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SEHSC SILICONES ENVIRONMENTAL, HEALTH AND SAFETY COUNCIL of North America

May 27, 2003

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TSCA Document Control Center (7407)
Office of Pollution Prevention and Toxics
US Environmental Protection Agency
Attn: TSCA Section 8(e) Coordinator
Ariel Rios Building
1200 Pennsylvania Avenue, NW
Washington, DC 20004

8903000196

Re: Final Report of Study, Subject of May 24, 2003, SEHSC Letter on TSCA Section 8(e) Notification of Substantial Risk: Hexamethylcyclotrisiloxane

On May 24, 2003, the Silicones Environmental, Health and Safety Council (SEHSC) submitted to the US EPA, on behalf of its member companies, information concerning a then ongoing study with hexamethylcyclotrisiloxane (CAS No. 541-05-9), in accordance with the provisions of Section 8(e) of the Toxic Substances and Control Act (TSCA). This study was conducted as part of SEHSC's commitment to the ICCA Global Initiative on HPV Chemicals.

SEHSC is a not-for-profit trade association whose mission is to promote the safe use and stewardship of silicones. The Council is comprised of North American silicone chemical producers and importers. SEHSC's members represent over 95 percent of silicone chemical manufacturing capacity in North America and include: Clariant LSM (Florida), Inc.; Degussa Corporation; Dow Corning Corporation; General Electric Silicones; Crompton Corporation, Osi Specialties business; Rhodia Inc.; Shin-Etsu Silicones of America; and Wacker Silicones, A Division of Wacker Chemical Corporation.

In the May 24, 2003, TSCA Section 8(e) Notification of Substantial Risk for Hexamethylcyclotrisiloxane, SEHSC agreed to provide the US EPA with a copy of the final report for the study, Combined Repeated Dose Toxicity Study with Reproduction/Developmental Toxicity Screening Test for Hexamethylcyclotrisiloxane (D3) in Sprague-Dawley Rats, Dow Corning Corporation, Dow Corning Study No. 9658. The final report for the aforementioned study is enclosed for the Agency's reference.

If you have any questions, please contact me at (703) 904-4322, rmenning@sehsc.com, or at the address provided herein.

Sincerely,

Contain NO CBI

Reo Menning
Executive Director

267282

11921 FREEDOM DRIVE, SUITE 550, RESTON, VA 20190 • (703) 904-4322 • Fax (703) 925-5955

Study Title:	Combined Repeated Dose Toxicity Study with the Reproductive/Developmental Toxicity Screening Test for Hexamethylcyclotrisiloxane (D3) in Sprague-Dawley Rats
Study Number:	9658
Test Substance:	Hexamethylcyclotrisiloxane
Study Director:	Paul A. Jean, Ph.D.
Sponsor:	Silicones Environmental, Health and Safety Council 11921 Freedom Drive Suite 550 Reston, VA 20190
Sponsor Representative:	Edward J. Hobbs
Testing Facility:	Health and Environmental Sciences Dow Corning Corporation 2200 West Salzburg Road Midland, MI 48686-0994
Study Completion Date:	December 13, 2002

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I. ABSTRACT:

The objective of this study was to evaluate the potential toxicity of hexamethylcyclotrisiloxane (D3) in a combined repeated-exposure toxicity study that included reproductive/developmental toxicity screening in Sprague-Dawley rats by whole-body vapor inhalation following enhanced test guidelines (Organisation for Economic Co-Operation and Development (OECD) Test Guideline number: 422, (1996) and United States Environmental Protection Agency (U.S.E.P.A.) Health Effects Test Guideline Office of Prevention, Pesticides, and Toxic Substances (OPPTS) 870.3650 (2000)). Exposure levels were 0 (control), 100, 500 and 2500 ppm D3. Rats were exposed to the test atmosphere for six hours per day for up to 39 consecutive days. Females in each exposure level were divided into a toxicity group (10 animals/group) and a reproductive group (10 animals/group). A single group of males (10 animals/group) was used for both the toxicity and reproductive phases of the study. Males and toxicity group females were treated for 28 and 29 days, respectively. Reproductive group females were treated for 14 days prior to the mating period, during the mating period and then up to and including gestation day 19.

Animals were observed twice daily (weekdays) or once daily (weekends) for mortality or morbidity. Clinical examinations were performed daily immediately following exposure. Detailed physical examinations and body weight measurements were performed weekly. Additional body weights on reproductive group females were obtained on gestational day (GD) 0, 7, 14, and 20, within 24 hours of parturition, and on postnatal day (PND) four. Individual food consumption was recorded at least weekly, except during the cohabitation period. Functional observational battery (FOB) and motor activity evaluations were performed on males and toxicity group females once prior to initiation of exposures and during the fourth week of exposure. Blood samples for hematology and serum chemistry evaluations were collected at the scheduled necropsy from males and toxicity group females. Complete necropsies were performed on the males and the toxicity group females and selected organs were weighed. Microscopic examination was performed on protocol-specified tissues from all animals in the control and 2500 ppm D3 exposure group males and toxicity group females. Microscopic examination was extended to include the liver (males and females), kidneys (males) and seminal vesicles (males) from the 100 and 500 ppm exposure groups.

A mating phase was initiated after the first two weeks of exposure by pairing reproductive group females with males of the same treatment group until positive evidence of mating was obtained. Reproductive and developmental parameters evaluated included evidence of mating, duration of gestation, mean litter size, mean live litter size, mean litter weight, mean ratio live births/litter size, corpora lutea count, and the number of uterine implantation sites. Dams and pups were euthanized on PND 4 and examined for external gross lesions.

In general, whole-body inhalation exposure of male and female Sprague-Dawley rats to D3 vapor at concentrations of up to 2500 ppm for 28-29 consecutive days was well tolerated, as suggested by minimal effects on body weight and food consumption, the lack of significant clinical signs of toxicity, and minimal effects on clinical pathology parameters.

Increased liver weight and centrilobular hepatocellular hypertrophy were demonstrated in both sexes exposed to 2500 ppm D3. These effects are considered to be an adaptive response characteristic of this class of cyclic siloxanes. No liver effects were identified at lower exposure levels.

Protein droplet nephropathy was observed by histopathology in the kidneys of males exposed to 500 and 2500 ppm D3. One male rat exposed to 100 ppm D3 was identified with a very slight lesion of the same nature. The finding was corroborated for males in the 2500 ppm exposure group by organ weight and clinical chemistry changes. Females were unaffected. This nephropathy, affecting only male rats, is consistent in incidence and histomorphology with that caused by excessive accumulation of α_{2U}-globulin in renal cortical tubules. This mechanism is specific to male rats and not associated with human risk.

In males exposed to 2500 ppm D3, there were statistically significant decreases in the absolute weight of epididymides (10%) and in the absolute (30%) and relative (27%) weight of the seminal vesicles. Slight atrophy of the seminal vesicles was identified microscopically in four of the 10 males in the 2500 ppm exposure group. The epididymides, testes, and prostate from D3-exposed rats were histologically normal.

Negative effects on litter size parameters were observed for female rats exposed to 2500 ppm D3. There was a 33% decrease in litter size and in the number of implantation sites. Gestation length, pup sex ratio, pup weight and viability, and corpora lutea counts were unaffected. No effects were observed at lower exposure concentrations.

At the 2500 ppm exposure level there was reduced motor activity in both sexes and a reduced "reaction to handling" in females. There was no histomorphologic evidence of neurotoxicity at any exposure level.

Within the context of the 28-day sub-acute toxicity portion of this study no adverse effects attributable to treatment were noted for female rats. Adverse effects attributed to treatment were noted for males and included an increased incidence of protein droplet nephropathy (100 ppm "no observable adverse effect level" (NOAEL)) and decreased seminal vesicle weight (500 ppm NOAEL). Within the reproductive/developmental toxicity screening test portion of this study a NOAEL of 500 ppm was identified for effects on reproductive parameters (decreased litter size and number of implantation sites).

II. GLP COMPLIANCE STATEMENT:

This study was conducted in compliance with current U.S.E.P.A. Toxic Substances Control Act Good Laboratory Practice Standards (GLP) 40 CFR Part 792, except feed and water analyses which were not conducted according to GLP. In addition, stability of the neat test substance under the conditions of storage at the testing facility was based upon information provided by the supplier. Protocol deviations are summarized in Table 46.

Paul A. Jean, Ph.D.
Study Director

13 DEC 2002
Date

Steven D. Crofoot, M.S.
Team Leader, Toxicology
Health and Environmental Sciences

12/13/2002
Date

Edward J. Hobbs
Sponsor Representative
Silicones Environmental, Health and Safety Council

12-5-02
Date

III. QUALITY ASSURANCE STATEMENT:

Title: Combined Repeated Dose Toxicity Study with the Reproductive/Developmental Toxicity Screening Test for Hexamethylcyclotrisiloxane (D3) in Sprague-Dawley Rats

This study has been audited by the Dow Corning Corporation Health and Environmental Sciences Quality Assurance Unit according to approved Standard Operating Procedures to assure that the raw data are accurately reflected within this final report. The following are the inspection dates and the dates inspection findings were reported.

Dates of Inspection	Phase Inspected	Findings Reported to Study Director	Findings Reported to Management
03-05 Dec 01	Draft Protocol Review	05 Dec 01	05 Dec 01
12 Mar 02	Chamber Homogeneity	12 Mar 02	13 Mar 02
21 Mar 02	Functional Observational Battery (FOB)	21 Mar 02	08 Apr 02
28 Mar 02	Daily Bag Standard and Exposure	28 Mar 02	09 Apr 02
15 Apr 02	Body Weight and Feeder Weight Collection	15 Apr 02	16 Apr 02
18 Apr 02	Detailed Physical Observations	19 Apr 02	23 Apr 02
22-23 Apr 02	Blood Collection, Prosection, Necropsy and Clinical Pathology Analyses	23 Apr 02	25 Apr 02
30 May 02	Sectioning Tissue Blocks	30 May 02	03 Jun 02
11 Jun 02	Microscopic Examination of Slides	12 Jun 02	12 Jun 02
5, 14, 28 Aug 02	Data – Test System	29 Aug 02	25 Sep 02
29 July, 01-09 and 22-30 Aug 02	Draft Final Report Tables and Associated Raw Data – Clinical Observations, Body Weights, Feed Consumption, Clinical Chemistry	03 Sep 02	19 Sep 02
02-06 Sep 02	Draft Final Report Tables and Associated Raw Data - FOB	09 Sep 02	19 Sep 02
10-18 Sep 02	Draft Report Tables and Associated Raw Data – Post-Partum Observations, Study Events and Histology Processing	18 Sep 02	19 Sep 02
25 Sep 02	Wet Specimen Verification	25 Sep 02	29 Sep 02
21-23 Aug and 19-25 Sep 02	Draft Final Report and Associated Raw Data Review	25 Sep 02	08 Oct 02

Joyce L. Henry
Manager, Quality Assurance
Health and Environmental Sciences
Dow Corning Corporation

12 Dec 02
Date

IV. APPROVAL SIGNATURES:

The undersigned have read and approved this report:

Paul A. Jean, Ph.D.
Study Director

13 DEC 2002
Date

Steven D. Crofoot, M.S.
Team Leader, Toxicology
Health and Environmental Sciences

12/13/2002
Date

James W. Crissman, D.V.M., Ph.D., D.A.C.V.P.
Attending Veterinarian and Pathologist

12/12/02
Date

Edward J. Hobbs
Sponsor Representative
Silicones Environmental, Health and Safety Council

12-5-02
Date

V. STUDY INFORMATION:

Study Initiation Date:	December 6, 2001
Experimental Start Date:	March 25, 2002
Experimental Termination Date:	November 7, 2002
Study Completion Date:	December 13, 2002
Study Number:	9658
Study Title:	Combined Repeated Dose Toxicity Study with the Reproductive/Developmental Toxicity Screening Test for Hexamethylcyclotrisiloxane (D3) in Sprague-Dawley Rats
Testing Facility:	Health and Environmental Sciences (HES) Dow Corning Corporation 2200 West Salzburg Road Midland, MI 48686-0994
Sponsor:	Silicones Environmental, Health and Safety Council 11921 Freedom Drive Suite 550 Reston, VA 20190
Study Director:	Paul A. Jean, Ph.D.
Team Leader, Toxicology:	Steven D. Crofoot, M.S.
Study Coordinator:	Matthew C. Long, B.S.
Manager, Quality Assurance:	Joyce L. Henry, B.S.
Attending Veterinarian/Pathologist:	James W. Crissman, D.V.M., Ph.D., D.A.C.V.P.
Manager, Analytical Chemistry and Operations:	Roy A. Campbell, B.S.
Supervisor, HES Technical Staff:	Joseph M. Tobin, B.S.

VI. OBJECTIVE:

The objective of this study was to evaluate D3 in a combined repeated-exposure toxicity study that included reproductive/developmental toxicity screening in Sprague-Dawley rats by whole-body vapor inhalation. The study was designed to meet the requirements of OECD Test Guideline number: 422 (1996) and U.S.E.P.A. Health Effects Test Guideline OPPTS 870.3650 (2000) and included enhancements to these guidelines.

VII. TEST SUBSTANCE SPECIFICATIONS:

The results of test substance characterization were reviewed by the study director and documented in the study file.

Identification:	Hexamethylcyclotrisiloxane
Batch Number:	1N44
CAS Number:	541-05-9
Physical Description:	Flammable white solid
Source:	GE Silicones 260 Hudson River Road Waterford, NY 12188
Handling Precautions:	Avoid contact with skin and eyes. Use only in a well ventilated area.
Chemical Stability:	Stable
Storage Conditions:	Store away from heat.
Expiration Date:	September 19, 2002
Solubility:	Common organic solvents.
Test Substance Characterization:	A characterization study was performed by Dow Corning Corporation (Study Number 9651). Report entitled *Characterization of Hexamethylcyclotrisiloxane (D3) (Lot/Batch/Reference No. 1N44).*
Purity:	> 99%

VIII. CARRIER:

J-tube Vaporization:

Identification:	Air
Source:	Nash® Air Compressor (Model/Size: AL-574)
Conditioning:	The air was passed through a series of purification filters (Matheson® model 460/461 and Balston® model 100-18-DX and 100-8-BX) prior to use in test atmosphere generation.

Chamber Make-up:

Identification:	Air
Source:	Building Air Supply
Conditioning:	Conditioned building air was passed through HEPA and activated charcoal filters before delivery to the chamber.

IX. JUSTIFICATION OF ROUTE OF EXPOSURE AND DOSE LEVELS:

Route: Test substance was administered by whole-body vapor inhalation.

Rationale: Whole-body vapor inhalation is an accepted method of administration of test substances. In particular, vapor inhalation exposure is appropriate for this test substance as it represents the most likely route of potential human exposure. Exposure levels were selected based on results from a previous inhalation toxicity study (Appendix T: 90-Day Study Abstract).

X. TEST SYSTEM:

Species:	*Rattus norvegicus*
Strain:	Crl:CD®(SD)IGS BR VAF/Plus®
Source:	Charles River Canada 188 LaSalle St. Constant, Canada
Age:	≥ 8 weeks at experimental start
Body Weight Range:	Females: 187 - 238 grams at experimental start Males: 250 - 313 grams at experimental start
Sex and Number:	80 females and 40 males Females were nulliparous and nonpregnant at experimental start.
Number of Groups:	4
Identification Method:	Upon receipt individual cage labels displayed a temporary quarantine (Q) number
Permanent Identification:	Ear tags and individual cage labels
Justification for Selection of Test System:	The rat is recognized as appropriate for toxicity studies and is recommended in the OECD test guideline for this study. CD® rats have been used in repeated-dose studies and are routinely utilized to evaluate reproductive toxicity.

XI. HOUSING AND MAINTENANCE:

Animal Receipt and Quarantine

Upon receipt, each animal was inspected by animal resource personnel and animals judged to be in good health and suitable as test animals were quarantined for five days. During the quarantine period animal resource personnel observed each animal at least once daily. The attending veterinarian examined all animals before release from quarantine and documented the general state of animal health.

Animal Housing

Animals were individually housed in suspended wire-mesh cages (7"x10"x7") elevated over Bed-O'Cobs® Alf-a' cobs bedding, during quarantine and throughout the course of the study, with the following exceptions. During cohabitation, reproductive group females were paired 1 to 1 with a male from the same treatment group in a suspended wire-mesh cage elevated over Bed-O'Cobs® Alf-a' cobs bedding. The animals were paired in the home cage of the male. On day 14 of gestation, Reproductive group females were moved into shoebox cages (8"x16.5"x8") containing Enviro-dri and Bed-O'Cobs® combination bedding for the remainder of the study.

Environmental Conditions

Animals were housed in an environmentally controlled animal room (12-hour fluorescent light/dark cycle, 66 - 77° Fahrenheit (19 - 25° Celsius), 30 - 70% relative humidity, 10 - 15 air changes per hour) throughout the in-life phase of the study. Temperature and humidity were continuously monitored via a seven-day chart recorder and hand recorded, twice a day on weekdays and once a day during weekends and holidays. One exception occurred in the 12-hour fluorescent light/dark cycle during the study. On study day one the light portion of the cycle was extended by approximately five minutes to complete the daily clinical observations.

Basal Diet

PMI® Certified Rodent Chow #5002, manufactured by PMI Nutrition International, St. Louis, MO, was offered *ad libitum* except during the daily inhalation exposure period and, for males and toxicity group females, the time between the last exposure and necropsy. The results of the manufacturer's periodic analysis of the certified feed were reviewed to ensure that heavy metals and pesticides were not present in concentrations that would be expected to affect the outcome of the study.

Drinking Water

Municipal water, further purified by reverse osmosis, was available *ad libitum* except during the daily exposure period. The water has been monitored on at least a semi-annual basis to determine compliance with the U.S.E.P.A. drinking water standards. The most recent analysis was reviewed and there were no contaminants in the water known to be present at levels expected to interfere with the integrity of the study.

XII. ANIMAL WELFARE ACT COMPLIANCE:

This study complied with all applicable sections of the final rules of the Animal Welfare Act regulations (9 CFR, Parts 1, 2, and 3) and was approved by the Institutional Laboratory Animal Care and Use Committee before animals were ordered.

XIII. EXPERIMENTAL DESIGN:

Randomization of Test System

Upon release from quarantine, rats were assigned to treatment groups based on a weight stratified randomization process. Animals outside ± 20% the mean body weight for each sex were excluded.

Organization of Test Groups and Exposure Levels

Group	Group ID	Sex	Number of Animals/Group	Exposure Level (ppm)	Frequency of Dosing	Exposure Duration
1	Controls	F	10/Toxicity Group & 10/ Reproductive Group	0	Daily	29 days minimum*
1	Controls	M	10	0	Daily	28 days
2	100 ppm	F	10/Toxicity Group & 10/ Reproductive Group	100 ppm	Daily	29 days minimum*
2	100 ppm	M	10	100 ppm	Daily	28day
3	500 ppm	F	10/Toxicity Group & 10/ Reproductive Group	500 ppm	Daily	29 days minimum*
3	500 ppm	M	10	500 ppm	Daily	28 days
4	2500 ppm	F	10/Toxicity Group & 10/ Reproductive Group	2500 ppm	Daily	29 days minimum*
4	2500 ppm	M	10	2500 ppm	Daily	28 days

*Reproductive group females were treated up to and including gestation day 19.

XIV. EXPOSURE SYSTEM:

Vapor Generation

Test substance was placed in a warming chamber (heated to approximately 75°C) to maintain it in a liquid state. The test substance was then metered into a heated metal J-tube for vaporization. The J-tube contained a column of 3/8" diameter stainless steel ball bearings to aid in the vaporization process. Air was used as a carrier in which the test substance was evaporated. The air was metered into the J-tube at a controlled rate of up to 50 lpm. The vapor exiting the J-tube was directed into the chamber inlet manifold for dilution in the chamber make-up air stream, which was then delivered into the exposure chamber.

Inhalation Chambers

Exposures were conducted in 450-liter stainless steel and glass Rochester-style inhalation chambers and stainless steel exposure caging (two layers of 10 compartments). Individual

animal cage position assignments within each chamber were rotated daily. Acclimation to the exposure system involved placing the animals into the exposure caging/chambers for 3 hours two days before the first exposure and for 6 hours the day before the first exposure. Chamber temperature, relative humidity, and airflow was maintained and recorded during the acclimation period similar to that anticipated for the exposure periods. Vapor generation systems were not in operation during acclimation. Daily test substance exposures were conducted such that chamber airflows were adjusted to maintain at least 10 exchanges of chamber volume per hour (minimum flow rate of 75 lpm). Chamber temperature was maintained within the range of 21.7 – 27.7°C. Chamber relative humidity was maintained within the range of 26.3 – 67.0%. Chamber airflow, temperature and relative humidity were monitored continuously and values were recorded at least once every 30 minutes. Chamber oxygen content was monitored once during the first day of exposure in each chamber to demonstrate that oxygen content was above 19% under applied experimental conditions.

The test atmosphere from each chamber was sampled by an automated sampling system. The system was designed such that test atmosphere was continuously pulled from the chamber and delivered to the analyzer through individual chamber sample lines and a stream selector valve. The sample lines were continuously purged with fresh test atmosphere during the entire exposure period.

Analysis of the test atmosphere was conducted by gas chromatography with flame ionization detection (GC/FID) and each chamber was evaluated at least once per hour during the exposure period. A standard curve, prepared by analyzing at least one bag standard from each of five calibration levels bracketing the expected range of exposure concentrations, was used to calculate the concentration of D3 in each chamber sample. Acceptance criteria for each standard curve required that the linear regression analysis correlation coefficient be at least 0.98 and that the peak area response for each bag standard used in the generation of the standard curve yield a calculated concentration within 10% of the nominal bag standard concentration. At least one bag standard was prepared and analyzed daily prior to the exposure period to verify the standard curve. The standard curve was considered valid if the actual concentration (calculated from the standard curve) was within 10% of the nominal bag standard concentration. A new standard curve was generated after the exposure period on study day 9.

For each day of exposure the mean test atmosphere D3 concentration (actual), as determined from the GC/FID analysis, was compared with the theoretical concentration (nominal), derived from the estimated chamber air flow rate and test substance consumption, as a quality control measure to evaluate exposure system performance.

Test atmosphere homogeneity was evaluated once prior to initiation of animal exposures. All chambers met acceptance criteria, which required that the mean values for each of the chamber zones be within 10% from the reference zone (approximate sampling location that was used during animal exposures). The stability of D3 vapor in air and sample line loss was evaluated using prepared bag standards. Method development activities that involved test running of the chambers to demonstrate vapor generation and chamber exposure system performance were conducted prior to initiating animal exposures. These activities were regarded as method development and the data generated are not presented. All data generated during these activities will be maintained in a method development section of the study file.

