3.20	2.01	14.75	0.63	0.30	0.00
S.E.	0.154	1.17	1.43	0.90	6.60	0.28	0.14	0.00
t'	0.7842							

INDIVIDUAL DATA 7-1-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of administration

Animal No.	Reticu- locyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
401	2.42	22.1	25.3	14.0	82.7	3.4	1.6	0.0
402	2.81	22.0	23.9	15.6	76.1	2.9	1.1	0.0
403	3.01	22.9	27.1	13.0	110.7	2.7	0.7	0.1
404	2.88	21.3	24.5	15.5	108.1	4.5	2.0	0.0
407	2.97	18.1	19.4	19.0	75.2	2.1	0.8	0.0
N	5	5	5	5	5	5	5	5
MEAN	2.818	21.28	24.04	15.42	90.56	3.12	1.24	0.02
S.D.	0.236	1.87	2.86	2.28	17.46	0.90	0.55	0.04
S.E.	0.105	0.83	1.28	1.02	7.81	0.40	0.25	0.02
t'	1.7737							

INDIVIDUAL DATA 7-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 6 of lactation

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
152	129.7	815	15.4	42.9	52.6	18.9	35.9	136.7
153	79.9	811	15.3	43.6	53.8	18.9	35.1	106.6
155	88.7	816	15.1	44.0	53.9	18.5	34.3	115.7
156	111.6	839	15.6	44.4	52.9	18.6	35.1	122.3
162	76.6	745	14.6	42.9	57.6	19.6	34.0	123.9
N	5	5	5	5	5	5	5	5
MEAN	97.30	805.2	15.20	43.56	54.16	18.90	34.88	121.04
S.D.	22.69	35.4	0.38	0.67	2.00	0.43	0.75	11.09
S.E.	10.15	15.8	0.17	0.30	0.90	0.19	0.34	4.96
M/C	2.4175	2.9871	3.7092	6.9547	1.1323	2.9565	2.7686	3.9270
F	0.3232	0.1098	0.3517	0.8736	1.5211	1.4430	1.1463	4.6661 *

INDIVIDUAL DATA 7-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 6 of lactation

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
251	94.2	838	15.3	43.2	51.6	18.3	35.4	123.1
253	101.7	813	15.1	43.6	53.6	18.6	34.6	137.0
255	103.0	800	15.3	44.2	55.3	19.1	34.6	136.4
256	71.3	841	16.1	46.6	55.4	19.1	34.5	122.6
262	129.3	798	14.8	42.5	53.3	18.5	34.8	131.0
N	5	5	5	5	5	5	5	5
MEAN	99.90	818.0	15.32	44.02	53.84	18.72	34.78	130.02
S.D.	20.78	20.5	0.48	1.57	1.58	0.36	0.36	6.95
S.E.	9.29	9.2	0.22	0.70	0.70	0.16	0.16	3.11
t'								1.3387

INDIVIDUAL DATA 7-2-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 6 of lactation

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
351	90.6	862	16.4	47.6	55.2	19.0	34.5	102.3
352	81.6	854	16.3	47.0	55.0	19.1	34.7	96.4
353	108.6	760	15.2	45.2	59.5	20.0	33.6	104.3
355	97.4	786	15.1	44.0	56.0	19.2	34.3	98.9
361	95.4	776	15.1	43.6	56.2	19.5	34.6	134.5
N	5	5	5	5	5	5	5	5
MEAN	94.72	807.6	15.62	45.48	56.38	19.36	34.34	107.28
S.D.	9.87	47.0	0.67	1.78	1.82	0.40	0.44	15.52
S.E.	4.41	21.0	0.30	0.79	0.81	0.18	0.20	6.94
t'								2.0513

INDIVIDUAL DATA 7-2-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 6 of lactation

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
451	128.6	841	15.7	45.1	53.6	18.7	34.8	125.5
453	85.0	776	14.7	42.3	54.5	18.9	34.8	121.4
455	114.3	788	15.8	46.6	59.1	20.1	33.9	126.9
457	114.5	892	16.5	47.8	53.6	18.5	34.5	138.0
462	86.9	776	14.0	40.4	52.1	18.0	34.7	127.3
N	5	5	5	5	5	5	5	5
MEAN	105.86	814.6	15.34	44.44	54.58	18.84	34.54	127.82
S.D.	19.09	50.9	0.99	3.05	2.67	0.78	0.38	6.15
S.E.	8.54	22.8	0.44	1.37	1.19	0.35	0.17	2.75
t'								1.0107

INDIVIDUAL DATA 7-2-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 6 of lactation

Animal No.	Reticuloocyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
152	7.97	19.4	17.9	30.1	86.1	11.0	2.4	0.1
153	9.30	18.4	21.5	11.0	62.6	4.9	1.4	0.0
155	7.47	17.7	18.6	18.2	63.7	5.4	1.4	0.0
156	8.05	18.2	20.4	19.8	84.2	5.3	2.3	0.0
162	11.42	17.4	20.7	11.1	61.7	3.1	0.7	0.0
N	5	5	5	5	5	5	5	5
MEAN	8.842	18.22	19.82	18.04	71.66	5.94	1.64	0.02
S.D.	1.591	0.77	1.51	7.85	12.35	2.98	0.71	0.04
S.E.	0.712	0.34	0.67	3.51	5.52	1.33	0.32	0.02
M/C	1.1895	4.2322	5.9851	3.3757	2.2695	5.2711	2.9411	∞ **
F	0.1489	3.7760 *	0.5290	0.9513	0.4621	0.4843	0.8297	
H							1.1176	

INDIVIDUAL DATA 7-2-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 6 of lactation

Animal No.	Reticuloocyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
251	7.96	18.6	19.4	19.8	69.9	3.5	1.0	0.0
253	11.80	19.0	20.9	33.5	58.1	8.9	1.2	0.0
255	8.90	18.5	21.7	30.2	64.4	6.8	1.6	0.0
256	10.65	19.2	19.9	9.6	55.3	4.7	1.7	0.0
262	7.79	19.3	19.3	45.6	73.7	8.7	1.2	0.1
N	5	5	5	5	5	5	5	5
MEAN	9.420	18.92	20.24	27.74	64.28	6.52	1.34	0.02
S.D.	1.749	0.36	1.03	13.70	7.73	2.39	0.30	0.04
S.E.	0.782	0.16	0.46	6.12	3.46	1.07	0.13	0.02
t'			1.5315					

INDIVIDUAL DATA 7-2-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 6 of lactation

Animal No.	Reticulocyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
351	8.35	18.2	20.0	25.1	61.4	3.2	0.9	0.0
352	8.63	17.8	19.8	22.8	52.8	4.5	1.5	0.0
353	8.55	20.3	20.2	16.8	83.7	7.0	1.0	0.1
355	9.15	18.3	19.4	16.0	75.6	5.3	0.5	0.0
361	11.24	19.5	19.4	27.8	60.5	5.1	2.0	0.0
N	5	5	5	5	5	5	5	5
MEAN	9.184	18.82	19.76	21.70	66.80	5.02	1.18	0.02
S.D.	1.187	1.04	0.36	5.16	12.53	1.38	0.58	0.04
S.E.	0.531	0.47	0.16	2.31	5.60	0.62	0.26	0.02
t'				1.3128				

INDIVIDUAL DATA 7-2-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 6 of lactation

Animal No.	Reticu- locyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
451	9.01	18.2	20.4	24.3	95.3	6.1	2.9	0.0
453	9.26	17.8	19.3	16.3	62.9	4.9	0.9	0.0
455	8.60	17.3	19.2	42.2	64.0	6.4	1.7	0.0
457	7.66	17.7	20.1	18.6	86.9	7.5	1.5	0.0
462	10.57	16.8	18.1	25.3	53.9	6.2	1.5	0.0
N	5	5	5	5	5	5	5	5
MEAN	9.020	17.56	19.42	25.34	72.60	6.22	1.70	0.00
S.D.	1.059	0.53	0.90	10.15	17.59	0.93	0.73	0.00
S.E.	0.474	0.24	0.40	4.54	7.87	0.41	0.33	0.00
t'				1.4440				

INDIVIDUAL DATA 7-3-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
102	66.5	920	16.6	47.0	51.1	18.0	35.3	120.5
105	120.2	956	16.4	46.2	48.3	17.2	35.5	125.3
107	116.2	955	17.0	48.5	50.8	17.8	35.1	137.2
108	112.9	911	16.0	45.1	49.5	17.6	35.5	137.4
111	127.5	914	16.3	46.4	50.8	17.8	35.1	104.4
N	5	5	5	5	5	5	5	5
MEAN	108.66	931.2	16.46	46.64	50.10	17.68	35.30	124.96
S.D.	24.19	22.4	0.37	1.25	1.18	0.30	0.20	13.67
S.E.	10.82	10.0	0.17	0.56	0.53	0.14	0.09	6.11
M/C	0.8876	1.4265	1.2091	0.6334	0.5871	0.0375	0.0054	0.0052
F	1.7350	1.6339	2.2685	2.3953	0.1206	0.0098	0.4156	0.3228

INDIVIDUAL DATA 7-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
405	127.2	859	15.6	44.0	51.2	18.2	35.5	130.3
406	117.3	943	16.5	46.6	49.4	17.5	35.4	129.2
409	144.0	956	16.8	47.6	49.8	17.6	35.3	124.1
410	132.1	880	15.2	43.3	49.2	17.3	35.1	119.1
411	105.8	879	15.6	43.8	49.8	17.7	35.6	98.0
N	5	5	5	5	5	5	5	5
MEAN	125.28	903.4	15.94	45.06	49.88	17.66	35.38	120.14
S.D.	14.52	43.2	0.68	1.91	0.78	0.34	0.19	13.16
S.E.	6.50	19.3	0.30	0.86	0.35	0.15	0.09	5.88

INDIVIDUAL DATA 7-3-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	Reticulocyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
102	3.01	19.8	28.0	11.8	51.8	2.1	0.8	0.0
105	2.76	19.3	30.3	32.2	81.5	4.6	1.9	0.0
107	3.19	20.6	28.5	10.8	100.4	1.9	3.0	0.1
108	3.35	17.9	26.4	18.0	87.7	4.8	2.4	0.0
111	3.01	17.2	24.8	13.6	109.1	3.5	1.2	0.1
N	5	5	5	5	5	5	5	5
MEAN	3.064	18.96	27.60	17.28	86.10	3.38	1.86	0.04
S.D.	0.221	1.39	2.09	8.78	21.99	1.36	0.89	0.05
S.E.	0.099	0.62	0.94	3.93	9.83	0.61	0.40	0.02
M/C	0.0299	0.0651	3.7171	4.4768 *	1.1943	0.7464	0.0716	∞ **
F	5.8767 *	2.6540	0.7262		1.0529	1.8000	0.5793	
H				1.8436				2.2500

INDIVIDUAL DATA 7-3-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	Reticulocyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
405	3.55	20.4	27.7	23.8	98.2	3.6	1.6	0.0
406	3.01	20.5	27.0	23.0	88.5	5.3	0.5	0.0
409	3.62	18.0	25.8	23.4	114.4	4.7	1.5	0.0
410	3.52	21.3	26.6	21.5	103.2	4.8	2.6	0.0
411	3.40	22.3	26.7	17.6	83.8	3.3	1.1	0.0
N	5	5	5	5	5	5	5	5
MEAN	3.420	20.50	26.76	21.86	97.62	4.34	1.46	0.00
S.D.	0.243	1.59	0.69	2.54	12.12	0.85	0.77	0.00
S.E.	0.108	0.71	0.31	1.13	5.42	0.38	0.34	0.00

t' 2.4242 *

INDIVIDUAL DATA 7-4-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
163	90.3	897	15.3	42.8	47.7	17.1	35.7	103.8
164	68.0	854	15.8	44.8	52.5	18.5	35.3	115.8
165	68.5	885	15.8	44.5	50.3	17.9	35.5	97.6
166	75.9	805	15.3	43.6	54.2	19.0	35.1	120.4
167	72.7	894	16.4	45.9	51.3	18.3	35.7	99.6
N	5	5	5	5	5	5	5	5
MEAN	75.08	867.0	15.72	44.32	51.20	18.16	35.46	107.44
S.D.	9.10	38.6	0.45	1.18	2.44	0.71	0.26	10.12
S.E.	4.07	17.3	0.20	0.53	1.09	0.32	0.12	4.52
M/C	1.1627	1.5339	0.0752	0.0933	0.8756	0.1933	1.2707	1.4165
F	0.3617	0.1461	0.9298	0.6541	0.0892	0.2930	0.2392	0.1602

INDIVIDUAL DATA 7-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	WBC $10^3/\mu\text{L}$	RBC $10^6/\mu\text{L}$	HGB g/dL	HCT %	MCV fL	MCH pg	MCHC g/dL	Platelet $10^3/\mu\text{L}$
463	105.6	878	16.3	45.7	52.1	18.6	35.7	112.4
464	69.0	873	15.4	43.4	49.7	17.6	35.5	124.6
465	84.3	857	14.9	41.9	48.9	17.4	35.6	119.8
466	78.3	862	15.2	44.1	51.2	17.6	34.5	122.4
467	63.4	828	15.3	43.2	52.2	18.5	35.4	77.6
N	5	5	5	5	5	5	5	5
MEAN	80.12	859.6	15.42	43.66	50.82	17.94	35.34	111.36
S.D.	16.38	19.6	0.53	1.39	1.47	0.56	0.48	19.42
S.E.	7.33	8.7	0.24	0.62	0.66	0.25	0.22	8.69

INDIVIDUAL DATA 7-4-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	Reticu- locyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
163	2.68	15.7	18.9	8.2	78.9	2.2	1.0	0.0
164	2.69	18.1	19.4	4.5	60.1	2.4	1.0	0.0
165	2.62	15.9	21.1	7.5	57.1	2.8	1.1	0.0
166	2.80	17.7	18.3	8.2	65.2	1.8	0.7	0.0
167	2.63	18.5	19.7	9.0	59.9	2.5	1.3	0.0
N	5	5	5	5	5	5	5	5
MEAN	2.684	17.18	19.48	7.48	64.24	2.34	1.02	0.00
S.D.	0.072	1.29	1.05	1.75	8.70	0.37	0.22	0.00
S.E.	0.032	0.58	0.47	0.78	3.89	0.17	0.10	0.00
M/C	6.1399 *	2.4634	0.1362	0.6219	0.5778	6.8859 **	3.7905	0.0000
F		2.0674	0.0358	0.9067	0.2312		2.1417	0.0000
H	5.7709 *				2.4545			

INDIVIDUAL DATA 7-4-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Hematological findings, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	Reticu- locyte %	PT sec	APTT sec	Differential count of WBC ($10^2/\mu\text{L}$)				
				Neutrophil	Lymphocyte	Monocyte	Eosinophil	Basophil
463	2.71	16.8	20.6	13.2	84.7	5.5	2.2	0.0
464	3.51	16.6	20.4	7.1	60.4	0.9	0.6	0.0
465	2.84	15.7	19.1	7.8	73.6	1.7	1.2	0.0
466	3.28	15.7	20.4	6.6	69.2	1.2	1.3	0.0
467	3.07	16.6	17.6	9.5	50.2	1.6	2.1	0.0
N	5	5	5	5	5	5	5	5
MEAN	3.082	16.28	19.62	8.84	67.62	2.18	1.48	0.00
S.D.	0.324	0.54	1.28	2.67	13.09	1.88	0.67	0.00
S.E.	0.145	0.24	0.57	1.20	5.85	0.84	0.30	0.00
U	1.0000 *							

INDIVIDUAL DATA 8-1-1

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L				
				Albumin	Globulin											
					α_1	α_2	β	γ								
101	5.6	2.91	1.09	52.1	17.1	8.2	17.8	4.8	79	36	428	0.5				
103	5.4	2.73	1.03	50.7	22.9	7.5	15.4	3.5	57	25	306	0.4				
104	5.4	2.80	1.07	51.7	19.0	8.2	16.2	4.9	68	28	367	0.2				
106	5.8	2.84	0.96	49.1	23.6	7.4	15.5	4.4	58	33	369	0.4				
109	5.7	2.88	1.02	50.6	22.2	7.8	15.1	4.3	58	27	316	0.3				
N	5	5	5	5	5	5	5	5	5	5	5	5				
MEAN	5.58	2.832	1.034	50.84	20.96	7.82	16.00	4.38	64.0	29.8	357.2	0.36				
S.D.	0.18	0.070	0.050	1.17	2.78	0.38	1.08	0.55	9.5	4.5	48.9	0.11				
S.E.	0.08	0.032	0.022	0.52	1.25	0.17	0.48	0.25	4.3	2.0	21.9	0.05				
M/C	1.8995	3.0038	5.0896	4.9758	2.3171	2.5836	1.9033	5.0535	0.3083	0.2855	3.7105	0.9268				
F	1.6296	2.2500	0.3663	0.3003	0.6818	1.2668	0.5724	2.1553	1.3557	0.5359	2.7700 †	1.5595				

INDIVIDUAL DATA 8-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L
				Albumin	α_1	α_2	β	γ				
201	5.9	2.95	1.00	50.1	23.7	6.9	15.0	4.3	74	36	317	0.4
203	5.8	2.89	0.99	49.7	20.5	7.8	15.9	6.1	80	43	471	0.2
204	5.4	2.93	1.19	54.2	18.8	6.9	16.5	3.6	75	29	356	0.4
205	5.7	2.81	0.97	49.3	24.8	6.9	15.4	3.6	58	30	267	0.3
206	5.7	2.78	0.95	48.7	23.0	7.7	16.0	4.6	77	31	414	0.4
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	5.70	2.872	1.020	50.40	22.16	7.24	15.76	4.44	72.8	33.8	365.0	0.34
S.D.	0.19	0.074	0.097	2.19	2.45	0.47	0.58	1.03	8.6	5.8	80.0	0.09
S.E.	0.08	0.033	0.043	0.98	1.10	0.21	0.26	0.46	3.8	2.6	35.8	0.04

t'

0.1383

INDIVIDUAL DATA 8-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L
				Albumin	α_1	α_2	β	γ				
302	5.8	2.77	0.91	47.7	24.1	7.8	16.2	4.2	63	32	365	0.4
303	5.8	2.82	0.95	48.7	20.2	7.5	18.2	5.4	71	32	531	0.5
305	5.5	2.82	1.04	51.1	21.6	6.6	15.9	4.8	88	38	475	0.4
306	5.4	2.66	0.97	49.3	20.4	8.3	16.8	5.2	80	31	384	0.6
308	5.4	2.78	1.06	51.6	20.9	6.8	15.7	5.0	61	25	379	0.3
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	5.58	2.770	0.986	49.68	21.44	7.40	16.56	4.92	72.6	31.6	426.8	0.44
S.D.	0.20	0.066	0.063	1.64	1.58	0.70	1.01	0.46	11.4	4.6	72.7	0.11
S.E.	0.09	0.029	0.028	0.73	0.71	0.31	0.45	0.21	5.1	2.1	32.5	0.05

t'

1.2344

INDIVIDUAL DATA 8-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of administration

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L
				Albumin	α_1	α_2	β	γ				
401	5.4	2.68	0.99	49.6	18.4	7.6	16.8	7.6	75	34	451	0.3
402	5.3	2.95	1.25	55.6	14.6	7.9	17.2	4.7	86	36	634	0.2
403	4.9	2.59	1.13	53.0	21.5	7.1	14.2	4.2	66	26	302	0.3
404	5.8	2.67	0.85	46.0	24.3	7.2	16.8	5.7	86	36	508	0.5
407	5.5	2.78	1.02	50.5	20.1	7.3	15.9	6.2	67	27	606	0.1
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	5.38	2.734	1.048	50.94	19.78	7.42	16.18	5.68	76.0	31.8	500.2	0.28
S.D.	0.33	0.138	0.151	3.62	3.61	0.33	1.20	1.33	9.8	4.9	133.1	0.15
S.E.	0.15	0.062	0.067	1.62	1.62	0.15	0.54	0.60	4.4	2.2	59.5	0.07

t'

2.5361

INDIVIDUAL DATA 8-1-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
101	0.07	154	59	33	13.8	0.52	144	4.56	106	9.4	7.1
103	0.07	146	74	27	15.2	0.53	143	5.01	107	9.6	7.6
104	0.05	133	70	26	15.0	0.56	147	4.33	109	9.4	6.8
106	0.06	148	67	39	13.4	0.57	144	4.97	106	9.8	7.6
109	0.06	160	63	68	15.3	0.55	143	5.54	106	9.8	7.8
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.062	148.2	66.6	38.6	14.54	0.546	144.2	4.882	106.8	9.60	7.38
S.D.	0.008	10.1	5.9	17.2	0.88	0.021	1.6	0.465	1.3	0.20	0.41
S.E.	0.004	4.5	2.6	7.7	0.39	0.009	0.7	0.208	0.6	0.09	0.19
M/C	1.1495	2.0416	5.6105	3.3852	4.6817	3.4679	1.8306	1.5764	3.1738	3.1574	0.4508
F	0.4848	1.6681	2.4019	4.5145 *	3.0351 †	1.4453	0.7719	0.2785	2.9333 †	0.3810	1.2335

INDIVIDUAL DATA 8-1-6

STUDY NO. SR07125 **TITLE :** PFHxD_A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Biochemical findings **ANIMAL :** Rat, Crl:CD(SD) **SEX :** Male **GROUP :** 4 mg/kg **PERIOD :** End of administration

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
201	0.06	168	67	51	14.0	0.62	143	5.00	106	10.0	6.5
203	0.07	143	39	43	15.2	0.63	147	4.27	108	9.7	7.4
204	0.06	179	56	78	12.2	0.48	145	5.15	112	9.2	6.8
205	0.04	160	46	79	14.9	0.58	143	5.05	107	9.6	6.2
206	0.05	153	39	35	16.3	0.58	144	4.75	108	9.4	7.2
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.056	160.6	49.4	57.2	14.52	0.578	144.4	4.844	108.2	9.58	6.82
S.D.	0.011	13.8	12.1	20.3	1.54	0.059	1.7	0.353	2.3	0.30	0.49
S.E.	0.005	6.2	5.4	9.1	0.69	0.027	0.7	0.158	1.0	0.14	0.22

t' 1.9115 0.0172 1.40000

INDIVIDUAL DATA 8-1-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
302	0.08	166	51	35	16.0	0.64	143	4.69	106	9.6	7.2
303	0.04	167	92	57	13.6	0.59	144	4.35	108	9.7	6.7
305	0.06	146	41	45	13.0	0.55	146	4.74	108	9.4	7.5
306	0.06	162	52	20	16.7	0.54	145	5.02	107	9.6	7.6
308	0.05	149	40	38	15.6	0.55	144	4.80	108	9.5	6.8
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.058	158.0	55.2	39.0	14.98	0.574	144.4	4.720	107.4	9.56	7.16
S.D.	0.015	9.8	21.3	13.6	1.60	0.042	1.1	0.242	0.9	0.11	0.40
S.E.	0.007	4.4	9.5	6.1	0.71	0.019	0.5	0.108	0.4	0.05	0.18
t'				0.0411	0.3775				0.6000		

INDIVIDUAL DATA 8-1-8

STUDY NO. SR07125 **TITLE :** PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Biochemical findings **ANIMAL :** Rat, Crl:CD(SD) **SEX :** Male **GROUP :** 100 mg/kg **PERIOD :** End of administration

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
401	0.08	158	45	22	20.4	0.57	144	4.93	108	9.2	6.4
402	0.06	148	59	33	13.7	0.51	146	4.63	111	9.4	6.6
403	0.07	155	30	14	16.5	0.47	145	5.40	111	9.3	7.2
404	0.06	142	49	22	20.2	0.58	146	4.94	108	9.8	7.7
407	0.05	148	40	16	16.8	0.51	146	4.64	110	9.6	7.4
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.064	150.2	44.6	21.4	17.52	0.528	145.4	4.908	109.6	9.46	7.06
S.D.	0.011	6.3	10.7	7.4	2.81	0.046	0.9	0.313	1.5	0.24	0.55
S.E.	0.005	2.8	4.8	3.3	1.26	0.021	0.4	0.140	0.7	0.11	0.24

t' 1.7677 2.5568 2.8000 *
t 1.7677 2.5568 2.8000 *

INDIVIDUAL DATA 8-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 6 of lactation