Administration of Doses

Test substance was administered by whole-body vapor inhalation. The exposure periods were defined as a 6 hrs/day exposure at target concentration sandwiched between two T_{99} periods (approximately 20 minutes each). Animals were removed from the chambers after the second T_{99} had expired. The T_{99} period was defined as the time required for the chamber atmosphere to reach 99% of the target concentration and was calculated based on chamber flow rate and size. For practical purposes, the $T_{99} \pm 2$ minutes was used on study.

The target exposure levels were: 0, 100, 500, and 2500 ppm D3.

XV. TEST SUBSTANCE ADMINISTRATION:

Treatment Regimen

Whole-body vapor inhalation exposure of rats to the test atmosphere was performed daily, seven days per week at approximately the same time each day. Treatment of both sexes began two weeks prior to initiating mating and continued through the mating period. Male animals were treated for 28 days and sacrificed on study day 29. Ten female animals per dose group were placed on study for the purpose of toxicological evaluation (toxicity group). These unmated females were treated for 29 days and sacrificed on study day 30. Daily exposure of reproductive group females continued up to and including day 19 of gestation. Dams were not exposed during the lactation period.

XVI. METHOD OF EUTHANASIA:

Scheduled Animal Death

Male and toxicity group females were euthanized by exsanguination (cutting of major abdominal vessels) following ketamine HCl / xylazine anesthesia. Reproductive group females were euthanized by CO_2 inhalation. Pups were deeply anesthetized by hypothermia and euthanized by decapitation.

Unscheduled Death

There were no unscheduled or premature deaths of adult animals.

XVII. MATING PROCEDURE:

A 1:1 mating ratio was used. After exposure on study day 14, the animals were paired by placing the lowest numbered ear tag reproductive group female within each group in the home cage of the male with the lowest numbered ear tag from the same group. Female animals were housed continuously, except during daily exposure, with the same male until evidence of copulation was obtained. Females were evaluated daily for evidence of copulation, as indicated by either a vaginal copulatory plug or sperm in the vaginal smear. Day 0 of gestation was defined as the day evidence of copulation was obtained, at which time the female was returned to her home cage. Females that delivered pups were euthanized on post-partum day four. One female in the reproductive group did not deliver and was euthanized on study day 44.

XVIII. TEST SYSTEM OBSERVATION:

Mortality/Morbidity

Animals were observed at least twice daily (once on weekends and holidays) in their cages for moribundity and mortality throughout the in-life phase of the study.

Clinical Observations

Daily Observations:

General clinical examinations were made at least once a day and were conducted immediately after exposure. The examinations included, but were not limited to changes in the skin, fur, eyes, mucous membranes; respiratory, circulatory, autonomic and central nervous systems functions; motor activity and behavior patterns. Findings were recorded for individual animals. The condition of animals without findings was documented in a general comment. General clinical examinations were not performed on days when detailed physical examinations were performed.

Detailed Physical Examinations:

All animals received a detailed physical examination once before the first exposure (to allow for within-subject comparisons), and weekly thereafter. Examinations were made outside the home cage in a standard arena at approximately the same time each day. Observations were detailed and carefully recorded. Examinations included, but were not limited to, changes in skin, fur, eyes, mucous membranes, occurrence of secretions and excretions, and autonomic activity. Changes in gait, posture and response to handling as well as the presence of clonic or tonic movement, stereotypies, difficult or prolonged parturition or bizarre behavior were recorded. The presence or absence of findings was recorded for individual animals.

XIX. PARAMETERS EVALUATED:

Body Weight

Individual animal body weights were recorded weekly beginning approximately one week prior to test substance administration, on the first day of dosing, and just prior to the scheduled necropsy. In addition to the above, reproductive group females were weighed on gestation days: 0, 7, 14, 20, within 24 hours after parturition, and day four postpartum.

Food Consumption

Individual food consumption was recorded weekly for toxicity group females. Individual food consumption for males and reproductive group females was recorded during the first two weeks of treatment and not during the period of cohabitation. Food consumption was recorded for the reproductive group females during gestation (weeks 1-3) and lactation periods (postpartum day 0 to postpartum day 4).

Functional Observational Battery

Functional observational battery and motor activity evaluations were performed on male and toxicity group female animals prior to the start of dosing (baseline evaluations) and during the fourth week of the exposure period. These assessments were conducted prior to the daily exposure period. Functional observational battery evaluations were performed under red light and motor activity evaluations were performed under reduced lighting (150 - 184 LUX) and white noise (60.1 - 64.0 db). A further description of the FOB components and the motor activity testing is provided below:

Cage Side Observations:

Unusual body movements, abnormal behavior and/or posture, and resistance to removal.

Hand-held Observations:

Palpebral closure, lacrimation, pupil size, pupil reactivity, salivation, muscle tone, extensor thrust response, and reactivity to handling.

Open Field Observations:

Level of ambulatory activity including rearing (one minute evaluation), responsiveness to sharp noise, touch, tail pinch, gait evaluation, and quantity of urine and fecal pellets voided.

Categorical Observations (made any time during the FOB):

Skin or hair coat, behavior, respiration, muscle movements, eyes, urine or feces, soiling, general abnormalities, and posture.

Measurements/Counts:

Rectal temperature, hind limb/forelimb grip strength, and landing foot splay.

Motor Activity

Motor activity was determined using a Cage Rack Activity System (San Diego Instruments, San Diego, CA). Each test session was 80 minutes in length.

Motor activity assessment involved obtaining baseline motor activity data prior to initiating exposures to assist in evaluating potential treatment-related effects by allowing each rat to act as its own control. The data collected reflects the movement of individual rats about the test cage and is characterized by the sum of events (beam breaks) per 8-minute interval (epoch) during the test session, total counts for the session (sum of the ten epochs), and habituation (decreasing activity with time in the test cage).

Clinical Pathology

Blood samples were collected from male and toxicity group females on the day of scheduled sacrifice. Food was removed from the cages of these animals the day prior to the scheduled sacrifice. After being anesthetized with Ketamine HCL/Xylazine, blood samples for hematological and clinical chemistry evaluations were collected from the abdominal vena cava

using a syringe and needle. Blood samples were then distributed to each of three sample collection tubes, which were either empty or contained sodium citrate or ethylenediaminetetraacetic acid (EDTA).

Hematology:

Blood samples were collected for the following evaluations: hematocrit, hemoglobin, erythrocyte count, total and differential leukocyte count, platelet count, and prothrombin time. Additionally, the erythrocyte indices: mean cell volume (MCV), mean cell hemoglobin, and mean cell hemoglobin concentration (MCHC) were assessed. These blood samples, with the exception of the blood samples collected for prothrombin time, were collected into test tubes containing EDTA and analyzed within six hours of collection. Whole blood samples were analyzed on the Baker 9000® Hematology analyzer (ABX Diagnostics, Irvine, CA).

White blood cell differentials and red blood cell morphology were determined from blood smears by manual counts and reported as a percentage based on a count of 100 cells.

Blood samples for prothrombin time (protime) were collected into test tubes containing sodium citrate, centrifuged within two hours of collection, and plasma collected for analysis. A semi-automated method using the BBL Fibrosystem (Becton Dickenson, Cockeysville, MD) and dried rabbit brain thromboplastin with calcium (Thromboplastin C Plus, Dade Behring, Newark, DE) was used to activate clotting. This method uses the detection of fibrin clot formation as the end point, with results reported in seconds.

Serum Chemistry:

Blood serum was collected for the following evaluations: total protein, albumin, glucose, cholesterol, total bilirubin, urea nitrogen, creatinine, alanine aminotransferase (ALT), aspartate aminotransferase (AST), alkaline phosphatase, calcium, phosphorus, sodium, potassium; and chloride. Serum was derived from blood samples collected into anticoagulant-free collection tubes, allowed to clot, and centrifuged. Serum samples were analyzed on the Monarch® 2000 Chemistry analyzer (Instrumentation Laboratory, Lexington, MA).

Morphologic Pathology

Macroscopic Examination:

Males and toxicity group females were subjected to a complete gross necropsy that included examination of the cranial, thoracic, and abdominal cavities and their contents. Reproductive group females were not subject to macroscopic examination at necropsy, however the number of corpora lutea and implantation sites was recorded. Pups found dead and pups sacrificed on day four postpartum were examined for external gross abnormalities only. No pup tissues were collected and the carcasses were discarded.

Organ Weights:

At necropsy, the following organs from males and toxicity group females were trimmed and weighed: adrenal glands, brain, heart, kidneys, liver, lungs, spleen and thymus. Testes, epididymides, seminal vesicles and prostate weights were recorded for all male adult animals. Ovary and uterine weights were recorded for toxicity group females.

Microscopic Examination:

The following tissues were collected from adult males and toxicity group females and preserved in 10% neutral buffered formalin: all gross lesions, adrenals, brain, heart, kidneys, liver, lymph nodes (mediastinal and mesenteric), lungs (infused with fixative as close to normal inspiratory volume as practical), nose (nares, nasal cavity and nasopharynx regions), spinal cord (cervical, thoracic, and lumbar), spleen, duodenum, jejunum, ileum, cecum, colon, stomach, peripheral nerve (sciatic), thymus, thyroid, trachea, uterus, urinary bladder, bone marrow (sternum), ovaries, prostate, and seminal vesicles. The testes and epididymides were preserved in Bouin's solution for 24 - 36 hours, and then transferred to 70% ethanol until processing. Tissues were trimmed and processed routinely, embedded in paraffin, sectioned at approximately five microns and stained with hematoxylin and eosin for histological examination. The nose was decalcified prior to sectioning. All remaining tissues were placed in 10% neutral buffered formalin for long-term storage.

All collected tissues were examined by light microscopy for high exposure group and control males and toxicity group females. Target tissues examined from the 100 and 500 ppm groups included liver from both sexes and kidneys and seminal vesicles from males. Histopathology findings were recorded using PathData® System v.6.1a1.

Reproductive Parameters

The duration of gestation was calculated for each female. Beginning on gestation day 20, pregnant animals were checked at least three times daily (twice daily on weekends and holidays) for evidence of parturition. Each litter was examined following delivery to determine the number and sex of the pups, stillbirths, live births, runts, sex ratio, and the presence of any gross abnormalities. Litter weights were taken within 24 hours of parturition and on day four postpartum. The day parturition began was considered postpartum day 0.

XX. STATISTICAL ANALYSIS:

All data analysis, with exception to histomorphologic findings, was carried out using SAS®(V.8.2). In all comparisons, the family wise error rate was held at 5% ($\alpha = 0.05$). Bartlett's test was used to check for heterogeneity of variances and a Kolmogorov-Smirnov test was used to test normality of the data. Parametric data was then tested using one-way Analysis of Variance (ANOVA) followed by Dunnett's Test (if significant); nonparametric data was tested by Kruskal Wallis Test followed by Wilcoxon (if significant) (Gad and Chengelis, 1998). For variables that had multiple measurements across time repeated-measures ANOVA was used to analyze the data. Categorical data was tested for equal proportions using the Fisher's Exact Test (Gad and Chengelis, 1998).

Fisher's exact test was used to determine if differences in the incidence of certain histomorphologic findings were statistically significant. The analysis was conducted within the PathData® System (v.6.1a1).

Statistically significant probabilities are reported at either the $p \leq 0.05$ or $p \leq 0.01$ levels.

Parameters Evaluated

The parameters evaluated for statistical significance were: body weight, body weight gain, food consumption, hematology and clinical chemistry values, organ weight, organ/body weight ratios, gestation length, litter size, pup viability, litter weight, live births/litter size, uterine implantation sites, number of corpora lutea, FOB and Motor Activity parameters, and histomorphology findings.

Data Analysis

Numerical data obtained during the conduct of the study was processed using Microsoft Excel™. Statistical analysis of the data was carried out using SAS®, V.8.2, (Appendix S) or the PathData® System (v.6.1a1).

XXI. RESULTS:

Exposure

Exposure data are summarized in Tables 1 - 2. Individual chamber concentrations and conditions can be found in Appendices A-B.

With few exceptions (see protocol deviations section), test groups were exposed to target vapor concentrations for 6 hrs/day for 28-39 days as required by group/subgroup designations. The overall mean daily exposure concentrations for male rats were below the limit of quantitation (BLQ), 99, 496, and 2505 ppm D3 for target exposure concentrations of 0, 100, 500 and 2500 ppm D3, respectively. Overall mean daily exposure concentrations for toxicity group females were BLQ, 99, 515, and 2499 ppm D3 for target exposure concentrations of 0, 100, 500 and 2500 ppm D3, respectively. Overall mean daily exposure concentrations for reproductive group females were BLQ, 99, 512, and 2492 ppm D3 for target exposure concentrations of 0, 100, 500, and 2500 ppm D3, respectively.

Mortality and Morbidity

All adult animals survived to their scheduled necropsy. At the scheduled necropsy, a 2500 ppm group male (D0072) was anesthetized by CO_2 inhalation following its poor response to anesthesia with ketamine HCl / xylazine.

During the parturition process and postnatal period pups found dead were removed from their cage and those found moribund or severely injured were euthanized.

Clinical Observations

The incidence of clinical observations is summarized in Tables 3 - 5. Individual clinical examination data are presented in Appendix C. Observations recorded during the weekly detailed physical examinations were not different from those recorded during the daily clinical examinations. Therefore, the findings were combined and are presented in total.

Clinical observations indicative of an adverse effect of treatment were minimal and involved an increased incidence of anogenital staining and probable porphyrin staining (head). Anogenital

staining was observed in 30% of males, 40% of toxicity group females and 100% of reproductive group females in the 2500 ppm groups. A single occurrence was observed in one reproductive group female in the 100 ppm group. Probable porphyrin staining (head) was not observed in male and female control groups and was noted once in two male rats, one each from the 100 and 2500 ppm exposure groups. In contrast, the incidence of probable porphyrin staining (head) in female rats was increased for females in the 2500 ppm group. Ninety percent of the toxicity group females and 100% of the reproductive group females in the 2500 ppm groups exhibited this finding at least once during the in-life phase of the study.

All other findings were typical for rats of this age, strain and husbandry and/or did not occur in a dose-related fashion.

Body Weight

Summaries of weekly mean body weights are shown in Tables 6 - 8. Mean body weight gain summaries are shown in Tables 9 - 11. Individual animal body weight data are given in Appendix D. Individual animal body weight gain data can be found in Appendix E.

Group mean body weights for 100, 500, and 2500 ppm D3-exposed males were not statistically different from the control group at any time; however, group mean body weight for males in the 2500 ppm exposure group was consistently lower than any other group after day 1. The difference in mean body weight between the control and 2500 ppm exposure group grew larger with time and resulted in a decrease of 6% relative to control on day 29. Similarly, group mean body weight gain was consistently lower for males in the 2500 ppm exposure group relative to controls for each time period. The decreases were statistically significant at week 1 (35% decrease) and for total weight gain (39% decrease).

Group mean body weights for 100, 500 and 2500 ppm D3-exposed toxicity group females were consistently lower than the control group. However, the decreases (approximately 6% relative to control) reached statistical significance only for the 2500 ppm exposure group on day 8 and 30. Group mean body weight gain was not statistically different in D3-exposed females relative to control. However, the mean value for total body weight gain was 25% lower for the 2500 ppm exposure group relative to control. These data suggest that exposure to 2500 ppm D3 results in a slight decrease in body weight/body weight gain.

Group mean body weight for 500 and 2500 ppm D3-exposed reproductive group females were consistently lower ($\leq$ 8%) than the control group. However, the decreases were not statistically significant except for the 2500 ppm exposure group on gestation day 20 (8% lower relative to control). Similarly, group mean body weight gain for D3–exposed reproductive group females in the 500 and 2500 ppm exposure groups were typically lower ($\leq$ 30%), though not statistically significant, than controls prior to week 3 of gestation. Females in the 2500 ppm exposure group demonstrated a statistically significant 21% decrease in body weight gain at gestation week 3. These changes in body weight and body weight gain are consistent with those observed for the toxicity group females (pre-mating) and with the effects on litter size (gestation and post-partum periods).

Food Consumption

Weekly mean food consumption data are presented in Tables 12 - 14. Individual animal food consumption values are given in Appendix F.

Mean weekly food consumption values were generally lower than control for male and toxicity group females exposed to D3. Statistically significant decreases (<15% relative to control values) were demonstrated for 2500 ppm group males (weeks 1 and 2), 2500 ppm group females (weeks 1, 2, and 3), and 500 ppm group females (weeks 2 and 3). There were no statistically significant differences in mean weekly food consumption values for D3-exposed reproductive group females relative to control. However, values were generally lower than control (2-12%). The one exception being a greater consumption of food (21%) during the parturition period (gestation day 20 –postpartum day 0/1) that likely represents differences in parturition (litter size and care).

Functional Observation Battery

Summaries of FOB data are presented in Tables 15 -18 and individual animal data are presented in Appendix G. The summary data for rectal temperature, hindlimb grip strength, forelimb grip strength or landing foot splay assessment are presented in Tables 19 -22 and individual animal data are provided in Appendix H.

1. Cage Side Observations: There were no treatment-related changes noted in unusual body movements, abnormal behavior, posture or resistance to removal.

2. Handling Observations: Palpebral closure, lacrimation, pupil size, and reactivity, salivation, muscle tone, and extensor thrust response were unaffected by treatment. There was a statistically significant change in the "reactivity to handling" categorical observation for 2500 ppm toxicity group females. This change reflects a greater proportion of rats in this exposure group exhibiting "minimal struggling" as compared to "moderate struggling with little or no vocalization" for the control group.

3. Open Field Observations: No differences were apparent between the control and treated groups in open field observations (level of ambulatory activity including rearing, responsiveness to sharp noise, touch, tail pinch, gait evaluation, quantity of urine and fecal pellets voided).

4. Categorical Observations: No differences were apparent between the control and treated groups for skin or hair coat, respiration, muscle movements, eyes, urine or feces, soiling, posture or general abnormalities.

5. Measurements/Counts: There were no treatment-related changes noted for rectal temperature, hindlimb grip strength, forelimb grip strength or landing foot splay.

Motor Activity

Motor activity data are presented in Tables 23 - 29 and individual animal data are presented in Appendix I.

Habituation, the characteristic of decreasing motor activity with time in the observation cage, was analyzed with the data from males and females combined and with the sexes separate. When combined there was no statistical difference between D3-exposed and control rats. When analyzed separately, there was no statistical difference between D3-exposed female rats and the controls. However, analysis of the data for males showed a statistically significant difference in habituation in the 500 ppm exposure group relative to control. Because this effect was not dose-responsive (i.e. only present in the 500 ppm exposure group and not in the 100 and 2500 ppm exposure groups), was not present in D3-exposed females, and was not detected in the combined data set, it was not considered treatment-related.

The effect of D3-exposure on motor activity (i.e. total number of locomotor counts per session) was evaluated statistically as a sexes-combined data set and then again with the sexes separated. The combined evaluation demonstrated reduced motor activity at the 2500 ppm exposure level relative to control. When analyzed separately, a statistically significant decrease in motor activity was demonstrated for 2500 ppm exposure group males and not females. This effect on motor activity is presented in Table 23. As shown, motor activity for males exposed to 2500 ppm D3 was markedly less than that for the other exposure groups. Similarly the total motor activity counts for 2500 ppm D3-exposed females were lower than that of the other exposure groups however the difference (baseline verses treated) was not as great as that for the males.

Clinical Pathology

Hematology:

Summaries of hematology parameters can be found in Tables 30-34 and Appendices J-L.

Hematology measures for D3-exposed animals were similar to controls. Differences were slight and the mean values for each group were within published ranges for animals of this strain, age and sex (Moore, 2000). Statistically significant differences were demonstrated for hemoglobin concentration in males (5% decrease at 2500 ppm) and females (4% increase at 100 ppm) and for platelet content in males (17% increase at 2500 ppm). In males, the modest elevation of platelets may represent a secondary thrombocytosis in response to the slight decrease in hemoglobin (Smith et al., 2002). The slight decrease in hemoglobin at 2500 ppm D3 for males may be related to the observed decrease in food consumption, decrease in body weight and/or effects on the kidney. The slight increase in hemoglobin for females in the 100 ppm exposure group is not considered treatment-related.

Clinical Chemistry:

Clinical chemistry data are presented in Tables 35 and 36, and Appendix M.

Clinical chemistry measures for D3-exposed rats were generally similar to those for control groups. Exceptions include a statistically significant decrease in alkaline phosphatase activity, decreased sodium concentration, and an increase in urea nitrogen in the 2500 ppm group males. Statistically significant changes in D3-exposed females included decreased calcium and glucose concentrations (500 and 2500 ppm females) and increased cholesterol (2500 ppm females).

In males, the 28% increase in urea nitrogen and 2% decrease in sodium in the 2500 ppm groups correlate with the observed kidney histopathology (see below). The 27% decrease in alkaline phosphatase activity observed for males in the 2500 ppm group is consistent with a reduction in food consumption; decreases in this parameter are not indicative of organ toxicity.

In females, the 3-4% decrease in calcium in the 500 and 2500 ppm exposure groups was not considered treatment-related based on the magnitude of the changes and that the values were within the normal range for this strain and age of rat (Levine, 1995). The 18-19% decrease in serum glucose levels for females in the 500 and 2500 ppm exposure groups represent moderate changes that may be related to the general health condition of the animals (decreased food consumption, slight decrease in body weight). Similarly the 30% increase in blood cholesterol may be related to the D3-induced liver effects.

Morphological Pathology

Macroscopic Examination:

Macroscopic pathology data are presented in Table 37 and individual animal data is presented in Appendix N.

There were no treatment-related macroscopic findings.

Organ Weights:

Organ weight and organ to body weight ratios are summarized in Tables 38 - 41 and individual animal data are presented in Appendices O-P.

Statistically significant increases in group mean liver weight and liver-to-body weight ratio were observed for males and toxicity group females in the 2500 ppm groups. Liver weight (liver-to-body weight ratio) was increased 22% (31%) for males and 32% (41%) for females. A statistically significant increase (6% relative to control) in liver-to-body weight ratio was observed for toxicity group females in the 500 ppm exposure group. However, because absolute organ weight was not affected the increase in relative organ weight change has been attributed to decreased body weight. The liver weight increases at 2500 ppm for males and toxicity group females are consistent with the liver hypertrophy identified by microscopic examination.

A statistically significant increase (13%) in kidney-to-body weight ratio was demonstrated for male rats at the 2500 ppm exposure level. Absolute kidney weight for this group was increased 6% relative to controls however this change was not statistically significant. The noted increase in serum urea nitrogen and histomorphological findings are consistent with the slight increase in kidney weight. The differential between absolute and relative kidney weight is attributed to decreased body weight.

Statistically identified decreases in the absolute weight of epididymides (10%), and seminal vesicles (30%) were observed for males in the 2500 ppm exposure level. When expressed relative to body weight only the decrease in seminal vesicle weight (27%) remained statistically significant. The decrease in seminal vesicle weight is consistent with the histomorphologic findings noted below.

In addition to the noted liver weight increase for D3-exposed toxicity group females, there were statistical increases in the relative (but not absolute) weights of the adrenal glands (16%) in the 2500 ppm group, and of the lungs (8%) in the 500 and 2500 ppm exposure groups. The adrenal gland effect may be a non-specific effect of stress at the highest exposure level. The modest increases in relative lung weight may be an effect of exposure; however, no histomorphologic correlate was observed, thus the finding is of minimal toxicologic significance. There was also a statistically identified decrease in heart weight (9% at 500 ppm) and adrenal weight (13% at 100 ppm) on an absolute organ weight basis but not when expressed relative to body weight; these findings are not considered treatment-related.

Microscopic Examination:

Histopathology findings are summarized in Tables 42 and 43. The individual animal reports and individual histopathology findings are given in Appendices N and Q.

Histomorphologic effects attributable to treatment were observed in the liver of rats of both sexes, and in the kidneys and seminal vesicles of males. Other findings were of an incidence and character common to rats of this age and stock and not attributable to treatment.

In the liver there was centrilobular hepatocellular hypertrophy in all rats of both sexes exposed to 2500 ppm D3. The change was uniformly graded as very slight in males and slight in females. The liver weight increases at the highest exposure in both sexes correlate with this finding. This is considered an adaptive change related to induction of cellular processes for increased xenobiotic metabolism/excretion. Treatment-related histomorphologic effects on the liver were not detected at lower exposure levels in either sex.

In the kidney, changes characterized as protein droplet nephropathy (PDN) were observed in male rats only. The severity and incidence were exposure related; all males exposed to 2500 ppm had slight to moderate PDN and there was a modest increase in the incidence and severity of tubular degeneration/regeneration (observed as small foci of basophilic tubules). At the 500 ppm D3 exposure level, nine of the 10 rats had very slight PDN with no increase in basophilic tubules. At 100 ppm D3, one of the 10 rats had very slight PDN changes, a finding that could be due to biologic variation since protein droplets are normally visible in the kidneys of most control male rats. PDN was characterized by an increase in the portion of renal cortical tubules (P2 segment) with visible eosinophilic droplets, often with angular droplet shapes suggesting crystallized material. With this increase in droplets there were increases in renal tubular cell necrosis, as indicated by nuclear pyknosis and karyorrhexis. Grade one, very slight, indicated a subtle increase in these findings over the background change in control males. Grades two

and three, slight and moderate, were associated with several fold increases in the portion of tubules with visible droplets, angular droplet shapes, and cortical tubular cell degeneration. These changes correlated with a modest increase (6%, not statistically significant) in kidney weight as well as a 28% increase in serum urea nitrogen and a very small decrease in serum sodium in the high exposure group males. These histomorphologic and clinical chemistry kidney findings are consistent with a nephropathy caused by α_{2U}-globulin retention and accumulation in the renal cortex, a finding not associated with human risk (USEPA, 1991).

The seminal vesicles were slightly atrophied in four of 10 males in the 2500 ppm exposure group. The change was characterized by decreased fluid in the gland lumen and decreased cell height and epithelial folding of the luminal epithelium. This finding correlated well with the noted decrease in organ weight.

Reproductive Parameters

The reproductive parameters are presented in Tables 44 and 45 and individual animal data are presented in Appendix R.

The mating phase began with pairing of reproductive group females with males after the exposure period on study day 14. All of the reproductive group females presented with positive evidence of mating on or before the seventh day of pairing and all but one of the reproductive group females became pregnant. The exception was a female from the 2500 ppm D3 exposure group (Animal D0033). The uterus from this female was negative for implantation sites.

Exposure to D3 had no affect on gestation length, pup sex ratio, pup weight, pup viability, and corpora lutea counts (PND 4). Statistically significant decreases were noted for litter size(number of pups per litter on PND 0 (33% decrease relative to control) and 4 (41% decrease relative to control)), litter weight (27% decrease relative to control on PND 0 and 4), and number of uterine implantation sites (33% decrease relative to control) in the 2500 ppm exposure group only.

Decreased litter size is considered responsible for lower mean body weight, body weight gain, and food consumption observed during gestation for this exposure group.

One reproductive group female (Animal D0035) in the 2500 ppm exposure group was observed squealing, shaking and with "bulging" eyes for approximately 10 minutes during the exposure period on study day 31. This same rat was observed to have an abnormal posture during the exposure period on study day 33. Physical and/or behavioral abnormalities were not observed at the end of the exposure period during return to the housing caging. No other rats in the exposure chamber exhibited these findings. This rat had a normal gestation period (22 days) and gave birth to 14 pups, none of which survived to PND 4. The loss of this litter represents the only complete litter loss for this exposure group. The only other rat on study to experience a complete litter loss was from the 100 ppm exposure group. The significance of this relationship, abnormal behaviors observed during exposure and litter loss, is not known.

XXII. CONCLUSIONS:

Whole-body inhalation exposure of male and female Sprague-Dawley rats to D3 vapor at concentrations of up to 2500 ppm for 28-29 consecutive days was generally well tolerated as suggested by minimal effects on body weight and food consumption, the lack of significant clinical signs of toxicity, and minimal effects on clinical pathology parameters.

Increased liver weight and centrilobular hepatocellular hypertrophy were observed in both sexes exposed to 2500 ppm D3. These effects are considered to be an adaptive response characteristic of this class of cyclic siloxanes (McKim *et al.* 1998, 1999; Burns-Naas *et al.,* 1998; Klykken *et al.,* 1999). No liver effects were identified at lower exposure levels.

Protein droplet nephropathy was observed by histopathology in the kidneys of males exposed to 500 or 2500 ppm D3. One male rat exposed to 100 ppm D3 was identified with a very slight lesion of the same nature. For males in the 2500 ppm exposure group, the finding was corroborated by organ weight and clinical chemistry changes. Females were unaffected. This nephropathy, affecting only male rats, is consistent in incidence and histomorphology with that caused by excessive accumulation of α_{2U}-globulin in renal cortical tubules. This mechanism is specific to male rats and not associated with human risk.

In males exposed to 2500 ppm D3, there were statistically significant decreases in the absolute weight of epididymides (10%) and in the absolute (30%) and relative (27%) weight of the seminal vesicles. Slight atrophy of the seminal vesicles was identified in four of the 10 males in the 2500 ppm exposure group. The epididymides, testes, and prostate from D3-exposed rats were histologically normal.

Negative effects on litter size parameters were observed for female rats exposed to 2500 ppm D3. There was a 33% decrease in litter size and in the number of implantation sites. Gestation length, pup sex ratio, pup weight and viability, and corpora lutea counts were unaffected. No effects were observed at lower exposure concentrations.

At the 2500 ppm exposure level there was reduced motor activity in both sexes, and reduced "reaction to handling" in females. There was no histopathological evidence of neurotoxicity.

Within the context of the 28-day sub-acute toxicity portion of this study no adverse effects attributable to treatment were noted for female rats. Adverse effects attributed to treatment were noted for males and included an increased incidence of protein droplet nephropathy (100 ppm "no observable adverse effect level" (NOAEL)) and decreased seminal vesicle weight (500 ppm NOAEL). Within the reproductive/developmental toxicity screening test portion of this study a NOAEL of 500 ppm was identified for effects on reproductive parameters (decreased litter size and number of implantation sites).

XXIII. ARCHIVE:

The protocol, amendments, deviations, all raw data, documentation, pathology tissue specimens, sample of the test substance, and the final report are retained by EPL Archives, Inc., 45610 Terminal Drive, Sterling, Virginia, 20166, as required by GLP regulations. Serum and plasma samples were discarded after wet specimen verification.

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XXV. DEVIATIONS FROM THE PROTOCOL:

Deviations from the protocol that occurred during the course of the study are summarized in Table 46. None of these deviations are believed to have impacted the outcome or validity of the study.

Table 1 - Exposure Concentration Summary

		Mean Daily Exposure	Std Dev
Males	**Controls**	BLQ*	1.4
	100 ppm	99	3.0
	500 ppm	496	30.8
	2500 ppm	2505	85.9
Tox Females	**Controls**	BLQ	1.4
	100 ppm	99	2.5
	500 ppm	515	14.9
	2500 ppm	2499	121.1
Repro Females	**Controls**		
	M1	BLQ	1.3
	M2	BLQ	1.3
	M3	BLQ	1.3
	M4	BLQ	1.2
	M6	BLQ	1.2
Repro Females	**100 ppm**		
	M1	99	2.3
	M2	99	2.3
	M3	99	2.3
	M4	99	2.3
Repro Females	**500 ppm**		
	M1	512	15.8
	M2	512	15.6
	M3	511	15.5
	M4	511	15.8
Repro Females	**2500 ppm**		
	M1	2493	113.2
	M2	2492	111.6
	M3	2492	110.0
	M4	2492	108.5

* Below Limit of Quantitation (0.01 - 3 ppm)
M1 = Animals that showed positive evidence of mating on Day 1 of mating period
M2 = Animals that showed positive evidence of mating on Day 2 of mating period
M3 = Animals that showed positive evidence of mating on Day 3 of mating period
M4 = Animals that showed positive evidence of mating on Day 4 of mating period
M6 = Animals that showed positive evidence of mating on Day 6 of mating period

Table 2 - Chamber Environmental Conditions Summary

Males		Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Dose Group	Exposure Period	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
Control	28 days	25.4	0.41	26.8	22.2	45.8	3.04	62.8	33.7	106	1.7	114	100
100 ppm	28 days	25.3	0.52	27.3	22.1	42.4	3.49	63.6	30.8	106	5.3	117	95
500 ppm	28 days	25.8	0.52	27.1	22.5	43.5	2.81	66.3	32.5	104	2.5	109	98
2500 ppm	28 days	25.8	0.53	27.7	22.2	43.5	2.85	64.1	33.9	106	1.9	112	99
Toxicity Group Females													
Control	29 days	25.8	0.25	26.9	22.0	42.7	3.67	62.1	32.3	106	1.4	110	102
100 ppm	29 days	25.2	0.36	26.3	22.1	48.6	3.44	67.0	37.4	103	2.5	113	96
500 ppm	29 days	26.2	0.18	27.2	21.9	39.3	3.71	60.5	26.3	106	2.0	113	98
2500 ppm	29 days	25.7	0.39	27.5	21.7	44.1	3.51	62.6	33.6	107	1.9	114	97

Table 2 - continued

Dose Group	Exposure Period	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
		Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
Control													
Subgroup M1	34 days	25.7	0.39	26.9	22.0	43.9	4.57	62.1	32.3	106	1.4	110	102
Subgroup M2	35 days	25.7	0.40	26.9	22.0	44.1	4.66	62.1	32.3	106	1.4	110	102
Subgroup M3	36 days	25.6	0.43	26.9	22.0	44.2	4.60	62.1	32.3	106	1.4	110	102
Subgroup M4	37 days	25.6	0.48	26.9	22.0	44.3	4.58	62.1	32.3	106	1.4	110	102
Subgroup M6	39 days	25.5	0.60	26.9	22.0	44.2	4.48	62.1	31.7	106	1.4	110	102
100 ppm													
Subgroup M1	34 days	25.2	0.38	26.6	22.1	48.5	3.28	67.0	37.4	103	2.8	113	96
Subgroup M2	35 days	25.2	0.39	26.6	22.1	48.4	3.23	67.0	37.4	103	2.8	113	96
Subgroup M3	36 days	25.2	0.40	26.6	22.1	48.3	3.36	67.0	29.8	103	2.8	113	96
Subgroup M4	37 days	25.1	0.42	26.6	22.1	48.2	3.38	67.0	29.8	103	2.8	113	96
500 ppm													
Subgroup M1	34 days	26.2	0.26	27.2	21.9	40.3	4.39	60.5	26.3	105	3.2	113	98
Subgroup M2	35 days	26.1	0.30	27.2	21.9	40.5	4.42	60.5	26.3	105	3.3	113	98
Subgroup M3	36 days	26.1	0.32	27.2	21.9	40.5	4.35	60.5	26.3	105	3.4	113	98
Subgroup M4	37 days	26.1	0.39	27.2	21.9	40.6	4.34	60.5	26.3	105	3.4	113	98
2500 ppm													
Subgroup M1	34 days	25.7	0.40	27.5	21.7	44.7	3.71	62.6	33.6	107	2.7	114	97
Subgroup M2	35 days	25.7	0.39	27.5	21.7	44.7	3.66	62.6	33.6	106	2.7	114	97
Subgroup M3	36 days	25.7	0.41	27.5	21.7	44.6	3.68	62.6	29.4	106	2.7	114	97
Subgroup M4	37 days	25.7	0.45	27.5	21.7	44.6	3.63	62.6	29.4	106	2.8	114	97

M1 = Animals that showed positive evidence of mating on Day 1 of mating period
M2 = Animals that showed positive evidence of mating on Day 2 of mating period
M3 = Animals that showed positive evidence of mating on Day 3 of mating period
M4 = Animals that showed positive evidence of mating on Day 4 of mating period
M6 = Animals that showed positive evidence of mating on Day 6 of mating period

Table 3 - Combined Clinical Observations and Detailed Physical Examination Summary: Male Rats

Total Occurrences/ No. of Animals

	Group 1 Controls	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Normal				
- No significant clinical observations	300/10	299/10	300/10	269/10
Disposition				
Found Dead	0/0	0/0	0/0	0/0
Moribund Sacrifice	0/0	0/0	0/0	0/0
Terminal Necropsy	10/10	10/10	10/10	10/10
Body/Integument				
Anogenital Staining	0/0	0/0	0/0	30/3
Probable Porphyrin Staining, head	0/0	1/1	0/0	1/1
Salivation	0/0	1/1	0/0	0/0

Table 4 - Combined Clinical Observations and Detailed Physical Examination Summary: Toxicity Group Female Rats

Total Occurrences/ No. of Animals

	Group 1 Controls	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Normal				
- No significant clinical observations	310/10	308/10	303/10	234/10
Disposition				
Found Dead	0/0	0/0	0/0	0/0
Moribund Sacrifice	0/0	0/0	0/0	0/0
Terminal Necropsy	10/10	10/10	10/10	10/10
Body/Integument				
Bloody Knuckle	0/0	1/1	0/0	0/0
Anogenital Staining	0/0	0/0	0/0	23/4
Probable Porphyrin Staining, head	0/0	1/1	6/2	63/9*
Toenail Bleeding	0/0	0/0	1/1	0/0

*$p \leq 0.01$ by Fisher's Exact Test

Table 5 - Combined Clinical Observations and Detailed Physical Examination Summary: Reproductive Group Female Rats

	Total Occurrences/ No. of Animals			
	Group 1 Controls	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Normal				
- No significant clinical observations	412/10	423/10	414/10	316/10
Disposition				
Found Dead	0/0	0/0	0/0	0/0
Moribund Sacrifice	0/0	0/0	0/0	0/0
Terminal Necropsy	10/10	10/10	10/10	10/10
Body/Integument				
Anogenital Staining	0/0	1/1	0/0	40/10*
Bleeding, ear	1/1	0/0	0/0	0/0
Generalized Hairloss, Anogenital Region	17/1	0/0	0/0	0/0
Toenail Bleeding	2/1	0/0	0/0	1/1
Probable Porphyrin Staining, head	0/0	0/0	14/3	89/10*
Salivation	0/0	0/0	0/0	2/1

*$p \leq 0.01$ by Fisher's Exact Test

Table 6 - Summary of Weekly Mean Body Weights (grams): Male Rats

	Day 1 25-Mar	Day 8 1-Apr	Day 15 8-Apr	Day 22 15-Apr	Day 29 22-Apr
Group 1 - Controls					
Mean	288.0	321.6	340.7	372.2	361.1
Standard Deviation	17.9	22.9	27.6	32.3	35.9
N	10	10	10	10	10
Group 2 - 100 ppm					
Mean	293.2	325.0	341.4	373.8	366.7
Standard Deviation	11.2	19.4	27.6	29.4	28.7
N	10	10	10	10	10
Group 3 - 500 ppm					
Mean	291.7	327.8	345.7	377.0	366.9
Standard Deviation	10.3	10.2	12.2	10.1	12.5
N	10	10	10	10	10
Group 4 - 2500 ppm					
Mean	293.3	315.1	329.0	352.2	337.7
Standard Deviation	12.4	16.0	23.7	26.3	28.2
N	10	10	10	10	10

Table 7 - Summary of Weekly Mean Body Weights (grams): Toxicity Group Female Rats

	Day 1 25-Mar	Day 8 1-Apr	Day 15 8-Apr	Day 22 15-Apr	Day 30 23-Apr
Group 1 - Controls					
Mean	214.5	234.5	248.3	256.8	241.9
Standard Deviation	6.2	7.3	6.3	10.4	8.2
N	10	10	10	10	10
Group 2 - 100 ppm					
Mean	212.4	231.5	243.8	252.5	238.0
Standard Deviation	9.5	9.1	9.6	9.1	10.5
N	10	10	10	10	10
Group 3 - 500 ppm					
Mean	210.2	227.6	238.6	243.9	229.9
Standard Deviation	7.2	11.8	14.8	15.5	13.0
N	10	10	10	10	10
Group 4 - 2500 ppm					
Mean	206.4	221.4*	235.4	244.5	227.0*
Standard Deviation	9.1	11.1	19.2	14.7	12.5
N	10	10	10	10	10

*$p \leq 0.05$ by Dunnett's Test

Table 8 - Summary of Weekly Mean Body Weights (grams): Reproductive Group Female Rats

	Day 1 25-Mar	Day 8 1-Apr	Day 15 8-Apr	Gestation Day 0	Gestation Day 7	Gestation Day 14	Gestation Day 20	Post Partum Day 0/1	Post Partum Day 4
Group 1 - Controls									
Mean	213.6	229.6	245.0	243.7	273.9	309.7	376.9	285.7	304.5
Standard Deviation	11.6	14.2	17.6	17.3	17.1	22.2	22.1	17.2	17.9
N	10	10	10	10	10	10	10	10	10
Group 2 - 100 ppm									
Mean	214.6	232.5	248.2	247.1	278.0	317.0	391.8	297.6	309.3
Standard Deviation	11.6	15.4	15.5	14.8	20.4	21.8	19.1	21.1	23.2
N	10	10	10	10	10	10	10	10	10
Group 3 - 500 ppm									
Mean	212.4	228.1	238.9	239.2	264.8	297.8	371.3	280.0	300.6
Standard Deviation	13.2	15.8	20.1	20.2	19.7	20.4	28.3	19.3	21.7
N	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm									
Mean	211.3	223.8	237.9	228.9	258.7	291.8	345.0*	280.4	290.4
Standard Deviation	7.4	9.3	11.0	16.0	10.7	14.5	21.6	17.0	19.5
N	10	10	10	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]

*$p \leq 0.05$ by Dunnett's Test

[a] Animal D0033 was not pregnant, values not used in calculations

Table 9 - Summary of Weekly Mean Body Weight Gain (grams): Male Rats

	Week 1	Week 2	Week 3	Week 4	Terminal Sac minus Day 1
Group 1 - Controls					
Mean	33.6	19.1	31.5	-11.1	73.1
Standard Deviation	7.9	7.1	10.2	11.8	21.4
N	10	10	10	10	10
Group 2 - 100 ppm					
Mean	31.8	16.4	32.4	-7.2	73.5
Standard Deviation	10.6	11.6	4.8	4.7	20.1
N	10	10	10	10	10
Group 3 - 500 ppm					
Mean	36.1	18.0	31.2	-10.1	75.2
Standard Deviation	5.7	10.9	8.6	5.5	7.2
N	10	10	10	10	10
Group 4 - 2500 ppm					
Mean	21.8*	14.0	23.2	-14.5	44.4*
Standard Deviation	6.2	9.0	5.5	5.5	17.4
N	10	10	10	10	10

* $p \leq 0.05$ by Dunnett's Test

Table 10 - Summary of Weekly Mean Body Weight Gain (grams): Toxicity Group Female Rats

	Week 1	Week 2	Week 3	Week 4	Terminal Sac minus Day 1
Group 1 - Controls					
Mean	20.0	13.8	8.5	-14.9	27.4
Standard Deviation	6.4	6.6	7.8	4.7	6.7
N	10	10	10	10	10
Group 2 - 100 ppm					
Mean	19.1	12.3	8.7	-14.6	25.6
Standard Deviation	3.1	6.1	6.7	4.5	4.7
N	10	10	10	10	10
Group 3 - 500 ppm					
Mean	17.4	11.0	5.3	-14.0	19.7
Standard Deviation	7.3	5.8	4.9	3.6	8.6
N	10	10	10	10	10
Group 4 - 2500 ppm					
Mean	14.9	14.0	9.2	-17.5	20.6
Standard Deviation	6.7	10.4	7.8	5.6	8.3
N	10	10	10	10	10

Table 11 - Summary of Weekly Mean Body Weight Gain (grams): Reproductive Group Female Rats

	Week 1 Pre Mating	Week 2 Pre Mating	Gestation Week 1	Gestation Week 2	Gestation Week 3	4 days Post Partum	Overall Weight Gain
Group 1 - Controls							
Mean	16.0	15.5	30.1	35.8	67.2	18.8	90.9
Standard Deviation	5.8	5.2	5.8	7.1	9.1	6.0	9.0
N	10	10	10	10	10	10	10
Group 2 - 100 ppm							
Mean	17.9	15.7	31.0	38.9	74.8	11.7	94.7
Standard Deviation	4.8	7.9	11.5	4.5	9.1	19.2	14.4
N	10	10	10	10	10	10	10
Group 3 - 500 ppm							
Mean	15.7	10.8	25.6	33.1	73.5	20.6	88.2
Standard Deviation	4.9	6.9	9.8	6.3	15.6	12.7	15.7
N	10	10	10	10	10	10	10
Group 4 - 2500 ppm							
Mean	12.5	14.1	29.8	33.1	53.2*	10.0	79.1
Standard Deviation	5.4	6.7	12.0	8.5	13.8	12.8	15.7
N	10	10	9[a]	9[a]	9[a]	9[a]	9[a]

*$p \leq 0.05$ by Dunnett's Test

[a] Animal D0033 not pregnant, values not used in calculations.