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L
				Albumin	α_1	α_2	β	γ				
152	6.4	3.34	1.10	52.3	17.3	7.9	18.7	3.8	111	41	258	0.8
153	6.7	3.40	1.03	50.8	21.9	6.5	15.7	5.1	76	27	220	0.7
155	6.4	3.44	1.16	53.7	17.0	6.4	17.2	5.7	85	24	179	0.8
156	6.7	3.34	0.99	49.9	20.9	6.8	17.3	5.1	74	27	190	0.9
162	6.3	3.32	1.11	52.6	20.3	7.8	16.1	3.2	66	25	223	1.0
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	6.50	3.368	1.078	51.86	19.48	7.08	17.00	4.58	82.4	28.8	214.0	0.84
S.D.	0.19	0.050	0.068	1.51	2.20	0.72	1.17	1.04	17.4	6.9	31.0	0.11
S.E.	0.08	0.022	0.030	0.67	0.99	0.32	0.53	0.46	7.8	3.1	13.9	0.05
M/C	10.7576 *	10.2196 *	1.7570	2.1699	3.0273	5.3521	0.5588	1.8340	7.5040	7.7470	7.3574	4.4651
F			0.8411	0.8578	0.4538	0.4848	0.4201	0.4083	0.6410	1.0674	0.1318	0.1813
H	3.1499	0.9328										

INDIVIDUAL DATA 8-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 6 of lactation

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L
				Albumin	α_1	α_2	β	γ				
251	6.7	3.49	1.08	52.0	19.6	6.9	17.1	4.4	74	33	200	1.0
253	6.5	3.22	0.98	49.4	21.9	7.9	17.2	3.6	73	25	196	0.7
255	6.6	3.36	1.04	50.9	20.6	8.0	16.8	3.7	77	30	220	1.0
256	6.4	3.20	1.00	50.0	20.0	7.2	16.7	6.1	88	33	232	0.6
262	6.3	2.95	0.88	46.9	20.5	8.2	19.4	5.0	84	29	201	0.6
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	6.50	3.244	0.996	49.84	20.52	7.64	17.44	4.56	79.2	30.0	209.8	0.78
S.D.	0.16	0.202	0.075	1.91	0.87	0.56	1.11	1.03	6.5	3.3	15.5	0.20
S.E.	0.07	0.090	0.034	0.86	0.39	0.25	0.50	0.46	2.9	1.5	6.9	0.09

INDIVIDUAL DATA 8-2-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 6 of lactation

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L
				Albumin	α_1	α_2	β	γ				
351	6.3	3.44	1.20	54.6	17.3	7.2	16.3	4.6	75	23	218	0.6
352	6.3	3.33	1.12	52.8	19.9	7.2	16.2	3.9	68	24	325	0.7
353	5.8	2.94	1.02	50.7	19.2	8.8	17.9	3.4	95	25	179	0.8
355	7.1	3.65	1.06	51.4	21.2	5.9	16.8	4.7	92	29	159	0.5
361	6.1	2.82	0.86	46.1	20.4	10.4	19.1	4.0	89	30	235	1.4
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	6.32	3.236	1.052	51.12	19.60	7.90	17.26	4.12	83.8	26.2	223.2	0.80
S.D.	0.48	0.347	0.127	3.18	1.48	1.73	1.23	0.54	11.7	3.1	64.4	0.35
S.E.	0.22	0.155	0.057	1.42	0.66	0.78	0.55	0.24	5.2	1.4	28.8	0.16

INDIVIDUAL DATA 8-2-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 6 of lactation

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L
				Albumin	α_1	α_2	β	γ				
451	6.5	3.51	1.17	53.9	20.8	6.0	15.9	3.4	77	29	164	0.9
453	6.5	3.23	0.99	49.8	21.8	8.1	16.6	3.7	72	33	162	1.1
455	6.3	3.36	1.14	53.4	19.0	7.7	14.8	5.1	81	30	313	0.6
457	6.5	3.44	1.13	53.0	18.6	7.0	16.5	4.9	73	30	190	1.0
462	6.4	3.14	0.96	49.1	19.5	8.6	19.1	3.7	71	31	195	0.8
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	6.44	3.336	1.078	51.84	19.94	7.48	16.58	4.16	74.8	30.6	204.8	0.88
S.D.	0.09	0.151	0.096	2.22	1.33	1.01	1.58	0.78	4.1	1.5	62.3	0.19
S.E.	0.04	0.068	0.043	0.99	0.59	0.45	0.71	0.35	1.9	0.7	27.9	0.09

INDIVIDUAL DATA 8-2-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 6 of lactation

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
152	0.07	127	60	50	26.4	0.58	141	4.79	104	11.0	8.7
153	0.09	136	63	19	22.4	0.59	140	4.94	104	11.0	7.9
155	0.08	177	43	16	24.5	0.72	142	4.19	105	10.8	8.8
156	0.08	159	69	38	28.3	0.69	140	5.30	103	11.0	9.6
162	0.08	182	53	57	27.3	0.66	141	5.19	104	10.9	8.6
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.080	156.2	57.6	36.0	25.78	0.648	140.8	4.882	104.0	10.94	8.72
S.D.	0.007	24.3	10.0	18.2	2.35	0.061	0.8	0.436	0.7	0.09	0.61
S.E.	0.003	10.9	4.5	8.2	1.05	0.027	0.4	0.195	0.3	0.04	0.27
M/C	∞ **	3.5321	0.3652	3.6081	4.6129	5.1168	1.9933	1.4195	3.8503	7.5746	9.9196 *
F		1.1793	0.3487	3.0440 †	3.1352 †	0.2196	3.0123 †	0.5675	5.7778 **	0.1575	
H		10.3957 *									4.8398

INDIVIDUAL DATA 8-2-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 6 of lactation

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
251	0.07	151	66	89	25.6	0.62	142	5.42	105	10.5	8.6
253	0.08	132	46	39	31.0	0.69	143	4.78	105	10.7	9.0
255	0.07	150	64	52	28.0	0.62	143	4.91	105	11.4	8.3
256	0.08	124	55	49	28.0	0.69	141	5.41	106	11.0	9.1
262	0.08	138	33	56	26.5	0.74	142	4.70	104	10.7	8.7
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.076	139.0	52.8	57.0	27.82	0.672	142.2	5.044	105.0	10.86	8.74
S.D.	0.005	11.6	13.6	19.0	2.05	0.052	0.8	0.347	0.7	0.35	0.32
S.E.	0.002	5.2	6.1	8.5	0.92	0.023	0.4	0.155	0.3	0.16	0.14
t' U	8.5000			2.2736	1.1472		1.9052		1.3608		

INDIVIDUAL DATA 8-2-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 6 of lactation

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
351	0.06	149	40	23	25.4	0.61	143	4.37	108	10.5	8.0
352	0.08	167	55	24	23.7	0.68	141	4.86	106	10.7	7.6
353	0.06	132	51	39	29.8	0.61	145	4.22	107	10.6	7.9
355	0.09	147	73	38	27.2	0.66	142	4.68	105	11.1	8.1
361	0.07	122	65	44	35.0	0.82	142	5.56	105	11.7	10.8
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.072	143.4	56.8	33.6	28.22	0.676	142.6	4.738	106.2	10.92	8.48
S.D.	0.013	17.2	12.7	9.5	4.41	0.086	1.5	0.524	1.3	0.49	1.31
S.E.	0.006	7.7	5.7	4.2	1.97	0.039	0.7	0.234	0.6	0.22	0.59
t' U	7.5000			0.2598	1.3721		2.4495		2.9938 *		

INDIVIDUAL DATA 8-2-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 6 of lactation

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
451	0.06	122	34	22	31.6	0.67	145	4.80	108	10.5	7.8
453	0.06	147	45	34	32.8	0.71	142	5.28	107	11.2	8.2
455	0.06	139	57	29	30.4	0.66	143	4.98	107	10.6	8.2
457	0.06	144	51	44	32.2	0.65	142	4.65	104	10.5	8.7
462	0.06	143	67	38	28.9	0.66	142	5.31	108	11.2	8.4
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.060	139.0	50.8	33.4	31.18	0.670	142.8	5.004	106.8	10.80	8.26
S.D.	0.000	9.9	12.4	8.4	1.55	0.023	1.3	0.290	1.6	0.37	0.33
S.E.	0.000	4.4	5.6	3.8	0.69	0.010	0.6	0.130	0.7	0.16	0.15
t U				0.2815	3.0366 *		2.7217 *		3.8103 **		

INDIVIDUAL DATA 8-3-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L
				Albumin	α_1	α_2	β	γ				
102	5.9	2.84	0.93	48.1	23.9	7.2	15.9	4.9	78	39	267	0.2
105	5.6	2.79	0.99	49.8	24.1	7.0	15.5	3.6	75	30	369	0.7
107	6.0	3.09	1.06	51.5	21.5	6.8	14.3	5.9	102	62	352	0.4
108	5.8	2.75	0.90	47.4	20.4	8.1	18.3	5.8	65	34	227	0.6
111	5.9	2.82	0.91	47.6	22.4	7.5	17.0	5.5	68	23	213	0.7
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	5.84	2.858	0.958	48.88	22.46	7.32	16.20	5.14	77.6	37.6	285.6	0.52
S.D.	0.15	0.134	0.067	1.74	1.58	0.51	1.52	0.94	14.6	14.8	71.4	0.22
S.E.	0.07	0.060	0.030	0.78	0.70	0.23	0.68	0.42	6.5	6.6	31.9	0.10
M/C	0.9971	0.0053	0.6745	0.6341	1.2198	1.7598	0.2952	6.5922 *	5.5128 *	12.4211 **	0.1210	0.3687
F	2.1978	1.2231	0.0118	0.0096	2.3294	1.0642	1.8673				5.1466 †	1.1825
H								0.1006	4.8109 *	2.4695		

INDIVIDUAL DATA 8-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L
				Albumin	α_1	α_2	β	γ				
405	5.5	2.86	1.08	51.9	19.7	7.3	15.5	5.6	67	28	279	0.4
406	5.9	2.77	0.88	47.0	21.7	7.3	18.4	5.6	59	27	459	0.4
409	5.9	2.90	0.97	49.2	20.6	7.8	17.2	5.2	60	29	502	0.8
410	5.6	2.57	0.84	45.7	23.4	7.2	18.1	5.6	64	29	374	1.1
411	5.3	2.73	1.05	51.3	15.7	9.7	17.6	5.7	66	26	381	0.8
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	5.64	2.766	0.964	49.02	20.22	7.86	17.36	5.54	63.2	27.8	399.0	0.70
S.D.	0.26	0.129	0.104	2.68	2.88	1.05	1.14	0.19	3.6	1.3	86.0	0.30
S.E.	0.12	0.058	0.047	1.20	1.29	0.47	0.51	0.09	1.6	0.6	38.4	0.13

2.2686

2.0000 *

INDIVIDUAL DATA 8-3-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Chol mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
102	0.05	179	66	38	15.5	0.58	144	4.38	105	10.0	7.0
105	0.08	142	79	13	17.4	0.56	144	4.33	106	9.7	6.6
107	0.07	167	43	15	15.2	0.55	146	4.61	106	9.6	7.5
108	0.05	190	88	50	15.2	0.62	142	5.29	106	9.7	7.6
111	0.07	139	50	12	17.5	0.55	145	4.39	108	9.8	6.7
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.064	163.4	65.2	25.6	16.16	0.572	144.2	4.600	106.2	9.76	7.08
S.D.	0.013	22.5	18.9	17.4	1.18	0.029	1.5	0.400	1.1	0.15	0.45
S.E.	0.006	10.0	8.5	7.8	0.53	0.013	0.7	0.179	0.5	0.07	0.20
M/C	2.5378	0.9673	0.7763	0.1552	0.0341	0.0020	3.0642	2.9259	0.2295	0.0626	0.0049
F	9.5238 *	0.0853	3.3311	1.0785	1.7389	2.3059	0.0800	1.7864	5.0625 †	0.1509	2.4605

INDIVIDUAL DATA 8-3-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Chol mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
405	0.04	150	38	32	14.4	0.58	144	5.05	107	9.5	8.1
406	0.05	182	63	35	14.6	0.54	145	4.88	107	9.9	7.7
409	0.05	151	44	74	17.3	0.55	145	4.68	109	9.9	7.1
410	0.04	162	55	16	14.0	0.50	144	4.73	107	9.8	7.8
411	0.04	155	35	35	15.3	0.55	144	4.94	110	9.9	7.0
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.044	160.0	47.0	38.4	15.12	0.544	144.4	4.856	108.0	9.80	7.54
S.D.	0.005	13.2	11.8	21.4	1.31	0.029	0.5	0.152	1.4	0.17	0.47
S.E.	0.002	5.9	5.3	9.6	0.58	0.013	0.2	0.068	0.6	0.08	0.21

t' 3.0861 *

2.2500

INDIVIDUAL DATA 8-4-1

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Recovery group ANIMAL : Rat, CrI:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L
				Albumin	α_1	α_2	β	γ				
163	6.5	3.80	1.41	58.5	16.4	6.1	13.4	5.6	66	26	145	1.0
164	6.6	3.91	1.45	59.1	19.0	5.3	12.8	3.8	62	37	130	0.7
165	6.3	3.49	1.24	55.4	16.7	6.5	13.6	7.8	71	24	156	0.6
166	6.6	3.95	1.50	59.9	17.4	5.6	12.5	4.6	60	30	151	0.5
167	6.4	3.61	1.30	56.5	19.4	5.6	11.8	6.7	59	24	92	0.6
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	6.48	3.752	1.380	57.88	17.78	5.82	12.82	5.70	63.6	28.2	134.8	0.68
S.D.	0.13	0.197	0.107	1.87	1.35	0.48	0.72	1.60	4.9	5.5	25.8	0.19
S.E.	0.06	0.088	0.048	0.84	0.61	0.21	0.32	0.72	2.2	2.5	11.6	0.09
M/C	1.3947	0.0405	0.1766	0.1806	0.9956	1.3260	1.8899	1.6551	2.1422	0.3662	1.7768	0.0803
F	4.2785 †	1.4368	0.0243	0.0222	2.6516	4.6758 †	2.8084	0.3931	0.0054	0.9101	1.4837	0.0230

INDIVIDUAL DATA 8-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	TP g/dL	Albumin g/dL	A/G ratio	Protein fraction (%)					AST IU/L	ALT IU/L	ALP IU/L	γ -GTP IU/L				
				Albumin	Globulin											
					α_1	α_2	β	γ								
463	6.4	3.82	1.48	59.7	16.8	5.8	11.9	5.8	57	19	138	0.7				
464	5.9	3.59	1.55	60.8	12.6	8.0	14.1	4.5	57	23	105	0.4				
465	6.0	3.37	1.28	56.2	14.2	7.4	16.1	6.1	53	18	186	0.6				
466	6.4	3.74	1.42	58.5	17.4	6.2	13.6	4.3	69	24	248	1.0				
467	6.4	3.53	1.23	55.2	18.1	6.6	14.8	5.3	80	37	160	0.8				
N	5	5	5	5	5	5	5	5	5	5	5	5				
MEAN	6.22	3.610	1.392	58.08	15.82	6.80	14.10	5.20	63.2	24.2	167.4	0.70				
S.D.	0.25	0.177	0.134	2.35	2.33	0.89	1.55	0.79	11.1	7.6	54.0	0.22				
S.E.	0.11	0.079	0.060	1.05	1.04	0.40	0.69	0.35	5.0	3.4	24.1	0.10				

t' 2.0684

2.1623

INDIVIDUAL DATA 8-4-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Recovery group ANIMAL : Rat, CrI:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Cho mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
163	0.08	148	85	12	17.2	0.63	143	4.41	109	10.3	5.7
164	0.08	157	70	13	16.2	0.57	146	4.44	110	10.3	6.4
165	0.08	141	65	11	15.0	0.67	145	4.18	108	10.3	6.6
166	0.11	150	57	11	17.2	0.63	145	3.88	109	10.3	6.1
167	0.07	148	69	10	17.7	0.60	144	4.39	108	10.4	6.8
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.084	148.8	69.2	11.4	16.66	0.620	144.6	4.260	108.8	10.32	6.32
S.D.	0.015	5.7	10.2	1.1	1.08	0.037	1.1	0.236	0.8	0.04	0.43
S.E.	0.007	2.6	4.6	0.5	0.48	0.017	0.5	0.105	0.4	0.02	0.19
M/C	0.2855	1.6756	0.0278	6.8149 **	0.0005	0.4957	0.7824	2.8284	1.3123	8.2334 **	0.0516
F	12.5000 **	0.0959	4.9723 †		2.8715	2.4155	0.4444	0.1567	5.5556 *		2.0753
H				4.0613 *						0.4208	

INDIVIDUAL DATA 8-4-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Biochemical findings, Recovery group ANIMAL : Rat, CrI:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	T-Bil mg/dL	Glucose mg/dL	T-Chol mg/dL	TG mg/dL	UN mg/dL	Crea mg/dL	Na mEq/L	K mEq/L	Cl mEq/L	Ca mg/dL	IP mg/dL
463	0.05	141	41	18	16.7	0.62	145	4.97	110	10.7	7.4
464	0.06	149	56	26	16.2	0.54	145	4.17	110	10.0	7.0
465	0.04	151	62	18	14.3	0.50	144	4.47	110	10.2	6.6
466	0.05	142	65	11	15.9	0.63	146	3.63	109	10.1	6.6
467	0.07	170	53	23	14.4	0.58	145	3.48	110	10.4	6.1
N	5	5	5	5	5	5	5	5	5	5	5
MEAN	0.054	150.6	55.4	19.2	15.50	0.574	145.0	4.144	109.8	10.28	6.74
S.D.	0.011	11.7	9.3	5.7	1.09	0.055	0.7	0.611	0.4	0.28	0.49
S.E.	0.005	5.2	4.2	2.6	0.49	0.024	0.3	0.273	0.2	0.12	0.22
t [†]	3.5355 **		2.2299						2.3570 *		
U				3.0000							

INDIVIDUAL DATA 9-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Serum hormone levels ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
101	0.41	75.482	6.07
103	0.51	79.707	2.42
104	0.41	68.949	2.52
106	0.38	44.288	3.49
109	0.54	80.136	4.16
N	5	5	5
MEAN	0.450	69.7124	3.732
S.D.	0.070	14.9065	1.491
S.E.	0.031	6.6664	0.667
M/C	2.0503	5.5537	7.0948
F	0.7482	0.9636	1.5943

INDIVIDUAL DATA 9-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Serum hormone levels ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
201	0.37	59.484	5.04
203	0.46	82.150	6.44
204	0.58	76.445	5.29
205	0.44	86.976	4.85
206	0.48	68.598	11.31
N	5	5	5
MEAN	0.466	74.7306	6.586
S.D.	0.076	10.9317	2.712
S.E.	0.034	4.8888	1.213

INDIVIDUAL DATA 9-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Serum hormone levels ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
302	0.45	74.635	4.95
303	0.42	71.314	16.11
305	0.39	93.549	2.58
306	0.29	78.032	7.44
308	0.40	82.700	4.24
N	5	5	5
MEAN	0.390	80.0460	7.064
S.D.	0.060	8.6452	5.351
S.E.	0.027	3.8662	2.393

INDIVIDUAL DATA 9-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Serum hormone levels ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of administration

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
401	0.28	77.710	20.10
402	0.54	67.620	7.03
403	0.41	71.566	9.09
404	0.38	69.410	7.46
407	0.57	70.609	4.73
N	5	5	5
MEAN	0.436	71.3830	9.682
S.D.	0.119	3.8313	6.029
S.E.	0.053	1.7134	2.696

INDIVIDUAL DATA 9-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Serum hormone levels, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 6 of lactation

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
152	0.73	57.401	5.27
153	0.74	60.341	2.54
155	0.72	81.219	3.47
156	0.77	65.540	4.95
162	0.71	62.915	6.16
N	5	5	5
MEAN	0.734	65.4832	4.478
S.D.	0.023	9.3002	1.454
S.E.	0.010	4.1592	0.650
M/C	4.3331	2.5929	6.2984
F	17.2796 **	0.1313	1.2221

INDIVIDUAL DATA 9-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Serum hormone levels, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 6 of lactation

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
251	0.58	82.545	3.59
253	0.56	39.907	2.37
255	0.65	63.610	14.55
256	0.62	71.082	2.82
262	0.62	70.636	3.84
N	5	5	5
MEAN	0.606	65.5560	5.434
S.D.	0.036	15.8637	5.130
S.E.	0.016	7.0945	2.294

t' 4.5086 **

INDIVIDUAL DATA 9-2-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Serum hormone levels, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 6 of lactation

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
351	0.59	50.032	5.12
352	0.62	67.786	2.10
353	0.62	68.392	3.24
355	0.74	59.182	8.10
361	0.56	63.895	3.48
N	5	5	5
MEAN	0.626	61.8574	4.408
S.D.	0.068	7.5664	2.329
S.E.	0.031	3.3838	1.041

t' 3.8041 **

INDIVIDUAL DATA 9-2-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Serum hormone levels, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 6 of lactation

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
451	0.48	83.947	7.17
453	0.56	74.079	3.05
455	0.58	53.806	5.53
457	0.53	71.697	14.81
462	0.51	48.274	11.13
N	5	5	5
MEAN	0.532	66.3606	8.338
S.D.	0.040	14.8499	4.661
S.E.	0.018	6.6411	2.084

t' 7.1151 **

INDIVIDUAL DATA 9-3-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Serum hormone levels ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
102	0.48	133.630	7.84
105	0.68	117.833	12.39
107	0.44	131.122	9.21
108	0.36	101.492	15.44
111	0.41	103.412	21.69
N	5	5	5
MEAN	0.474	117.4978	13.314
S.D.	0.123	15.0048	5.530
S.E.	0.055	6.7103	2.473
M/C	1.6589	0.1935	0.9831
F	0.1284	10.8958 *	0.0076

INDIVIDUAL DATA 9-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Serum hormone levels ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
405	0.52	90.366	10.96
406	0.39	87.444	10.98
409	0.50	102.157	17.77
410	0.46	95.766	11.81
411	0.39	70.540	16.30
N	5	5	5
MEAN	0.452	89.2546	13.564
S.D.	0.061	11.8703	3.229
S.E.	0.027	5.3086	1.444
t'		3.3009 *	

INDIVIDUAL DATA 9-4-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Serum hormone levels, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
163	0.96	35.046	2.82
164	0.91	47.784	4.50
165	0.74	74.292	3.49
166	0.66	33.043	4.81
167	0.65	41.118	3.17
N	5	5	5
MEAN	0.784	46.2566	3.758
S.D.	0.143	16.6958	0.859
S.E.	0.064	7.4666	0.384
M/C	6.0585 *	2.3288	24.6447 **
F		2.2691	
H	1.3613		0.8836

INDIVIDUAL DATA 9-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Serum hormone levels, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	T3 ng/mL	T4 ng/mL	TSH ng/mL
463	0.70	57.966	4.63
464	0.70	65.049	2.49
465	0.65	66.254	3.61
466	0.72	50.432	127.11
467	0.65	52.701	6.02
N	5	5	5
MEAN	0.684	58.4804	28.772
S.D.	0.032	7.1067	54.988
S.E.	0.014	3.1782	24.591

INDIVIDUAL DATA 10-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

Animal No.	Findings	Day of euthanasia
101	No abnormal findings	Next day of day 42 of administration
103	Liver: Atrophy, papillary process of caudate lobe Yellowish green discoloration, papillary process of caudate lobe	Next day of day 42 of administration
104	No abnormal findings	Next day of day 42 of administration
106	Kidney (right and left): Dilatation, renal pelvis	Next day of day 42 of administration
109	No abnormal findings	Next day of day 42 of administration
110	No abnormal findings	Next day of day 42 of administration
112	No abnormal findings	Next day of day 42 of administration

INDIVIDUAL DATA 10-1-2

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	Findings	Day of euthanasia
201	No abnormal findings	Next day of day 42 of administration
202	No abnormal findings	Next day of day 42 of administration
203	No abnormal findings	Next day of day 42 of administration
204	No abnormal findings	Next day of day 42 of administration
205	No abnormal findings	Next day of day 42 of administration
206	No abnormal findings	Next day of day 42 of administration
207	No abnormal findings	Next day of day 42 of administration
208	No abnormal findings	Next day of day 42 of administration
209	No abnormal findings	Next day of day 42 of administration
210	No abnormal findings	Next day of day 42 of administration
211	No abnormal findings	Next day of day 42 of administration
212	No abnormal findings	Next day of day 42 of administration

INDIVIDUAL DATA 10-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	Findings	Day of euthanasia
301	No abnormal findings	Next day of day 42 of administration
302	No abnormal findings	Next day of day 42 of administration
303	No abnormal findings	Next day of day 42 of administration
304	No abnormal findings	Next day of day 42 of administration
305	No abnormal findings	Next day of day 42 of administration
306	No abnormal findings	Next day of day 42 of administration
307	No abnormal findings	Next day of day 42 of administration
308	No abnormal findings	Next day of day 42 of administration
309	Epididymis (left): Yellowish white patch, cauda (3 x 3, mm)	Next day of day 42 of administration
310	No abnormal findings	Next day of day 42 of administration
311	No abnormal findings	Next day of day 42 of administration
312	No abnormal findings	Next day of day 42 of administration

INDIVIDUAL DATA 10-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of administration

Animal No.	Findings	Day of euthanasia
401	No abnormal findings	Next day of day 42 of administration
402	No abnormal findings	Next day of day 42 of administration
403	No abnormal findings	Next day of day 42 of administration
404	No abnormal findings	Next day of day 42 of administration
407	No abnormal findings	Next day of day 42 of administration
408	No abnormal findings	Next day of day 42 of administration
412	No abnormal findings	Next day of day 42 of administration

INDIVIDUAL DATA 10-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 6 of lactation

Animal No.	Findings	Day of euthanasia
151	No abnormal findings	Day 6 of lactation
152	No abnormal findings	Day 6 of lactation
153	No abnormal findings	Day 6 of lactation
154	No abnormal findings	Day 6 of lactation
155	No abnormal findings	Day 6 of lactation
156	No abnormal findings	Day 6 of lactation
157	No abnormal findings	Day 6 of lactation
158	No abnormal findings	Day 6 of lactation
159	No abnormal findings	Day 6 of lactation
160	No abnormal findings	Day 6 of lactation
161 ^a	No abnormal findings	Day 26 of gestation
162	No abnormal findings	Day 6 of lactation

a: Non-pregnancy.