Table 12 - Summary of Weekly Mean Food Consumption (grams/animal/day): Male Rats

	Week 1	Week 2
Group 1 - Controls		
Mean	24.7	25.0
Standard Deviation	2.5	2.3
N	10	10
Group 2 - 100 ppm		
Mean	24.1	24.6
Standard Deviation	2.5	2.7
N	10	10
Group 3 - 500 ppm		
Mean	24.8	25.2
Standard Deviation	1.5	1.3
N	10	10
Group 4 - 2500 ppm		
Mean	22.2*	21.9*
Standard Deviation	2.4	2.2
N	10	10

*$p \leq 0.05$ by Dunnett's Test

Table 13 - Summary of Weekly Mean Food Consumption (grams/animal/day): Toxicity Group Female Rats

	Week 1	Week 2	Week 3	Week 4
Group 1 - Controls				
Mean	20.2	20.6	21.1	20.2
Standard Deviation	1.4	1.7	1.8	1.4
N	10	10	10	10
Group 2 - 100 ppm				
Mean	19.2	19.1	20.2	19.0
Standard Deviation	1.0	1.4	1.6	1.7
N	10	10	10	10
Group 3 - 500 ppm				
Mean	19.1	18.7*	19.5*	18.8
Standard Deviation	1.8	1.4	1.3	1.2
N	10	10	10	10
Group 4 - 2500 ppm				
Mean	17.8*	18.0*	18.7*	18.9
Standard Deviation	1.3	2.0	1.2	1.2
N	10	10	10	10

*$p \leq 0.05$ by Dunnett's Test

Table 14 - Summary of Weekly Mean Food Consumption (grams/animal/day): Reproductive Group Female Rats

	Pre-mating Week 1	Pre-mating Week 2	Gestation Week 1	Gestation Week 2	Gestation Week 3	Gestation Day 20 to PostPartum Day 0/1	Periparturiant Week
Group 1 - Controls							
Mean	19.3	20.0	23.5	25.6	24.5	20.9	30.4
Standard Deviation	1.3	1.5	1.6	2.0	4.6	3.6	5.1
N	10	10	10	10	10	10	10
Group 2 - 100 ppm							
Mean	19.4	19.8	23.7	25.9	24.3	19.8	29.0
Standard Deviation	2.6	2.2	3.7	3.5	3.4	5.0	8.0
N	10	10	10	10	10	10	10
Group 3 - 500 ppm							
Mean	18.6	18.8	22.0	23.9	23.5	21.8	30.3
Standard Deviation	2.4	2.4	2.8	2.1	2.5	4.6	5.6
N	10	10	10	10	10	10	10
Group 4 - 2500 ppm							
Mean	17.5	18.2	20.8	23.6	23.9	25.3	26.8
Standard Deviation	1.7	1.5	1.9	2.7	2.6	2.9	7.0
N	10	10	9[a]	9[a]	9[a]	9[a]	9[a]

[a] Animal D0033 not pregnant, values not used in calculations.

Table 15 - Summary of Functional Observation Battery Ranked/Categorical Incidence: Male Rats (Baseline)

Number of Animals Showing Each Observation

	Group 1 0 ppm	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Number of animals tested:	10	10	10	10
Resistance to removal: minimally avoids capture but is not aggressive	8	7	7	10
Resistance to removal: moderate; actively avoids capture but is not aggressive	2	3	3	0
Palpebral closure: open	10	10	10	10
Pupil reactivity: dilated, briskly reactive pupil	10	10	10	10
Pupil size: normal pupil size	10	10	10	10
Lacrimation: glistening cornea	10	10	10	10
Salivation: glistening oral cavity/lips, no wetness	10	10	10	10
Muscle tone: moderate; animal is neither very relaxed nor tense	10	10	10	10
Extensor thrust: moderate; clearly detectable	10	10	10	10
Reactivity to handling: minimal struggling	4	2	5	4
Reactivity to handling: moderate struggling with little or no vocalization	6	7	4	6
Reactivity to handling: pronounced struggling, animal may vocalize	0	1	1	0
Level of activity: moderate; inactive less than 50% of the time	10	10	10	10

Table 15 - continued

Number of Animals Showing Each Observation

	Group 1 0 ppm	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Number of animals tested:	10	10	10	10
Responsiveness to sharp noise: minimal; only the ears move	1	0	0	1
Responsiveness to sharp noise: moderate; clear movement of the head, neck and ears	9	9	10	9
Responsiveness to sharp noise: pronounced; clear movement of head, neck, ears, and body	0	1	0	0
Responsiveness to touch: moderate; spontaneously lifts tail when touched	10	10	10	10
Responsiveness to tail pinch: minimal; animal flinches but does not vocalize or move rapidly	0	1	0	1
Responsiveness to tail pinch: moderate; animal flinches, moves rapidly and/or vocalizes	10	9	10	9
Gait evaluation: smooth and coordinated gait	10	10	10	10
Urination: no drops of urine	1	0	1	3
Urination: minimal; a few (1-4) small drops or a small pool (~1cm)	6	6	7	1
Urination: moderate; more than one small pool, or a large pool (>1cm)	2	4	1	6
Urination: pronounced; clearly a large amount of urine (more than moderate)	1	0	1	0
Defecation: no pellets voided	4	1	4	3
Defecation: minimal; 1 or 2 pellets voided	5	8	4	6
Defecation: moderate; 3 pellets voided	1	1	2	1
Categorical observations:				
None	10	10	9	9
Feces: soft	0	0	1	1

Table 16 - Summary of Functional Observation Battery Ranked/Categorical Incidence: Male Rats (Treated)

Number of Animals Showing Each Observation

	Group 1 0 ppm	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Number of animals tested:	10	10	10	10
Resistance to removal: minimally avoids capture but is not aggressive	9	10	10	10
Resistance to removal: moderate; actively avoids capture but is not aggressive	1	0	0	0
Palpebral closure: open	10	10	10	10
Pupil reactivity: dilated, briskly reactive pupil	10	10	10	10
Pupil size: normal pupil size	10	10	10	10
Lacrimation: glistening cornea	10	10	10	10
Salivation: glistening oral cavity/lips, no wetness	10	10	10	10
Muscle tone: moderate; animal is neither very relaxed nor tense	10	10	10	10
Extensor thrust: slight; barely detectable response	0	0	1	0
Extensor thrust: moderate; clearly detectable	10	10	9	10
Reactivity to handling: minimal struggling	6	4	5	5
Reactivity to handling: moderate; struggling with little or no vocalization	4	6	5	5
Level of activity: minimal; inactive more than 50% of the time	1	2	1	2
Level of activity: moderate; inactive less than 50% of the time	9	8	9	8

Table 16 - continued

Number of Animals Showing Each Observation

	Group 1 0 ppm	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Number of animals tested:	10	10	10	10
Responsiveness to sharp noise: minimal; only the ears move	1	0	0	0
Responsiveness to sharp noise: moderate; clear movement of head, neck and ears	7	10	10	10
Responsiveness to sharp noise: pronounced; clear movement of head, neck, ears, and body	2	0	0	0
Responsiveness to touch: moderate; spontaneously lifts tail when touched	10	10	10	10
Responsiveness to tail pinch: moderate; animal flinches, moves rapidly and/or vocalizes	10	9	10	10
Responsiveness to tail pinch: pronounced; animal moves very quickly and bends towards the tail and vocalizes	0	1	0	0
Gait evaluation: smooth and coordinated gait	10	10	10	10
Urination: no drops of urine	0	0	0	3
Urination: minimal; a few (1-4) small drops or a small pool (~ 1cm)	4	3	1	1
Urination: moderate; more than one small pool, or a large pool (>1cm)	4	6	5	5
Urination: pronounced; clearly a large amount of urine (more than moderate)	2	1	4	1
Defecation: no pellets voided	0	0	1	2
Defecation: minimal; 1 or 2 pellets voided	6	4	7	6
Defecation: moderate; 3 pellets voided	2	4	1	1
Defecation: pronounced; 4 pellets voided	1	1	0	1
Defecation: exaggerated; 5 or more pellets voided	1	1	1	0
Categorical observations:				
None	10	9	10	10
Soiling: perineal, urine	0	1	0	0

Table 17 - Summary of Functional Observation Battery Ranked/Categorical Incidence: Toxicity Group Female Rats (Baseline)

	Number of Animals Showing Each Observation			
	Group 1 0 ppm	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Number of animals tested:	10	10	10	10
Resistance to removal: minimally avoids capture but is not aggressive	5	6	8	4
Resistance to removal: moderate; actively avoids capture but is not aggressive	5	4	2	6
Palpebral closure: open	10	10	10	10
Pupil reactivity: dilated, briskly reactive pupil	10	9	10	9
Pupil reactivity: constricted, cannot assess	0	1	0	1
Pupil size: normal pupil size	10	10	10	10
Lacrimation: glistening cornea	10	10	10	10
Salivation: glistening oral cavity/lips, no wetness	10	10	10	10
Muscle tone: moderate; animal is neither very relaxed nor tense	10	10	10	10
Extensor thrust: moderate; clearly detectable	10	10	10	10
Reactivity to handling: minimal struggling	0	1	2	1
Reactivity to handling: moderate struggling with little or no vocalization	9	9	6	9
Reactivity to handling: pronounced struggling, animal may vocalize	1	0	2	0
Level of activity: moderate; inactive less than 50% of the time	10	10	10	9
Level of activity: pronounced; leaves circle, occasional inactivity	0	0	0	1

Table 17 - continued

Number of Animals Showing Each Observation

	Group 1 0 ppm	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Number of animals tested:	10	10	10	10
Responsiveness to sharp noise: minimal; only the ears move	0	1	0	1
Responsiveness to sharp noise: moderate; clear movement of the head, neck and ears	10	9	10	9
Responsiveness to touch: moderate; spontaneously lifts tail when touched	10	10	10	10
Responsiveness to tail pinch: minimal; animal flinches but does not vocalize or move rapidly	1	2	0	0
Responsiveness to tail pinch: moderate; animal flinches, moves rapidly and/or vocalizes	9	8	10	10
Gait evaluation: smooth and coordinated gait	10	10	10	10
Urination: no drops of urine	5	4	3	7
Urination: minimal; a few (1-4) small drops or a small pool (~ 1cm)	4	4	7	2
Urination: moderate; more than one small pool, or a large pool (>1cm)	1	2	0	1
Defecation: no pellets voided	5	3	7	7
Defecation: minimal; 1 or 2 pellets voided	3	4	3	3
Defecation: moderate; 3 pellets voided	2	3	0	0
Categorical observations:				
None	10	10	10	10

Table 18 - Summary of Functional Observation Battery Ranked/Categorical Incidence: Toxicity Group Female Rats (Treated)

Number of Animals Showing Each Observation

	Group 1 0 ppm	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Number of animals tested:	10	10	10	9[a]
Resistance to removal: minimally avoids capture but is not aggressive	10	10	10	9
Palpebral closure: open	10	10	10	9
Pupil reactivity: dilated, briskly reactive pupil	10	10	10	9
Pupil size: normal pupil size	10	10	10	9
Lacrimation: glistening cornea	10	10	10	9
Salivation: glistening oral cavity/lips, no wetness	10	10	10	9
Muscle tone: moderate; animal is neither very relaxed nor tense	10	10	10	9
Extensor thrust: moderate; clearly detectable	10	10	10	9
Reactivity to handling: minimal struggling	0	2	1	6*
Reactivity to handling: moderate struggling with little or no vocalization	9	8	9	3
Reactivity to handling: pronounced struggling, animal may vocalize	1	0	0	0
Level of activity: none, rat doesn't leave the center circle (20 cm dia.)	0	0	1	0
Level of activity: minimal; inactive more than 50% of the time	2	2	2	0
Level of activity: moderate; inactive less than 50% of the time	8	8	7	9

[a]Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

*"Reactivity to Handling" as a category was statistically different ($p \leq 0.01$) by Fisher's Exact Test from control.

Table 18 - continued
Number of Animals Showing Each Observation

	Group 1 0 ppm	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Number of animals tested:	10	10	10	9[a]
Responsiveness to sharp noise: none, no response to click	0	1	0	0
Responsiveness to sharp noise: minimal; only the ears move	0	1	1	1
Responsiveness to sharp noise: moderate; clear movement of the head, neck and ears	8	8	8	7
Responsiveness to sharp noise: pronounced; clear movement of head, neck, ears, and body	1	0	1	1
Responsiveness to sharp noise: exaggerated; whole body response to stimulus, more intense than 'pronounced'	1	0	0	0
Responsiveness to touch: moderate; spontaneously lifts tail when touched	10	10	10	9
Responsiveness to tail pinch: moderate; animal flinches, moves rapidly and/or vocalizes	10	9	10	9
Responsiveness to tail pinch: pronounced; animal moves very quickly, bends towards the tail and vocalizes	0	1	0	0
Gait evaluation: smooth and coordinated gait	10	10	10	9
Urination: no drops of urine	0	1	1	1
Urination: minimal; a few (1-4) small drops or a small pool (~ 1cm)	2	4	5	6
Urination: moderate; more than one small pool, or a large pool (>1cm)	8	4	4	2
Urination: pronounced; clearly a large amount of urine (more than moderate)	0	1	0	0
Defecation: no pellets voided	4	4	3	8
Defecation: minimal; 1 or 2 pellets voided	2	1	4	0
Defecation: moderate; 3 pellets voided	1	0	2	1
Defecation: pronounced; 4 pellets voided	1	2	1	0
Defecation: exaggerated; 5 or more pellets voided	2	3	0	0
Categorical observations:				
None	10	10	10	9

[a]Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Table 19 - Summary of Mean Rectal Temperatures (°C) in Male and Female Rats

	Baseline	Treated
Group 1 - Controls		
Mean	37.6	37.7
Standard Deviation	0.4	0.6
N	20	20
Group 2 - 100 ppm		
Mean	37.4	37.8
Standard Deviation	0.4	0.5
N	20	20
Group 3 - 500 ppm		
Mean	37.7	37.9
Standard Deviation	0.4	0.5
N	20	20
Group 4 - 2500 ppm		
Mean	37.6	37.7
Standard Deviation	0.4	0.7
N	19[a]	19[a]

[a] Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Table 20 - Summary of Mean Hind limb Grip Strength (gram-force) in Male and Female Rats

	Baseline	Treated
Group 1 - Controls		
Mean	741	1173
Standard Deviation	142	234
N	20	20
Group 2 - 100 ppm		
Mean	771	1160
Standard Deviation	157	140
N	20	20
Group 3 - 500 ppm		
Mean	766	1202
Standard Deviation	154	228
N	20	20
Group 4 - 2500 ppm		
Mean	758	1074
Standard Deviation	158	201
N	19[a]	19[a]

[a]Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Table 21 - Summary of Mean Forelimb Grip Strength (gram-force) in Male and Female Rats

	Baseline	Treated
Group 1 - Controls		
Mean	878	1145
Standard Deviation	173	216
N	20	20
Group 2 - 100 ppm		
Mean	862	973
Standard Deviation	155	183
N	20	20
Group 3 - 500 ppm		
Mean	907	1075
Standard Deviation	129	256
N	20	20
Group 4 - 2500 ppm		
Mean	901	1085
Standard Deviation	118	194
N	19[a]	19[a]

[a]Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Table 22 - Summary of Mean Landing Foot Splay (centimeters) in Male and Female Rats

	Baseline	Treated
Group 1 - Controls		
Mean	12.1	10.9
Standard Deviation	1.9	1.7
N	20	20
Group 2 - 100 ppm		
Mean	11.7	10.8
Standard Deviation	1.7	1.7
N	20	20
Group 3 - 500 ppm		
Mean	11.8	10.8
Standard Deviation	1.5	1.7
N	20	20
Group 4 - 2500 ppm		
Mean	11.4	10.5
Standard Deviation	2.0	1.8
N	19[a]	19[a]

[a]Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Table 23 - Motor Activity Assessment Summary of Total Counts (Transformed[1])

	Males		Females		Combined	
	Baseline	Treated	Baseline	Treated	Baseline	Treated
Group 1 - Controls						
Mean	61.76	65.19	90.08	76.01	75.92	70.60
Standard Deviation	15.58	10.61	25.24	15.11	25.06	13.86
N	10	10	10	10	20	20
Group 2 - 100 ppm						
Mean	73.10	77.85	72.35	70.84	72.73	74.35
Standard Deviation	26.93	17.17	23.66	18.97	24.68	17.97
N	10	10	10	10	20	20
Group 3 - 500 ppm						
Mean	69.63	81.90	80.80	74.39	75.21	78.14
Standard Deviation	25.41	17.26	21.81	21.03	23.75	19.12
N	10	10	10	10	20	20
Group 4 - 2500 ppm						
Mean	73.35	42.72*	82.79	60.92	77.82	51.34*
Standard Deviation	22.48	9.37	22.77	9.60	22.51	13.11
N	10	10	9[a]	9[a]	19[a]	19[a]

[1]Square Root transformation of raw data counts.

[a]Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

*$p \leq 0.05$ by Repeated Measures ANOVA/Linear Contrasts

Table 24 - Motor Activity Assessment Summary by Interval (Transformed[1]): Baseline Males

Interval[2]	1	2	3	4	5	6	7	8	9	10
Group 1 - Controls										
Mean	15.09	11.87	8.27	7.66	5.85	4.02	2.20	1.30	2.89	2.60
Standard Deviation	1.69	2.12	3.59	3.82	2.42	4.15	2.82	3.43	4.46	3.44
N	10	10	10	10	10	10	10	10	10	10
Group 2 - 100 ppm										
Mean	15.12	11.97	8.85	7.07	7.41	7.96	5.82	4.45	3.00	1.47
Standard Deviation	1.83	2.42	4.61	4.89	4.80	4.82	4.08	5.24	4.18	3.41
N	10	10	10	10	10	10	10	10	10	10
Group 3 - 500 ppm										
Mean	14.49	11.45	8.27	8.37	7.78	1.53	3.42	4.96	4.92	4.43
Standard Deviation	1.97	2.99	2.82	3.24	3.64	2.86	3.78	5.01	5.73	5.45
N	10	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm										
Mean	15.28	11.59	10.94	8.64	7.32	5.15	4.84	3.47	2.65	3.46
Standard Deviation	1.30	1.93	1.00	1.28	2.14	3.94	4.54	4.58	4.20	4.96
N	10	10	10	10	10	10	10	10	10	10

[1]Square Root transformation of raw data counts.
[2]Session consisted of 10 eight minute intervals.

Table 25 - Motor Activity Assessment by Interval (Transformed[1]): Treated Males

Interval[2]	1	2	3	4	5	6	7	8	9	10
Group 1 - Controls										
Mean	14.78	10.65	7.97	5.96	6.97	4.59	2.61	3.09	4.80	3.77
Standard Deviation	2.00	1.47	2.31	3.21	2.73	3.63	2.96	2.92	3.21	3.70
N	10	10	10	10	10	10	10	10	10	10
Group 2 - 100 ppm										
Mean	14.11	11.26	9.39	7.86	9.16	5.84	4.99	6.03	4.82	4.38
Standard Deviation	1.77	1.81	1.95	1.79	2.15	3.06	3.68	4.30	3.64	3.83
N	10	10	10	10	10	10	10	10	10	10
Group 3 - 500 ppm										
Mean	14.46	11.13	9.37	7.52	7.15	7.14	8.06	6.85	5.61	4.60
Standard Deviation	1.36	2.17	3.36	4.55	3.88	3.00	2.75	3.20	3.66	4.16
N	10	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm										
Mean	11.93	9.20	6.84	4.31	2.71	2.09	1.57	0.94	1.58	1.57
Standard Deviation	1.15	1.37	1.99	1.99	2.13	1.50	1.35	1.42	2.41	2.12
N	10	10	10	10	10	10	10	10	10	10

[1]Square Root transformation of raw data counts.
[2]Session consisted of 10 eight minute intervals.

Table 26 - Motor Activity Assessment Summary by Interval (Transformed[1]): Baseline Females

Interval[2]	1	2	3	4	5	6	7	8	9	10
Group 1 - Controls										
Mean	15.55	12.62	10.79	10.79	9.92	8.67	6.14	4.72	5.60	5.28
Standard Deviation	1.18	1.77	1.40	1.48	2.69	3.95	4.93	4.89	4.75	4.91
N	10	10	10	10	10	10	10	10	10	10
Group 2 - 100 ppm										
Mean	14.05	10.92	9.74	9.63	8.43	4.25	3.32	3.46	3.60	4.96
Standard Deviation	2.12	3.61	3.01	3.66	3.48	3.72	4.07	4.46	3.91	3.88
N	10	10	10	10	10	10	10	10	10	10
Group 3 - 500 ppm										
Mean	15.57	12.49	10.96	9.54	8.17	7.80	5.54	3.38	3.60	3.76
Standard Deviation	1.48	1.51	2.60	3.80	3.27	3.30	4.71	4.92	4.75	4.31
N	10	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm										
Mean	15.80	12.00	11.89	10.37	8.28	3.11	3.26	4.06	6.19	7.85
Standard Deviation	1.28	3.87	2.29	2.50	4.51	4.30	4.28	4.83	5.08	4.51
N	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]

[1]Square Root transformation of raw data counts.
[2]Session consisted of 10 eight minute intervals.
[a]Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Table 27 - Motor Activity Summary by Interval (Transformed[1]): Treated Females

Interval[2]	1	2	3	4	5	6	7	8	9	10
Group 1 - Controls										
Mean	15.27	11.78	8.56	5.50	5.16	5.62	6.02	7.28	6.46	4.35
Standard Deviation	1.44	1.88	2.63	2.97	3.80	4.46	4.14	4.42	3.02	3.07
N	10	10	10	10	10	10	10	10	10	10
Group 2 - 100 ppm										
Mean	13.89	9.71	7.55	7.07	5.73	5.70	5.24	6.25	5.51	4.19
Standard Deviation	1.09	2.09	3.05	3.79	3.61	2.70	2.80	2.98	4.46	2.91
N	10	10	10	10	10	10	10	10	10	10
Group 3 - 500 ppm										
Mean	14.46	11.14	7.65	5.69	5.03	5.61	7.02	6.71	6.02	5.06
Standard Deviation	1.21	1.93	2.81	4.03	3.98	4.85	3.32	4.55	4.15	4.27
N	10	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm										
Mean	12.91	9.83	7.07	6.09	5.48	4.50	1.82	2.64	4.02	6.56
Standard Deviation	1.33	2.35	3.13	3.82	3.62	2.28	2.04	3.52	3.26	4.03
N	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]

[1]Square Root transformation of raw data counts.
[2]Session consisted of 10 eight minute intervals.
[a]Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Table 28 - Summary of Mean Motor Activity Assessment by Interval in Rats (Sexes Combined [Baseline]) Transformed[1]

Interval[2]	1	2	3	4	5	6	7	8	9	10
Group 1 - Controls										
Mean	15.32	12.25	9.53	9.23	7.88	6.34	4.17	3.01	4.25	3.94
Standard Deviation	1.44	1.93	2.95	3.24	3.25	4.61	4.40	4.47	4.69	4.35
N	20	20	20	20	20	20	20	20	20	20
Group 2 - 100 ppm										
Mean	14.58	11.44	9.29	8.35	7.92	6.10	4.57	3.96	3.30	3.21
Standard Deviation	2.00	3.04	3.82	4.40	4.12	4.60	4.17	4.76	3.95	3.98
N	20	20	20	20	20	20	20	20	20	20
Group 3 - 500 ppm										
Mean	15.03	11.97	9.61	8.95	7.97	4.66	4.48	4.17	4.26	4.10
Standard Deviation	1.78	2.36	2.98	3.49	3.37	4.40	4.29	4.90	5.17	4.80
N	20	20	20	20	20	20	20	20	20	20
Group 4 - 2500 ppm										
Mean	15.52	11.78	11.39	9.46	7.77	4.18	4.09	3.75	4.33	5.54
Standard Deviation	1.28	2.93	1.75	2.09	3.40	4.13	4.37	4.57	4.85	5.14
N	19[a]	19	19	19	19	19	19	19	19	19

[1]Square root transformation of raw data counts

[2]Session consisted of 10 eight minute intervals.

[a]Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Table 29 - Summary of Mean Motor Activity Assessment by Interval in Rats (Sexes Combined [Treated]) Transformed[1]

Interval[2]	1	2	3	4	5	6	7	8	9	10
Group 1 - Controls										
Mean	15.02	11.22	8.26	5.73	6.07	5.10	4.32	5.19	5.63	4.06
Standard Deviation	1.72	1.74	2.43	3.02	3.35	3.99	3.92	4.23	3.15	3.32
N	20	20	20	20	20	20	20	20	20	20
Group 2 - 100 ppm										
Mean	14.00	10.49	8.47	7.47	7.45	5.77	5.12	6.14	5.17	4.29
Standard Deviation	1.44	2.06	2.67	2.91	3.39	2.81	3.18	3.60	3.98	3.31
N	20	20	20	20	20	20	20	20	20	20
Group 3 - 500 ppm										
Mean	14.46	11.14	8.51	6.61	6.09	6.38	7.54	6.78	5.82	4.83
Standard Deviation	1.25	2.00	3.14	4.29	3.98	4.00	3.02	3.83	3.82	4.11
N	20	20	20	20	20	20	20	20	20	20
Group 4 - 2500 ppm*										
Mean	12.40	9.50	6.95	5.15	4.02	3.23	1.69	1.75	2.73	3.94
Standard Deviation	1.30	1.87	2.52	3.05	3.18	2.23	1.67	2.70	3.03	4.00
N	19[a]	19	19	19	19	19	19	19	19	19

[1]Square root transformation of raw data counts.

[2]Session consisted of 10 eight minute intervals.

[a]Animal D0120 was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

*Total Motor Activity was statistically significant ($p \leq 0.05$) by Repeated Measures ANOVA/Linear Contrasts (Sexes Combined).

Table 30 - Summary of Mean Hematology: Male Rats

	White Blood Cells (X 10^3/cu mm)	Red Blood Cells (X 10^6/cu mm)	Hemoglobin (g/dL)	Hematocrit (%)	Mean Cell Volume (cubic microns)	Mean Cell Hemoglobin (picograms)	Mean Cell Hemaglobin Concentration (%)	Platlets (X 10^3/cu mm)
Group 1 - Controls								
Mean	12.2	8.0	16.9	44.4	55.5	21.1	38.0	1091.6
Standard Deviation	4.0	0.5	0.6	2.2	1.4	0.9	0.9	68.3
N	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]
Group 2 - 100 ppm								
Mean	13.0	8.0	16.6	43.7	54.8	20.9	38.1	1122.6
Standard Deviation	5.1	0.4	0.5	1.7	1.5	0.8	0.7	67.7
N	10	10	10	10	10	10	10	10
Group 3 - 500 ppm								
Mean	12.1	7.8	16.4	43.0	55.4	21.1	38.1	1125.0
Standard Deviation	3.5	0.4	0.5	1.8	1.2	0.7	0.8	83.8
N	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm								
Mean	11.3	7.9	16.0*	42.6	53.9	20.2	37.5	1279.4*
Standard Deviation	2.2	0.4	0.4	1.3	1.8	1.0	0.8	123.8
N	10	10	10	10	10	10	10	10

*$p \leq 0.05$ by Dunnett's Test

[a] Sample D0041 clotted

Table 31 - Summary of Mean Hematology Data: Toxicity Group Female Rats

	White Blood Cells (X 10^3/cu mm)	Red Blood Cells (X 10^6/cu mm)	Hemoglobin (g/dL)	Hematocrit (%)	Mean Cell Volume (cubic microns)	Mean Cell Hemoglobin (picograms)	Mean Cell Hemaglobin Concentration (%)	Platlets (X 10^3/cu mm)
Group 1 - Controls								
Mean	8.0	7.1	15.3	39.8	55.8	21.4	38.4	1103.1
Standard Deviation	3.2	0.3	0.9	2.4	3.6	1.5	0.5	101.0
N	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]
Group 2 - 100 ppm								
Mean	8.7	7.4	16.0*	42.2	57.2	21.7	37.9	1109.8
Standard Deviation	4.0	0.5	0.7	2.3	1.3	0.7	0.7	126.0
N	9[b]	9[b]	9[b]	9[b]	9[b]	9[b]	9[b]	9[b]
Group 3 - 500 ppm								
Mean	7.3	7.2	15.6	41.2	57.2	21.7	37.9	1083.4
Standard Deviation	2.4	0.4	0.6	1.7	1.7	0.7	0.5	111.2
N	9[c]	9[c]	9[c]	9[c]	9[c]	9[c]	9[c]	9[c]
Group 4 - 2500 ppm								
Mean	7.7	7.0	15.2	40.5	58.0	21.8	37.6	1136.5
Standard Deviation	1.2	0.3	0.4	1.5	1.7	0.8	0.8	92.1
N	10	10	10	10	10	10	10	10

[a] Sample D0088 clotted
[b] Sample D0095 clotted
[c] Sample D0108 clotted

Table 32 - Summary of Mean Hematology Differential Data: Male Rats

	Segmented Neutrophils (%)	Band Neutrophils (%)	Lymphocytes (%)	Monocytes (%)	Eosinophils (%)	Basophils (%)
Group 1 - Controls						
Mean	13.4	0.2	85.3	0.9	0.1	0.0
Standard Deviation	5.5	0.4	6.2	0.9	0.3	0.0
N	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]
Group 2 - 100 ppm						
Mean	10.8	0.0	88.3	0.9	0.0	0.0
Standard Deviation	3.0	0.0	3.2	1.4	0.0	0.0
N	10	10	10	10	10	10
Group 3 - 500 ppm						
Mean	17.9	0.0	81.7	0.4	0.0	0.0
Standard Deviation	4.5	0.0	4.6	0.5	0.0	0.0
N	10	10	10	10	10	10
Group 4 - 2500 ppm						
Mean	13.0	0.1	86.2	0.7	0.0	0.0
Standard Deviation	3.4	0.3	3.4	0.8	0.0	0.0
N	10	10	10	10	10	10

[a] Sample D0041 clotted

Table 33 - Summary of Mean Hematology Differential Data: Toxicity Group Female Rats

	Segmented Neutrophils (%)	Band Neutrophils (%)	Lymphocytes (%)	Monocytes (%)	Eosinophils (%)	Basophils (%)
Group 1 - Controls						
Mean	9.1	0.0	90.7	0.2	0.0	0.0
Standard Deviation	4.3	0.0	4.4	0.4	0.0	0.0
N	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]
Group 2 - 100 ppm						
Mean	12.7	0.0	86.9	0.3	0.0	0.0
Standard Deviation	5.6	0.0	5.6	0.7	0.0	0.0
N	9[b]	9[b]	9[b]	9[b]	9[b]	9[b]
Group 3 - 500 ppm						
Mean	14.7	0.0	84.9	0.3	0.1	0.0
Standard Deviation	8.0	0.0	7.9	0.7	0.3	0.0
N	9[c]	9[c]	9[c]	9[c]	9[c]	9[c]
Group 4 - 2500 ppm						
Mean	10.3	0.0	89.0	0.7	0.0	0.0
Standard Deviation	2.8	0.0	2.7	0.8	0.0	0.0
N	10	10	10	10	10	10

[a] Sample D0088 clotted
[b] Sample D0095 clotted
[c] Sample D0108 clotted

Table 34 - Summary of Mean Prothrombin Time in Rats

		Males Results (seconds)	Females Results (seconds)
Group 1 - Controls			
	Mean	16.6	15.9
	Standard Deviation	1.1	0.7
	N	9[a]	9[b]
Group 2 - 100 ppm			
	Mean	16.1	15.9
	Standard Deviation	0.9	0.8
	N	10	7[c]
Group 3 - 500 ppm			
	Mean	17.5	16.3
	Standard Deviation	1.7	0.4
	N	10	8[d]
Group 4 - 2500 ppm			
	Mean	16.4	16.5
	Standard Deviation	0.9	0.7
	N	10	10

[a] Sample D0041 clotted
[b] Sample D0088 clotted
[c] Samples D0095, D0097 and D0099 clotted
[d] Sample D0104 and D0108 clotted

Table 35 - Summary of Mean Clinical Chemistry Data: Male Rats

	Albumin g/dL	Alkaline Phosphatase U/L	ALT U/L	AST U/L	Total Bilirubin mg/dL	Calcium mg/dL	Creatinine mg/dL	Cholesterol mg/dL	Glucose mg/dL	Phosphorus mg/dL	Total Protein g/dL	Urea Nitrogen mg/dL	Sodium mmol/L	Potassium mmol/L	Chloride mmol/L
Group 1 - Controls															
Mean	3.7	135.4	26.8	98.6	0.2	10.0	0.7	39.8	124.1	7.9	6.1	14.9	146.7	4.6	109.1
Std Dev.	0.5	32.8	7.2	17.9	0.1	0.3	0.1	5.5	15.5	0.8	0.4	4.0	2.3	0.7	2.1
N	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
Group 2 - 100 ppm															
Mean	3.9	140.0	28.2	92.4	0.2	9.9	0.7	44.6	120.3	8.1	6.0	16.0	145.6	5.1	108.5
Std Dev.	0.1	20.1	6.5	7.7	0.1	0.5	0.1	15.0	23.1	1.5	0.4	2.1	1.3	1.7	1.4
N	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
Group 3 - 500 ppm															
Mean	3.8	135.0	30.4	94.2	0.1	9.8	0.6	42.2	110.9	7.6	6.0	15.9	145.8	4.4	109.7
Std Dev.	0.2	24.1	5.8	8.6	0.1	0.3	0.1	5.8	14.0	0.5	0.3	1.7	1.5	0.3	1.7
N	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]
Group 4 - 2500 ppm															
Mean	3.8	98.5**	36.6	87.0	0.1	9.8	0.6	41.8	127.5	7.8	6.2	19.1*	144.4**	4.8	107.8
Std Dev.	0.2	10.1	12.1	8.3	0.1	0.3	0.1	5.2	36.1	0.7	0.4	3.7	1.5	0.6	2.0
N	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10

[a] Sample D0061 not taken

*$p \leq 0.05$ by Dunnett's Test

**$p \leq 0.01$ by Wilcoxon Test

Table 36 - Summary of Mean Clinical Chemistry Data: Toxicity Group Female Rats

	Albumin g/dL	Alkaline Phosphatase U/L	ALT U/L	AST U/L	Total Bilirubin mg/dL	Calcium mg/dL	Creatinine mg/dL	Cholesterol mg/dL	Glucose mg/dL	Phosphorus mg/dL	Total Protein g/dL	Urea Nitrogen mg/dL	Sodium mmol/L	Potassium mmol/L	Chloride mmol/L
Group 1 - Controls															
Mean	4.4	56.9	23.9	85.6	0.2	10.1	0.7	51.2	137.4	6.3	6.6	14.1	143.8	4.0	109.4
Std Dev.	0.3	10.7	7.0	6.7	0.0	0.2	0.1	11.7	20.0	1.0	0.5	2.0	1.2	0.3	1.3
N	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
Group 2 - 100 ppm															
Mean	4.5	64.8	24.2	84.2	0.2	9.9	0.7	52.9	138.3	6.1	6.7	15.9	144.1	4.6	110.0
Std Dev.	0.2	13.4	3.3	10.3	0.0	0.8	0.1	10.9	29.2	1.2	0.6	2.3	1.3	1.2	1.3
N	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]	9[a]
Group 3 - 500 ppm															
Mean	4.2	65.4	24.2	90.1	0.2	9.7**	0.7	53.0	112.8*	6.7	6.5	15.6	144.9	4.2	110.7
Std Dev.	0.2	14.9	3.5	16.8	0.0	0.3	0.1	10.6	17.4	1.1	0.3	2.3	1.8	0.3	2.0
N	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm															
Mean	4.0	52.8	22.1	75.8	0.2	9.8**	0.6	66.5*	110.7*	6.2	6.4	14.6	144.2	4.2	109.8
Std Dev.	0.3	11.7	4.4	9.5	0.1	0.2	0.1	13.4	16.5	0.6	0.5	3.3	1.4	0.2	1.1
N	10	10	10	10	10	10	10	10	10	10	10	10	10	10	10

[a] Sample D0099 not taken

*$p \leq 0.05$ by Dunnett's Test

**$p \leq 0.01$ by Wilcoxon Test

Table 37 - Summary of Macroscopic Observations in Rats

		Group 1 Controls	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Males	No. Examined	10	10	10	10
- Lung - dark foci		-	1	-	-
- Seminal Vesicles - diminished in size		-	-	-	1
Females	- No Findings				

Table 38 - Summary of Mean Organ Weights (grams): Male Rats

	Brain	Heart	Liver	Lungs	Kidneys	Adrenals	Thymus	Spleen	Testes	Epididymides	Seminal Vesicles	Prostate	Body Weight
Group 1 - Controls													
Mean	2.036	1.181	9.738	1.279	2.543	0.056	0.364	0.705	3.052	1.105	1.375	0.870	361.1
Standard Deviation	0.126	0.162	1.412	0.134	0.268	0.009	0.081	0.143	0.206	0.131	0.217	0.174	35.9
N	10	10	10	10	10	10	10	9[a]	10	10	10	10	10
Group 2 - 100 ppm													
Mean	1.993	1.160	9.670	1.347	2.612	0.057	0.366	0.714	3.062	1.113	1.376	0.906	366.7
Standard Deviation	0.129	0.164	1.376	0.085	0.312	0.014	0.111	0.102	0.349	0.096	0.372	0.257	28.7
N	10	10	10	10	10	10	10	10	10	10	10	10	10
Group 3 - 500 ppm													
Mean	2.005	1.118	9.911	1.319	2.569	0.057	0.355	0.682	3.079	1.077	1.533	0.925	366.9
Standard Deviation	0.053	0.135	0.745	0.116	0.091	0.014	0.111	0.059	0.188	0.058	0.221	0.118	12.5
N	10	10	10	10	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm													
Mean	1.989	1.091	11.864*	1.260	2.706	0.056	0.304	0.666	2.997	0.989*	0.956*	0.685	337.7
Standard Deviation	0.114	0.101	1.343	0.075	0.435	0.012	0.081	0.158	0.199	0.082	0.265	0.130	28.2
N	10	10	10	10	10	10	10	10	10	10	10	10	10

*$p \leq 0.05$ by Dunnett's Test

[a] Spleen weight for animal D0047 was inadvertently not taken at necropsy.

Table 39 - Summary of Mean Organ Weights (grams): Toxicity Group Female Rats

	Brain	Heart	Liver	Lungs	Kidneys	Adrenals	Thymus	Spleen	Uterus	Ovaries	Body Weight
Group 1 - Controls											
Mean	1.902	0.839	6.821	1.097	1.643	0.075	0.310	0.536	0.727	0.104	241.9
Standard Deviation	0.063	0.073	0.364	0.067	0.095	0.008	0.072	0.060	0.272	0.021	8.2
N	10	10	10	10	10	10	10	10	10	10	10
Group 2 - 100 ppm											
Mean	1.857	0.783	7.150	1.126	1.582	0.065*	0.342	0.488	0.685	0.101	238.0
Standard Deviation	0.046	0.030	0.977	0.117	0.160	0.005	0.048	0.063	0.325	0.014	10.5
N	10	10	10	10	10	10	10	10	10	10	10
Group 3 - 500 ppm											
Mean	1.927	0.760*	6.851	1.118	1.627	0.075	0.307	0.504	0.749	0.114	229.9
Standard Deviation	0.064	0.050	0.417	0.056	0.126	0.008	0.066	0.087	0.279	0.033	13.0
N	10	10	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm											
Mean	1.855	0.793	9.036**	1.106	1.621	0.081	0.284	0.538	0.594	0.102	227.0*
Standard Deviation	0.062	0.072	0.501	0.067	0.125	0.012	0.066	0.081	0.219	0.012	12.5
N	10	10	10	10	10	10	10	10	10	10	10

*$p \leq 0.05$ by Dunnett's Test
**$p \leq 0.01$ by Wilcoxon Test

Table 40 - Summary of Mean Organ to Body Weight Ratios: Male Rats

	Brain/ Body Weight Ratio	Heart/ Body Weight Ratio	Liver/ Body Weight Ratio	Lung/ Body Weight Ratio	Kidney/ Body Weight Ratio	Adrenal/ Body Weight Ratio	Thymus/ Body Weight Ratio	Spleen/ Body Weight Ratio	Testes/ Body Weight Ratio	Epididymide/ Body Weight Ratio	Seminal Vesicle/ Body Weight Ratio	Prostate/ Body Weight Ratio
Group 1 - Control												
Mean	0.00567	0.00327	0.02696	0.00354	0.00704	0.00015	0.00100	0.00193	0.00851	0.00307	0.00383	0.00240
Standard Deviation	0.00047	0.00036	0.00290	0.00018	0.00024	0.00002	0.00018	0.00024	0.00078	0.00030	0.00064	0.00035
N	10	10	10	10	10	10	10	9[a]	10	10	10	10
Group 2 - 100 ppm												
Mean	0.00545	0.00315	0.02629	0.00368	0.00711	0.00016	0.00100	0.00194	0.00838	0.00305	0.00374	0.00246
Standard Deviation	0.00035	0.00028	0.00202	0.00015	0.00049	0.00003	0.00030	0.00020	0.00101	0.00031	0.00089	0.00065
N	10	10	10	10	10	10	10	10	10	10	10	10
Group 3 - 500 ppm												
Mean	0.00547	0.00304	0.02706	0.00359	0.00701	0.00015	0.00097	0.00186	0.00840	0.00294	0.00418	0.00252
Standard Deviation	0.00022	0.00034	0.00231	0.00023	0.00032	0.00003	0.00030	0.00015	0.00054	0.00017	0.00059	0.00028
N	10	10	10	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm												
Mean	0.00591	0.00325	0.03528**	0.00375	0.00799**	0.00017	0.00090	0.00196	0.00895	0.00294	0.00280*	0.00202
Standard Deviation	0.00034	0.00040	0.00411	0.00026	0.00084	0.00003	0.00022	0.00037	0.00109	0.00024	0.00071	0.00028
N	10	10	10	10	10	10	10	10	10	10	10	10

*$p \leq 0.05$ by Dunnett's Test

**$p \leq 0.01$ by Wilcoxon Test

[a] Spleen weight for animal D0047 was inadvertently not taken at necropsy.

Table 41 - Summary of Mean Organ to Body Weight Ratios: Toxicity Group Female Rats

	Brain/ Body Weight Ratio	Heart/ Body Weight Ratio	Liver/ Body Weight Ratio	Lung/ Body Weight Ratio	Kidney/ Body Weight Ratio	Adrenal/ Body Weight Ratio	Thymus/ Body Weight Ratio	Spleen/ Body Weight Ratio	Uterus/ Body Weight Ratio	Ovary/ Body Weight Ratio
Group 1 - Controls										
Mean	0.00787	0.00347	0.02819	0.00453	0.00679	0.00031	0.00128	0.00221	0.00300	0.00043
Standard Deviation	0.00039	0.00027	0.00124	0.00022	0.00031	0.00003	0.00028	0.00020	0.00109	0.00010
N	10	10	10	10	10	10	10	10	10	10
Group 2 - 100 ppm										
Mean	0.00782	0.00330	0.03007	0.00473	0.00665	0.00027	0.00144	0.00206	0.00285	0.00043
Standard Deviation	0.00043	0.00021	0.00394	0.00037	0.00067	0.00002	0.00023	0.00028	0.00127	0.00007
N	10	10	10	10	10	10	10	10	10	10
Group 3 - 500 ppm										
Mean	0.00840*	0.00331	0.02981**	0.00487*	0.00708	0.00033	0.00135	0.00219	0.00323	0.00050
Standard Deviation	0.00043	0.00020	0.00082	0.00024	0.00045	0.00004	0.00032	0.00034	0.00106	0.00016
N	10	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm										
Mean	0.00819	0.00350	0.03986**	0.00488*	0.00717	0.00036*	0.00124	0.00236	0.00261	0.00045
Standard Deviation	0.00050	0.00034	0.00230	0.00031	0.00075	0.00006	0.00024	0.00027	0.00088	0.00004
N	10	10	10	10	10	10	10	10	10	10

*$p \leq 0.05$ by Dunnett's Test
**$p \leq 0.01$ by Wilcoxon Test

Table 42 - Summary of Histopathology Findings in Male Rats

Tissue		Group 1 Controls	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Lung					
- Inflammation	No. Examined	10	-	-	10
	Grade 1	1	-	-	1
	Grade 2	-	-	-	1
- Alveolar histiocytosis		1	-	-	-
	Grade 1	3	-	-	7
	Grade 2	2	-	-	6
- Granuloma		1	-	-	1
	Grade 2	1	-	-	-
		1	-	-	-
Seminal Vesicles					
- Atrophy	No. Examined	10	10	10	10
	Grade 2	-	-	-	4*
		-	-	-	4
Brain					
- Neuronal degeneration	No. Examined	10	-	-	10
	Grade 1	-	-	-	1
		-	-	-	1
Nose					
- Hemorrhage	No. Examined	10	-	-	10
	Grade 2	-	-	-	1
		-	-	-	1
Stomach					
- Mucosal gland dilation	No. Examined	10	-	-	10
	Grade 2	-	-	-	1
		-	-	-	1
Liver					
- Focal necrosis	No. Examined	10	10	10	10
	Grade 1	1	1	2	-
-Lymphoid cell infiltration		1	1	2	-
	Grade 1	1	2	3	-
	Grade 2	-	2	3	-
-Centrilobular hepatocellular hypertrophy		1	-	-	-
	Grade 1	-	-	-	10**
		-	-	-	10
Kidneys					
- Protein droplet nephropathy	No. Examined	10	10	10	10
	Grade 1	-	1	9**	10**
	Grade 2	-	1	9	-
	Grade 3	-	-	-	5
-Tubular degeneration/regeneration		-	-	-	5
	Grade 1	6	4	6	8
	Grade 2	6	4	6	4
- Lymphoid cell infiltration		-	-	-	4
	Grade 1	2	4	1	3
- Inflammation		2	4	1	3
	Grade 3	-	-	1	-
		-	-	1	-