INDIVIDUAL DATA 10-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 6 of lactation

Animal No.	Findings	Day of euthanasia
251	No abnormal findings	Day 6 of lactation
252 ^a	No abnormal findings	Day 6 of lactation
253	No abnormal findings	Day 6 of lactation
254 ^a	No abnormal findings	Day 6 of lactation
255	No abnormal findings	Day 6 of lactation
256	No abnormal findings	Day 6 of lactation
257	No abnormal findings	Day 6 of lactation
258	No abnormal findings	Day 6 of lactation
259	No abnormal findings	Day 6 of lactation
260	Stomach: White mass, limiting ridge (1 x 1 x 1, mm)	Day 6 of lactation
261	No abnormal findings	Day 6 of lactation
262	No abnormal findings	Day 6 of lactation

a: All pups died by day 3 of lactation .

INDIVIDUAL DATA 10-2-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 6 of lactation

Animal No.	Findings	Day of cuthanasia
351	No abnormal findings	Day 6 of lactation
352	No abnormal findings	Day 6 of lactation
353	No abnormal findings	Day 6 of lactation
354	No abnormal findings	Day 6 of lactation
355	No abnormal findings	Day 6 of lactation
356	No abnormal findings	Day 6 of lactation
357	No abnormal findings	Day 6 of lactation
358	No abnormal findings	Day 6 of lactation
359	No abnormal findings	Day 6 of lactation
360	No abnormal findings	Day 6 of lactation
361	No abnormal findings	Day 6 of lactation
362	No abnormal findings	Day 6 of lactation

INDIVIDUAL DATA 10-2-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 6 of lactation

Animal No.	Findings	Day of euthanasia
451	No abnormal findings	Day 6 of lactation
452	No abnormal findings	Day 6 of lactation
453	No abnormal findings	Day 6 of lactation
454	No abnormal findings	Day 6 of lactation
455	No abnormal findings	Day 6 of lactation
456	No abnormal findings	Day 6 of lactation
457	No abnormal findings	Day 6 of lactation
458	No abnormal findings	Day 6 of lactation
459	No abnormal findings	Day 6 of lactation
460	Skin of right posterior abdominal region: Subcutaneous yellow mass (20 x 10 x 5, mm)	Day 6 of lactation
461	No abnormal findings	Day 6 of lactation
462	No abnormal findings	Day 6 of lactation

INDIVIDUAL DATA 10-3-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	Findings	Day of euthanasia
102	No abnormal findings	Next day of day 14 of recovery
105	No abnormal findings	Next day of day 14 of recovery
107	No abnormal findings	Next day of day 14 of recovery
108	No abnormal findings	Next day of day 14 of recovery
111	Ileum : Diverticulum (5 x 3 x 3, mm)	Next day of day 14 of recovery

INDIVIDUAL DATA 10-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	Findings	Day of euthanasia
405	No abnormal findings	Next day of day 14 of recovery
406	No abnormal findings	Next day of day 14 of recovery
409	No abnormal findings	Next day of day 14 of recovery
410	No abnormal findings	Next day of day 14 of recovery
411	No abnormal findings	Next day of day 14 of recovery

INDIVIDUAL DATA 10-4-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	Findings	Day of euthanasia
163	No abnormal findings	Next day of day 14 of recovery
164	No abnormal findings	Next day of day 14 of recovery
165	No abnormal findings	Next day of day 14 of recovery
166	No abnormal findings	Next day of day 14 of recovery
167	No abnormal findings	Next day of day 14 of recovery

INDIVIDUAL DATA 10-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Gross findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	Findings	Day of euthanasia
463	No abnormal findings	Next day of day 14 of recovery
464	No abnormal findings	Next day of day 14 of recovery
465	No abnormal findings	Next day of day 14 of recovery
466	No abnormal findings	Next day of day 14 of recovery
467	No abnormal findings	Next day of day 14 of recovery

INDIVIDUAL DATA 11-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain		Pituitary gland	
		g	g	%	g	%	g	%	g	%	g	%	mg	10 ⁻³ %
101	466	11.72	2.52		2.94	0.63	0.67	0.14	1.31	0.28	2.25	0.48	14.9	3.20
103	480	11.72	2.44		3.85	0.80	0.93	0.19	1.48	0.31	2.24	0.47	15.8	3.29
104	474	11.81	2.49		3.00	0.63	0.82	0.17	1.59	0.34	2.16	0.46	13.8	2.91
106	446	11.14	2.50		3.16	0.71	0.87	0.20	1.54	0.35	2.11	0.47	12.2	2.74
109	561	14.37	2.56		3.29	0.59	0.74	0.13	1.47	0.26	2.24	0.40	12.8	2.28
N		5	5	5	5	5	5	5	5	5	5	5	5	5
MEAN		485.4	12.152	2.502	3.248	0.672	0.806	0.166	1.478	0.308	2.200	0.456	13.90	2.884
S.D.		44.2	1.268	0.044	0.363	0.084	0.103	0.030	0.106	0.038	0.062	0.032	1.48	0.403
S.E.		19.8	0.567	0.020	0.163	0.037	0.046	0.014	0.047	0.017	0.028	0.014	0.66	0.180
M/C		9.5196 *	3.2686	5.5965	3.5892	6.5587	0.3163	0.8188	2.1104	2.6270	2.5325	2.5131	1.3291	3.0256
F		10.4025 **	79.3461 **		1.8642	2.8639 †	0.3619	0.4436	1.8638	1.2100	0.6259	1.8448	3.1453 †	1.7605
H		9.4614 *												

INDIVIDUAL DATA 11-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain		Pituitary gland	
		g	g	%	g	%	g	%	g	%	g	%	mg	10 ⁻³ %
201	468	11.86	2.53		2.75	0.59	0.65	0.14	1.15	0.25	2.00	0.43	13.4	2.86
203	479	11.21	2.34		2.96	0.62	0.77	0.16	1.46	0.30	2.20	0.46	12.9	2.69
204	488	11.82	2.42		3.25	0.67	0.65	0.13	1.50	0.31	2.21	0.45	11.0	2.25
205	495	12.66	2.56		2.83	0.57	0.82	0.17	1.29	0.26	2.12	0.43	10.1	2.04
206	483	11.49	2.38		2.94	0.61	0.85	0.18	1.36	0.28	2.09	0.43	11.6	2.40
N		5	5	5	5	5	5	5	5	5	5	5	5	5
MEAN		482.6	11.808	2.446	2.946	0.612	0.748	0.156	1.352	0.280	2.124	0.440	11.80	2.448
S.D.		10.1	0.545	0.095	0.190	0.038	0.094	0.021	0.140	0.025	0.086	0.014	1.35	0.330
S.E.		4.5	0.244	0.043	0.085	0.017	0.042	0.009	0.063	0.011	0.039	0.006	0.61	0.148
t' U			0.6282	0.9019		1.7011							2.7277 *	
		8.0000												

INDIVIDUAL DATA 11-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain		Pituitary gland	
		g	g	%	g	%	g	%	g	%	g	%	mg	10 ⁻³ %
302	451	11.34	2.51		2.79	0.62	0.57	0.13	1.34	0.30	2.19	0.49	12.9	2.86
303	495	13.53	2.73		3.00	0.61	0.92	0.19	1.32	0.27	2.02	0.41	14.5	2.93
305	485	12.09	2.49		2.76	0.57	0.74	0.15	1.43	0.29	2.16	0.45	12.6	2.60
306	491	11.59	2.36		3.06	0.62	0.74	0.15	1.30	0.26	2.28	0.46	12.7	2.59
308	509	12.03	2.36		3.04	0.60	0.77	0.15	1.44	0.28	2.28	0.45	13.8	2.71
N		5	5	5	5	5	5	5	5	5	5	5	5	5
MEAN		486.2	12.116	2.490	2.930	0.604	0.748	0.154	1.366	0.280	2.186	0.452	13.30	2.738
S.D.		21.6	0.849	0.151	0.144	0.021	0.124	0.022	0.065	0.016	0.107	0.029	0.82	0.153
S.E.		9.6	0.380	0.068	0.064	0.009	0.056	0.010	0.029	0.007	0.048	0.013	0.37	0.068
t' U			0.0657	0.1933		1.9280							0.7794	
		8.0000												

INDIVIDUAL DATA 11-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of administration

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain		Pituitary gland	
		g	g	%	g	%	g	%	g	%	g	%	mg	10 ⁻³ %
401	438	14.34	3.27		3.20	0.73	0.71	0.16	1.45	0.33	2.22	0.51	13.9	3.17
402	459	15.48	3.37		3.04	0.66	0.67	0.15	1.45	0.32	2.27	0.49	11.4	2.48
403	437	13.98	3.20		3.28	0.75	0.90	0.21	1.30	0.30	1.90	0.43	11.3	2.59
404	457	14.67	3.21		2.75	0.60	0.66	0.14	1.19	0.26	2.15	0.47	11.7	2.56
407	432	14.05	3.25		3.02	0.70	0.81	0.19	1.21	0.28	2.15	0.50	12.8	2.96
N		5	5	5	5	5	5	5	5	5	5	5	5	5
MEAN		444.6	14.504	3.260	3.058	0.688	0.750	0.170	1.320	0.298	2.138	0.480	12.22	2.752
S.D.		12.5	0.610	0.068	0.204	0.060	0.103	0.029	0.126	0.029	0.142	0.032	1.11	0.298
S.E.		5.6	0.273	0.030	0.091	0.027	0.046	0.013	0.056	0.013	0.064	0.014	0.50	0.133
t' U			4.2950 **	12.2084 **		0.4536								2.1822
			2.0000 *											

INDIVIDUAL DATA 11-1-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

Animal No.	Thymus		Thyroid		Adrenal		Prostate		Seminal vesicle	
	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	g	%
101	278	59.66	18.1	3.88	70	15.02	1085	232.83	2.54	0.55
103	342	71.25	20.7	4.31	67	13.96	835	173.96	2.14	0.45
104	284	59.92	19.7	4.16	70	14.77	822	173.42	1.90	0.40
106	335	75.11	19.6	4.39	73	16.37	757	169.73	2.27	0.51
109	349	62.21	16.6	2.96	70	12.48	750	133.69	2.55	0.45
N	5	5	5	5	5	5	5	5	5	5
MEAN	317.6	65.630	18.94	3.940	70.0	14.520	849.8	176.726	2.280	0.472
S.D.	33.8	7.096	1.60	0.581	2.1	1.433	136.8	35.592	0.276	0.058
S.E.	15.1	3.173	0.72	0.260	0.9	0.641	61.2	15.917	0.123	0.026
M/C	5.7964	3.7318	5.3622	2.3336	7.8468 *	1.2958	3.1436	2.0584	0.5533	0.4054
F	1.2406	1.5004	3.0436 †	3.6607 *		1.1033	1.7976	0.9502	0.5046	0.5066
H					5.6971					

INDIVIDUAL DATA 11-1-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	Thymus		Thyroid		Adrenal		Prostate		Seminal vesicle	
	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	g	%
201	207	44.23	20.2	4.32	53	11.32	1029	219.87	2.25	0.48
203	249	51.98	22.8	4.76	72	15.03	691	144.26	2.56	0.53
204	312	63.93	21.2	4.34	52	10.66	656	134.43	2.58	0.53
205	389	78.59	20.0	4.04	70	14.14	496	100.20	2.07	0.42
206	360	74.53	18.7	3.87	45	9.32	895	185.30	2.22	0.46
N	5	5	5	5	5	5	5	5	5	5
MEAN	303.4	62.652	20.58	4.266	58.4	12.094	753.4	156.812	2.336	0.484
S.D.	75.6	14.577	1.53	0.339	11.9	2.406	209.5	46.506	0.224	0.047
S.E.	33.8	6.519	0.68	0.152	5.3	1.076	93.7	20.798	0.100	0.021
t'			0.8914	0.8406						

INDIVIDUAL DATA 11-1-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	Thymus		Thyroid		Adrenal		Prostate		Seminal vesicle	
	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	g	%
302	238	52.77	23.4	5.19	77	17.07	1031	228.60	2.23	0.49
303	434	87.68	23.4	4.73	50	10.10	613	123.84	2.03	0.41
305	375	77.32	19.0	3.92	58	11.96	828	170.72	2.03	0.42
306	481	97.96	24.6	5.01	62	12.63	1302	265.17	2.53	0.52
308	356	69.94	30.9	6.07	65	12.77	771	151.47	2.04	0.40
N	5	5	5	5	5	5	5	5	5	5
MEAN	376.8	77.134	24.26	4.984	62.4	12.906	909.0	187.960	2.172	0.448
S.D.	92.0	17.251	4.28	0.778	9.9	2.559	265.8	57.778	0.218	0.054
S.E.	41.1	7.715	1.92	0.348	4.4	1.145	118.9	25.839	0.097	0.024

t' 2.8917 * 2.6921 *

INDIVIDUAL DATA 11-1-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of administration

Animal No.	Thymus		Thyroid		Adrenal		Prostate		Seminal vesicle	
	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	g	%
401	343	78.31	17.5	4.00	58	13.24	826	188.58	2.21	0.50
402	306	66.67	22.0	4.79	63	13.73	586	127.67	2.37	0.52
403	291	66.59	22.4	5.13	66	15.10	680	155.61	2.50	0.57
404	335	73.30	26.7	5.84	44	9.63	530	115.97	2.06	0.45
407	372	86.11	22.2	5.14	58	13.43	615	142.36	1.72	0.40
N	5	5	5	5	5	5	5	5	5	5
MEAN	329.4	74.196	22.16	4.980	57.8	13.026	647.4	146.038	2.172	0.488
S.D.	31.8	8.279	3.26	0.668	8.4	2.033	113.5	28.091	0.302	0.065
S.E.	14.2	3.702	1.46	0.299	3.8	0.909	50.8	12.563	0.135	0.029
t'			1.7503	2.6818 *						

INDIVIDUAL DATA 11-1-9

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

Animal No.	Body weight g	Testis		Epididymis	
		g	%	g	%
101	466	3.84	0.82	1.54	0.33
103	480	3.99	0.83	1.51	0.31
104	474	3.30	0.70	1.21	0.26
106	446	3.21	0.72	1.47	0.33
109	561	3.49	0.62	1.39	0.25
110	526	3.55	0.67	1.38	0.26
112	550	3.40	0.62	1.40	0.25
N	7	7	7	7	7
MEAN	500.4	3.540	0.711	1.414	0.284
S.D.	44.8	0.283	0.086	0.109	0.037
S.E.	16.9	0.107	0.033	0.041	0.014
M/C	6.2357	0.7787	2.7424	0.2120	3.2020
F	2.3344 †	1.2568	1.1292	2.5909 †	3.2424 *

INDIVIDUAL DATA 11-1-10

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	Body weight		Testis		Epididymis	
		g	g	%	g	%
201	468	3.17	0.68	1.21	0.26	
202	502	2.90	0.58	1.13	0.23	
203	479	3.07	0.64	1.16	0.24	
204	488	3.55	0.73	1.48	0.30	
205	495	3.23	0.65	1.25	0.25	
206	483	3.64	0.75	1.23	0.25	
207	486	3.14	0.65	1.31	0.27	
208	490	3.63	0.74	1.37	0.28	
209	497	2.88	0.58	1.14	0.23	
210	517	3.34	0.65	1.25	0.24	
211	536	3.42	0.64	1.41	0.26	
212	472	3.36	0.71	1.25	0.26	
N		12	12	12	12	12
MEAN	492.8	3.278	0.667	1.266	0.256	
S.D.	19.0	0.259	0.057	0.109	0.021	
S.E.	5.5	0.075	0.016	0.031	0.006	
t'	0.5325		2.6997 *	2.1180		

INDIVIDUAL DATA 11-1-11

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	Body weight g	Testis		Epididymis	
		g	%	g	%
301	487	3.35	0.69	1.31	0.27
302	451	3.01	0.67	1.27	0.28
303	495	3.18	0.64	1.22	0.25
304	500	2.84	0.57	1.11	0.22
305	485	3.40	0.70	1.28	0.26
306	491	3.22	0.66	1.38	0.28
307	488	3.81	0.78	1.54	0.32
308	509	3.83	0.75	1.31	0.26
309	473	3.49	0.74	1.35	0.29
310	496	3.48	0.70	1.43	0.29
311	513	3.31	0.65	1.21	0.24
312	564	3.64	0.65	1.50	0.27
N	12	12	12	12	12
MEAN	496.0	3.380	0.683	1.326	0.269
S.D.	26.9	0.298	0.057	0.123	0.026
S.E.	7.8	0.086	0.016	0.036	0.008
t'	0.3071		1.6085	1.1255	

INDIVIDUAL DATA 11-1-12

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of administration

Animal No.	Body weight		Testis		Epididymis	
		g		g		%
401	438	3.95	0.90	1.53	0.35	
402	459	2.96	0.64	1.21	0.26	
403	437	3.24	0.74	1.19	0.27	
404	457	3.05	0.67	1.40	0.31	
407	432	3.10	0.72	1.39	0.32	
408	488	3.27	0.67	1.36	0.28	
412	529	3.63	0.69	1.41	0.27	
N	7	7	7	7	7	
MEAN	462.9	3.314	0.719	1.356	0.294	
S.D.	34.9	0.354	0.087	0.119	0.033	
S.E.	13.2	0.134	0.033	0.045	0.013	
t'	2.3184		0.9477	0.6623		

INDIVIDUAL DATA 11-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 6 of lactation

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain	
		g	g	%	g	%	g	%	g	%	g	%
152	292	9.99	3.42		2.05	0.70	0.85	0.29	1.07	0.37	2.21	0.76
153	295	9.59	3.25		1.93	0.65	0.74	0.25	1.00	0.34	2.01	0.68
155	290	10.34	3.57		1.92	0.66	0.58	0.20	1.05	0.36	2.11	0.73
156	301	10.03	3.33		2.01	0.67	0.70	0.23	1.04	0.35	2.05	0.68
162	324	10.88	3.36		2.23	0.69	0.74	0.23	1.18	0.36	2.02	0.62
N		5	5	5	5	5	5	5	5	5	5	5
MEAN		300.4	10.166	3.386	2.028	0.674	0.722	0.240	1.068	0.356	2.080	0.694
S.D.		13.8	0.480	0.120	0.125	0.021	0.097	0.033	0.068	0.011	0.082	0.054
S.E.		6.2	0.215	0.054	0.056	0.009	0.043	0.015	0.030	0.005	0.037	0.024
M/C		0.4251	1.0980	1.5133	2.8206	2.1863	2.0617	1.7705	1.9462	0.8969	0.5044	6.4960
F		0.0771	1.2985	2.3273	0.9125	1.6271	0.9779	1.2263	2.0687	2.6584 †	0.1499	0.0385

INDIVIDUAL DATA 11-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 6 of lactation

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain		
		g		g	%		g	%		g	%		%
251	290	9.88	3.41	1.89	0.65	0.57	0.20	0.99	0.34	1.97	0.68		
253	284	9.10	3.20	1.63	0.57	0.68	0.24	0.99	0.35	1.98	0.70		
255	298	10.62	3.56	2.05	0.69	0.75	0.25	1.04	0.35	2.07	0.69		
256	301	9.19	3.05	2.02	0.67	0.77	0.26	0.96	0.32	2.12	0.70		
262	313	9.73	3.11	1.97	0.63	0.97	0.31	1.08	0.35	2.09	0.67		
N		5	5	5	5	5	5	5	5	5	5	5	5
MEAN		297.2	9.704	3.266	1.912	0.642	0.748	0.252	1.012	0.342	2.046	0.688	
S.D.		11.1	0.612	0.214	0.169	0.046	0.147	0.040	0.048	0.013	0.067	0.013	
S.E.		5.0	0.274	0.096	0.076	0.021	0.066	0.018	0.021	0.006	0.030	0.006	

1.5556

t'

INDIVIDUAL DATA 11-2-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 6 of lactation

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain		
		g		g	%		g	%		g	%		%
351	282	8.80	3.12	2.01	0.71	0.63	0.22	1.00	0.35	2.01	0.71		
352	292	9.95	3.41	1.93	0.66	0.77	0.26	0.99	0.34	2.06	0.71		
353	294	9.77	3.32	2.12	0.72	0.86	0.29	0.95	0.32	1.92	0.65		
355	302	10.59	3.51	1.93	0.64	0.69	0.23	1.03	0.34	2.10	0.70		
361	322	10.87	3.38	2.03	0.63	0.94	0.29	1.03	0.32	2.18	0.68		
N		5	5	5	5	5	5	5	5	5	5	5	5
MEAN		298.4	9.996	3.348	2.004	0.672	0.778	0.258	1.000	0.334	2.054	0.690	
S.D.		15.0	0.806	0.145	0.079	0.041	0.125	0.033	0.033	0.013	0.097	0.025	
S.E.		6.7	0.361	0.065	0.035	0.018	0.056	0.015	0.015	0.006	0.044	0.011	

2.4444

INDIVIDUAL DATA 11-2-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 6 of lactation

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain		
		g		g	%		g	%		g	%		%
451	295	9.58	3.25	1.97	0.67	0.61	0.21	0.97	0.33	2.09	0.71		
453	290	11.00	3.79	2.03	0.70	0.59	0.20	0.96	0.33	2.16	0.74		
455	282	9.84	3.49	1.82	0.65	0.66	0.23	0.96	0.34	1.92	0.68		
457	311	11.12	3.58	2.33	0.75	0.67	0.22	0.96	0.31	2.06	0.66		
462	306	11.10	3.63	2.14	0.70	0.77	0.25	1.10	0.36	2.09	0.68		
N		5	5	5	5	5	5	5	5	5	5	5	5
MEAN		296.8	10.528	3.548	2.058	0.694	0.660	0.222	0.990	0.334	2.064	0.694	
S.D.		11.8	0.754	0.199	0.191	0.038	0.070	0.019	0.062	0.018	0.088	0.031	
S.E.		5.3	0.337	0.089	0.085	0.017	0.031	0.009	0.028	0.008	0.040	0.014	

2.4444

INDIVIDUAL DATA 11-2-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : Day 6 of lactation

Animal No.	Pituitary gland		Thymus		Thyroid		Adrenal		Ovary	
	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %
152	17.1	5.86	150	51.37	20.6	7.05	67	22.95	111	38.01
153	13.8	4.68	358	121.36	17.6	5.97	77	26.10	131	44.41
155	15.5	5.34	197	67.93	15.8	5.45	80	27.59	107	36.90
156	15.3	5.08	314	104.32	18.6	6.18	76	25.25	144	47.84
162	19.4	5.99	234	72.22	23.8	7.35	82	25.31	123	37.96
N	5	5	5	5	5	5	5	5	5	5
MEAN	16.22	5.390	250.6	83.440	19.28	6.400	76.4	25.440	123.2	41.024
S.D.	2.13	0.544	84.9	28.587	3.06	0.784	5.8	1.682	15.0	4.832
S.E.	0.95	0.243	38.0	12.784	1.37	0.351	2.6	0.752	6.7	2.161
M/C	12.7933 **	2.8980	1.8708	1.8098	1.0509	2.5588	0.6127	0.9677	2.4431	0.6739
F		1.4073	1.8093	1.7648	1.3256	1.3659	0.4628	0.5657	1.1916	1.2303
H		4.6613								

INDIVIDUAL DATA 11-2-6

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : Day 6 of lactation

Animal No.	Pituitary gland		Thymus		Thyroid		Adrenal		Ovary	
	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %
251	14.3	4.93	217	74.83	16.5	5.69	81	27.93	101	34.83
253	16.7	5.88	268	94.37	14.6	5.14	81	28.52	84	29.58
255	18.1	6.07	135	45.30	12.8	4.30	76	25.50	127	42.62
256	18.0	5.98	325	107.97	14.5	4.82	72	23.92	111	36.88
262	15.5	4.95	216	69.01	20.5	6.55	88	28.12	125	39.94
N	5	5	5	5	5	5	5	5	5	5
MEAN	16.52	5.562	232.2	78.296	15.78	5.300	79.6	26.798	109.6	36.770
S.D.	1.63	0.572	70.4	24.114	2.95	0.862	6.0	1.998	17.8	4.994
S.E.	0.73	0.256	31.5	10.784	1.32	0.385	2.7	0.893	8.0	2.233

INDIVIDUAL DATA 11-2-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : Day 6 of lactation

Animal No.	Pituitary gland		Thymus		Thyroid		Adrenal		Ovary	
	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %
351	17.0	6.03	269	95.39	16.0	5.67	68	24.11	113	40.07
352	16.9	5.79	306	104.79	22.1	7.57	79	27.05	125	42.81
353	17.6	5.99	381	129.59	14.7	5.00	78	26.53	105	35.71
355	17.4	5.76	277	91.72	18.5	6.13	90	29.80	116	38.41
361	16.6	5.16	379	117.70	13.5	4.19	82	25.47	109	33.85
N	5	5	5	5	5	5	5	5	5	5
MEAN	17.10	5.746	322.4	107.838	16.96	5.712	79.4	26.592	113.6	38.170
S.D.	0.40	0.348	54.4	15.767	3.42	1.270	7.9	2.117	7.6	3.532
S.E.	0.18	0.156	24.3	7.051	1.53	0.568	3.5	0.947	3.4	1.580

INDIVIDUAL DATA 11-2-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : Day 6 of lactation

Animal No.	Pituitary gland		Thymus		Thyroid		Adrenal		Ovary	
	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %
451	15.8	5.36	243	82.37	18.0	6.10	84	28.47	114	38.64
453	15.4	5.31	237	81.72	19.4	6.69	82	28.28	148	51.03
455	15.1	5.35	344	121.99	14.7	5.21	66	23.40	111	39.36
457	14.8	4.76	280	90.03	19.7	6.33	76	24.44	130	41.80
462	15.8	5.16	280	91.50	18.4	6.01	69	22.55	116	37.91
N	5	5	5	5	5	5	5	5	5	5
MEAN	15.38	5.188	276.8	93.522	18.04	6.068	75.4	25.428	123.8	41.748
S.D.	0.44	0.252	42.6	16.510	1.99	0.547	7.9	2.773	15.4	5.391
S.E.	0.20	0.113	19.1	7.384	0.89	0.245	3.5	1.240	6.9	2.411

INDIVIDUAL DATA 11-3-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain		Pituitary gland		Thymus	
		g	g	%	g	%	g	%	g	%	g	%	mg	10 ⁻³ %	mg	10 ⁻³ %
102	492	11.45	2.33		2.85	0.58	0.62	0.13	1.50	0.30	2.09	0.42	12.6	2.56	272	55.28
105	498	11.99	2.41		2.96	0.59	0.73	0.15	1.44	0.29	2.24	0.45	12.5	2.51	221	44.38
107	487	11.09	2.28		3.16	0.65	0.78	0.16	1.57	0.32	2.08	0.43	11.1	2.28	284	58.32
108	544	14.62	2.69		3.21	0.59	0.79	0.15	1.55	0.28	2.13	0.39	11.4	2.10	235	43.20
111	553	12.73	2.30		3.11	0.56	1.04	0.19	1.57	0.28	2.22	0.40	17.0	3.07	331	59.86
N	5	5	5		5	5	5	5	5	5	5	5	5	5	5	5
MEAN	514.8	12.376	2.402		3.058	0.594	0.792	0.156	1.526	0.294	2.152	0.418	12.92	2.504	268.6	52.208
S.D.	31.2	1.398	0.168		0.149	0.034	0.154	0.022	0.056	0.017	0.074	0.024	2.37	0.366	43.4	7.870
S.E.	13.9	0.625	0.075		0.067	0.015	0.069	0.010	0.025	0.007	0.033	0.011	1.06	0.164	19.4	3.520
M/C	1.4056	0.0047	1.4608		1.7801	1.3198	1.3785	0.9176	2.8705	0.6127	1.7477	0.5874	0.0000	0.1088	0.2422	0.4985
F	1.7492	6.6954 *	11.8440 **		0.6637	2.2588	1.6480	1.5077	1.8824	0.0860	0.1899	2.9512	0.1494	0.0006	0.0158	0.2316

INDIVIDUAL DATA 11-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain		Pituitary gland		Thymus	
		g	g	%	g	%	g	%	g	%	g	%	mg	10 ⁻³ %	mg	10 ⁻³ %
405	510	12.23	2.40		3.23	0.63	0.64	0.13	1.65	0.32	2.14	0.42	16.5	3.24	225	44.12
406	469	15.01	3.20		3.21	0.68	0.59	0.13	1.34	0.29	2.18	0.46	11.9	2.54	220	46.91
409	487	15.43	3.17		2.84	0.58	0.73	0.15	1.51	0.31	2.17	0.45	10.6	2.18	350	71.87
410	503	15.12	3.01		3.66	0.73	0.80	0.16	1.31	0.26	2.22	0.44	11.3	2.25	311	61.83
411	501	15.33	3.06		2.98	0.59	0.70	0.14	1.34	0.27	2.13	0.43	11.4	2.28	257	51.30
N	5	5	5		5	5	5	5	5	5	5	5	5	5	5	5
MEAN	494.0	14.624	2.968		3.184	0.642	0.692	0.142	1.430	0.290	2.168	0.440	12.34	2.498	272.6	55.206
S.D.	16.3	1.349	0.327		0.312	0.063	0.081	0.013	0.146	0.025	0.036	0.016	2.37	0.437	56.4	11.495
S.E.	7.3	0.603	0.146		0.140	0.028	0.036	0.006	0.065	0.011	0.016	0.007	1.06	0.195	25.2	5.141

t' 2.5875 * 3.4415 **

INDIVIDUAL DATA 11-3-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	Thyroid		Adrenal		Testis		Epididymis		Prostate		Seminal vesicle	
	mg	10 ⁻³ %	mg	10 ⁻³ %	g	%	g	%	mg	10 ⁻³ %	g	%
102	18.9	3.84	70	14.23	2.74	0.56	1.10	0.22	876	178.05	2.56	0.52
105	28.4	5.70	73	14.66	3.61	0.72	1.47	0.30	806	161.85	2.01	0.40
107	18.7	3.84	64	13.14	3.49	0.72	1.35	0.28	528	108.42	1.96	0.40
108	22.7	4.17	73	13.42	2.97	0.55	1.33	0.24	771	141.73	2.05	0.38
111	20.8	3.76	72	13.02	3.65	0.66	1.58	0.29	887	160.40	2.64	0.48
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	21.90	4.262	70.4	13.694	3.292	0.642	1.366	0.266	773.6	150.090	2.244	0.436
S.D.	3.98	0.819	3.8	0.717	0.411	0.083	0.180	0.034	145.5	26.611	0.328	0.061
S.E.	1.78	0.366	1.7	0.321	0.184	0.037	0.080	0.015	65.1	11.901	0.147	0.027
M/C	0.0032	0.0019	0.7620	0.7488	0.7434	0.5080	0.1134	0.0015	0.1867	0.2014	0.2354	0.1875
F	0.0383	0.2853	22.6510 **	17.4763 **	0.6269	1.7751	0.7396	1.6266	0.0195	0.2181	0.0467	0.5633

INDIVIDUAL DATA 11-3-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	Thyroid		Adrenal		Testis		Epididymis		Prostate		Seminal vesicle	
	mg	10 ⁻³ %	mg	10 ⁻³ %	g	%	g	%	mg	10 ⁻³ %	g	%
405	15.9	3.12	60	11.76	3.46	0.68	1.48	0.29	938	183.92	2.61	0.51
406	21.6	4.61	49	10.45	3.29	0.70	1.43	0.30	638	136.03	2.30	0.49
409	22.9	4.70	62	12.73	3.91	0.80	1.70	0.35	676	138.81	2.01	0.41
410	25.1	4.99	56	11.13	3.36	0.67	1.33	0.26	657	130.62	2.44	0.49
411	26.5	5.29	49	9.78	3.30	0.66	1.34	0.27	1032	205.99	2.06	0.41
N	5	5	5	5	5	5	5	5	5	5	5	5
MEAN	22.40	4.542	55.2	11.170	3.464	0.702	1.456	0.294	788.2	159.074	2.284	0.462
S.D.	4.10	0.838	6.1	1.144	0.258	0.057	0.150	0.035	183.2	33.800	0.253	0.048
S.E.	1.83	0.375	2.7	0.512	0.116	0.025	0.067	0.016	81.9	15.116	0.113	0.022

t' 4.7593 ** 4.1805 **

INDIVIDUAL DATA 11-4-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain	
		g	g	%	g	%	g	%	g	%	g	%
163	276	7.18	2.60		1.67	0.61	0.52	0.19	0.96	0.35	1.85	0.67
164	270	6.38	2.36		1.79	0.66	0.43	0.16	1.04	0.39	1.98	0.73
165	294	7.03	2.39		1.73	0.59	0.49	0.17	0.85	0.29	2.14	0.73
166	273	6.83	2.50		1.74	0.64	0.39	0.14	1.01	0.37	2.20	0.81
167	284	7.35	2.59		1.86	0.65	0.57	0.20	0.92	0.32	2.09	0.74
N		5	5	5	5	5	5	5	5	5	5	5
MEAN		279.4	6.954	2.488	1.758	0.630	0.480	0.172	0.956	0.344	2.052	0.736
S.D.		9.7	0.374	0.111	0.071	0.029	0.071	0.024	0.075	0.040	0.139	0.050
S.E.		4.3	0.167	0.050	0.032	0.013	0.032	0.011	0.034	0.018	0.062	0.022
M/C		0.7378	3.0156	2.7633	2.2664	1.9883	0.0809	0.0153	2.0815	1.8530	1.7497	0.0648
F		0.1747	1.2171	2.6605	4.3623 †	4.6992 †	0.9944	1.3279	3.8343 †	1.4922	0.4078	0.0314

INDIVIDUAL DATA 11-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	Body weight		Liver		Kidney		Spleen		Heart		Brain	
		g	g	%	g	%	g	%	g	%	g	%
463	273	7.36	2.70		1.88	0.69	0.48	0.18	0.86	0.32	2.06	0.75
464	275	6.21	2.26		2.09	0.76	0.59	0.21	0.94	0.34	2.05	0.75
465	275	8.21	2.99		2.10	0.76	0.58	0.21	0.89	0.32	1.93	0.70
466	257	6.91	2.69		1.72	0.67	0.51	0.20	0.86	0.33	2.06	0.80
467	300	8.72	2.91		1.84	0.61	0.45	0.15	0.87	0.29	1.94	0.65
N		5	5	5	5	5	5	5	5	5	5	5
MEAN		276.0	7.482	2.710	1.926	0.698	0.522	0.190	0.884	0.320	2.008	0.730
S.D.		15.4	1.003	0.283	0.165	0.064	0.061	0.025	0.034	0.019	0.067	0.057
S.E.		6.9	0.448	0.127	0.074	0.029	0.027	0.011	0.015	0.008	0.030	0.025
t'				2.0886	2.1678			1.9581				

INDIVIDUAL DATA 11-4-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Absolute and relative organ weights, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.	Pituitary gland		Thymus		Thyroid		Adrenal		Ovary	
	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %
163	15.7	5.69	198	71.74	16.3	5.91	71	25.72	98	35.51
164	13.0	4.81	335	124.07	21.1	7.81	62	22.96	127	47.04
165	16.8	5.71	281	95.58	17.6	5.99	68	23.13	126	42.86
166	16.1	5.90	264	96.70	17.3	6.34	65	23.81	78	28.57
167	16.7	5.88	305	107.39	19.8	6.97	69	24.30	85	29.93
N	5	5	5	5	5	5	5	5	5	5
MEAN	15.66	5.598	276.6	99.096	18.42	6.604	67.0	23.984	102.8	36.782
S.D.	1.55	0.451	51.4	19.098	1.97	0.793	3.5	1.109	22.8	8.034
S.E.	0.69	0.202	23.0	8.541	0.88	0.355	1.6	0.496	10.2	3.593
M/C	1.4495	2.2975	0.5025	0.1251	1.0748	0.2014	4.5684 *	5.7293 *	0.1778	0.2541
F	0.0566	0.0111	1.9208	1.9625	0.7482	0.8160			0.1013	0.0507
H						0.3951	1.3200			

INDIVIDUAL DATA 11-4-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Absolute and relative organ weights, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	Pituitary gland		Thymus		Thyroid		Adrenal		Ovary	
	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %	mg	10 ⁻³ %
463	11.0	4.03	281	102.93	13.2	4.84	67	24.54	101	37.00
464	19.2	6.98	286	104.00	17.6	6.40	94	34.18	145	52.73
465	14.8	5.38	363	132.00	19.4	7.05	79	28.73	90	32.73
466	14.8	5.76	339	131.91	13.4	5.21	65	25.29	74	28.79
467	16.7	5.57	307	102.33	20.8	6.93	66	22.00	78	26.00
N	5	5	5	5	5	5	5	5	5	5
MEAN	15.30	5.544	315.2	114.634	16.88	6.086	74.2	26.948	97.6	35.450
S.D.	3.01	1.052	35.1	15.823	3.46	1.007	12.4	4.704	28.5	10.513
S.E.	1.34	0.471	15.7	7.076	1.55	0.451	5.6	2.104	12.8	4.701

INDIVIDUAL DATA 12-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Male GROUP: 0 mg/kg PERIOD: End of administration

Animal No.	101	103	104	106	109	110	112
Organ: Findings							
Lung: Aggregation, macrophage, alveolar Metaplasia, osscous, alvcoli	-	+	-	-	+	*	*
Trachea:	-	-	+	-	-	*	*
Esophagus:	N	N	N	N	N	*	*
Forestomach:	N	N	N	N	N	*	*
Stomach, limiting ridge:	N	N	N	N	N	*	*
Glandular stomach:	N	N	N	N	N	*	*
Duodenum:	N	N	N	N	N	*	*
Jejunum:	N	N	N	N	N	*	*
Ileum (including Peyer's patch):	N	N	N	N	N	*	*
Cecum:	N	N	N	N	N	*	*
Colon:	N	N	N	N	N	*	*
Rectum:	N	N	N	N	N	*	*
Pancreas:	N	N	N	N	N	*	*
Liver: Fatty change, periportal Microgranuloma Necrosis, massive Deposit, hemosiderin/ hematoidin Fibrosis Mineralization	-	-	-	-	+	-	-
Heart:	N	N	N	N	N	*	*
Right kidney: Regeneration, tubular epithelium Dilatation, renal pelvis	-	+	-	-	-	*	*
Left kidney: Regeneration, tubular epithelium Dilatation, renal pelvis	-	+	-	-	-	*	*
Urinary bladder:	N	N	N	N	N	*	*
Right testis:	N	N	N	N	N	*	*
Left testis:	N	N	N	N	N	*	*
Right epididymis:	N	N	N	N	N	*	*
Left epididymis:	N	N	N	N	N	*	*
Prostate: Cellular infiltration, inflammatory cell	-	-	-	+	-	-	+
Right seminal vesicle:	N	N	N	N	N	*	*
Left seminal vesicle:	N	N	N	N	N	*	*
Right coagulating gland:	N	N	N	N	N	*	*
Left coagulating gland:	N	N	N	N	N	*	*

N: No abnormal findings, -: normal, +: slight change, ++: moderate change, +++: severe change.

*: Not examined.

(to be continued)

INDIVIDUAL DATA 12-1-1 (continued)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Male GROUP: 0 mg/kg PERIOD: End of administration

Animal No.	101	103	104	106	109	110	112
Organ: Findings							
Cerebrum:	N	N	N	N	N	*	*
Cerebellum:	N	N	N	N	N	*	*
Pons:	N	N	N	N	N	*	*
Spinal cord:	N	N	N	N	N	*	*
Sciatic nerve:	N	N	N	N	N	*	*
Spleen:	N	N	N	N	N	*	*
Thymus:	N	N	N	N	N	*	*
Bone marrow of right femur:	N	N	N	N	N	*	*
Right submandibular lymph node:	N	N	N	N	N	*	*
Left submandibular lymph node:	N	N	N	N	N	*	*
Mesenteric lymph node:	N	N	N	N	N	*	*
Pituitary gland:	N	N	N	N	N	*	*
Right thyroid:	N	N	N	N	N	*	*
Left thyroid:	N	N	N	N	N	*	*
Right parathyroid:	N	N	N	N	N	*	*
Left parathyroid:	N	N	N	N	#	*	*
Right adrenal:	N	N	N	N	N	*	*
Left adrenal:	N	N	N	N	N	*	*
Right eyeball:	N	N	N	N	N	*	*
Left eyeball:	N	N	N	N	N	*	*
Right Harderian gland:	N	N	N	N	N	*	*
Left Harderian gland:	N	N	N	N	N	*	*
Right femur:	N	N	N	N	N	*	*

N: No abnormal findings.

*: Not examined.

#: Not examined because of missing.

INDIVIDUAL DATA 12-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Male GROUP: 4 mg/kg PERIOD: End of administration

Animal No.	201	202	203	204	205	206	207	208	209	210	211	212
Organ: Findings												
Liver: Microgranuloma	+	+	-	-	+	+	+	+	+	+	-	-
Prostate: Cellular infiltration, inflammatory cell	+	+	+	+	+	-	-	+	-	+	-	+

-: Normal, +: slight change.

INDIVIDUAL DATA 12-1-3

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Male GROUP: 20 mg/kg PERIOD: End of administration

Animal No.	301	302	303	304	305	306	307	308	309	310	311	312
Organ: Findings												
Liver: Hypertrophy, hepatocyte, centrilobular	-	+	-	+	+	-	+	-	-	-	+	-
Fatty change, centrilobular	-	-	-	+	-	-	+	-	-	-	-	-
Microgranuloma	-	+	+	-	+	+	+	+	-	+	+	-
Left epididymis: Granuloma, spermatic	*	*	*	*	*	*	*	*	+	*	*	*
Prostate: Cellular infiltration, inflammatory cell	-	-	+	-	-	+	-	-	-	-	-	-

-: Normal, +: slight change.

*: Not examined.

INDIVIDUAL DATA 12-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Male GROUP: 100 mg/kg PERIOD: End of administration

Animal No.	401	402	403	404	407	408	412
Organ: Findings							
Lung: Aggregation, macrophage, alveolar Mineralization, artery	+	++	-	-	-	*	*
Trachea:	+	-	-	+	-	*	*
Esophagus:	N	N	N	N	N	*	*
Forestomach:	N	N	N	N	N	*	*
Stomach, limiting ridge:	N	N	N	N	N	*	*
Glandular stomach:	N	N	N	N	N	*	*
Duodenum:	N	N	N	N	N	*	*
Jejunum:	N	N	N	N	N	*	*
Ileum (including Peyer's patch):	N	N	N	N	N	*	*
Cecum:	N	N	N	N	N	*	*
Colon:	N	N	N	N	N	*	*
Rectum:	N	N	N	N	N	*	*
Pancreas:	N	N	N	N	N	*	*
Liver: Hypertrophy, hepatocyte, centrilobular Fatty change, centrilobular Microgranuloma	++	++	++	++	++	++	++
Heart: Myocardial degeneration, focal	-	+	-	-	-	*	*
Right kidney:	N	N	N	N	N	*	*
Left kidney:	N	N	N	N	N	*	*
Urinary bladder:	N	N	N	N	N	*	*
Right testis:	N	N	N	N	N	*	*
Left testis:	N	N	N	N	N	*	*
Right epididymis:	N	N	N	N	N	*	*
Left epididymis:	N	N	N	N	N	*	*
Prostate: Cellular infiltration, inflammatory cell	-	+	+	+	+	+	+
Right seminal vesicle:	N	N	N	N	N	*	*
Left seminal vesicle:	N	N	N	N	N	*	*
Right coagulating gland:	N	N	N	N	N	*	*
Left coagulating gland:	N	N	N	N	N	*	*

N: No abnormal findings, -: normal, +: slight change, ++: moderate change.

*: Not examined.

(to be continued)

INDIVIDUAL DATA 12-1-4 (continued)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Male GROUP: 100 mg/kg PERIOD: End of administration

Animal No.	401	402	403	404	407	408	412
Organ: Findings							
Cerebrum:	N	N	N	N	N	*	*
Cerebellum:	N	N	N	N	N	*	*
Pons:	N	N	N	N	N	*	*
Spinal cord:	N	N	N	N	N	*	*
Sciatic nerve:	N	N	N	N	N	*	*
Spleen:	N	N	N	N	N	*	*
Thymus:	N	N	N	N	N	*	*
Bone marrow of right femur:	N	N	N	N	N	*	*
Right submandibular lymph node:	N	N	N	N	N	*	*
Left submandibular lymph node:	N	N	N	N	N	*	*
Mesenteric lymph node:	N	N	N	N	N	*	*
Pituitary gland:	N	N	N	N	N	*	*
Right thyroid:	N	N	N	N	N	*	*
Left thyroid:	N	N	N	N	N	*	*
Right parathyroid:	N	N	N	N	N	*	*
Left parathyroid:	N	#	N	N	N	*	*
Right adrenal:	N	N	N	N	N	*	*
Left adrenal:	N	N	N	N	N	*	*
Right eyeball:	N	N	N	N	N	*	*
Left eyeball:	N	N	N	N	N	*	*
Right Harderian gland:	N	N	N	N	N	*	*
Left Harderian gland:	N	N	N	N	N	*	*
Right femur:	N	N	N	N	N	*	*

N: No abnormal findings.

*: Not examined.

#: Not examined because of missing.