*p≤0.05 by One-Sided Fisher's Exact Test
**p≤0.01 by One-Sided Fisher's Exact Test

Table 42 - continued

Tissue		Group 1 Controls	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Urinary Bladder	No. Examined	10	-	-	10
- Seminal Coagulum		1	-	-	1
	Grade 2	1	-	-	1
- Lymphoid cell infiltration		-	-	-	2
	Grade 1	-	-	-	1
	Grade 2	-	-	-	1
Testes	No. Examined	10	-	-	10
- Tubular atrophy		-	-	-	1
	Grade 1	-	-	-	1
- Multinucleated spermatid giant cells		-	-	-	1
	Grade 1	-	-	-	1
Epididymides	No. Examined	10	-	-	10
- Lymphoid cell infiltration		-	-	-	1
	Grade 1	-	-	-	1
- Intratubular cellular debris		3	-	-	1
	Grade 1	3	-	-	1
Prostate Gland	No. Examined	10	-	-	10
- Lymphoid cell infiltration		5	-	-	3
	Grade 1	2	-	-	1
	Grade 2	1	-	-	1
	Grade 3	2	-	-	1
Adrenal Glands	No. Examined	10	-	-	10
- Hypertrophy		1	-	-	1
	Grade 1	-	-	-	1
	Grade 2	1	-	-	-
Spleen	No. Examined	10	-	-	10
- Lymphoid hyperplasia		1	-	-	-
	Grade 3	1	-	-	-
Mesenteric Lymph Node	No. Examined	10	-	-	10
- Lymphoid hyperplasia		1	-	-	-
	Grade 3	1	-	-	-

Table 43 - Summary of Histopathology Findings in Toxicity Group Female Rats

Tissue		Group 1 Controls	Group 2 100 ppm	Group 3 500 ppm	Group 4 2500 ppm
Lung	No. Examined	10	-	-	10
- Alveolar histiocytosis		3	-	-	3
	Grade 1	3	-	-	3
Stomach	No. Examined	10	-	-	10
- Inflammation		1	-	-	2
	Grade 1	-	-	-	1
	Grade 2	1	-	-	1
- Chief cell hyperplasia		1	-	-	-
	Grade 2	1	-	-	-
- Glandular hyperplasia		-	-	-	1
	Grade 1	-	-	-	1
Liver	No. Examined	10	10	10	10
- Tigroid basophilic focus		-	-	-	1
	Grade 1	-	-	-	1
- Centrilobular hepatocellular hypertrophy		-	-	-	10**
	Grade 2	-	-	-	10
Kidneys	No. Examined	10	-	-	10
- Tubular degeneration / regeneration		4	-	-	2
	Grade 1	4	-	-	2
- Lymphoid cell infiltration		-	-	-	1
	Grade 1	-	-	-	1
Urinary Bladder	No. Examined	10	-	-	10
- Lymphoid cell infiltration		-	-	-	1
	Grade 1	-	-	-	1
Ovaries	No. Examined	10	-	-	10
- Cyst(s)		2	-	-	-
Uterus	No. Examined	10	-	-	10
- Dilation		1	-	-	-
	Grade 3	1	-	-	-
Adrenal Glands	No. Examined	10	-	-	10
- Anomaly		-	-	-	1
Spleen	No. Examined	10	-	-	10
- Lymphoid hyperplasia		1	-	-	1
	Grade 2	1	-	-	1

**$p \leq 0.01$ by One-Sided Fisher's Exact Test

Table 44 - Summary of Mean Reproductive Parameters for Reproductive Group Female Rats

			Litter						PND 0 Initial		PND 4 Final			
							PND 4							
	Days Gestation	Undetermined Sex	Male Pups	Female Pups	Males/ Females	Total Pups	Viable (Live) Pups	Viable/ Total	Litter Weight (Grams)	Average Pup Weight (Grams)	Litter Weight (Grams)	Average Pup Weight (Grams)	Total Implants	Corpora Lutea Counts
Group 1 - Controls														
Mean	21.7	0.1	7.5	7.7	1.2	15.3	14.6	0.96	92.7	6.1	138.9	9.5	16.5	18.5
Std Dev.	0.5	0.3	2.1	2.7	0.8	2.1	1.6	0.07	12.4	0.6	19.4	0.8	1.7	3.1
N	10	10	10	10	10	10	10	10	10	10	10	10	10	10
Group 2 - 100 ppm														
Mean	21.8	0.1	7.7	7.6	1.5	15.4	13.8	0.88	93.2	6.1	144.7	9.5	15.9	17.0
Std Dev.	0.4	0.3	2.3	3.0	1.5	1.7	5.0	0.31	8.4	0.3	11.5	1.2	1.3	1.6
N	10	10	10	10	10	10	10	10	10	10	9[a]	9[a]	10	10
Group 3 - 500 ppm														
Mean	21.9	0.0	6.9	8.1	0.9	15.0	14.2	0.93	93.1	6.3	136.3	9.9	15.2	17.7
Std Dev.	0.3	0.0	2.0	2.5	0.3	3.7	3.6	0.07	18.9	0.7	24.9	1.5	3.8	3.8
N	10	10	10	10	10	10	10	10	10	10	10	10	10	10
Group 4 - 2500 ppm														
Mean	22.1	0.2	5.1	5.0	1.3	10.3**	8.6**	0.87	67.5*	6.6	101.3*	10.8	11.1**	16.6
Std Dev.	0.3	0.7	1.9	2.4	0.6	3.8	4.7	0.33	23.6	0.4	33.3	1.0	3.9	5.2
N	9[b]	9[b]	9[b]	9[b]	9[b]	9[b]	9[b]	9[b]	9[b]	9[b]	8[c]	8[c]	9[b]	9[b]

*p≤0.05 by Dunnett's Test

**p≤0.01 by Wilcoxon Test

[a] Animal D0012 cannabilized pups, values not used in calculations.

[b] Animal D0033 not pregnant, values not used in calculations.

[c] Animal D0033 not pregnant, animal D0035 cannabilized pups, values not used in calculations.

Table 45 - Summary of Reproductive Performance for Reproductive Group Female Rats

	Group 1 - Controls	Group 2 - 100 ppm	Group 3 - 500 ppm	Group 4 - 2500 ppm
Females on Study	10	10	10	10
Females that died during study	0	0	0	0
Females euthanized in extremis	0	0	0	0
Females Allowed to Deliver				
NonGravid	0	0	0	1
Gravid	10	10	10	9
Females with Evidence of Copulation				
Number which delivered	10	10	10	9
Number which did not deliver	0	0	0	1
Females with No Evidence of Copulation				
Number which delivered	0	0	0	0
Number which did not deliver	0	0	0	0

Table 46 - Summary of Protocol Deviations and Additions

Protocol Section	Protocol Requirement	Deviation or Addition	Impact
Section X.G. Page 14	Functional observation battery (FOB) will be performed on all adult male and all Toxicity group females prior to the start of dosing (baseline evaluations).	Animal D0019 (reproductive group female) was inadvertently included in the baseline FOB evaluation.	None.
Section X.G. Page 14	Functional observation battery (FOB) will be performed on all adult male and all Toxicity group females prior to the start of dosing (baseline evaluations).	The extra animals (i.e. not assigned to a dose group) were evaluated for FOB and motor activity. The animals were evaluated to obtain baseline data should they be needed as replacement animals.	None.
Section X.B.2 Page 8	Temperature and relative humidity will be maintained within the range of 22 ± 3°C and 50 ± 20%, respectively.	Chamber temperatures increased during the daily exposures. Excursions above the stated range were typical for each chamber. Chamber temperatures ranged from 21.7 to 27.7°C.	None.
Section VII.C. Page 5	Animals will be housed in an environmentally controlled animal room (12-hour fluorescent-light/dark cycle)	On March 25, 2002 the light portion of the light cycle was extended approximately 5 minutes to complete the recording of clinical observations.	None.
Section X.G. Page 14	Functional observation battery (FOB) will be performed on all adult male and all Toxicity group females prior to the start of dosing (baseline evaluations).	FOB data was not available for one toxicity group female (D0040). This rat, originally a reproductive group female, was substituted for toxicity group female after initiation of exposures and after conducting baseline assessments.	None.
Section X.B.2. Page 8	Temperature and relative humidity will be maintained within the range of 22 ± 3°C and 50 ± 20%, respectively.	On April 1, 2002, one of the relative humidity readings for Chamber 30903-E was below the minimum protocol specification. The reading was 29.6%	None.
Section X.B.3. Page 9	The exposure periods will be defined as a 6hr/day exposure at target concentration.	On April 3, 2002 the exposure at target concentration in chamber 30903-E (Group 2 males) was 5 hours and 29 minutes.	None.
Section X.B.3. Page 9	The exposure periods will be defined as a 6hr/day exposure at target concentration.	On April 4, 2002 the exposure to target concentration was inadvertently extended 13 minutes for rats in chamber 30903-E (group 3 toxicity and reproductive females) and reduced by 13 minutes for rats in chamber 30908-E (group 3 males).	None.

Table 46 (continued)

Protocol Section	Protocol Requirement	Deviation or Addition	Impact
Section X.B.3. Page 9	Animals will be removed from the chambers after the second T_{99} has expired.	On study days 1 - 11 control animals were removed from inhalation chambers prior to the end of the second T_{99} time.	None.
Section X.B.3. Page 9	The exposure periods will be defined as a 6hr/day exposure at target concentration. This will involve a daily exposure routine whereby the 6 hr exposure at target concentration is sandwiched between two T_{99} periods.	On April 5, 2002, a reproductive group female (D0010, control) was inadvertently not placed into the exposure cage during the loading process. This rat was added to the exposure cage approximately 3 minutes after initiating the exposure start. As a control animal, this did not involve test article exposure.	None.
Section X.B.3. Page 9	The exposure periods will be defined as a 6hr/day exposure at target concentration.	On April 17, 2002, the chamber analysis data suggests that the exposure duration for chamber 30904-E (2500 ppm) was decreased by 33 minutes.	None.
Section X.G. Page 15	At necropsy, from all males and toxicity group females the following organs will be taken, trimmed and weighed as soon as possible after dissection: adrenals, brain, heart, kidneys, liver, lungs, spleen and thymus.	The spleen from one male rat (D0047) in the control exposure group was inadvertently not weighed.	None.
Section X. E. Page 11	All animals assigned to toxicity groups will be euthanized by exsanguination brought about by the cutting of major abdominal vessels following Ketamine HCL/Xylazine anesthesia.	One male rat (D0072) from the 2500 ppm exposure level was not responding to Ketamine HCL/Xylazine anesthesia and as a consequence was anesthetized via carbon dioxide inhalation.	None.
Section VII. C. Page 5	Documentation (of temperature and humidity) will include the recorder chart and hand recording of values twice a day on weekdays.	Temperature and humidity values were not hand recorded during the a.m. observations for the animal room. Values were within acceptable ranges as documented on the chart recorder.	None.

Table 46 (continued)

Protocol Section	Protocol Requirement	Deviation or Addition	Impact
Section X. H. Page 16	Blood samples will be collected from all males and all toxicity group females on the day of scheduled sacrifice as a terminal procedure. While under Ketamine/xylazine anesthesia a syringe and needle will be used to collect blood samples for hematological and clinical chemistry evaluations.	On April 22, 2002, insufficient blood was collected from a 500 ppm group rat (D0061) for analysis of clinical chemistry parameters. Additionally, clotting of blood samples collected from control group rat (D0041) prevented WBC differentials, prothrombin time, and hematology for that animal.	None.
Section X. H. Page 16	Blood samples will be collected from all males and all toxicity group females on the day of scheduled sacrifice as a terminal procedure. While under Ketamine/xylazine anesthesia a syringe and needle will be used to collect blood samples for hematological and clinical chemistry evaluations.	On April 23, 2002, insufficient blood was collected from a 100 ppm group rat (D0099) for analysis of clinical chemistry parameters. Additionally, clotting of blood samples from the following list of animals prevented analysis of the indicated parameters. D0088 (Control) – WBC differentials, prothrombin time, hematology D0095 (100 ppm) – WBC differentials, prothrombin time, hematology D0097 (100 ppm) – prothrombin time D0099 (100 ppm) – prothrombin time D0104 (500 ppm) – prothrombin time D0108 (500 ppm) – WBC differentials, prothrombin time, hematology	None.
Section X.G. Page 15	The following organs/tissues will be taken and preserved in fixative: lungs (preserved by inflation with fixative and then immersion).	On April 23, 2002, the left lung from a toxicity group female (D0110) failed to inflate upon infusion of formalin.	None.
Section X.B.2. Page 8	Temperature and relative humidity will be maintained within the range of 22 ± 3°C and 50 ± 20%, respectively.	On April 29, 2002, the humidity in chambers 30908-E, 30909-E, and 30910-E was below the protocol specified range (28.3%, 29.4% and 29.8% respectively).	None.
Section X.G. Page 14	Feeder weights will be taken on lactation day 0 and 4.	Reproductive females D0002 (control) and D0039 (2500 ppm) had their feeders weighed on lactation day 1.	None.

Appendix A

Group Daily Mean Exposure Levels and Nominal Concentrations

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Males

Controls

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	BLQ	NA	NA	NA
Day 2	BLQ	NA	NA	NA
Day 3	BLQ	NA	NA	NA
Day 4	BLQ	NA	NA	NA
Day 5	BLQ	NA	NA	NA
Day 6	BLQ	NA	NA	NA
Day 7	BLQ	NA	NA	NA
Day 8	BLQ	NA	NA	NA
Day 9	BLQ	NA	NA	NA
Day 10	BLQ	NA	NA	NA
Day 11	BLQ	NA	NA	NA
Day 12	BLQ	NA	NA	NA
Day 13	BLQ	NA	NA	NA
Day 14	BLQ	NA	NA	NA
Day 15	BLQ	NA	NA	NA
Day 16	BLQ	NA	NA	NA
Day 17	BLQ	NA	NA	NA
Day 18	BLQ	NA	NA	NA
Day 19	BLQ	NA	NA	NA
Day 20	BLQ	NA	NA	NA
Day 21	BLQ	NA	NA	NA
Day 22	BLQ	NA	NA	NA
Day 23	BLQ	NA	NA	NA
Day 24	BLQ	NA	NA	NA
Day 25	BLQ	NA	NA	NA
Day 26	BLQ	NA	NA	NA
Day 27	BLQ	NA	NA	NA
Day 28	BLQ	NA	NA	NA
Mean	**BLQ**			
Std Dev.	**NA**			

NA = Not Applicable
[a] Percent difference actual to nominal
BLQ = Below Limit of Quantitation (0.01 - 3 ppm)

100 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	105	11.8	95	110.5
Day 2	98	1.6	90	108.9
Day 3	100	1.3	92	108.7
Day 4	96	1.8	90	106.7
Day 5	97	1.1	89	109.0
Day 6	97	1.3	91	106.6
Day 7	95	1.3	89	106.7
Day 8	105	1.4	89	118.0
Day 9	101	1.0	91	111.0
Day 10	96	2.1	89	107.9
Day 11	99	14.0	97	102.1
Day 12	103	2.5	104	99.0
Day 13	100	0.9	102	98.0
Day 14	103	1.7	103	100.0
Day 15	103	2.6	100	103.0
Day 16	99	1.3	100	99.0
Day 17	98	1.4	99	99.0
Day 18	97	1.1	99	98.0
Day 19	97	1.6	98	99.0
Day 20	98	1.2	100	98.0
Day 21	98	0.9	98	100.0
Day 22	97	1.8	99	98.0
Day 23	96	1.4	98	98.0
Day 24	98	1.0	101	97.0
Day 25	98	2.0	99	99.0
Day 26	96	1.7	101	95.0
Day 27	93	1.6	98	94.9
Day 28	95	2.6	104	91.3
Mean	**99**			
Std Dev.	**3.0**			

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Males

500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	490	4.4	481	101.9
Day 2	477	3.8	472	101.1
Day 3	504	10.8	497	101.4
Day 4	498	16.8	499	99.8
Day 5	508	6.9	502	101.2
Day 6	358	206.0	431	83.1
Day 7	494	3.1	507	97.4
Day 8	529	9.3	484	109.3
Day 9	538	11.0	483	111.4
Day 10	520	6.1	486	107.0
Day 11	472	138.3	457	103.3
Day 12	514	10.5	477	107.8
Day 13	510	7.7	479	106.5
Day 14	515	19.7	483	106.6
Day 15	507	17.8	463	109.5
Day 16	496	12.1	467	106.2
Day 17	492	17.6	478	102.9
Day 18	517	19.9	486	106.4
Day 19	501	19.3	469	106.8
Day 20	506	12.5	477	106.1
Day 21	495	4.2	476	104.0
Day 22	481	10.7	476	101.1
Day 23	491	13.9	474	103.6
Day 24	496	14.9	484	102.5
Day 25	489	8.5	473	103.4
Day 26	485	7.0	474	102.3
Day 27	499	14.4	497	100.4
Day 28	494	14.2	489	101.0
Mean	**496**			
Std Dev.	**30.8**			

[a] Percent difference actual to nominal

2500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	2174	868.2	2156	100.8
Day 2	2483	14.3	2524	98.4
Day 3	2482	26.2	2485	99.9
Day 4	2485	25.3	2479	100.2
Day 5	2465	18.6	2473	99.7
Day 6	2465	19.8	2530	97.4
Day 7	2478	53.2	2532	97.9
Day 8	2542	52.9	2380	106.8
Day 9	2624	21.2	2498	105.0
Day 10	2548	31.5	2406	105.9
Day 11	2568	21.9	2458	104.5
Day 12	2616	36.6	2455	106.6
Day 13	2585	20.9	2486	104.0
Day 14	2582	24.0	2458	105.0
Day 15	2605	11.7	2472	105.4
Day 16	2609	9.5	2458	106.1
Day 17	2481	167.2	2423	102.4
Day 18	2499	23.9	2403	104.0
Day 19	2545	31.0	2472	103.0
Day 20	2480	31.3	2364	104.9
Day 21	2532	24.1	2467	102.6
Day 22	2440	19.0	2384	102.3
Day 23	2435	12.5	2402	101.4
Day 24	2511	43.7	2301	109.1
Day 25	2460	14.7	2441	100.8
Day 26	2460	23.0	2449	100.4
Day 27	2490	35.0	2505	99.4
Day 28	2483	87.9	2548	97.4
Mean	**2505**			
Std Dev.	**85.9**			

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Toxicity Females

Controls

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	BLQ	NA	NA	NA
Day 2	BLQ	NA	NA	NA
Day 3	BLQ	NA	NA	NA
Day 4	BLQ	NA	NA	NA
Day 5	BLQ	NA	NA	NA
Day 6	BLQ	NA	NA	NA
Day 7	BLQ	NA	NA	NA
Day 8	BLQ	NA	NA	NA
Day 9	BLQ	NA	NA	NA
Day 10	BLQ	NA	NA	NA
Day 11	BLQ	NA	NA	NA
Day 12	BLQ	NA	NA	NA
Day 13	BLQ	NA	NA	NA
Day 14	BLQ	NA	NA	NA
Day 15	BLQ	NA	NA	NA
Day 16	BLQ	NA	NA	NA
Day 17	BLQ	NA	NA	NA
Day 18	BLQ	NA	NA	NA
Day 19	BLQ	NA	NA	NA
Day 20	BLQ	NA	NA	NA
Day 21	BLQ	NA	NA	NA
Day 22	BLQ	NA	NA	NA
Day 23	BLQ	NA	NA	NA
Day 24	BLQ	NA	NA	NA
Day 25	BLQ	NA	NA	NA
Day 26	BLQ	NA	NA	NA
Day 27	BLQ	NA	NA	NA
Day 28	BLQ	NA	NA	NA
Day 29	BLQ	NA	NA	NA
Mean	**BLQ**			
Std Dev.	**NA**			

NA = Not Applicable
[a] Percent difference actual to nominal
BLQ = Below Limit of Quantitation (0.01 - 3 ppm)

100 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	100	10.8	98	102.0
Day 2	100	1.3	98	102.0
Day 3	98	1.5	98	100.0
Day 4	98	10.2	97	101.0
Day 5	99	1.2	98	101.0
Day 6	96	1.2	97	99.0
Day 7	98	0.9	98	100.0
Day 8	103	1.9	95	108.4
Day 9	99	1.0	97	102.1
Day 10	97	1.5	98	99.0
Day 11	96	1.2	96	100.0
Day 12	101	1.7	101	100.0
Day 13	96	2.8	95	101.1
Day 14	98	1.3	97	101.0
Day 15	99	3.7	97	102.1
Day 16	100	7.4	100	100.0
Day 17	101	1.2	100	101.0
Day 18	100	2.0	99	101.0
Day 19	100	1.3	99	101.0
Day 20	100	1.9	101	99.0
Day 21	102	2.2	101	101.0
Day 22	102	1.4	103	99.0
Day 23	99	1.1	99	100.0
Day 24	101	1.2	102	99.0
Day 25	101	1.9	101	100.0
Day 26	101	1.6	103	98.1
Day 27	96	1.8	98	98.0
Day 28	97	2.0	100	97.0
Day 29	91	16.3	94	96.8
Mean	**99**			
Std Dev.	**2.5**			

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Toxicity Females

500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	527	8.3	534	98.7
Day 2	511	20.6	487	104.9
Day 3	525	7.2	530	99.1
Day 4	508	7.1	501	101.4
Day 5	523	2.2	523	100.0
Day 6	508	2.6	504	100.8
Day 7	514	2.4	515	99.8
Day 8	539	11.0	491	109.8
Day 9	552	3.6	508	108.7
Day 10	530	42.5	460	115.2
Day 11	532	8.5	477	111.5
Day 12	539	7.0	508	106.1
Day 13	513	6.3	492	104.3
Day 14	517	3.6	485	106.6
Day 15	514	3.4	482	106.6
Day 16	513	3.6	485	105.8
Day 17	499	3.2	478	104.4
Day 18	506	6.9	476	106.3
Day 19	507	5.4	480	105.6
Day 20	508	6.8	481	105.6
Day 21	512	10.9	490	104.5
Day 22	503	2.5	492	102.2
Day 23	494	2.2	476	103.8
Day 24	510	2.4	492	103.7
Day 25	516	4.3	492	104.9
Day 26	522	5.6	508	102.8
Day 27	516	6.7	513	100.6
Day 28	499	7.1	498	100.2
Day 29	478	10.5	493	97.0
Mean	**515**			
Std Dev.	**14.9**			