INDIVIDUAL DATA 12-2-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings	ANIMAL: Rat, Crl: CD (SD)	SEX: Female	GROUP: 0 mg/kg	PERIOD: End of administration									
Animal No.		151	152	153	154	155	156	157	158	159	160	161	162
Organ: Findings													
Lung: Aggregation, macrophage, alveolar Mineralization, artery		*	-	-	*	+	-	*	*	*	*	*	-
Trachea:		*	-	-	*	-	-	*	*	*	*	*	+
Esophagus:		*	N	N	*	N	N	*	*	*	*	*	N
Forestomach:		*	N	N	*	N	N	*	*	*	*	*	N
Stomach, limiting ridge:		*	N	N	*	N	N	*	*	*	*	*	N
Glandular stomach:		*	N	N	*	N	N	*	*	*	*	*	N
Duodenum:		*	N	N	*	N	N	*	*	*	*	*	N
Jejunum:		*	N	N	*	N	N	*	*	*	*	*	N
Ileum (including Peyer's patch):		*	N	N	*	N	N	*	*	*	*	*	N
Cecum:		*	N	N	*	N	N	*	*	*	*	*	N
Colon:		*	N	N	*	N	N	*	*	*	*	*	N
Rectum:		*	N	N	*	N	N	*	*	*	*	*	N
Pancreas:		*	N	N	*	N	N	*	*	*	*	*	N
Liver: Microgranuloma		+	+	+	-	+	+	+	-	-	-	-	+
Heart:		*	N	N	*	N	N	*	*	*	*	*	N
Right kidney:		*	N	N	*	N	N	*	*	*	*	*	N
Left kidney: Cellular infiltration, inflammatory cell, renal pelvic mucosa		*	-	-	*	-	+	*	*	*	*	*	-
Urinary bladder:		*	N	N	*	N	N	*	*	*	*	*	N
Right ovary:		*	N	N	*	N	N	*	*	*	*	*	N
Left ovary:		*	N	N	*	N	N	*	*	*	*	*	N
Right uterine horn:		*	N	N	*	N	N	*	*	*	*	*	N
Left uterine horn:		*	N	N	*	N	N	*	*	*	*	*	N
Uterine cervix:		*	N	N	*	N	N	*	*	*	*	*	N

N: No abnormal findings, -: normal, +: slight change.

(to be continued)

*: Not examined.

INDIVIDUAL DATA 12-2-1 (continued)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings	ANIMAL: Rat, Crl: CD (SD)	SEX: Female	GROUP: 0 mg/kg	PERIOD: End of administration									
Animal No.		151	152	153	154	155	156	157	158	159	160	161	162
Organ: Findings													
Cerebrum:		*	N	N	*	N	N	*	*	*	*	*	N
Cerebellum:		*	N	N	*	N	N	*	*	*	*	*	N
Pons:		*	N	N	*	N	N	*	*	*	*	*	N
Spinal cord:		*	N	N	*	N	N	*	*	*	*	*	N
Sciatic nerve:		*	N	N	*	N	N	*	*	*	*	*	N
Spleen:		*	N	N	*	N	N	*	*	*	*	*	N
Thymus:		*	N	N	*	N	N	*	*	*	*	*	N
Bone marrow of right femur:		*	N	N	*	N	N	*	*	*	*	*	N
Right submandibular lymph node:		*	N	N	*	N	N	*	*	*	*	*	N
Left submandibular lymph node:		*	N	N	*	N	N	*	*	*	*	*	N
Mesenteric lymph node:		*	N	N	*	N	N	*	*	*	*	*	N
Pituitary gland: Tubular hyperplasia, pars nervosa		*	-	-	*	+	-	*	*	*	*	*	-
Right thyroid:		*	N	N	*	N	N	*	*	*	*	*	N
Left thyroid:		*	N	N	*	N	N	*	*	*	*	*	N
Right parathyroid:		*	#	N	*	N	N	*	*	*	*	*	N
Left parathyroid:		*	N	N	*	N	N	*	*	*	*	*	N
Right adrenal:		*	N	N	*	N	N	*	*	*	*	*	N
Left adrenal:		*	N	N	*	N	N	*	*	*	*	*	N
Right eyeball:		*	N	N	*	N	N	*	*	*	*	*	N
Left eyeball:		*	N	N	*	N	N	*	*	*	*	*	N
Right Harderian gland:		*	N	N	*	N	N	*	*	*	*	*	N
Left Harderian gland:		*	N	N	*	N	N	*	*	*	*	*	N
Right femur:		*	N	N	*	N	N	*	*	*	*	*	N
Mammary gland:		*	N	N	*	N	N	*	*	*	*	*	N

N: No abnormal findings, -: normal, +: slight change.

*: Not examined.

#: Not examined because of missing.

INDIVIDUAL DATA 12-2-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Female GROUP: 4 mg/kg PERIOD: End of administration

Animal No.	251	252	253	254	255	256	257	258	259	260	261	262
Organ: Findings												
Stomach, limiting ridge: Cyst, squamous cell	*	*	*	*	*	*	*	*	*	+	*	*
Liver: Fatty change, periportal	-	-	-	+	-	-	-	-	-	-	-	-
Microgranuloma	+	+	-	+	+	-	+	-	+	+	-	-
Necrosis, focal	-	-	-	-	-	-	-	-	+	-	-	-

-: Normal, +: slight change.

*: Not examined.

INDIVIDUAL DATA 12-2-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Female GROUP: 20 mg/kg PERIOD: End of administration

Animal No.	351	352	353	354	355	356	357	358	359	360	361	362
Organ: Findings												
Liver: Fatty change, periportal Microgranuloma	-	-	+	-	-	-	-	-	-	-	-	-
	+	+	+	-	+	-	+	+	+	-	+	-

-: Normal, +: slight change.

INDIVIDUAL DATA 12-2-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings	ANIMAL: Rat, Crl: CD (SD)	SEX: Female	GROUP: 100 mg/kg	PERIOD: End of administration									
Animal No.		451	452	453	454	455	456	457	458	459	460	461	462
Organ: Findings													
Lung: Aggregation, macrophage, alveolar		+	*	-	*	-	*	-	*	*	*	*	*
Trachea:		N	*	N	*	N	*	N	*	*	*	*	*
Esophagus:		N	*	N	*	N	*	N	*	*	*	*	*
Forestomach:		N	*	N	*	N	*	N	*	*	*	*	*
Stomach, limiting ridge:		N	*	N	*	N	*	N	*	*	*	*	*
Glandular stomach:		N	*	N	*	N	*	N	*	*	*	*	*
Duodenum:		N	*	N	*	N	*	N	*	*	*	*	*
Jejunum:		N	*	N	*	N	*	N	*	*	*	*	*
Ileum (including Peyer's patch):		N	*	N	*	N	*	N	*	*	*	*	*
Cecum:		N	*	N	*	N	*	N	*	*	*	*	*
Colon:		N	*	N	*	N	*	N	*	*	*	*	*
Rectum:		N	*	N	*	N	*	N	*	*	*	*	*
Pancreas:		N	*	N	*	N	*	N	*	*	*	*	*
Liver: Hypertrophy, hepatocyte, centrilobular		+	-	+	+	+	+	+	+	+	-	-	-
Fatty change, periportal		-	-	-	-	-	-	-	-	-	-	-	+
Microgranuloma		-	-	+	+	+	+	-	-	-	+	+	+
Necrosis, focal		-	-	-	-	+	-	-	-	-	-	-	-
Heart:		N	*	N	*	N	*	N	*	*	*	*	*
Right kidney:		N	*	N	*	N	*	N	*	*	*	*	*
Left kidney: Mineralization, cortico-medullary junction		+	*	-	*	-	*	-	*	*	*	*	-
Mineralization, papilla		-	*	+	*	-	*	-	*	*	*	*	-
Urinary bladder:		N	*	N	*	N	*	N	*	*	*	*	*
Right ovary:		N	*	N	*	N	*	N	*	*	*	*	*
Left ovary:		N	*	N	*	N	*	N	*	*	*	*	*
Right uterine horn:		N	*	N	*	N	*	N	*	*	*	*	*
Left uterine horn:		N	*	N	*	N	*	N	*	*	*	*	N
Uterine cervix:		N	*	N	*	N	*	N	*	*	*	*	N

N: No abnormal findings, -: normal, +: slight change.

*: Not examined.

(to be continued)

INDIVIDUAL DATA 12-2-4 (continued)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Histopathological findings	ANIMAL: Rat, Crl: CD (SD)	SEX: Female	GROUP: 100 mg/kg	PERIOD: End of administration									
Animal No.		451	452	453	454	455	456	457	458	459	460	461	462
Organ: Findings													
Cerebrum:		N	*	N	*	N	*	N	*	*	*	*	*
Cerebellum:		N	*	N	*	N	*	N	*	*	*	*	N
Pons:		N	*	N	*	N	*	N	*	*	*	*	N
Spinal cord:		N	*	N	*	N	*	N	*	*	*	*	N
Sciatic nerve:		N	*	N	*	N	*	N	*	*	*	*	N
Spleen:		N	*	N	*	N	*	N	*	*	*	*	N
Thymus:		N	*	N	*	N	*	N	*	*	*	*	N
Bone marrow of right femur:		N	*	N	*	N	*	N	*	*	*	*	N
Right submandibular lymph node:		N	*	N	*	N	*	N	*	*	*	*	N
Left submandibular lymph node:		N	*	N	*	N	*	N	*	*	*	*	N
Mesenteric lymph node:		N	*	N	*	N	*	N	*	*	*	*	N
Pituitary gland:		N	*	N	*	N	*	N	*	*	*	*	N
Right thyroid:		N	*	N	*	N	*	N	*	*	*	*	N
Left thyroid:		N	*	N	*	N	*	N	*	*	*	*	N
Right parathyroid:		N	*	N	*	N	*	N	*	*	*	*	N
Left parathyroid:		N	*	N	*	N	*	N	*	*	*	*	N
Right adrenal:		N	*	N	*	N	*	N	*	*	*	*	N
Left adrenal:		N	*	N	*	N	*	N	*	*	*	*	N
Right eyeball:		N	*	N	*	N	*	N	*	*	*	*	N
Left eyeball:		N	*	N	*	N	*	N	*	*	*	*	N
Right Harderian gland:		N	*	N	*	N	*	N	*	*	*	*	N
Left Harderian gland:		N	*	N	*	N	*	N	*	*	*	*	N
Right femur:		N	*	N	*	N	*	N	*	*	*	*	N
Mammary gland: Fibroadenoma		-	*	-	*	-	*	-	*	*	<+>	*	-

N: No abnormal findings, -: normal, <+>: presence in "presence or not" basis.

*: Not examined.

INDIVIDUAL DATA 12-3-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Male GROUP: 0 mg/kg PERIOD: End of recovery

Animal No.	102	105	107	108	111
Organ: Findings					
Lung: Aggregation, macrophage, alveolar Mineralization, artery	-	-	-	+	-
Trachea:	-	-	-	-	+
Esophagus:	N	N	N	N	N
Forestomach:	N	N	N	N	N
Stomach, limiting ridge:	N	N	N	N	N
Glandular stomach:	N	N	N	N	N
Duodenum:	N	N	N	N	N
Jejunum:	N	N	N	N	N
Ileum (including Peyer's patch): Diverticulum	-	-	-	-	+
Cecum:	N	N	N	N	N
Colon:	N	N	N	N	N
Rectum:	N	N	N	N	N
Pancreas:	N	N	N	N	N
Liver: Fatty change, periportal Microgranuloma	-	+	-	+	+
Heart:	N	N	N	N	N
Right kidney:	N	N	N	N	N
Left kidney:	N	N	N	N	N
Urinary bladder:	N	N	N	N	N
Right testis:	N	N	N	N	N
Left testis:	N	N	N	N	N
Right epididymis:	N	N	N	N	N
Left epididymis:	N	N	N	N	N
Prostate: Cellular infiltration, inflammatory cell	-	-	+	-	-
Right seminal vesicle:	N	N	N	N	N
Left seminal vesicle:	N	N	N	N	N
Right coagulating gland:	N	N	N	N	N
Left coagulating gland:	N	N	N	N	N

N: No abnormal findings, -: normal, +: slight change.

(to be continued)

INDIVIDUAL DATA 12-3-1 (continued)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Male GROUP: 0 mg/kg PERIOD: End of recovery

Animal No.	102	105	107	108	111
Organ: Findings					
Cerebrum:	N	N	N	N	N
Cerebellum:	N	N	N	N	N
Pons:	N	N	N	N	N
Spinal cord:	N	N	N	N	N
Sciatic nerve:	N	N	N	N	N
Spleen:	N	N	N	N	N
Thymus:	N	N	N	N	N
Bone marrow of right femur:	N	N	N	N	N
Right submandibular lymph node:	N	N	N	N	N
Left submandibular lymph node:	N	N	N	N	N
Mesenteric lymph node:	N	N	N	N	N
Pituitary gland:	N	N	N	N	N
Right thyroid:	N	N	N	N	N
Left thyroid:	N	N	N	N	N
Right parathyroid:	N	N	N	N	N
Left parathyroid:	N	N	N	N	N
Right adrenal:	N	N	N	N	N
Left adrenal:	N	N	N	N	N
Right eyeball:	N	N	N	N	N
Left eyeball:	N	N	N	N	N
Right Harderian gland:	N	N	N	N	N
Left Harderian gland:	N	N	N	N	N
Right femur:	N	N	N	N	N

N: No abnormal findings.

INDIVIDUAL DATA 12-3-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Male GROUP: 100 mg/kg PERIOD: End of recovery

Animal No.	405	406	409	410	411
Organ: Findings					
Lung: Aggregation, macrophage, alveolar	+	+	+	+	+
Trachea:	N	N	N	N	N
Esophagus:	N	N	N	N	N
Forestomach:	N	N	N	N	N
Stomach, limiting ridge:	N	N	N	N	N
Glandular stomach:	N	N	N	N	N
Duodenum:	N	N	N	N	N
Jejunum:	N	N	N	N	N
Ileum (including Peyer's patch):	N	N	N	N	N
Cecum:	N	N	N	N	N
Colon:	N	N	N	N	N
Rectum:	N	N	N	N	N
Pancreas:	N	N	N	N	N
Liver: Hypertrophy, hepatocyte, centrilobular	+	+	+	+	+
Fatty change, centrilobular	-	-	+	-	-
Microgranuloma	-	+	+	-	-
Heart:	N	N	N	N	N
Right kidney:	N	N	N	N	N
Left kidney:	N	N	N	N	N
Urinary bladder:	N	N	N	N	N
Right testis:	N	N	N	N	N
Left testis:	N	N	N	N	N
Right epididymis: Granuloma, spermatic	-	-	+	-	-
Left epididymis:	N	N	N	N	N
Prostate: Cellular infiltration, inflammatory cell	-	-	+	-	+
Right seminal vesicle:	N	N	N	N	N
Left seminal vesicle:	N	N	N	N	N
Right coagulating gland:	N	N	N	N	N
Left coagulating gland:	N	N	N	N	N

N: No abnormal findings, -: normal, +: slight change.

(to be continued)

INDIVIDUAL DATA 12-3-2 (continued)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Male GROUP: 100 mg/kg PERIOD: End of recovery

Animal No.	405	406	409	410	411
Organ: Findings					
Cerebrum:	N	N	N	N	N
Cerebellum:	N	N	N	N	N
Pons:	N	N	N	N	N
Spinal cord:	N	N	N	N	N
Sciatic nerve:	N	N	N	N	N
Spleen:	N	N	N	N	N
Thymus:	N	N	N	N	N
Bone marrow of right femur:	N	N	N	N	N
Right submandibular lymph node:	N	N	N	N	N
Left submandibular lymph node:	N	N	N	N	N
Mesenteric lymph node:	N	N	N	N	N
Pituitary gland: Cyst, pars distalis	-	-	-	+	-
Right thyroid:	N	N	N	N	N
Left thyroid:	N	N	N	N	N
Right parathyroid:	N	N	#	N	N
Left parathyroid:	N	N	N	N	N
Right adrenal:	N	N	N	N	N
Left adrenal:	N	N	N	N	N
Right eyeball:	N	N	N	N	N
Left eyeball:	N	N	N	N	N
Right Harderian gland:	N	N	N	N	N
Left Harderian gland:	N	N	N	N	N
Right femur:	N	N	N	N	N

N: No abnormal findings, -: normal, +: slight change.

#: Not examined because of missing.

INDIVIDUAL DATA 12-4-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Female GROUP: 0 mg/kg PERIOD: End of recovery

Animal No.	163	164	165	166	167
Organ: Findings					
Lung: Aggregation, macrophage, alveolar Metaplasia, osseous, alcocoli	-	-	-	+	-
Trachea:	-	-	-	+	-
Esophagus:	N	N	N	N	N
Forestomach:	N	N	N	N	N
Stomach, limiting ridge:	N	N	N	N	N
Glandular stomach:	N	N	N	N	N
Duodenum:	N	N	N	N	N
Jejunum:	N	N	N	N	N
Ileum (including Peyer's patch):	N	N	N	N	N
Cecum:	N	N	N	N	N
Colon:	N	N	N	N	N
Rectum:	N	N	N	N	N
Pancreas:	N	N	N	N	N
Liver: Microgranuloma	-	-	+	-	-
Heart:	N	N	N	N	N
Right kidney: Cellular infiltration, inflammatory cell, renal pelvic mucosa	-	+	-	-	-
Left kidney:	N	N	N	N	N
Urinary bladder:	N	N	N	N	N
Right ovary:	N	N	N	N	N
Left ovary:	N	N	N	N	N
Right uterine horn:	N	N	N	N	N
Left uterine horn:	N	N	N	N	N
Uterine cervix:	N	N	N	N	N

N: No abnormal findings, -: normal, +: slight change.

(to be continued)

INDIVIDUAL DATA 12-4-1 (continued)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Female GROUP: 0 mg/kg PERIOD: End of recovery

Animal No.	163	164	165	166	167
Organ: Findings					
Cerebrum:	N	N	N	N	N
Cerebellum:	N	N	N	N	N
Pons:	N	N	N	N	N
Spinal cord:	N	N	N	N	N
Sciatic nerve:	N	N	N	N	N
Spleen:	N	N	N	N	N
Thymus:	N	N	N	N	N
Bone marrow of right femur:	N	N	N	N	N
Right submandibular lymph node:	N	N	N	N	N
Left submandibular lymph node:	N	N	N	N	N
Mesenteric lymph node:	N	N	N	N	N
Pituitary gland:	N	N	N	N	N
Right thyroid:	N	N	N	N	N
Left thyroid:	N	N	N	N	N
Right parathyroid:	N	N	N	N	N
Left parathyroid:	N	N	N	N	N
Right adrenal:	N	N	N	N	N
Left adrenal:	N	N	N	N	N
Right eyeball:	N	N	N	N	N
Left eyeball:	N	N	N	N	N
Right Harderian gland:	N	N	N	N	N
Left Harderian gland:	N	N	N	N	N
Right femur:	N	N	N	N	N
Mammary gland:	N	N	N	N	N

N: No abnormal findings.

INDIVIDUAL DATA 12-4-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Female GROUP: 100 mg/kg PERIOD: End of recovery

Animal No.	463	464	465	466	467
Organ: Findings					
Lung: Mineralization, artery	-	+	-	-	-
Metaplasia, osseous, alveoli	+	-	-	-	-
Trachea:	N	N	N	N	N
Esophagus:	N	N	N	N	N
Forestomach:	N	N	N	N	N
Stomach, limiting ridge:	N	N	N	N	N
Glandular stomach:	N	N	N	N	N
Duodenum:	N	N	N	N	N
Jejunum:	N	N	N	N	N
Ileum (including Peyer's patch):	N	N	N	N	N
Cecum:	N	N	N	N	N
Colon:	N	N	N	N	N
Rectum:	N	N	N	N	N
Pancreas:	N	N	N	N	N
Liver: Hypertrophy, hepatocyte, centrilobular	-	-	-	+	-
Microgranuloma	+	+	-	+	+
Heart:	N	N	N	N	N
Right kidney: Mineralization, cortex	-	+	-	-	-
Left kidney:	N	N	N	N	N
Urinary bladder:	N	N	N	N	N
Right ovary:	N	N	N	N	N
Left ovary:	N	N	N	N	N
Right uterine horn:	N	N	N	N	N
Left uterine horn:	N	N	N	N	N
Uterine cervix:	N	N	N	N	N

N: No abnormal findings, -: normal, +: slight change.

(to be continued)

INDIVIDUAL DATA 12-4-2 (continued)

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Histopathological findings ANIMAL: Rat, Crl: CD (SD) SEX: Female GROUP: 100 mg/kg PERIOD: End of recovery

Animal No.	463	464	465	466	467
Organ: Findings					
Cerebrum:	N	N	N	N	N
Cerebellum:	N	N	N	N	N
Pons:	N	N	N	N	N
Spinal cord:	N	N	N	N	N
Sciatic nerve:	N	N	N	N	N
Spleen:	N	N	N	N	N
Thymus:	N	N	N	N	N
Bone marrow of right femur:	N	N	N	N	N
Right submandibular lymph node:	N	N	N	N	N
Left submandibular lymph node:	N	N	N	N	N
Mesenteric lymph node:	N	N	N	N	N
Pituitary gland:	N	N	N	N	N
Right thyroid:	N	N	N	N	N
Left thyroid:	N	N	N	N	N
Right parathyroid:	N	N	N	N	N
Left parathyroid:	N	N	#	N	#
Right adrenal:	N	N	N	N	N
Left adrenal:	N	N	N	N	N
Right eyeball:	N	N	N	N	N
Left eyeball:	N	N	N	N	N
Right Harderian gland:	N	N	N	N	N
Left Harderian gland:	N	N	N	N	N
Right femur:	N	N	N	N	N
Mammary gland:	N	N	N	N	N

N: No abnormal findings.

#: Not examined because of missing.

INDIVIDUAL DATA 12-5-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

Animal No.	Animal										
	A	B	C	D	E	F	G	H	I	J	K
101	—	—	—	—	—	—	+	—	—	—	—
103	+	—	—	—	—	—	—	3+	2+	2+	2+
104	—	—	+	—	—	—	+	—	—	—	—
106	—	—	—	—	—	—	+	—	—	—	—
109	+	—	—	—	—	—	+	—	—	—	—
110	#	#	#	—	—	—	—	—	—	—	—
112	#	#	#	—	—	—	+	—	—	—	—

N	5	5	5	7	7	7	7	7	7	7	7
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H	0.0571	28.5520 **	4.4286	4.4286	4.4286	4.4286
X ²						

- A: Lung: Aggregation, macrophage, alveolar.
- B: Lung: Mineralization, artery.
- C: Lung: Metaplasia, osseous, alveoli.
- D: Liver: Hypertrophy, hepatocyte, centrilobular.
- E: Liver: Fatty change, centrilobular.
- F: Liver: Fatty change, periportal.
- G: Liver: Microgranuloma.
- H: Liver: Necrosis, massive.
- I: Liver: Deposit, hemosiderin/ hematoidin.
- J: Liver: Fibrosis.
- K: Liver: Mineralization.

INDIVIDUAL DATA 12-5-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	A	B	C	D	E	F	G	H	I	J	K
201	#	#	#	—	—	—	+	—	—	—	—
202	#	#	#	—	—	—	+	—	—	—	—
203	#	#	#	—	—	—	—	—	—	—	—
204	#	#	#	—	—	—	—	—	—	—	—
205	#	#	#	—	—	—	+	—	—	—	—
206	#	#	#	—	—	—	+	—	—	—	—
207	#	#	#	—	—	—	+	—	—	—	—
208	#	#	#	—	—	—	+	—	—	—	—
209	#	#	#	—	—	—	+	—	—	—	—
210	#	#	#	—	—	—	+	—	—	—	—
211	#	#	#	—	—	—	—	—	—	—	—
212	#	#	#	—	—	—	—	—	—	—	—

N	0	0	0	12	12	12	12	12	12	12	12
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U	42.0000					
p		1.0000	0.3684	0.6215		

- A: Lung: Aggregation, macrophage, alveolar.
- B: Lung: Mineralization, artery.
- C: Lung: Metaplasia, osseous, alveoli.
- D: Liver: Hypertrophy, hepatocyte, centrilobular.
- E: Liver: Fatty change, centrilobular.
- F: Liver: Fatty change, periportal.
- G: Liver: Microgranuloma.
- H: Liver: Necrosis, massive.
- I: Liver: Deposit, hemosiderin/ hematoidin.
- J: Liver: Fibrosis.
- K: Liver: Mineralization.

INDIVIDUAL DATA 12-5-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	A	B	C	D	E	F	G	H	I	J	K
301	#	#	#	—	—	—	—	—	—	—	—
302	#	#	#	+	—	—	+	—	—	—	—
303	#	#	#	—	—	—	+	—	—	—	—
304	#	#	#	+	+	—	—	—	—	—	—
305	#	#	#	+	—	—	+	—	—	—	—
306	#	#	#	—	—	—	+	—	—	—	—
307	#	#	#	+	+	—	+	—	—	—	—
308	#	#	#	—	—	—	+	—	—	—	—
309	#	#	#	—	—	—	—	—	—	—	—
310	#	#	#	—	—	—	—	—	—	—	—
311	#	#	#	+	—	—	+	—	—	—	—
312	#	#	#	—	—	—	—	—	—	—	—

N	0	0	0	12	12	12	12	12	12	12	12
---	---	---	---	----	----	----	----	----	----	----	----

U	24.5000										
p		0.3860	0.3684	0.6215							

A: Lung: Aggregation, macrophage, alveolar.

B: Lung: Mineralization, artery.

C: Lung: Metaplasia, osseous, alveoli.

D: Liver: Hypertrophy, hepatocyte, centrilobular.

E: Liver: Fatty change, centrilobular.

F: Liver: Fatty change, periportal.

G: Liver: Microgranuloma.

H: Liver: Necrosis, massive.

I: Liver: Deposit, hemosiderin/ hematoidin.

J: Liver: Fibrosis.

K: Liver: Mineralization.

INDIVIDUAL DATA 12-5-4

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of administration

U 0.0000 **
p 0.2222 0.5000 0.0003 ** 0.5000 0.2960

- A: Lung: Aggregation, macrophage, alveolar.
- B: Lung: Mineralization, artery.
- C: Lung: Metaplasia, osscous, alveoli.
- D: Liver: Hypertrophy, hepatocyte, centrilobular.
- E: Liver: Fatty change, centrilobular.
- F: Liver: Fatty change, periportal.
- G: Liver: Microgranuloma.
- H: Liver: Necrosis, massive.
- I: Liver: Deposit, hemosiderin/ hematoidin.
- J: Liver: Fibrosis.
- K: Liver: Mineralization.

INDIVIDUAL DATA 12-5-5

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of administration

No.	L	M	N	O	P
101	—	—	—	—	—
103	—	+	—	—	—
104	—	—	—	—	—
106	—	—	+	—	+
109	—	—	—	—	—
110	#	#	#	#	—
112	#	#	#	#	+

N	5	5	5	5	7
---	---	---	---	---	---

 χ^2

- L: Heart: Myocardial degeneration, focal.
 M: Kidney: Regeneration, tubular epithelium.
 N: Kidney: Dilatation, renal pelvis.
 O: Epididymis: Granuloma, spermatic.
 P: Prostate: Cellular infiltration, inflammatory cell.

INDIVIDUAL DATA 12-5-6

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Statistical analysis in histopathological findings ANIMAL : Rat, CrI:CD(SD) SEX : Male GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	L	M	N	O	P
201	#	#	#	#	+
202	#	#	#	#	+
203	#	#	#	#	+
204	#	#	#	#	+
205	#	#	#	#	+
206	#	#	#	#	-
207	#	#	#	#	-
208	#	#	#	#	+
209	#	#	#	#	-
210	#	#	#	#	+
211	#	#	#	#	-
212	#	#	#	#	+

- p**
- L: Heart: Myocardial degeneration, focal.
- M: Kidney: Regeneration, tubular epithelium.
- N: Kidney: Dilatation, renal pelvis.
- O: Epididymis: Granuloma, spermatic.
- P: Prostate: Cellular infiltration, inflammatory cell,

0.1299

- 584 -

INDIVIDUAL DATA 12-5-7

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 20 mg/kg PERIOD : End of administration

Animal					
No.	L	M	N	O	P
301	#	#	#	#	-
302	#	#	#	#	-
303	#	#	#	#	+
304	#	#	#	#	-
305	#	#	#	#	-
306	#	#	#	#	+
307	#	#	#	#	-
308	#	#	#	#	-
309	#	#	#	+	-
310	#	#	#	#	-
311	#	#	#	#	-
312	#	#	#	#	-

N	0	0	0	1	12
---	---	---	---	---	----

p	0.1667	0.4750
---	--------	--------

L: Heart: Myocardial degeneration, focal.

M: Kidney: Regeneration, tubular epithelium.

N: Kidney: Dilatation, renal pelvis.

O: Epididymis: Granuloma, spermatic.

P: Prostate: Cellular infiltration, inflammatory cell.

INDIVIDUAL DATA 12-5-8

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of administration

No.	L	M	N	O	P
401	—	—	—	—	—
402	+	—	—	—	+
403	—	—	—	—	+
404	—	—	—	—	+
407	—	—	—	—	+
408	#	#	#	#	+
412	#	#	#	#	+
N	5	5	5	5	7

p 0.5000 0.5000 0.5000 1.0000 0.0513

L: Heart: Myocardial degeneration, focal.

M: Kidney: Regeneration, tubular epithelium.

N: Kidney: Dilatation, renal pelvis.

O: Epididymis: Granuloma, spermatic.

P: Prostate: Cellular infiltration, inflammatory cell.

INDIVIDUAL DATA 12-6-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : End of administration

No.	Animal											
	A	B	C	D	E	F	G	H	I	J	K	L
151	#	#	#	—	—	+	—	#	#	#	#	#
152	—	—	—	—	—	+	—	—	—	—	—	—
153	—	—	—	—	—	+	—	—	—	—	—	—
154	#	#	#	—	—	—	—	#	#	#	#	#
155	+	—	—	—	—	+	—	—	—	—	+	—
156	—	—	—	—	—	+	—	+	—	—	—	—
157	#	#	#	—	—	+	—	#	#	#	#	#
158	#	#	#	—	—	—	—	#	#	#	#	#
159	#	#	#	—	—	—	—	#	#	#	#	#
160	#	#	#	—	—	—	—	#	#	#	#	#
161	#	#	#	—	—	—	—	#	#	#	#	#
162	—	+	—	—	—	+	—	—	—	—	—	—

N	5	5	5	12	12	12	12	5	5	5	5	5
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X²

- A: Lung: Aggregation, macrophage, alveolar.
- B: Lung: Mineralization, artery.
- C: Stomach, limiting ridge: Cyst, squamous cell.
- D: Liver: Hypertrophy, hepatocyte, centrilobular.
- E: Liver: Fatty change, periportal.
- F: Liver: Microgranuloma.
- G: Liver: Necrosis, focal.
- H: Kidney: Cellular infiltration, inflammatory cell, renal pelvic mucosa.
- I: Kidney: Mineralization, cortico-medullary junction.
- J: Kidney: Mineralization, papilla.
- K: Pituitary gland: Tubular hyperplasia, pars nervosa.
- L: Mammary gland: Fibroadenoma.

INDIVIDUAL DATA 12-6-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg PERIOD : End of administration

Animal No.	A	B	C	D	E	F	G	H	I	J	K	L
251	#	#	#	—	—	+	—	#	#	#	#	#
252	#	#	#	—	—	+	—	#	#	#	#	#
253	#	#	#	—	—	—	—	#	#	#	#	#
254	#	#	#	—	+	+	—	#	#	#	#	#
255	#	#	#	—	—	+	—	#	#	#	#	#
256	#	#	#	—	—	—	—	#	#	#	#	#
257	#	#	#	—	—	—	+	—	#	#	#	#
258	#	#	#	—	—	—	—	#	#	#	#	#
259	#	#	#	—	—	—	+	+	#	#	#	#
260	#	#	+	—	—	—	+	—	#	#	#	#
261	#	#	#	—	—	—	—	#	#	#	#	#
262	#	#	#	—	—	—	—	#	#	#	#	#

N	0	0	1	12	12	12	12	0	0	0	0	0
p			0.1667	1.0000	0.5000	0.6599	0.5000					

A: Lung: Aggregation, macrophage, alveolar.

B: Lung: Mineralization, artery.

C: Stomach, limiting ridge: Cyst, squamous cell.

D: Liver: Hypertrophy, hepatocyte, centrilobular.

E: Liver: Fatty change, periportal.

F: Liver: Microgranuloma.

G: Liver: Necrosis, focal.

H: Kidney: Cellular infiltration, inflammatory cell, renal pelvic mucosa.

I: Kidney: Mineralization, cortico-medullary junction.

J: Kidney: Mineralization, papilla.

K: Pituitary gland: Tubular hyperplasia, pars nervosa.

L: Mammary gland: Fibroadenoma.

INDIVIDUAL DATA 12-6-3

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg PERIOD : End of administration

Animal No.	A	B	C	D	E	F	G	H	I	J	K	L
351	#	#	#	—	—	+	—	#	#	#	#	#
352	#	#	#	—	—	+	—	#	#	#	#	#
353	#	#	#	—	+	+	—	#	#	#	#	#
354	#	#	#	—	—	—	—	#	#	#	#	#
355	#	#	#	—	—	+	—	#	#	#	#	#
356	#	#	#	—	—	—	—	#	#	#	#	#
357	#	#	#	—	—	—	+	—	#	#	#	#
358	#	#	#	—	—	—	+	—	#	#	#	#
359	#	#	#	—	—	—	+	—	#	#	#	#
360	#	#	#	—	—	—	—	—	#	#	#	#
361	#	#	#	—	—	—	+	—	#	#	#	#
362	#	#	#	—	—	—	—	—	#	#	#	#

N	0	0	0	12	12	12	12	0	0	0	0	0
---	---	---	---	----	----	----	----	---	---	---	---	---

p	1.0000	0.5000	0.5000	1.0000
---	--------	--------	--------	--------

A: Lung: Aggregation, macrophage, alveolar.

B: Lung: Mineralization, artery.

C: Stomach, limiting ridge: Cyst, squamous cell.

D: Liver: Hypertrophy, hepatocyte, centrilobular.

E: Liver: Fatty change, periportal.

F: Liver: Microgranuloma.

G: Liver: Necrosis, focal.

H: Kidney: Cellular infiltration, inflammatory cell, renal pelvic mucosa.

I: Kidney: Mineralization, cortico-medullary junction.

J: Kidney: Mineralization, papilla.

K: Pituitary gland: Tubular hyperplasia, pars nervosa.

L: Mammary gland: Fibroadenoma.

INDIVIDUAL DATA 12-6-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : End of administration

Animal No.	A	B	C	D	E	F	G	H	I	J	K	L
451	+	-	-	+	-	-	-	-	+	-	-	-
452	#	#	#	-	-	-	-	#	#	#	#	#
453	-	-	-	+	-	+	-	-	-	+	-	-
454	#	#	#	+	-	+	-	#	#	#	#	#
455	-	-	-	+	-	+	+	-	-	-	-	-
456	#	#	#	+	-	+	-	#	#	#	#	#
457	-	-	-	+	-	-	-	-	-	-	-	-
458	#	#	#	+	-	-	-	#	#	#	#	#
459	#	#	#	+	-	-	-	#	#	#	#	#
460	#	#	#	-	-	+	-	#	#	#	#	<+>
461	#	#	#	-	-	+	-	#	#	#	#	#
462	-	-	-	-	+	+	-	-	-	-	-	-
N	5	5	5	12	12	12	12	5	5	5	5	6
p	0.7778	0.5000	1.0000	0.0007 **	0.5000	0.6599	0.5000	0.5000	0.5000	0.5000	0.5000	0.5455

A: Lung: Aggregation, macrophage, alveolar.

B: Lung: Mineralization, artery.

C: Stomach, limiting ridge: Cyst, squamous cell.

D: Liver: Hypertrophy, hepatocyte, centrilobular.

E: Liver: Fatty change, periportal.

F: Liver: Microgranuloma.

G: Liver: Necrosis, focal.

H: Kidney: Cellular infiltration, inflammatory cell, renal pelvic mucosa.

I: Kidney: Mineralization, cortico-medullary junction.

J: Kidney: Mineralization, papilla.

K: Pituitary gland: Tubular hyperplasia, pars nervosa.

L: Mammary gland: Fibroadenoma.

INDIVIDUAL DATA 12-7-1

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 0 mg/kg PERIOD : End of recovery

Animal No.										
	A	B	C	D	E	F	G	H	I	J
102	—	—	—	—	—	—	—	—	—	—
105	—	—	—	—	—	—	+	—	—	—
107	—	—	—	—	—	—	—	—	+	—
108	+	—	—	—	—	+	+	—	—	—
111	—	+	+	—	—	—	+	—	—	—
N	5	5	5	5	5	5	5	5	5	5

 χ^2

- A: Lung: Aggregation, macrophage, alveolar.
- B: Lung: Mineralization, artery.
- C: Ileum: Diverticulum.
- D: Liver: Hypertrophy, hepatocyte, centrilobular.
- E: Liver: Fatty change, centrilobular.
- F: Liver: Fatty change, periportal.
- G: Liver: Microgranuloma.
- H: Epididymis: Granuloma, spermatic.
- I: Prostate: Cellular infiltration, inflammatory cell.
- J: Pituitary gland: Cyst, pars distalis.

INDIVIDUAL DATA 12-7-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)

Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Male GROUP : 100 mg/kg PERIOD : End of recovery

Animal No.	A	B	C	D	E	F	G	H	I	J
405	+	-	-	+	-	-	-	-	-	-
406	+	-	-	+	-	-	+	-	-	-
409	+	-	-	+	+	-	+	+	+	-
410	+	-	-	+	-	-	-	-	-	+
411	+	-	-	+	-	-	-	-	+	-
N	5	5	5	5	5	5	5	5	5	5

p	0.0238 *	0.5000	0.5000	0.0040 **	0.5000	0.5000	0.5000	0.5000	0.5000	0.5000
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A: Lung: Aggregation, macrophage, alveolar.

B: Lung: Mineralization, artery.

C: Ileum: Diverticulum.

D: Liver: Hypertrophy, hepatocyte, centrilobular.

E: Liver: Fatty change, centrilobular.

F: Liver: Fatty change, periportal.

G: Liver: Microgranuloma.

H: Epididymis: Granuloma, spermatic.

I: Prostate: Cellular infiltration, inflammatory cell.

J: Pituitary gland: Cyst, pars distalis.

INDIVIDUAL DATA 12-8-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg PERIOD : End of recovery

No.	Animal						
	A	B	C	D	E	F	G
163	—	—	—	—	—	—	—
164	—	—	—	—	—	+	—
165	—	—	—	—	+	—	—
166	+	—	+	—	—	—	—
167	—	—	—	—	—	—	—
N	5	5	5	5	5	5	5

 χ^2

A: Lung: Aggregation, macrophage, alveolar.

B: Lung: Mineralization, artery.

C: Lung: Metaplasia, osseous, alveoli.

D: Liver: Hypertrophy, hepatocyte, centrilobular.

E: Liver: Microgranuloma.

F: Kidney: Cellular infiltration, inflammatory cell, renal pelvic mucosa.

G: Kidney: Mineralization, cortex.

INDIVIDUAL DATA 12-8-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Statistical analysis in histopathological findings ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg PERIOD : End of recovery

No.	Animal						
	A	B	C	D	E	F	G
463	—	—	+	—	+	—	—
464	—	+	—	—	+	—	+
465	—	—	—	—	—	—	—
466	—	—	—	+	+	—	—
467	—	—	—	—	+	—	—
N	5	5	5	5	5	5	5

p 0.5000 0.5000 0.7778 0.5000 0.1032 0.5000 0.5000

A: Lung: Aggregation, macrophage, alveolar.

B: Lung: Mineralization, artery.

C: Lung: Metaplasia, osseous, alveoli.

D: Liver: Hypertrophy, hepatocyte, centrilobular.

E: Liver: Microgranuloma.

F: Kidney: Cellular infiltration, inflammatory cell, renal pelvic mucosa.

G: Kidney: Mineralization, cortex.

INDIVIDUAL DATA 13-1-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Estrous cycle, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

Animal No.	Administration day														Mating day				
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	1	2	3	4	5
151	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC
152	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#	#	#	#
153	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#	#
154	III	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#
155	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#	#
156	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#	#
157	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#
158	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#	#
159	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#
160	V	II	III	IV	V	II	III	III	IV	V	II	III	IV	V	II	IIIIC	#	#	#
161	II	III	IV	V	II	III	IV	V	II	III	IV	V	V	III	IIIIC	#	#	#	#
162	II	III	IV	V	V	II	III	IV	V	II	III	IV	V	V	II	IIIIC	#	#	#

II: Proestrus. III: Estrus. IV: Metestrus. V: Diestrus. C: Copulated.

#: Blank.

INDIVIDUAL DATA 13-1-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Estrous cycle, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg

Animal No.	Administration day														Mating day						
	Before mating day																				
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	1	2	3	4	5	6	7
251	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#	#	#	#	#
252	III	IV	V	V	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	V	IIIC
253	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#	#	#	#	#
254	II	III	IV	V	II	III	IV	V	V	II	III	III	IV	V	II	IIIC	#	#	#	#	#
255	II	III	IV	V	II	III	III	IV	V	II	III	III	IV	V	II	IIIC	#	#	#	#	#
256	III	IV	V	II	III	IV	V	II	III	III	IV	V	II	II	IIIC	#	#	#	#	#	#
257	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#	#	#
258	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#	#	#
259	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#	#	#
260	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#	#	#
261	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#	#	#
262	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#	#	#	#

II: Proestrus. III: Estrus. IV: Metestrus. V: Diestrus. C: Copulated.

#: Blank.

INDIVIDUAL DATA 13-1-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Estrous cycle, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg

Animal No.	Administration day														Mating day		
	Before mating day																
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	1	2	3
351	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#
352	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#
353	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#
354	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC
355	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#	#
356	III	IV	V	V	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC
357	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC
358	III	IV	V	V	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC
359	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC
360	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC	#
361	II	III	IV	V	II	III	IV	V	II	III	III	IV	V	II	IIIC	#	#
362	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIC

II: Proestrus. III: Estrus. IV: Metestrus. V: Diestrus. C: Copulated.

#: Blank.

INDIVIDUAL DATA 13-1-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Estrous cycle, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Administration day														Mating day			
	Before mating day							Mating day							1	2	3	4
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	1	2	3	4
451	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#	#	#
452	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC
453	IV	V	V	III	IV	V	V	III	IV	V	II	III	IV	V	II	IIIIC	#	#
454	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC
455	III	IV	V	V	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#
456	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC
457	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#	#	#
458	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC
459	III	IV	V	V	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#
460	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#
461	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#
462	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	IIIIC	#	#

II: Proestrus. III: Estrus. IV: Metestrus. V: Diestrus. C: Copulated.

#: Blank.

INDIVIDUAL DATA 13-2-1

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Estrous cycle, Recovery group ANIMAL : Rat, CrI:CD(SD) SEX : Female GROUP : 0 mg/kg

Animal No.	Administration day																											
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23	24	25	26	27	28
163	V	II	III	IV	V	V	II	III	IV	V	II	III	IV	V	V	II	III	IV	V	V	III	IV	V	V	III	IV	V	V
164	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV
165	II	III	IV	V	V	II	III	IV	V	II	III	IV	V	V	II	III	IV	V	V	V	III	IV	V	V	II	III	IV	V
166	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III
167	II	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV

Animal No.	Administration day												Estrous cycle			
	29	30	31	32	33	34	35	36	37	38	39	40	41	42	Normality	Length (days)
163	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	Normal	4.2
164	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	Normal	4.0
165	V	II	III	IV	V	V	V	III	IV	V	V	II	III	IV	Normal	4.9
166	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	Normal	4.0
167	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	Normal	4.0

II: Proestrus. III: Estrus. IV: Metestrus. V: Diestrus.

Normal estrous	N	5
cycle (%)	MEAN	4.22
5/5 (100)	S.D.	0.39

Estrous cycle: Normal; Female rats cycling normally (defined as having a mean cycle length between 4 and 6 days).

Normal estrous cycle = (number of female with normal estrous cycle / number of females examined) x 100.

(to be continued)

INDIVIDUAL DATA 13-2-1 (continued)

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Estrous cycle, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

Animal No.	Recovery day													Estrous cycle			
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	A	Normality	Length (days)
163	V	II	III	IV	V	V	III	IV	V	II	III	IV	V	II	III	Normal	4.0
164	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	Normal	4.0
165	V	V	V	III	IV	V	V	V	III	IV	V	V	V	III	III	Normal	5.0
166	II	III	IV	V	II	III	IV	V	V	III	IV	V	II	III	IV	Normal	4.0
167	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	Normal	4.0

II: Proestrus. III: Estrus. IV: Metestrus. V: Diestrus.

A: Autopsy day.

Normal estrous cycle (%)	N	5
5/5 (100)	MEAN	4.20
	S.D.	0.45

Estrous cycle: Normal; Female rats cycling normally (defined as having a mean cycle length between 4 and 6 days).

Normal estrous cycle = (number of female with normal estrous cycle / number of females examined) x 100.

INDIVIDUAL DATA 13-2-2

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Estrous cycle, Recovery group ANIMAL : Rat, CrI:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Administration day																											
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23	24	25	26	27	28
463	V	II	III	IV	V	V	III	IV	V	II	III	IV	V	V	III	IV												
464	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II
465	V	II	III	III	IV	V	II	II	III	IV	V	II																
466	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	V	III	IV	V
467	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	V

Animal No.	Administration day												Estrous cycle			
	29	30	31	32	33	34	35	36	37	38	39	40	41	42	Normality	Length (days)
463	V	V	III	IV	V	II	III	IV	V	V	III	IV	V	V	Normal	4.0
464	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	Normal	4.0
465	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	Normal	4.1
466	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	Normal	4.0
467	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	Normal	4.0

II: Proestrus. III: Estrus. IV: Metestrus. V: Diestrus.

Normal estrous cycle (%)	N	5
	MEAN	4.02
5/5 (100)	S.D.	0.04

Estrous cycle: Normal; Female rats cycling normally (defined as having a mean cycle length between 4 and 6 days).

Normal estrous cycle = (number of female with normal estrous cycle / number of females examined) x 100. (to be continued)

INDIVIDUAL DATA 13-2-2 (continued)

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Estrous cycle, Recovery group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Recovery day														Estrous cycle		Length (days)
	1	2	3	4	5	6	7	8	9	10	11	12	13	14	A	Normality	
463	III	IV	V	V	III	IV	V	V	III	IV	V	V	III	IV	V	Normal	4.0
464	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	V	II	Normal	4.0
465	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	Normal	4.0
466	IV	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	Normal	4.0
467	V	II	III	IV	V	II	III	IV	V	II	III	IV	V	II	III	Normal	4.0

II: Proestrus. III: Estrus. IV: Metestrus. V: Diestrus.

A: Autopsy day.

Normal estrous cycle (%)	N	5
5/5 (100)	MEAN	4.00
	S.D.	0.00

Estrous cycle: Normal; Female rats cycling normally (defined as having a mean cycle length between 4 and 6 days).

Normal estrous cycle = (number of female with normal estrous cycle / number of females examined) x 100.

INDIVIDUAL DATA 14-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Reproduction performance, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

Female No.	Estrous cycle		Paired male No.	Copulation		Pregnancy	Parturition	Gestation length (days)	Nursing
	Normality	Length (days)		Mating week	1				
151	Normal	4.0	101	+	+	+	+	22	+
152	Normal	4.0	102	+	+	+	+	23	+
153	Normal	4.0	103	+	+	+	+	22	+
154	Normal	4.0	104	+	+	+	+	22	+
155	Normal	4.0	105	+	+	+	+	22	+
156	Normal	4.0	106	+	+	+	+	22	+
157	Normal	4.0	107	+	+	+	+	22	+
158	Normal	4.0	108	+	+	+	+	23	+
159	Normal	4.0	109	+	+	+	+	22	+
160	Normal	4.3	110	+	+	+	+	23	+
161	Normal	4.3	111	+	-	#	#	#	#
162	Normal	4.7	112	+	+	+	+	22	+

N	12	11
MEAN	4.11	22.3
S.D.	0.22	0.5
Normal estrous cycle (%)	Copulation index (%) Male Female	Fertility index (%) Gestation index (%) Nursing index (%)
12/12 (100)	12/12 (100) 12/12 (100)	11/12 (91.7) 11/11 (100) 11/11 (100)

Estrous cycle: Normal; Female rats cycling normally (defined as having a mean cycle length between 4 and 6 days).