2500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	2480	30.1	2486	99.8
Day 2	2522	22.1	2585	97.6
Day 3	2480	23.2	2570	96.5
Day 4	2279	659.7	2459	92.7
Day 5	2538	6.5	2620	96.9
Day 6	2468	12.5	2603	94.8
Day 7	2515	5.7	2668	94.3
Day 8	2555	49.1	2471	103.4
Day 9	2636	50.7	2579	102.2
Day 10	2544	15.2	2505	101.6
Day 11	2591	31.2	2577	100.5
Day 12	2559	44.4	2525	101.3
Day 13	2537	34.7	2547	99.6
Day 14	2614	30.5	2591	100.9
Day 15	2606	8.4	2564	101.6
Day 16	2548	9.3	2535	100.5
Day 17	2583	18.7	2609	99.0
Day 18	2498	29.7	2512	99.4
Day 19	2510	29.9	2520	99.6
Day 20	2518	24.7	2558	98.4
Day 21	2462	18.1	2525	97.5
Day 22	2000	942.0	2300	87.0
Day 23	2511	47.8	2584	97.2
Day 24	2354	751.1	2450	96.1
Day 25	2449	142.2	2821	86.8
Day 26	2565	29.7	2639	97.2
Day 27	2508	23.3	2672	93.9
Day 28	2573	19.7	2730	94.2
Day 29	2455	79.8	2601	94.4
Mean	**2499**			
Std Dev.	**121.1**			

[a] Percent difference actual to nominal

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Reproductive Females Subgroup M1

Controls

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	BLQ	NA	NA	NA
Day 2	BLQ	NA	NA	NA
Day 3	BLQ	NA	NA	NA
Day 4	BLQ	NA	NA	NA
Day 5	BLQ	NA	NA	NA
Day 6	BLQ	NA	NA	NA
Day 7	BLQ	NA	NA	NA
Day 8	BLQ	NA	NA	NA
Day 9	BLQ	NA	NA	NA
Day 10	BLQ	NA	NA	NA
Day 11	BLQ	NA	NA	NA
Day 12	BLQ	NA	NA	NA
Day 13	BLQ	NA	NA	NA
Day 14	BLQ	NA	NA	NA
Day 15	BLQ	NA	NA	NA
Day 16	BLQ	NA	NA	NA
Day 17	BLQ	NA	NA	NA
Day 18	BLQ	NA	NA	NA
Day 19	BLQ	NA	NA	NA
Day 20	BLQ	NA	NA	NA
Day 21	BLQ	NA	NA	NA
Day 22	BLQ	NA	NA	NA
Day 23	BLQ	NA	NA	NA
Day 24	BLQ	NA	NA	NA
Day 25	BLQ	NA	NA	NA
Day 26	BLQ	NA	NA	NA
Day 27	BLQ	NA	NA	NA
Day 28	BLQ	NA	NA	NA
Day 29	BLQ	NA	NA	NA
Day 30	BLQ	NA	NA	NA
Day 31	BLQ	NA	NA	NA
Day 32	BLQ	NA	NA	NA
Day 33	BLQ	NA	NA	NA
Day 34	BLQ	NA	NA	NA
Mean	**BLQ**			
Std Dev.	**NA**			

NA = Not Applicable
[a] Percent difference actual to nominal
BLQ = Below Limit of Quantitation (0.01 - 3 ppm)

100 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	100	10.8	98	102.0
Day 2	100	1.3	98	102.0
Day 3	98	1.5	98	100.0
Day 4	98	10.2	97	101.0
Day 5	99	1.2	98	101.0
Day 6	96	1.2	97	99.0
Day 7	98	0.9	98	100.0
Day 8	103	1.9	95	108.4
Day 9	99	1.0	97	102.1
Day 10	97	1.5	98	99.0
Day 11	96	1.2	96	100.0
Day 12	101	1.7	101	100.0
Day 13	96	2.8	95	101.1
Day 14	98	1.3	97	101.0
Day 15	99	3.7	97	102.1
Day 16	100	7.4	100	100.0
Day 17	101	1.2	100	101.0
Day 18	100	2.0	99	101.0
Day 19	100	1.3	99	101.0
Day 20	100	1.9	101	99.0
Day 21	102	2.2	101	101.0
Day 22	102	1.4	103	99.0
Day 23	99	1.1	99	100.0
Day 24	101	1.2	102	99.0
Day 25	101	1.9	101	100.0
Day 26	101	1.6	103	98.1
Day 27	96	1.8	98	98.0
Day 28	97	2.0	100	97.0
Day 29	91	16.3	94	96.8
Day 30	99	3.6	101	98.0
Day 31	98	1.9	101	97.0
Day 32	101	1.2	102	99.0
Day 33	100	1.8	103	97.1
Day 34	101	1.4	105	96.2
Mean	**99**			
Std Dev.	**2.3**			

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Reproductive Females Subgroup M1

500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	527	8.3	534	98.7
Day 2	511	20.6	487	104.9
Day 3	525	7.2	530	99.1
Day 4	508	7.1	501	101.4
Day 5	523	2.2	523	100.0
Day 6	508	2.6	504	100.8
Day 7	514	2.4	515	99.8
Day 8	539	11.0	491	109.8
Day 9	552	3.6	508	108.7
Day 10	530	42.5	460	115.2
Day 11	532	8.5	477	111.5
Day 12	539	7.0	508	106.1
Day 13	513	6.3	492	104.3
Day 14	517	3.6	485	106.6
Day 15	514	3.4	482	106.6
Day 16	513	3.6	485	105.8
Day 17	499	3.2	478	104.4
Day 18	506	6.9	476	106.3
Day 19	507	5.4	480	105.6
Day 20	508	6.8	481	105.6
Day 21	512	10.9	490	104.5
Day 22	503	2.5	492	102.2
Day 23	494	2.2	476	103.8
Day 24	510	2.4	492	103.7
Day 25	516	4.3	492	104.9
Day 26	522	5.6	508	102.8
Day 27	516	6.7	513	100.6
Day 28	499	7.1	498	100.2
Day 29	478	10.5	493	97.0
Day 30	489	8.5	491	99.6
Day 31	503	22.6	496	101.4
Day 32	493	11.8	507	97.2
Day 33	490	15.8	509	96.3
Day 34	495	12.6	517	95.7
Mean	**512**			
Std Dev.	**15.8**			

2500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	2480	30.1	2486	99.8
Day 2	2522	22.1	2585	97.6
Day 3	2480	23.2	2570	96.5
Day 4	2279	659.7	2459	92.7
Day 5	2538	6.5	2620	96.9
Day 6	2468	12.5	2603	94.8
Day 7	2515	5.7	2668	94.3
Day 8	2555	49.1	2471	103.4
Day 9	2636	50.7	2579	102.2
Day 10	2544	15.2	2505	101.6
Day 11	2591	31.2	2577	100.5
Day 12	2559	44.4	2525	101.3
Day 13	2537	34.7	2547	99.6
Day 14	2614	30.5	2591	100.9
Day 15	2606	8.4	2564	101.6
Day 16	2548	9.3	2535	100.5
Day 17	2583	18.7	2609	99.0
Day 18	2498	29.7	2512	99.4
Day 19	2510	29.9	2520	99.6
Day 20	2518	24.7	2558	98.4
Day 21	2462	18.1	2525	97.5
Day 22	2000	942.0	2300	87.0
Day 23	2511	47.8	2584	97.2
Day 24	2354	751.1	2450	96.1
Day 25	2449	142.2	2821	86.8
Day 26	2565	29.7	2639	97.2
Day 27	2508	23.3	2672	93.9
Day 28	2573	19.7	2730	94.2
Day 29	2455	79.8	2601	94.4
Day 30	2424	41.2	2522	96.1
Day 31	2478	73.4	2572	96.3
Day 32	2495	80.4	2624	95.1
Day 33	2413	101.1	2555	94.4
Day 34	2485	44.1	2652	93.7
Mean	**2493**			
Std Dev.	**113.2**			

[a] Percent difference actual to nominal

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Reproductive Females Subgroup M2

Controls

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	BLQ	NA	NA	NA
Day 2	BLQ	NA	NA	NA
Day 3	BLQ	NA	NA	NA
Day 4	BLQ	NA	NA	NA
Day 5	BLQ	NA	NA	NA
Day 6	BLQ	NA	NA	NA
Day 7	BLQ	NA	NA	NA
Day 8	BLQ	NA	NA	NA
Day 9	BLQ	NA	NA	NA
Day 10	BLQ	NA	NA	NA
Day 11	BLQ	NA	NA	NA
Day 12	BLQ	NA	NA	NA
Day 13	BLQ	NA	NA	NA
Day 14	BLQ	NA	NA	NA
Day 15	BLQ	NA	NA	NA
Day 16	BLQ	NA	NA	NA
Day 17	BLQ	NA	NA	NA
Day 18	BLQ	NA	NA	NA
Day 19	BLQ	NA	NA	NA
Day 20	BLQ	NA	NA	NA
Day 21	BLQ	NA	NA	NA
Day 22	BLQ	NA	NA	NA
Day 23	BLQ	NA	NA	NA
Day 24	BLQ	NA	NA	NA
Day 25	BLQ	NA	NA	NA
Day 26	BLQ	NA	NA	NA
Day 27	BLQ	NA	NA	NA
Day 28	BLQ	NA	NA	NA
Day 29	BLQ	NA	NA	NA
Day 30	BLQ	NA	NA	NA
Day 31	BLQ	NA	NA	NA
Day 32	BLQ	NA	NA	NA
Day 33	BLQ	NA	NA	NA
Day 34	BLQ	NA	NA	NA
Day 35	BLQ	NA	NA	NA
Mean	**BLQ**			
Std Dev.	**NA**			

NA = Not Applicable
[a] Percent difference actual to nominal
BLQ = Below Limit of Quantitation (0.01 - 3 ppm)

100 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	100	10.8	98	102.0
Day 2	100	1.3	98	102.0
Day 3	98	1.5	98	100.0
Day 4	98	10.2	97	101.0
Day 5	99	1.2	98	101.0
Day 6	96	1.2	97	99.0
Day 7	98	0.9	98	100.0
Day 8	103	1.9	95	108.4
Day 9	99	1.0	97	102.1
Day 10	97	1.5	98	99.0
Day 11	96	1.2	96	100.0
Day 12	101	1.7	101	100.0
Day 13	96	2.8	95	101.1
Day 14	98	1.3	97	101.0
Day 15	99	3.7	97	102.1
Day 16	100	7.4	100	100.0
Day 17	101	1.2	100	101.0
Day 18	100	2.0	99	101.0
Day 19	100	1.3	99	101.0
Day 20	100	1.9	101	99.0
Day 21	102	2.2	101	101.0
Day 22	102	1.4	103	99.0
Day 23	99	1.1	99	100.0
Day 24	101	1.2	102	99.0
Day 25	101	1.9	101	100.0
Day 26	101	1.6	103	98.1
Day 27	96	1.8	98	98.0
Day 28	97	2.0	100	97.0
Day 29	91	16.3	94	96.8
Day 30	99	3.6	101	98.0
Day 31	98	1.9	101	97.0
Day 32	101	1.2	102	99.0
Day 33	100	1.8	103	97.1
Day 34	101	1.4	105	96.2
Day 35	100	1.6	102	98.0
Mean	**99**			
Std Dev.	**2.3**			

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Reproductive Females Subgroup M2

500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	527	8.3	534	98.7
Day 2	511	20.6	487	104.9
Day 3	525	7.2	530	99.1
Day 4	508	7.1	501	101.4
Day 5	523	2.2	523	100.0
Day 6	508	2.6	504	100.8
Day 7	514	2.4	515	99.8
Day 8	539	11.0	491	109.8
Day 9	552	3.6	508	108.7
Day 10	530	42.5	460	115.2
Day 11	532	8.5	477	111.5
Day 12	539	7.0	508	106.1
Day 13	513	6.3	492	104.3
Day 14	517	3.6	485	106.6
Day 15	514	3.4	482	106.6
Day 16	513	3.6	485	105.8
Day 17	499	3.2	478	104.4
Day 18	506	6.9	476	106.3
Day 19	507	5.4	480	105.6
Day 20	508	6.8	481	105.6
Day 21	512	10.9	490	104.5
Day 22	503	2.5	492	102.2
Day 23	494	2.2	476	103.8
Day 24	510	2.4	492	103.7
Day 25	516	4.3	492	104.9
Day 26	522	5.6	508	102.8
Day 27	516	6.7	513	100.6
Day 28	499	7.1	498	100.2
Day 29	478	10.5	493	97.0
Day 30	489	8.5	491	99.6
Day 31	503	22.6	496	101.4
Day 32	493	11.8	507	97.2
Day 33	490	15.8	509	96.3
Day 34	495	12.6	517	95.7
Day 35	507	10.2	505	100.4
Mean	**512**			
Std Dev.	**15.6**			

2500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	2480	30.1	2486	99.8
Day 2	2522	22.1	2585	97.6
Day 3	2480	23.2	2570	96.5
Day 4	2279	659.7	2459	92.7
Day 5	2538	6.5	2620	96.9
Day 6	2468	12.5	2603	94.8
Day 7	2515	5.7	2668	94.3
Day 8	2555	49.1	2471	103.4
Day 9	2636	50.7	2579	102.2
Day 10	2544	15.2	2505	101.6
Day 11	2591	31.2	2577	100.5
Day 12	2559	44.4	2525	101.3
Day 13	2537	34.7	2547	99.6
Day 14	2614	30.5	2591	100.9
Day 15	2606	8.4	2564	101.6
Day 16	2548	9.3	2535	100.5
Day 17	2583	18.7	2609	99.0
Day 18	2498	29.7	2512	99.4
Day 19	2510	29.9	2520	99.6
Day 20	2518	24.7	2558	98.4
Day 21	2462	18.1	2525	97.5
Day 22	2000	942.0	2300	87.0
Day 23	2511	47.8	2584	97.2
Day 24	2354	751.1	2450	96.1
Day 25	2449	142.2	2821	86.8
Day 26	2565	29.7	2639	97.2
Day 27	2508	23.3	2672	93.9
Day 28	2573	19.7	2730	94.2
Day 29	2455	79.8	2601	94.4
Day 30	2424	41.2	2522	96.1
Day 31	2478	73.4	2572	96.3
Day 32	2495	80.4	2624	95.1
Day 33	2413	101.1	2555	94.4
Day 34	2485	44.1	2652	93.7
Day 35	2467	46.2	2522	97.8
Mean	**2492**			
Std Dev.	**111.6**			

[a] Percent difference actual to nominal

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Reproductive Females Subgroup M3

Controls

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	BLQ	NA	NA	NA
Day 2	BLQ	NA	NA	NA
Day 3	BLQ	NA	NA	NA
Day 4	BLQ	NA	NA	NA
Day 5	BLQ	NA	NA	NA
Day 6	BLQ	NA	NA	NA
Day 7	BLQ	NA	NA	NA
Day 8	BLQ	NA	NA	NA
Day 9	BLQ	NA	NA	NA
Day 10	BLQ	NA	NA	NA
Day 11	BLQ	NA	NA	NA
Day 12	BLQ	NA	NA	NA
Day 13	BLQ	NA	NA	NA
Day 14	BLQ	NA	NA	NA
Day 15	BLQ	NA	NA	NA
Day 16	BLQ	NA	NA	NA
Day 17	BLQ	NA	NA	NA
Day 18	BLQ	NA	NA	NA
Day 19	BLQ	NA	NA	NA
Day 20	BLQ	NA	NA	NA
Day 21	BLQ	NA	NA	NA
Day 22	BLQ	NA	NA	NA
Day 23	BLQ	NA	NA	NA
Day 24	BLQ	NA	NA	NA
Day 25	BLQ	NA	NA	NA
Day 26	BLQ	NA	NA	NA
Day 27	BLQ	NA	NA	NA
Day 28	BLQ	NA	NA	NA
Day 29	BLQ	NA	NA	NA
Day 30	BLQ	NA	NA	NA
Day 31	BLQ	NA	NA	NA
Day 32	BLQ	NA	NA	NA
Day 33	BLQ	NA	NA	NA
Day 34	BLQ	NA	NA	NA
Day 35	BLQ	NA	NA	NA
Day 36	BLQ	NA	NA	NA
Mean	**BLQ**			
Std Dev.	**NA**			

100 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	100	10.8	98	102.0
Day 2	100	1.3	98	102.0
Day 3	98	1.5	98	100.0
Day 4	98	10.2	97	101.0
Day 5	99	1.2	98	101.0
Day 6	96	1.2	97	99.0
Day 7	98	0.9	98	100.0
Day 8	103	1.9	95	108.4
Day 9	99	1.0	97	102.1
Day 10	97	1.5	98	99.0
Day 11	96	1.2	96	100.0
Day 12	101	1.7	101	100.0
Day 13	96	2.8	95	101.1
Day 14	98	1.3	97	101.0
Day 15	99	3.7	97	102.1
Day 16	100	7.4	100	100.0
Day 17	101	1.2	100	101.0
Day 18	100	2.0	99	101.0
Day 19	100	1.3	99	101.0
Day 20	100	1.9	101	99.0
Day 21	102	2.2	101	101.0
Day 22	102	1.4	103	99.0
Day 23	99	1.1	99	100.0
Day 24	101	1.2	102	99.0
Day 25	101	1.9	101	100.0
Day 26	101	1.6	103	98.1
Day 27	96	1.8	98	98.0
Day 28	97	2.0	100	97.0
Day 29	91	16.3	94	96.8
Day 30	99	3.6	101	98.0
Day 31	98	1.9	101	97.0
Day 32	101	1.2	102	99.0
Day 33	100	1.8	103	97.1
Day 34	101	1.4	105	96.2
Day 35	100	1.6	102	98.0
Day 36	99	1.2	102	97.1
Mean	**99**			
Std Dev.	**2.3**			

NA = Not Applicable
[a] Percent difference actual to nominal
BLQ = Below Limit of Quantitation (0.01 - 3 ppm)

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Reproductive Females Subgroup M3

500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	527	8.3	534	98.7
Day 2	511	20.6	487	104.9
Day 3	525	7.2	530	99.1
Day 4	508	7.1	501	101.4
Day 5	523	2.2	523	100.0
Day 6	508	2.6	504	100.8
Day 7	514	2.4	515	99.8
Day 8	539	11.0	491	109.8
Day 9	552	3.6	508	108.7
Day 10	530	42.5	460	115.2
Day 11	532	8.5	477	111.5
Day 12	539	7.0	508	106.1
Day 13	513	6.3	492	104.3
Day 14	517	3.6	485	106.6
Day 15	514	3.4	482	106.6
Day 16	513	3.6	485	105.8
Day 17	499	3.2	478	104.4
Day 18	506	6.9	476	106.3
Day 19	507	5.4	480	105.6
Day 20	508	6.8	481	105.6
Day 21	512	10.9	490	104.5
Day 22	503	2.5	492	102.2
Day 23	494	2.2	476	103.8
Day 24	510	2.4	492	103.7
Day 25	516	4.3	492	104.9
Day 26	522	5.6	508	102.8
Day 27	516	6.7	513	100.6
Day 28	499	7.1	498	100.2
Day 29	478	10.5	493	97.0
Day 30	489	8.5	491	99.6
Day 31	503	22.6	496	101.4
Day 32	493	11.8	507	97.2
Day 33	490	15.8	509	96.3
Day 34	495	12.6	517	95.7
Day 35	507	10.2	505	100.4
Day 36	497	12.5	507	98.0
Mean	**511**			
Std Dev.	**15.5**			

2500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	2480	30.1	2486	99.8
Day 2	2522	22.1	2585	97.6
Day 3	2480	23.2	2570	96.5
Day 4	2279	659.7	2459	92.7
Day 5	2538	6.5	2620	96.9
Day 6	2468	12.5	2603	94.8
Day 7	2515	5.7	2668	94.3
Day 8	2555	49.1	2471	103.4
Day 9	2636	50.7	2579	102.2
Day 10	2544	15.2	2505	101.6
Day 11	2591	31.2	2577	100.5
Day 12	2559	44.4	2525	101.3
Day 13	2537	34.7	2547	99.6
Day 14	2614	30.5	2591	100.9
Day 15	2606	8.4	2564	101.6
Day 16	2548	9.3	2535	100.5
Day 17	2583	18.7	2609	99.0
Day 18	2498	29.7	2512	99.4
Day 19	2510	29.9	2520	99.6
Day 20	2518	24.7	2558	98.4
Day 21	2462	18.1	2525	97.5
Day 22	2000	942.0	2300	87.0
Day 23	2511	47.8	2584	97.2
Day 24	2354	751.1	2450	96.1
Day 25	2449	142.2	2821	86.8
Day 26	2565	29.7	2639	97.2
Day 27	2508	23.3	2672	93.9
Day 28	2573	19.7	2730	94.2
Day 29	2455	79.8	2601	94.4
Day 30	2424	41.2	2522	96.1
Day 31	2478	73.4	2572	96.3
Day 32	2495	80.4	2624	95.1
Day 33	2413	101.1	2555	94.4
Day 34	2485	44.1	2652	93.7
Day 35	2467	46.2	2522	97.8
Day 36	2505	49.2	2598	96.4
Mean	**2492**			
Std Dev.	**110.0**			

[a] Percent difference actual to nominal

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Reproductive Females Subgroup M4

Controls

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	BLQ	NA	NA	NA
Day 2	BLQ	NA	NA	NA
Day 3	BLQ	NA	NA	NA
Day 4	BLQ	NA	NA	NA
Day 5	BLQ	NA	NA	NA
Day 6	BLQ	NA	NA	NA
Day 7	BLQ	NA	NA	NA
Day 8	BLQ	NA	NA	NA
Day 9	BLQ	NA	NA	NA
Day 10	BLQ	NA	NA	NA
Day 11	BLQ	NA	NA	NA
Day 12	BLQ	NA	NA	NA
Day 13	BLQ	NA	NA	NA
Day 14	BLQ	NA	NA	NA
Day 15	BLQ	NA	NA	NA
Day 16	BLQ	NA	NA	NA
Day 17	BLQ	NA	NA	NA
Day 18	BLQ	NA	NA	NA
Day 19	BLQ	NA	NA	NA
Day 20	BLQ	NA	NA	NA
Day 21	BLQ	NA	NA	NA
Day 22	BLQ	NA	NA	NA
Day 23	BLQ	NA	NA	NA
Day 24	BLQ	NA	NA	NA
Day 25	BLQ	NA	NA	NA
Day 26	BLQ	NA	NA	NA
Day 27	BLQ	NA	NA	NA
Day 28	BLQ	NA	NA	NA
Day 29	BLQ	NA	NA	NA
Day 30	BLQ	NA	NA	NA
Day 31	BLQ	NA	NA	NA
Day 32	BLQ	NA	NA	NA
Day 33	BLQ	NA	NA	NA
Day 34	BLQ	NA	NA	NA
Day 35	BLQ	NA	NA	NA
Day 36	BLQ	NA	NA	NA
Day 37	BLQ	NA	NA	NA
Mean	**BLQ**			
Std Dev.	**NA**			