Copulation and pregnancy : -, Negative; +, Positive.

Parturition and nursing : +, Normal.

Normal estrous cycle = (number of females with normal estrous cycle / number of females examined) x 100.

Copulation index = (number of animals with successful copulation / number of animals mated) x 100.

Fertility index = (number of pregnant females / number of pairs with successful copulation) x 100.

Gestation index = (number of females with live pups / number of pregnant females) x 100.

Nursing index = (number of females nursing live pups on lactation day 4 / number of females with live pups delivery) x 100.

#: Blank.

INDIVIDUAL DATA 14-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Reproduction performance, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg

Female No.	Estrous cycle		Paired male No.	Copulation		Pregnancy	Parturition	Gestation length (days)	Nursing
	Normality	Length (days)		Mating week	1				
251	Normal	4.0	201	+	+	+	+	22	+
252	Normal	4.0	202	+	+	+	+	23	-
253	Normal	4.0	203	+	+	+	+	22	+
254	Normal	4.7	204	+	+	+	+	23	-
255	Normal	4.7	205	+	+	+	+	22	+
256	Normal	4.7	206	+	+	+	+	22	+
257	Normal	4.0	207	+	+	+	+	23	+
258	Normal	4.0	208	+	+	+	+	22	+
259	Normal	4.0	209	+	+	+	+	22	+
260	Normal	4.0	210	+	+	+	+	22	+
261	Normal	4.0	211	+	+	+	+	23	+
262	Normal	4.0	212	+	+	+	+	22	+

N	12	12		
MEAN	4.18	22.3		
S.D.	0.32	0.5		
Normal estrous cycle (%)	Copulation index (%) Male 12/12 (100)	Fertility index (%) Female 12/12 (100)	Gestation index (%) 12/12 (100)	Nursing index (%) 10/12 (83.3)

Estrous cycle: Normal; Female rats cycling normally (defined as having a mean cycle length between 4 and 6 days).

Copulation and pregnancy : +, Positive.

Parturition and nursing : -, Abnormal; +, Normal.

Normal estrous cycle = (number of females with normal estrous cycle / number of females examined) x 100.

Copulation index = (number of animals with successful copulation / number of animals mated) x 100.

Fertility index = (number of pregnant females / number of pairs with successful copulation) x 100.

Gestation index = (number of females with live pups / number of pregnant females) x 100.

Nursing index = (number of females nursing live pups on lactation day 4 / number of females with live pups delivery) x 100.

INDIVIDUAL DATA 14-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Reproduction performance, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg

Female No.	Estrous cycle		Paired male No.	Copulation		Pregnancy	Parturition	Gestation length (days)	Nursing
	Normality	Length (days)		Mating week	1				
351	Normal	4.0	301	+	+	+	+	22	+
352	Normal	4.0	302	+	+	+	+	22	+
353	Normal	4.0	303	+	+	+	+	22	+
354	Normal	4.0	304	+	+	+	+	22	+
355	Normal	4.0	305	+	+	+	+	22	+
356	Normal	4.0	306	+	+	+	+	23	+
357	Normal	4.0	307	+	+	+	+	23	+
358	Normal	4.0	308	+	+	+	+	23	+
359	Normal	4.0	309	+	+	+	+	22	+
360	Normal	4.0	310	+	+	+	+	22	+
361	Normal	4.3	311	+	+	+	+	22	+
362	Normal	4.0	312	+	+	+	+	22	+
N		12						12	
MEAN		4.03						22.3	
S.D.		0.09						0.5	
Normal estrous cycle (%)				Copulation index (%)	Fertility index (%)	Gestation index (%)		Nursing index (%)	
				Male	Female				
12/12 (100)				12/12 (100)	12/12 (100)	12/12 (100)	12/12 (100)	12/12 (100)	

Estrous cycle: Normal; Female rats cycling normally (defined as having a mean cycle length between 4 and 6 days).

Copulation and pregnancy : +, Positive.

Parturition and nursing : +, Normal.

Normal estrous cycle = (number of females with normal estrous cycle / number of females examined) x 100.

Copulation index = (number of animals with successful copulation / number of animals mated) x 100.

Fertility index = (number of pregnant females / number of pairs with successful copulation) x 100.

Gestation index = (number of females with live pups / number of pregnant females) x 100.

Nursing index = (number of females nursing live pups on lactation day 4 / number of females with live pups delivery) x 100.

INDIVIDUAL DATA 14-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Reproduction performance, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Female No.	Estrous cycle		Paired male No.	Copulation		Pregnancy	Parturition	Gestation length (days)	Nursing
	Normality	Length (days)		Mating week	1				
451	Normal	4.0	401	+	+	+	+	22	+
452	Normal	4.0	402	+	+	+	+	22	+
453	Normal	4.0	403	+	+	+	+	22	+
454	Normal	4.0	404	+	+	+	+	23	+
455	Normal	4.0	405	+	+	+	+	22	+
456	Normal	4.0	406	+	+	+	+	23	+
457	Normal	4.0	407	+	+	+	+	22	+
458	Normal	4.0	408	+	+	+	+	22	+
459	Normal	4.0	409	+	+	+	+	22	+
460	Normal	4.0	410	+	+	+	+	22	+
461	Normal	4.0	411	+	+	+	+	22	+
462	Normal	4.0	412	+	+	+	+	22	+

N	12	12		
MEAN	4.00	22.2		
S.D.	0.00	0.4		
Normal estrous cycle (%)	Copulation index (%) Male 12/12 (100)	Fertility index (%) Female 12/12 (100)	Gestation index (%) 12/12 (100)	Nursing index (%) 12/12 (100)

Estrous cycle: Normal; Female rats cycling normally (defined as having a mean cycle length between 4 and 6 days).

Copulation and pregnancy : +, Positive.

Parturition and nursing : +, Normal.

Normal estrous cycle = (number of females with normal estrous cycle / number of females examined) x 100.

Copulation index = (number of animals with successful copulation / number of animals mated) x 100.

Fertility index = (number of pregnant females / number of pairs with successful copulation) x 100.

Gestation index = (number of females with live pups / number of pregnant females) x 100.

Nursing index = (number of females nursing live pups on lactation day 4 / number of females with live pups delivery) x 100.

INDIVIDUAL DATA 15-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Pregnancy and litter data, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 0 mg/kg

Animal No.	Number of corpora lutea	Number of implantation sites	Implantation index (%)	Delivery index (%)	Lactation day 0								Live birth index (%)	Lactation day 4				
					Number of pups delivered				Sex ratio					Number of live pups	Sex ratio		Viability index (%)	
					Total	Male	Female	Alive	Dead	All pups	Live pups	Live pups			Live pups	Live pups		
151	16	16	100.00	100.00	16	8	8	16	0	8/16	0.50	8/16	0.50	100.00	16	8/16	0.50	100.00
152	15	14	93.33	100.00	14	8	6	14	0	8/14	0.57	8/14	0.57	100.00	14	8/14	0.57	100.00
153	16	16	100.00	93.75	15	5	10	15	0	5/15	0.33	5/15	0.33	100.00	15	5/15	0.33	100.00
154	17	16	94.12	87.50	14	5	9	14	0	5/14	0.36	5/14	0.36	100.00	14	5/14	0.36	100.00
155	17	17	100.00	94.12	16	4	12	16	0	4/16	0.25	4/16	0.25	100.00	16	4/16	0.25	100.00
156	18	18	100.00	94.44	17	9	8	17	0	9/17	0.53	9/17	0.53	100.00	17	9/17	0.53	100.00
157	14	13	92.86	84.62	11	6	5	11	0	6/11	0.55	6/11	0.55	100.00	11	6/11	0.55	100.00
158	17	17	100.00	100.00	17	8	9	17	0	8/17	0.47	8/17	0.47	100.00	17	8/17	0.47	100.00
159	17	16	94.12	100.00	16	10	6	16	0	10/16	0.63	10/16	0.63	100.00	15	9/15	0.60	93.75
160	17	17	100.00	88.24	15	13	2	15	0	13/15	0.87	13/15	0.87	100.00	15	13/15	0.87	100.00
161 ^a	#	#	#	#	#	#	#	#	#	#/#	#/#	#/#	#/#	#	#	#/#	#	#
162	17	17	100.00	94.12	16	8	8	15	1	8/16	0.50	7/15	0.47	93.75	15	7/15	0.47	100.00
N	11	11	11	11	11	11	11	11	11	11	11	11	11	11	11	11	11	
MEAN	16.5	16.1	97.675	94.254	15.2	7.6	7.5	15.1	0.1	0.505	0.503	99.432	15.0	0.500	99.432			
S.D.	1.1	1.4	3.243	5.541	1.7	2.6	2.7	1.7	0.3	0.165	0.166	1.884	1.7	0.164	1.884			
S.E.	0.3	0.4	0.978	1.671	0.5	0.8	0.8	0.5	0.1	0.050	0.050	0.568	0.5	0.049	0.568			
M/C	3.9390	3.4459	1.8345	2.5155	1.1267	1.3625	0.5303	1.8435	∞ **	0.7828	0.6837	∞ **	21.6613 **	0.6316	92.5767 **			
F	1.3614	1.3471	0.8018	0.9995	2.0109	0.9807	1.1823	1.5047		0.9161	0.8663			1.1040				
H									1.0799				1.0756	1.9177			4.9394	

Implantation index = (number of implantation sites / number of corpora lutea) x 100.

Delivery index = (number of pups born / number of implantation sites) x 100.

Sex ratio on lactation day 0 = (number of male pups born / number of pups born) or (number of live male pups / number of live pups).

Sex ratio on lactation day 4 = number of live male pups / number of live pups.

Live birth index = (number of live pups on lactation day 0 / number of pups born) x 100.

Viability index on lactation day 4 = (number of live pups on lactation day 4 / number of live pups on lactation day 0) x 100.

a: Non-pregnancy.

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INDIVIDUAL DATA 15-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Pregnancy and litter data, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 4 mg/kg

Animal No.	Number of corpora lutea	Number of implantation sites	Implantation index (%)	Delivery index (%)	Lactation day 0								Live birth index (%)	Lactation day 4				
					Number of pups delivered				Sex ratio					Number of live pups	Sex ratio		Viability index (%)	
					Total	Male	Female	Alive	Dead	All pups	Live pups	Live pups			Live pups	Live pups		
251	16	15	93.75	100.00	15	5	10	15	0	5/15	0.33	5/15	0.33	100.00	15	5/15	0.33	100.00
252	18	18	100.00	100.00	18	6	12	15	3	6/18	0.33	5/15	0.33	83.33	0	#/#	#	0.00
253	18	17	94.44	94.12	16	10	6	16	0	10/16	0.63	10/16	0.63	100.00	16	10/16	0.63	100.00
254	16	16	100.00	100.00	16	11	5	16	0	11/16	0.69	11/16	0.69	100.00	0	#/#	#	0.00
255	17	17	100.00	94.12	16	5	11	16	0	5/16	0.31	5/16	0.31	100.00	16	5/16	0.31	100.00
256	17	16	94.12	100.00	16	3	13	16	0	3/16	0.19	3/16	0.19	100.00	16	3/16	0.19	100.00
257	18	16	88.89	93.75	15	7	8	15	0	7/15	0.47	7/15	0.47	100.00	15	7/15	0.47	100.00
258	18	18	100.00	100.00	18	7	11	18	0	7/18	0.39	7/18	0.39	100.00	17	7/17	0.41	94.44
259	17	17	100.00	88.24	15	7	8	15	0	7/15	0.47	7/15	0.47	100.00	15	7/15	0.47	100.00
260	14	14	100.00	85.71	12	7	5	12	0	7/12	0.58	7/12	0.58	100.00	12	7/12	0.58	100.00
261	18	18	100.00	100.00	18	4	14	18	0	4/18	0.22	4/18	0.22	100.00	18	4/18	0.22	100.00
262	17	17	100.00	100.00	17	6	11	17	0	6/17	0.35	6/17	0.35	100.00	17	6/17	0.35	100.00
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	10	12	
MEAN	17.0	16.6	97.600	96.328	16.0	6.5	9.5	15.8	0.3	0.413	0.413	98.611	13.1	0.396	82.870			
S.D.	1.2	1.2	3.800	5.125	1.7	2.3	3.1	1.6	0.9	0.158	0.158	4.812	6.3	0.144	38.741			
S.E.	0.3	0.4	1.097	1.480	0.5	0.7	0.9	0.5	0.3	0.046	0.046	1.389	1.8	0.046	11.184			

Implantation index = (number of implantation sites / number of corpora lutea) x 100.

Delivery index = (number of pups born / number of implantation sites) x 100.

Sex ratio on lactation day 0 = (number of male pups born / number of pups born) or (number of live male pups / number of live pups).

Sex ratio on lactation day 4 = number of live male pups / number of live pups.

Live birth index = (number of live pups on lactation day 0 / number of pups born) x 100.

Viability index on lactation day 4 = (number of live pups on lactation day 4 / number of live pups on lactation day 0) x 100.

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INDIVIDUAL DATA 15-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Pregnancy and litter data, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 20 mg/kg

Animal No.	Number of corpora lutea	Number of implantation sites	Implantation index (%)	Delivery index (%)	Lactation day 0								Live birth index (%)	Lactation day 4				
					Number of pups delivered				Sex ratio					Number of live pups	Sex ratio		Viability index (%)	
					Total	Male	Female	Alive	Dead	All pups	Live pups	Live pups			Live pups	Live pups		
351	12	11	91.67	81.82	9	4	5	9	0	4/9	0.44	4/9	0.44	100.00	9	4/9	0.44	100.00
352	16	16	100.00	100.00	16	7	9	16	0	7/16	0.44	7/16	0.44	100.00	16	7/16	0.44	100.00
353	15	14	93.33	92.86	13	5	8	13	0	5/13	0.38	5/13	0.38	100.00	13	5/13	0.38	100.00
354	17	16	94.12	100.00	16	4	12	16	0	4/16	0.25	4/16	0.25	100.00	16	4/16	0.25	100.00
355	16	16	100.00	100.00	16	8	8	16	0	8/16	0.50	8/16	0.50	100.00	16	8/16	0.50	100.00
356	15	15	100.00	80.00	12	5	7	12	0	5/12	0.42	5/12	0.42	100.00	12	5/12	0.42	100.00
357	17	15	88.24	86.67	13	9	4	13	0	9/13	0.69	9/13	0.69	100.00	13	9/13	0.69	100.00
358	15	14	93.33	100.00	14	9	5	14	0	9/14	0.64	9/14	0.64	100.00	13	9/13	0.69	92.86
359	14	14	100.00	100.00	14	6	8	14	0	6/14	0.43	6/14	0.43	100.00	14	6/14	0.43	100.00
360	16	15	93.75	93.33	14	5	9	14	0	5/14	0.36	5/14	0.36	100.00	14	5/14	0.36	100.00
361	17	17	100.00	94.12	16	3	13	16	0	3/16	0.19	3/16	0.19	100.00	16	3/16	0.19	100.00
362	20	20	100.00	85.00	17	7	10	17	0	7/17	0.41	7/17	0.41	100.00	17	7/17	0.41	100.00
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	
MEAN	15.8	15.3	96.203	92.817	14.2	6.0	8.2	14.2	0.0	0.429	0.429	100.000	14.1	0.433	99.405			
S.D.	1.9	2.1	4.235	7.652	2.2	2.0	2.7	2.2	0.0	0.140	0.140	0.000	2.3	0.147	2.061			
S.E.	0.6	0.6	1.223	2.209	0.6	0.6	0.8	0.6	0.0	0.040	0.040	0.000	0.7	0.042	0.595			

Implantation index = (number of implantation sites / number of corpora lutea) x 100.

Delivery index = (number of pups born / number of implantation sites) x 100.

Sex ratio on lactation day 0 = (number of male pups born / number of pups born) or (number of live male pups / number of live pups).

Sex ratio on lactation day 4 = number of live male pups / number of live pups.

Live birth index = (number of live pups on lactation day 0 / number of pups born) x 100.

Viability index on lactation day 4 = (number of live pups on lactation day 4 / number of live pups on lactation day 0) x 100.

INDIVIDUAL DATA 15-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Pregnancy and litter data, Main group ANIMAL : Rat, Crl:CD(SD) SEX : Female GROUP : 100 mg/kg

Animal No.	Number of corpora lutea	Number of implantation sites	Implantation index (%)	Delivery index (%)	Lactation day 0								Live birth index (%)	Lactation day 4				
					Number of pups delivered				Sex ratio					Number of live pups	Sex ratio		Viability index (%)	
					Total	Male	Female	Alive	Dead	All pups	Live pups	Live pups			Live pups	Live pups		
451	14	14	100.00	92.86	13	6	7	13	0	6/13	0.46	6/13	0.46	100.00	13	6/13	0.46	100.00
452	14	14	100.00	92.86	13	6	7	13	0	6/13	0.46	6/13	0.46	100.00	13	6/13	0.46	100.00
453	17	16	94.12	100.00	16	10	6	16	0	10/16	0.63	10/16	0.63	100.00	15	10/15	0.67	93.75
454	16	16	100.00	81.25	13	5	8	13	0	5/13	0.38	5/13	0.38	100.00	13	5/13	0.38	100.00
455	17	16	94.12	93.75	15	6	9	15	0	6/15	0.40	6/15	0.40	100.00	15	6/15	0.40	100.00
456	16	16	100.00	81.25	13	7	6	13	0	7/13	0.54	7/13	0.54	100.00	12	7/12	0.58	92.31
457	17	17	100.00	100.00	17	5	12	17	0	5/17	0.29	5/17	0.29	100.00	16	5/16	0.31	94.12
458	15	15	100.00	93.33	14	9	5	14	0	9/14	0.64	9/14	0.64	100.00	8	5/8	0.63	57.14
459	14	13	92.86	84.62	11	8	3	10	1	8/11	0.73	7/10	0.70	90.91	10	7/10	0.70	100.00
460	17	17	100.00	100.00	17	14	3	17	0	14/17	0.82	14/17	0.82	100.00	17	14/17	0.82	100.00
461	19	19	100.00	84.21	16	5	11	16	0	5/16	0.31	5/16	0.31	100.00	15	5/15	0.33	93.75
462	17	17	100.00	100.00	17	4	13	17	0	4/17	0.24	4/17	0.24	100.00	17	4/17	0.24	100.00
N	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	12	
MEAN	16.1	15.8	98.425	92.011	14.6	7.1	7.5	14.5	0.1	0.492	0.489	99.243	13.7	0.498	0.498	94.256		
S.D.	1.6	1.6	2.866	7.438	2.0	2.8	3.3	2.2	0.3	0.183	0.180	2.624	2.7	0.180	0.180	12.109		
S.E.	0.5	0.5	0.827	2.147	0.6	0.8	0.9	0.6	0.1	0.053	0.052	0.758	0.8	0.052	0.052	3.496		

Implantation index = (number of implantation sites / number of corpora lutea) x 100.

Delivery index = (number of pups born / number of implantation sites) x 100.

Sex ratio on lactation day 0 = (number of male pups born / number of pups born) or (number of live male pups / number of live pups).

Sex ratio on lactation day 4 = number of live male pups / number of live pups.

Live birth index = (number of live pups on lactation day 0 / number of pups born) x 100.

Viability index on lactation day 4 = (number of live pups on lactation day 4 / number of live pups on lactation day 0) x 100.

019 -

INDIVIDUAL DATA 16-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 General appearance, pups ANIMAL : Rat,Crl:CD(SD) SEX : Male, Female GROUP : 0 mg/kg

Dam No.	Day of lactation	Male			Female		
		Number of pups	Finding and number of pups with findings		Number of pups	Finding and number of pups with findings	
151	0-4	8	No abnormal findings, 8		8	No abnormal findings, 8	
152	0-4	8	No abnormal findings, 8		6	No abnormal findings, 6	
153	0-4	5	No abnormal findings, 5		10	No abnormal findings, 10	
154	0-4	5	No abnormal findings, 5		9	No abnormal findings, 9	
155	0-4	4	No abnormal findings, 4		12	No abnormal findings, 12	
156	0-4	9	No abnormal findings, 9		8	No abnormal findings, 8	
157	0-4	6	No abnormal findings, 6		5	No abnormal findings, 5	
158	0-4	8	No abnormal findings, 8		9	No abnormal findings, 9	
159	0	10	No abnormal findings, 10		6	No abnormal findings, 6	
	1	10	Missing, 1; No abnormal findings, 9		6	No abnormal findings, 6	
	2-4	9	No abnormal findings, 9		6	No abnormal findings, 6	
160	0-4	13	No abnormal findings, 13		2	No abnormal findings, 2	
161 ^a	#	#	#		#	#	
162	0	8	Death(milk-band not examined because of autolysis of abdominal cavity), 1; No abnormal findings, 7		8	No abnormal findings, 8	
	1-4	7	No abnormal findings, 7		8	No abnormal findings, 8	

a: Non-pregnancy.

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INDIVIDUAL DATA 16-2

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 General appearance, pups ANIMAL : Rat,Crl:CD(SD) SEX : Male, Female GROUP : 4 mg/kg

Dam No.	Day of lactation	Male			Female		
		Number of pups	Finding and number of pups with findings		Number of pups	Finding and number of pups with findings	
251	0-4	5	No abnormal findings, 5		10	No abnormal findings, 10	
252	0	6	Death (milk-band negative), 1; Milk-band negative, 5		12	Death (milk-band not examined because of autolysis of abdominal cavity), 2; Milk-band negative, 8; No abnormal findings, 2	
	1	5	Death (milk-band negative), 4; Milk-band negative, 1		10	Death (milk-band negative), 8; Milk-band negative, 2	
	2	1	Milk-band negative, 1		2	Milk-band negative, 2	
	3	1	Missing, 1		2	Death (milk-band not examined because of autolysis of abdominal cavity), 2;	
253	0-4	10	No abnormal findings, 10		6	No abnormal findings, 6	
254	0	11	No abnormal findings, 11		5	No abnormal findings, 5	
	1	11	Death (milk-band negative), 1; Milk-band negative, 10		5	Milk-band negative, 5	
	2	10	Death (milk-band negative), 5; Missing, 3; Milk-band negative, 2		5	Death (milk-band negative), 1; Milk-band negative, 4	
	3	2	Death (milk-band not examined because of autolysis of abdominal cavity), 1; Missing, 1		4	Missing, 4	
255	0-4	5	No abnormal findings, 5		11	No abnormal findings, 11	
256	0-4	3	No abnormal findings, 3		13	No abnormal findings, 13	
257	0-4	7	No abnormal findings, 7		8	No abnormal findings, 8	
258	0	7	No abnormal findings, 7		11	No abnormal findings, 11	
	1	7	No abnormal findings, 7		11	Missing, 1; No abnormal findings, 10	
	2-4	7	No abnormal findings, 7		10	No abnormal findings, 10	
259	0-4	7	No abnormal findings, 7		8	No abnormal findings, 8	
260	0-4	7	No abnormal findings, 7		5	No abnormal findings, 5	
261	0-4	4	No abnormal findings, 4		14	No abnormal findings, 14	
262	0-4	6	No abnormal findings, 6		11	No abnormal findings, 11	

INDIVIDUAL DATA 16-3

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 General appearance, pups ANIMAL : Rat,Crl:CD(SD) SEX : Male, Female GROUP : 20 mg/kg

Dam No.	Day of lactation	Male			Female		
		Number of pups	Finding and number of pups with findings		Number of pups	Finding and number of pups with findings	
351	0-4	4	No abnormal findings, 4		5	No abnormal findings, 5	
352	0-4	7	No abnormal findings, 7		9	No abnormal findings, 9	
353	0-4	5	No abnormal findings, 5		8	No abnormal findings, 8	
354	0-4	4	No abnormal findings, 4		12	No abnormal findings, 12	
355	0-4	8	No abnormal findings, 8		8	No abnormal findings, 8	
356	0-4	5	No abnormal findings, 5		7	No abnormal findings, 7	
357	0-4	9	No abnormal findings, 9		4	No abnormal findings, 4	
358	0-2	9	No abnormal findings, 9		5	No abnormal findings, 5	
	3	9	No abnormal findings, 9		5	Missing, 1; No abnormal findings, 4	
	4	9	No abnormal findings, 9		4	No abnormal findings, 4	
359	0-4	6	No abnormal findings, 6		8	No abnormal findings, 8	
360	0-4	5	No abnormal findings, 5		9	No abnormal findings, 9	
361	0-4	3	No abnormal findings, 3		13	No abnormal findings, 13	
362	0-4	7	No abnormal findings, 7		10	No abnormal findings, 10	

INDIVIDUAL DATA 16-4

STUDY NO. SR07125 TITLE : PFHxD A Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 General appearance, pups ANIMAL : Rat,Crl:CD(SD) SEX : Male, Female GROUP : 100mg/kg

Dam No.	Day of lactation	Male			Female		
		Number of pups	Finding and number of pups with findings	Number of pups	Finding and number of pups with findings		
451	0-4	6	No abnormal findings, 6	7	No abnormal findings, 7		
452	0-4	6	No abnormal findings, 6	7	No abnormal findings, 7		
453	0-1	10	No abnormal findings, 10	6	No abnormal findings, 6		
	2	10	No abnormal findings, 10	6	Missing, 1; No abnormal findings; 5		
	3-4	10	No abnormal findings, 10	5	No abnormal findings, 5		
454	0-4	5	No abnormal findings, 5	8	No abnormal findings, 8		
455	0-4	6	No abnormal findings, 6	9	No abnormal findings, 9		
456	0-1	7	No abnormal findings, 7	6	No abnormal findings, 6		
	2	7	No abnormal findings, 7	6	Death (milk-band positive), 1; No abnormal findings, 5		
	3-4	7	No abnormal findings, 7	5	No abnormal findings, 5		
457	0	5	No abnormal findings, 5	12	No abnormal findings, 12		
	1	5	No abnormal findings, 5	12	Death (milk-band negative), 1; No abnormal findings, 11		
	2-4	5	No abnormal findings, 5	11	No abnormal findings, 11		
458	0	9	Milk-band negative, 1; No abnormal findings, 8	5	Milk-band negative, 1; No abnormal findings, 4		
	1	9	Missing, 1; Milk-band negative, 2; No abnormal findings, 6	5	Death (milk-band not examined because of autolysis of abdominal cavity), 1; Milk-band negative, 1; No abnormal findings, 3		
	2	8	Death (milk-band not examined because of autolysis of abdominal cavity), 3; No abnormal findings, 5	4	Missing, 1; No abnormal findings, 3		
	3-4	5	No abnormal findings, 5	3	No abnormal findings, 3		
459	0	8	Death (milk-band positive), 1; No abnormal findings, 7	3	No abnormal findings, 3		
	1-4	7	No abnormal findings, 7	3	No abnormal findings, 3		
460	0-4	14	No abnormal findings, 14	3	No abnormal findings, 3		
461	0	5	No abnormal findings, 5	11	Milk-band negative, 1; No abnormal findings, 10		
	1	5	No abnormal findings, 5	11	Missing, 1; No abnormal findings, 10		
	2-4	5	No abnormal findings, 5	10	No abnormal findings, 10		
462	0-4	4	No abnormal findings, 4	13	No abnormal findings, 13		

INDIVIDUAL DATA 17-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, pups ANIMAL : Rat, Crl:CD(SD) SEX : Male, Female GROUP : 0 mg/kg

Dam No.	Male			Female		
	Body weight (g) on lactation day			Body weight (g) on lactation day		
	0	1	4	0	1	4
151	6.4	7.1	10.3	6.1	6.8	9.9
152	7.6	8.2	11.7	7.2	7.9	11.1
153	6.2	6.9	9.6	6.1	6.6	9.1
154	6.5	7.0	10.4	6.0	6.6	9.7
155	5.9	6.5	9.8	5.8	6.6	9.8
156	6.1	6.6	9.3	5.7	6.1	8.7
157	6.5	7.1	11.0	6.0	6.7	10.3
158	7.2	7.8	11.1	6.8	7.4	10.1
159	6.2	7.1	10.4	5.9	6.7	10.2
160	7.5	8.1	12.0	7.0	7.5	11.5
161 ^a	#	#	#	#	#	#
162	6.8	7.4	10.2	6.4	7.1	10.2
N	11	11	11	11	11	11
MEAN	6.63	7.25	10.53	6.27	6.91	10.05
S.D.	0.58	0.56	0.85	0.51	0.51	0.79
S.E.	0.17	0.17	0.26	0.15	0.16	0.24
M/C	1.9583	4.2738	3.2081	1.1413	4.9817	3.0921
F	0.5188	0.4573	0.9695	0.6639	0.7355	0.8507

a: Non-pregnancy.

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INDIVIDUAL DATA 17-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, pups ANIMAL : Rat, Crl:CD(SD) SEX : Male, Female GROUP : 4 mg/kg

Dam No.	Male			Female		
	Body weight (g) on lactation day			Body weight (g) on lactation day		
	0	1	4	0	1	4
251	6.5	6.9	10.5	6.1	6.7	9.9
252 ^a	6.0	5.8	#	5.6	4.8	#
253	6.3	6.8	9.8	5.9	6.4	9.5
254 ^a	6.1	5.5	#	5.5	4.8	#
255	6.7	7.3	10.7	6.2	6.9	9.7
256	6.5	6.9	9.6	6.1	6.5	9.5
257	8.2	9.1	12.7	7.5	8.2	11.9
258	6.0	6.3	8.9	5.9	6.2	8.7
259	6.5	7.0	9.0	6.2	6.6	8.7
260	7.5	8.6	13.5	7.1	8.2	12.7
261	7.3	8.3	11.6	6.7	7.4	10.4
262	6.4	6.9	10.0	5.8	6.3	8.7
N	12	12	10	12	12	10
MEAN	6.67	7.12	10.63	6.22	6.58	9.97
S.D.	0.67	1.08	1.54	0.60	1.07	1.36
S.E.	0.19	0.31	0.49	0.17	0.31	0.43

a : All pups died by day 3 of lactation.

#: Blank.

INDIVIDUAL DATA 17-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, pups ANIMAL : Rat, Crl:CD(SD) SEX : Male, Female GROUP : 20 mg/kg

Dam No.	Male			Female		
	Body weight (g) on lactation day			Body weight (g) on lactation day		
	0	1	4	0	1	4
351	7.0	8.0	12.0	6.9	7.6	12.0
352	6.6	7.1	10.3	5.9	6.1	8.4
353	7.0	7.7	11.2	6.7	7.6	11.4
354	6.1	6.9	10.2	5.6	6.2	9.2
355	6.0	6.3	9.7	5.9	6.2	9.2
356	8.3	8.5	12.1	7.6	8.0	11.3
357	7.5	8.2	11.3	7.4	8.2	10.8
358	7.2	8.2	12.5	6.8	7.9	11.8
359	6.4	6.9	10.4	6.2	6.8	10.1
360	6.5	6.8	9.2	6.0	6.3	8.8
361	6.5	6.9	10.0	6.0	6.5	9.5
362	5.9	6.5	9.1	5.8	6.3	9.3
N	12	12	12	12	12	12
MEAN	6.75	7.33	10.67	6.40	6.98	10.15
S.D.	0.69	0.75	1.14	0.66	0.82	1.25
S.E.	0.20	0.22	0.33	0.19	0.24	0.36

INDIVIDUAL DATA 17-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Body weight, pups ANIMAL : Rat, Crl:CD(SD) SEX : Male, Female GROUP : 100 mg/kg

Dam No.	Male			Female		
	Body weight (g) on lactation day			Body weight (g) on lactation day		
	0	1	4	0	1	4
451	6.4	7.0	10.3	5.9	6.5	9.6
452	6.3	7.1	9.7	5.8	6.4	9.2
453	6.4	6.6	9.4	6.1	6.4	8.9
454	7.1	8.1	11.7	6.8	7.8	11.5
455	6.9	7.5	10.0	6.3	6.8	9.3
456	7.3	8.2	11.8	7.2	8.1	11.8
457	6.2	6.9	9.8	5.8	6.6	9.3
458	5.6	5.2	7.6	5.4	4.9	6.8
459	6.6	7.2	11.0	6.1	6.5	10.0
460	6.2	6.7	8.8	5.9	6.3	8.6
461	6.2	6.8	10.4	5.6	6.4	9.5
462	6.2	6.3	8.6	6.0	6.4	8.7
N	12	12	12	12	12	12
MEAN	6.45	6.97	9.93	6.08	6.59	9.43
S.D.	0.46	0.80	1.24	0.50	0.79	1.31
S.E.	0.13	0.23	0.36	0.14	0.23	0.38

INDIVIDUAL DATA 18

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
 Gross findings of dead pups on lactation days 0-4 ANIMAL : Rat,Crl:CD(SD) SEX : Male, Female

Group	Dam No.	Day of death	Number of pups		
			Male	Female	Findings
0 mg/kg	162	0	1	#	No abnormal findings (autolysis in abdominal cavity)
4 mg/kg	252	0	1	#	No abnormal findings
		0	#	2	No abnormal findings (autolysis in abdominal cavity)
		1	4	8	No abnormal findings
	254	3	#	2	No abnormal findings (autolysis in abdominal cavity)
		1	1	#	No abnormal findings
		2	5	1	No abnormal findings
100 mg/kg	456	3	1	#	No abnormal findings (autolysis in abdominal cavity)
		2	#	1	No abnormal findings
		1	#	1	No abnormal findings
	458	1	#	1	No abnormal findings (autolysis in abdominal cavity)
		2	2	#	No abnormal findings
		2	1	#	No abnormal findings (autolysis in abdominal cavity)
	459	0	1	#	No abnormal findings

#: Blank.

INDIVIDUAL DATA 19-1

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Gross findings of pups euthanized on lactation day 4 ANIMAL : Rat,Crl:CD(SD) SEX : Male, Female GROUP : 0 mg/kg

Dam No.	Number of pups		
	Male	Female	Findings
151	8	8	No abnormal findings
152	8	6	No abnormal findings
153	5	10	No abnormal findings
154	5	9	No abnormal findings
155	4	12	No abnormal findings
156	9	8	No abnormal findings
157	6	5	No abnormal findings
158	8	9	No abnormal findings
159	9	6	No abnormal findings
160	13	2	No abnormal findings
161 ^a	#	#	#
162	7	8	No abnormal findings

a: Non-pregnancy.

#: Blank.

INDIVIDUAL DATA 19-2

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Gross findings of pups euthanized on lactation day 4 ANIMAL : Rat,Crl:CD(SD) SEX : Male, Female GROUP : 4 mg/kg

Dam No.	Number of pups		
	Male	Female	Findings
251	5	10	No abnormal findings
252 ^a	#	#	#
253	10	6	No abnormal findings
254 ^a	#	#	#
255	5	11	No abnormal findings
256	3	13	No abnormal findings
257	7	8	No abnormal findings
258	7	10	No abnormal findings
259	7	8	No abnormal findings
260	7	5	No abnormal findings
261	4	14	No abnormal findings
262	6	11	No abnormal findings

a: All pups died by day 3 of lactation .

#: Blank.

INDIVIDUAL DATA 19-3

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Gross findings of pups euthanized on lactation day 4 ANIMAL : Rat,Crl:CD(SD) SEX : Male, Female GROUP : 20 mg/kg

Dam No.	Number of pups		
	Male	Female	Findings
351	4	5	No abnormal findings
352	7	9	No abnormal findings
353	5	8	No abnormal findings
354	4	12	No abnormal findings
355	8	8	No abnormal findings
356	5	7	No abnormal findings
357	9	4	No abnormal findings
358	9	4	No abnormal findings
359	6	8	No abnormal findings
360	5	9	No abnormal findings
361	3	13	No abnormal findings
362	7	10	No abnormal findings

INDIVIDUAL DATA 19-4

STUDY NO. SR07125 TITLE : PFHxDA Combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (p.o.)
Gross findings of pups euthanized on lactation day 4 ANIMAL : Rat,Crl:CD(SD) SEX : Male, Female GROUP : 100 mg/kg

Dam No.	Number of pups		
	Male	Female	Findings
451	6	7	No abnormal findings
452	6	7	No abnormal findings
453	10	5	No abnormal findings
454	5	8	No abnormal findings
455	6	9	No abnormal findings
456	7	5	No abnormal findings
457	1	#	Liver: Yellowish white patch, middle lobe (5 x 3, mm)
	4	11	No abnormal findings
458	5	3	No abnormal findings
459	7	3	No abnormal findings
460	14	3	No abnormal findings
461	5	10	No abnormal findings
462	4	13	No abnormal findings

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EXFLUOR RESEARCH CORPORATION

2350 Double Creek Drive
Round Rock, Texas 78664
(512) 310-9044

C E R T I F I C A T E O F A N A L Y S I S

Issue Date: August 31, 2006

Product Name: Perfluorohexadecanoic acid

CAS # 67905-19-5

Chemical Formula: $\text{CF}_3(\text{CF}_2)_{14}\text{COOH}$

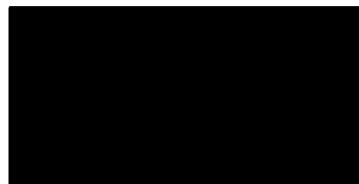
Lot # 1262

Physical State White solid

Products other than Perfluorohexadecanoic acid ≤ 5 %

Perfluorohexadecanoic acid 95 % min.

Test Results 95.3 %



Exflouor Research Corporation

被験物質原体の純度試験 報告書

被験物質名：ペルフルオロヘキサデカン酸
 製造会社名：Exfluor Research Corporation
 ロット番号：1262
 測定年月日：2011年3月31日～2011年4月1日
 保存条件：遮光容器にて室温保存
 測定方法：LC/MS/MS法(純度試験；定量法)
 試験成績：

項目	結果				
	測定の繰返し数	純度 (%)			
純度試験 (定量法)		測定値 (%)	平均値	標準偏差	変動係数 (%)
1	99.9				
2	97.8	98.8	1.06	1.07	
	3	98.6			

委託者：株式会社 化合物安全性研究所

試験施設：野外科学株式会社 環境計量部

〒065-0043 札幌市東区苗穂町12丁目2番39号

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分析担当者

分析証明書番号 : 1371

均一性試験 分析証明書

被 験 物 質 名 : ペルフルオロヘキサデカン酸

ロ ッ ト 番 号 : 1262

媒 体 : 0.5%カルボキシメチルセルロースナトリウム水溶液

調 製 年 月 日 : 2010年6月25日

分析試験実施時期 : 投与開始前

測 定 開 始 日 : 2010年6月25日

測 定 方 法 : LC-MS/MS 法

試 験 成 績 :

調製液 表示濃度	測定の 繰返し数	被験物質濃度 (mg/mL)	平均値 ±標準偏差	変動係数 (%)	
0.01 mg/mL	上層 1	0.00999	0.00970 ±0.000208	2.1	
	2	0.00980			
	中層 1	0.00977			
	2	0.00937			
100 mg/mL	下層 1	0.00965	97.4 ±2.26	2.3	
	2	0.00963			
	上層 1	99.0			
	2	98.9			
	中層 1	97.8	97.4 ±2.26		
	2	99.4			
	下層 1	95.0			
	2	94.1			

判 定 基 準 : 変動係数が 5%以下の場合を適とする。

合 否 判 定 : 適

備 考 :

実施試験名 ; ペルフルオロヘキサデカン酸のラットにおける反復投与毒性・生殖発生毒性併合試験 (試験番号 ; SR07125)

試験施設 株式会社 化合物安全性研究所

化学分析責任者 :

2010 年 7 月 8 日

分析証明書番号 : 1372

安定性試験 分析証明書

被 驗 物 質 名 : ペルフルオロヘキサデカン酸
 ロ ッ ト 番 号 : 1262
 媒 体 : 0.5%カルボキシメチルセルロースナトリウム水溶液
 調 製 年 月 日 : 2010年8月5日
 分析試験実施時期 : 投与開始前
 測 定 開 始 日 : 2010年8月5日(調製時)、
 2010年8月15日(冷蔵保存10日の室温保存4時間後)
 保 存 条 件 : 冷蔵保存後、室温保存
 測 定 方 法 : LC-MS/MS法
 試 験 成 績 :

調製液 表示濃度 mg/mL	測定の 繰返し数	被驗物質濃度 (mg/mL)	
		調製時	冷蔵保存10日の 室温保存4時間後
0.01	1	0.0103	0.0103
0.01	2	0.0102	0.00974
mg/mL	平均値	0.0103	0.0100
	残存率(%)	-	97.1
100	1	101	105
100	2	103	102
mg/mL	平均値	102	104
	残存率(%)	-	102.0

判 定 基 準 : 残存率が 85~115%の場合を適とする。
 合 否 判 定 : 0.01 および 100 mg/mL の被驗物質調製液について、冷蔵保存10日の室温保存4時間後まで安定性が認められた(調製日を0日として起算)。

備 考 :
 実施試験名 ; ペルフルオロヘキサデカン酸のラットにおける反復投与毒性・生殖発生毒性併合試験(試験番号 ; SR07125)

試験施設 株式会社 化合物安全性研究所

化学分析責任者 :

2010 年 8 月 17 日

分析証明書番号 : 1373

濃度確認試験 分析証明書

被 験 物 質 名 : ペルフルオロヘキサデカン酸
 ロ ッ ト 番 号 : 1262
 媒 体 : 0.5%カルボキシメチルセルロースナトリウム水溶液
 調 製 年 月 日 : 2010年12月15日
 分析試験実施時期 : 初回調製時
 測 定 年 月 日 : 2010年12月16日
 測 定 方 法 : LC-MS/MS 法
 試 験 成 績 :

調製液 表示濃度 (mg/mL)	測定の 繰返し数	調製液		
		被験物質濃度 (mg/mL)	平均値	含有率 (%)
0.4	1	0.428	0.418	104.5
	2	0.408	±0.0141	
2.0	1	2.04	2.06	103.0
	2	2.08	±0.028	
10	1	10.4	10.4	104.0
	2	10.4	±0.00	

判 定 基 準 : 含有率が 85~115%の場合を適とする。

合 否 判 定 : 適

備 考 :

実施試験名 ; ペルフルオロヘキサデカン酸のラットにおける反復投与毒性・生殖発生毒性併合試験(試験番号 SR07125)

試験施設 株式会社 化合物安全性研究所

化学分析責任者 :

2010 年 12 月 21 日

分析証明書番号 : 1374

濃度確認試験 分析証明書

被 験 物 質 名 : ペルフルオロヘキサデカン酸
 ロ ッ ト 番 号 : 1262
 媒 体 : 0.5%カルボキシメチルセルロースナトリウム水溶液
 調 製 年 月 日 : 2011年1月27日
 分析試験実施時期 : 最終回調製時
 測 定 年 月 日 : 2011年1月28日
 測 定 方 法 : LC-MS/MS法
 試 験 成 績 :

調製液 表示濃度 (mg/mL)	測定の 繰返し数	調製液		
		被験物質濃度 (mg/mL)	平均値	含有率 (%)
0.4	1	0.384	0.394	98.5
	2	0.404	±0.0141	
2.0	1	1.99	1.97	98.5
	2	1.94	±0.035	
10	1	9.79	9.85	98.5
	2	9.91	±0.085	

判 定 基 準 : 含有率が 85~115%の場合を適とする。

合 否 判 定 : 適

備 考 :

実施試験名 ; ペルフルオロヘキサデカン酸のラットにおける反復投与毒性・生殖発生毒性併合試験(試験番号 SR07125)

試験施設 株式会社 化合物安全性研究所

化学分析責任者 : [REDACTED]

2011 年 / 月 31 日

Symbols and process for statistical analysis in individual data

M/C: Values for Bartlett's test for homogeneity of variance,

$p \leq 0.05$ Kruskal-Wallis test

$p > 0.05$ One way analysis of variance

F : Values for one way analysis of variance,

$p \leq 0.10$ Dunnett's procedure

H : Values for Kruskal-Wallis test,

$p \leq 0.10$ Mann-Whitney's U-test

† : Significant difference, $p \leq 0.10$

* : Significant difference, $p \leq 0.05$

** : Significant difference, $p \leq 0.01$

t' : Values for Dunnett's procedure

U : Values for Mann-Whitney's U-test

* : Significant difference, $p \leq 0.05$

** : Significant difference, $p \leq 0.01$

χ^2 : Values for chi-square test

p : Values for Fisher's exact probability test

* : Significant difference, $p \leq 0.05$

** : Significant difference, $p \leq 0.01$

Definitions for detailed clinical and functional observations

Item	Category No.					
In the cage:						
Body position/Posture						
	1; Normal (sitting, etc.)	2; Sleeping	3; Crouching	4; Prone, lateral	5; Standing, jumping	6; Cataleptic
Respiratory pattern						
	1; Normal	2; Slightly abnormal, rapid or slow	3; Moderately abnormal, difficult to breath	4; Severe abnormal, labored	5; Dyspnea	
Tremor / Convulsion						
	1; Not present	2; Irregularly, only the legs	3; Durable, only the legs	4; Clonic, systemic	5; Tonic, systemic	
Stereotype						
Rolling	0; Not present	1; Sometimes	2; Frequently			
Repetitive circling						
	0; Not present	1; Sometimes	2; Frequently			
Bizarre behavior						
Biting/Selfmutilation						
	1; Not present	2; Present				
On the hand: while removing the animal from its cage						
Ease of removal						
	1; Very easy	2; Easy	3; Slightly difficult	4; Difficult	5; Very difficult	
Ease of handling						
	1; Very easy	2; Easy	3; Slightly difficult	4; Difficult	5; Very difficult	
Muscle tone	1; Low	2; Normal	3; High			
Piloerection	1; Not present	2; Slightly present, around head and back	3; Slightly present, systemic	4; Severely present, systemic		
Fur	1; Normal	2; Slightly stained	3; Stained			
Eyes	1; Not present, normal	2; Slight ptosis, half closed	3; Ptosis	4; Closed		
Mucous membranes						
	-1; Dark purplish appearance, cyanosis	0; Normal	1; Red appearance			

(to be continued)

(Continued 1)

Item	Category No.		
Skin	0; Pale appearance, cyanosis	1; Normal	2; Red appearance
Pupil size	1; Normal	2; Slightly mydriatic	3; Mydriatic
Lacration	1; Not present	2; Wet around the eye	3; Wet 4; Severely wet
Salivation	1; Not present	2; Wet around the mouth	3; Wet 4; Severely wet
Secretions/Excretions			
	0; Not present	1; Present	
In the open-field: when placed the animal in an open-field			
Gait	0; Not moved	1; Normal	2; Difficult to walk 3; Unable to walk, paralytic
Co-ordination of movement			
	0; staggering (ataxic)	1; Normal	
Reactivity to environmental stimuli			
	0; Not present	1; present (to noise, etc.)	
Searching	0; Not present	1; present (sniffing, standing, etc.)	
Urination	0; Not present	1; Present	
Defecation	0; Not present	1; Present	
Stereotype			
Excessive grooming			
	0; Not present	1; Sometimes	2; Frequently
Unusual head movement			
	0; Not present	1; Sometimes	2; Frequently
Bizarre behavior			
Walking backward			
	1; Not present	2; Present	
Vocalization	1; Not present, sometimes	2; Present, frequently	
Aggression	1; Not present	2; Present	

(to be continued)

(Continued 2)

Item	Category No.
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On the desk : functional observation, stimulus reactivity

Visual reactivity: approach response

1; Jumping	2; Turning away	3; No reaction	4; Approach	5; Attack
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Touch reactivity: touch response

1; Hyposensitive	2; Turning away	3; Hypersensitive
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Auditory reactivity: response to Galton's whistle

0; No reaction	1; Normal, moving the auricle	2; Sensitive, moving the body	3; Hypersensitive, surprising and jumping
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Pain reactivity: tail pinch response

0; No reaction	1; Dull, vocalizing	2; Normal, vocalizing and turning back	3; Hypersensitive attacking or jumping
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Proprioceptive reactivity: returning from enforced posture

0; No returning	1; Returning
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Righting reflex: landing performance from 30 cm above

1; Normal, landing by foot	2; Abnormal, landing by body
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Others

Grip strength: measuring by CPU gage[®]

expressed by mean values (g) from 3 trials; forelimb and hindlimb

Motor activity measurements: measuring by SUPERMEX and CompACT AMS

expressed by total counts for 60 minutes at 10 minutes' intervals