NA = Not Applicable
[a] Percent difference actual to nominal
BLQ = Below Limit of Quantitation (0.01 - 3 ppm)

100 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	100	10.8	98	102.0
Day 2	100	1.3	98	102.0
Day 3	98	1.5	98	100.0
Day 4	98	10.2	97	101.0
Day 5	99	1.2	98	101.0
Day 6	96	1.2	97	99.0
Day 7	98	0.9	98	100.0
Day 8	103	1.9	95	108.4
Day 9	99	1.0	97	102.1
Day 10	97	1.5	98	99.0
Day 11	96	1.2	96	100.0
Day 12	101	1.7	101	100.0
Day 13	96	2.8	95	101.1
Day 14	98	1.3	97	101.0
Day 15	99	3.7	97	102.1
Day 16	100	7.4	100	100.0
Day 17	101	1.2	100	101.0
Day 18	100	2.0	99	101.0
Day 19	100	1.3	99	101.0
Day 20	100	1.9	101	99.0
Day 21	102	2.2	101	101.0
Day 22	102	1.4	103	99.0
Day 23	99	1.1	99	100.0
Day 24	101	1.2	102	99.0
Day 25	101	1.9	101	100.0
Day 26	101	1.6	103	98.1
Day 27	96	1.8	98	98.0
Day 28	97	2.0	100	97.0
Day 29	91	16.3	94	96.8
Day 30	99	3.6	101	98.0
Day 31	98	1.9	101	97.0
Day 32	101	1.2	102	99.0
Day 33	100	1.8	103	97.1
Day 34	101	1.4	105	96.2
Day 35	100	1.6	102	98.0
Day 36	99	1.2	102	97.1
Day 37	101	0.9	104	97.1
Mean	**99**			
Std Dev.	**2.3**			

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Reproductive Females Subgroup M4

500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	527	8.3	534	98.7
Day 2	511	20.6	487	104.9
Day 3	525	7.2	530	99.1
Day 4	508	7.1	501	101.4
Day 5	523	2.2	523	100.0
Day 6	508	2.6	504	100.8
Day 7	514	2.4	515	99.8
Day 8	539	11.0	491	109.8
Day 9	552	3.6	508	108.7
Day 10	530	42.5	460	115.2
Day 11	532	8.5	477	111.5
Day 12	539	7.0	508	106.1
Day 13	513	6.3	492	104.3
Day 14	517	3.6	485	106.6
Day 15	514	3.4	482	106.6
Day 16	513	3.6	485	105.8
Day 17	499	3.2	478	104.4
Day 18	506	6.9	476	106.3
Day 19	507	5.4	480	105.6
Day 20	508	6.8	481	105.6
Day 21	512	10.9	490	104.5
Day 22	503	2.5	492	102.2
Day 23	494	2.2	476	103.8
Day 24	510	2.4	492	103.7
Day 25	516	4.3	492	104.9
Day 26	522	5.6	508	102.8
Day 27	516	6.7	513	100.6
Day 28	499	7.1	498	100.2
Day 29	478	10.5	493	97.0
Day 30	489	8.5	491	99.6
Day 31	503	22.6	496	101.4
Day 32	493	11.8	507	97.2
Day 33	490	15.8	509	96.3
Day 34	495	12.6	517	95.7
Day 35	507	10.2	505	100.4
Day 36	497	12.5	507	98.0
Day 37	487	16.0	498	97.8
Mean	**511**			
Std Dev.	**15.8**			

2500 ppm

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	2480	30.1	2486	99.8
Day 2	2522	22.1	2585	97.6
Day 3	2480	23.2	2570	96.5
Day 4	2279	659.7	2459	92.7
Day 5	2538	6.5	2620	96.9
Day 6	2468	12.5	2603	94.8
Day 7	2515	5.7	2668	94.3
Day 8	2555	49.1	2471	103.4
Day 9	2636	50.7	2579	102.2
Day 10	2544	15.2	2505	101.6
Day 11	2591	31.2	2577	100.5
Day 12	2559	44.4	2525	101.3
Day 13	2537	34.7	2547	99.6
Day 14	2614	30.5	2591	100.9
Day 15	2606	8.4	2564	101.6
Day 16	2548	9.3	2535	100.5
Day 17	2583	18.7	2609	99.0
Day 18	2498	29.7	2512	99.4
Day 19	2510	29.9	2520	99.6
Day 20	2518	24.7	2558	98.4
Day 21	2462	18.1	2525	97.5
Day 22	2000	942.0	2300	87.0
Day 23	2511	47.8	2584	97.2
Day 24	2354	751.1	2450	96.1
Day 25	2449	142.2	2821	86.8
Day 26	2565	29.7	2639	97.2
Day 27	2508	23.3	2672	93.9
Day 28	2573	19.7	2730	94.2
Day 29	2455	79.8	2601	94.4
Day 30	2424	41.2	2522	96.1
Day 31	2478	73.4	2572	96.3
Day 32	2495	80.4	2624	95.1
Day 33	2413	101.1	2555	94.4
Day 34	2485	44.1	2652	93.7
Day 35	2467	46.2	2522	97.8
Day 36	2505	49.2	2598	96.4
Day 37	2473	38.9	2573	96.1
Mean	**2492**			
Std Dev.	**108.5**			

[a] Percent difference actual to nominal

Appendix A
Group Daily Mean Exposure Levels and Nominal Concentrations

Reproductive Females Subgroup M6
Controls

	Mean Exposure Concentration (ppm D3)	Std Dev	Nominal Exposure Concentration (ppm D3)	%[a]
Day 1	BLQ	NA	NA	NA
Day 2	BLQ	NA	NA	NA
Day 3	BLQ	NA	NA	NA
Day 4	BLQ	NA	NA	NA
Day 5	BLQ	NA	NA	NA
Day 6	BLQ	NA	NA	NA
Day 7	BLQ	NA	NA	NA
Day 8	BLQ	NA	NA	NA
Day 9	BLQ	NA	NA	NA
Day 10	BLQ	NA	NA	NA
Day 11	BLQ	NA	NA	NA
Day 12	BLQ	NA	NA	NA
Day 13	BLQ	NA	NA	NA
Day 14	BLQ	NA	NA	NA
Day 15	BLQ	NA	NA	NA
Day 16	BLQ	NA	NA	NA
Day 17	BLQ	NA	NA	NA
Day 18	BLQ	NA	NA	NA
Day 19	BLQ	NA	NA	NA
Day 20	BLQ	NA	NA	NA
Day 21	BLQ	NA	NA	NA
Day 22	BLQ	NA	NA	NA
Day 23	BLQ	NA	NA	NA
Day 24	BLQ	NA	NA	NA
Day 25	BLQ	NA	NA	NA
Day 26	BLQ	NA	NA	NA
Day 27	BLQ	NA	NA	NA
Day 28	BLQ	NA	NA	NA
Day 29	BLQ	NA	NA	NA
Day 30	BLQ	NA	NA	NA
Day 31	BLQ	NA	NA	NA
Day 32	BLQ	NA	NA	NA
Day 33	BLQ	NA	NA	NA
Day 34	BLQ	NA	NA	NA
Day 35	BLQ	NA	NA	NA
Day 36	BLQ	NA	NA	NA
Day 37	BLQ	NA	NA	NA
Day 38	BLQ	NA	NA	NA
Day 39	BLQ	NA	NA	NA
Mean	**BLQ**			
Std Dev.	**NA**			

NA = Not Applicable
[a] Percent difference actual to nominal
BLQ = Below Limit of Quantitation (0.01 - 3 ppm)

Appendix B

Daily Exposure Chamber Environmental Conditions

Appendix B
Daily Exposure Chamber Environmental Conditions

Males Controls

Exposure	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Day	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	25.6	0.90	26.1	22.9	45.1	6.12	50.7	33.7	108	1.1	109	105
2	25.7	0.96	26.3	22.8	43.3	2.51	47.0	39.9	101	0.7	102	100
3	25.7	0.92	26.3	22.9	42.5	3.16	48.7	37.6	104	1.5	108	102
4	25.6	0.94	26.2	22.8	42.8	2.57	48.8	40.8	107	0.7	108	106
5	25.7	0.95	26.3	22.9	45.2	2.65	48.9	39.9	105	0.5	106	104
6	25.3	0.87	25.7	22.5	46.8	3.35	50.1	39.7	105	0.4	106	105
7	25.6	0.94	26.1	22.7	44.2	3.56	51.7	40.2	107	0.4	107	106
8	26.0	1.22	26.7	22.3	46.1	3.12	49.1	37.4	108	0.7	109	107
9	25.8	1.15	26.5	22.2	45.0	2.93	53.2	41.1	105	0.8	106	104
10	25.8	0.88	26.4	23.4	46.4	2.87	53.2	43.2	106	0.5	107	106
11	25.0	0.70	25.5	23.0	40.9	2.39	44.8	35.7	105	0.7	107	105
12	25.0	0.57	25.3	23.3	43.4	2.92	48.0	36.7	102	0.7	104	102
13	24.6	0.65	25.3	23.0	43.7	2.57	49.4	40.8	107	0.9	108	106
14	25.0	0.58	25.5	23.5	46.1	2.36	48.9	42.3	106	0.6	107	105
15	25.1	0.72	25.6	23.0	40.4	2.55	44.4	37.0	107	0.4	108	106
16	25.2	0.74	25.7	23.3	43.5	2.92	48.2	39.5	105	0.4	106	105
17	24.8	0.63	25.6	23.2	43.7	4.44	52.9	38.4	106	0.7	107	105
18	25.0	0.83	25.8	23.0	46.3	3.48	50.5	37.0	108	1.0	109	106
19	25.1	0.76	25.6	23.2	50.6	3.64	59.3	47.7	106	1.1	108	105
20	24.7	0.59	25.3	23.1	44.8	2.98	50.9	41.5	106	1.2	107	104
21	25.7	0.69	26.4	23.9	49.9	4.31	60.2	43.9	104	0.3	105	104
22	25.9	0.83	26.3	23.4	48.7	4.74	62.1	44.9	104	0.5	105	104
23	26.0	0.98	26.8	23.3	51.0	3.61	61.4	48.4	106	0.6	107	105
24	25.7	1.00	26.2	22.5	50.5	4.15	62.8	47.8	105	0.4	106	105
25	25.5	0.75	25.9	23.2	50.5	2.45	56.9	47.9	105	1.6	109	104
26	25.4	0.67	25.9	23.5	46.8	4.34	56.2	42.5	108	1.4	111	107
27	25.8	0.86	26.4	23.2	44.9	4.55	55.7	38.5	107	3.4	114	104
28	25.7	0.89	26.3	23.1	50.6	2.93	57.6	46.6	105	0.9	107	104

Appendix B
Daily Exposure Chamber Environmental Conditions

Males 100 ppm

Exposure Day	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	25.4	0.80	25.8	23.0	42.4	6.16	47.5	30.8	111	1.3	112	108
2	26.0	1.00	26.8	23.0	38.9	2.04	42.2	35.1	112	0.5	113	112
3	25.5	0.94	26.1	22.6	38.5	1.96	41.6	36.4	110	1.2	113	109
4	25.5	1.01	26.1	22.4	37.5	1.17	40.4	36.3	113	0.8	114	112
5	25.5	0.72	25.9	23.4	40.9	3.50	44.5	34.4	114	0.5	114	113
6	25.3	0.89	25.8	22.5	42.7	3.31	47.3	36.5	112	0.7	114	112
7	25.2	0.89	25.7	22.5	39.7	2.34	43.8	36.4	113	0.4	114	113
8	26.5	1.28	27.3	22.7	39.1	3.11	42.2	31.9	113	1.2	116	112
9	25.3	1.03	25.9	22.2	40.9	1.84	44.4	37.9	111	1.0	112	109
10	25.4	0.78	25.9	23.2	42.1	2.79	46.4	35.2	115	0.8	117	114
11	24.7	0.75	25.3	22.8	39.1	2.16	43.8	35.1	103	5.9	114	98
12	24.6	0.70	25.1	22.7	40.5	3.02	45.4	35.6	99	0.5	100	99
13	24.7	0.84	25.5	22.8	40.4	2.39	45.6	38.0	101	0.9	103	99
14	25.0	0.78	25.6	23.0	43.8	1.77	46.6	40.0	100	0.5	101	99
15	24.8	0.60	25.2	23.0	38.0	2.49	41.8	34.1	101	1.8	103	99
16	24.8	0.74	25.4	23.0	40.7	2.73	45.4	37.1	102	0.3	102	101
17	24.6	0.61	25.3	23.1	39.8	3.14	44.5	34.8	103	0.7	104	102
18	24.5	0.79	25.2	22.6	45.1	3.02	49.7	37.5	103	1.0	105	102
19	24.8	0.76	25.5	23.0	48.4	2.28	53.6	46.6	104	1.1	107	103
20	24.5	0.67	25.1	22.7	43.5	3.18	50.0	39.7	103	1.0	104	101
21	25.6	0.67	26.2	23.7	46.6	3.36	53.7	42.0	103	0.5	103	102
22	25.0	0.75	25.6	22.9	47.9	3.53	57.3	44.7	103	1.1	105	102
23	26.2	1.12	27.0	22.9	47.3	3.41	57.6	45.2	103	0.9	104	101
24	25.4	1.04	25.9	22.1	48.6	4.67	63.6	46.3	102	0.8	103	101
25	25.4	0.68	25.8	23.3	47.6	1.69	52.5	46.1	102	1.7	105	101
26	25.6	0.68	26.0	23.6	45.6	3.95	54.4	40.9	102	1.8	104	100
27	25.8	0.84	26.3	23.2	38.8	4.23	50.7	33.1	104	1.4	106	101
28	25.5	0.82	26.0	23.1	42.3	3.15	48.3	37.6	98	3.6	107	95

Appendix B
Daily Exposure Chamber Environmental Conditions

Males 500 ppm

Exposure	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Day	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	25.7	0.83	26.2	23.2	43.4	5.86	50.1	33.2	106	0.9	107	104
2	25.2	0.71	25.7	23.1	43.4	2.23	46.3	39.9	107	0.4	108	106
3	26.4	1.05	27.1	23.1	38.0	5.08	51.7	32.5	102	2.3	107	100
4	26.4	1.05	27.0	23.1	38.8	2.32	43.3	36.1	102	3.0	107	100
5	26.3	1.02	26.9	23.2	41.3	1.71	44.9	38.1	101	0.3	101	100
6	25.6	0.75	26.1	23.0	44.4	2.79	48.1	39.9	99	0.9	101	98
7	26.4	1.14	27.0	22.8	40.1	4.36	51.7	34.1	100	0.5	101	100
8	26.5	1.05	27.1	23.3	42.0	2.94	44.5	33.9	104	1.3	105	101
9	26.3	1.16	27.0	22.7	41.2	3.33	49.5	38.2	102	1.0	104	100
10	25.9	0.89	26.5	23.4	42.6	3.05	49.1	38.1	102	1.3	104	100
11	25.0	0.67	25.5	23.3	39.5	2.21	43.3	34.9	103	3.8	108	99
12	25.3	0.78	25.9	23.1	42.3	2.67	47.5	38.5	106	0.9	106	104
13	24.8	0.72	25.4	23.0	43.2	2.80	49.1	38.7	106	1.0	108	104
14	25.0	0.65	25.5	23.3	45.4	2.95	50.1	40.5	105	0.8	107	104
15	25.4	0.68	25.9	23.4	41.3	3.24	47.5	36.5	107	0.5	107	106
16	25.8	0.86	26.3	23.4	43.3	2.91	48.7	39.1	107	0.7	108	106
17	25.0	0.61	25.7	23.4	42.2	4.34	51.7	37.5	105	0.6	106	104
18	25.4	0.77	26.0	23.5	46.8	2.47	49.5	42.3	103	2.1	108	101
19	25.8	0.77	26.4	23.6	48.3	4.70	61.3	44.7	107	1.8	109	105
20	25.1	0.73	25.7	23.1	45.0	3.72	53.3	40.7	105	1.6	106	102
21	26.1	0.80	26.7	23.8	46.0	4.15	57.1	41.5	105	0.6	106	104
22	25.9	0.87	26.5	23.3	45.8	4.75	59.9	42.1	105	0.8	106	104
23	25.8	0.94	26.5	23.0	48.3	3.82	60.3	46.1	104	1.4	108	103
24	26.5	1.22	27.1	22.5	47.5	6.00	66.3	43.9	101	1.9	105	100
25	26.1	0.78	26.6	23.7	46.1	2.84	54.3	43.7	105	0.7	106	104
26	25.9	0.71	26.4	23.8	44.3	4.71	55.9	40.1	105	0.8	106	104
27	25.9	0.71	26.4	23.7	41.9	4.21	50.7	36.5	100	1.9	104	98
28	26.2	0.79	26.8	23.9	46.7	3.34	51.9	40.3	99	1.4	103	98

Appendix B
Daily Exposure Chamber Environmental Conditions

Males 2500 ppm

Exposure	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Day	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	25.5	0.97	26.5	22.9	46.1	5.15	52.0	35.6	105	1.5	107	102
2	25.7	1.11	26.4	22.5	44.3	2.32	47.4	39.9	104	0.6	105	103
3	25.5	0.85	26.1	22.9	40.1	2.63	44.8	36.8	105	0.9	106	104
4	26.0	1.14	26.7	22.6	41.7	1.84	46.1	39.8	105	1.6	108	103
5	26.0	0.88	26.5	23.4	42.5	2.82	47.9	36.0	105	1.1	107	104
6	25.7	0.92	26.2	22.8	46.8	3.71	50.9	39.9	103	1.4	105	102
7	25.7	0.99	26.1	22.6	42.9	1.75	45.4	39.0	103	2.8	106	99
8	26.5	1.16	27.3	23.1	43.5	2.02	45.6	37.7	109	2.4	111	104
9	26.0	1.12	26.8	22.8	42.3	2.43	46.6	39.4	104	1.1	105	102
10	26.2	0.91	26.9	23.9	42.2	3.20	48.8	36.6	108	0.5	108	107
11	25.1	0.79	25.6	23.0	38.4	2.85	44.2	33.9	106	1.0	108	105
12	25.0	0.84	25.7	22.8	41.1	3.64	47.7	34.6	106	0.8	106	104
13	24.8	0.80	25.5	22.9	41.6	2.25	47.5	39.7	105	1.3	108	103
14	25.1	0.67	25.6	23.5	44.3	2.48	48.3	40.1	106	0.5	107	105
15	25.3	0.80	25.9	23.1	39.6	3.68	48.3	36.2	106	0.6	107	105
16	25.5	0.82	26.2	23.7	40.5	3.76	46.4	35.4	106	0.5	106	105
17	25.0	0.65	25.7	23.4	39.6	3.00	44.0	35.0	107	0.6	108	106
18	25.3	0.88	26.1	23.5	41.8	3.42	45.8	34.0	108	1.1	109	106
19	25.6	0.87	26.2	23.5	46.3	2.44	51.4	43.5	107	1.2	109	106
20	25.3	0.90	26.1	23.3	42.3	3.43	48.0	38.6	110	1.4	112	107
21	26.2	0.92	26.7	23.7	46.6	3.98	55.9	41.9	107	0.6	108	106
22	26.1	1.11	26.7	22.9	46.4	4.45	58.1	41.9	109	0.7	110	108
23	27.0	1.21	27.7	23.3	47.0	5.38	64.1	43.3	109	0.8	111	108
24	26.1	1.23	26.8	22.2	49.7	4.02	62.1	46.7	107	1.8	111	106
25	26.0	0.87	26.7	23.7	46.0	2.72	52.9	43.0	108	0.5	109	108
26	26.1	0.98	26.8	23.5	44.0	4.89	54.8	39.8	107	1.0	109	106
27	26.2	0.90	27.0	23.7	42.5	4.70	54.0	38.3	106	1.0	107	104
28	26.5	1.04	27.2	23.6	47.7	3.92	56.5	41.9	103	2.8	108	99

Appendix B
Daily Exposure Chamber Environmental Conditions

Toxicity Females Control

Exposure	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Day	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	25.2	1.11	25.9	22.0	42.0	4.74	48.7	32.3	107	1.0	108	104
2	25.4	0.92	26.1	22.8	39.0	2.07	44.1	37.0	107	0.8	108	106
3	25.8	0.95	26.4	22.9	37.2	4.28	45.8	32.6	106	1.1	108	104
4	25.7	1.16	26.4	22.3	35.9	2.44	41.2	34.2	108	0.8	110	107
5	25.6	1.00	26.2	22.6	40.3	2.46	45.5	34.7	103	0.5	103	102
6	25.5	0.93	26.0	22.6	41.3	2.66	43.8	35.1	106	0.9	108	105
7	25.5	0.97	26.0	22.5	38.9	2.55	44.6	36.4	108	0.5	109	108
8	25.7	0.94	26.4	22.9	41.1	3.16	46.0	33.7	108	0.6	109	107
9	25.5	1.13	26.3	22.2	39.9	3.29	46.5	35.2	107	1.1	108	105
10	25.6	0.90	26.2	23.2	42.9	3.19	51.1	39.6	106	0.9	107	105
11	25.8	0.92	26.4	23.2	39.9	2.57	44.2	35.9	107	0.7	108	106
12	25.9	0.91	26.4	23.2	41.0	3.09	48.7	37.8	104	0.5	105	103
13	25.5	0.86	26.2	23.1	41.6	3.73	50.0	37.6	107	0.9	108	105
14	25.9	0.88	26.5	23.3	43.6	3.41	50.6	39.6	106	0.6	107	105
15	25.9	0.98	26.4	22.8	39.0	4.52	50.1	33.6	106	0.6	107	105
16	26.2	0.97	26.8	23.3	39.5	2.64	44.4	35.9	105	0.5	106	105
17	26.0	0.84	26.7	23.6	41.0	4.51	47.2	35.2	106	0.7	107	105
18	25.9	1.05	26.5	22.9	42.0	2.51	46.8	35.9	107	0.8	107	105
19	25.9	0.94	26.4	23.2	47.2	4.48	58.1	42.9	108	1.5	110	106
20	26.0	0.76	26.5	23.7	43.6	3.27	50.4	40.0	105	1.3	107	103
21	25.7	0.94	26.3	22.9	48.1	4.58	60.8	44.6	104	0.0	104	104
22	26.1	1.00	26.6	23.0	45.0	4.04	55.6	42.0	104	0.5	105	104
23	26.0	1.11	26.9	22.7	49.6	4.31	62.1	46.0	108	0.7	108	106
24	25.6	1.06	26.1	22.2	47.1	4.28	60.0	44.3	104	0.0	104	104
25	25.9	0.89	26.4	23.2	46.9	3.81	56.9	43.7	107	0.5	107	106
26	25.9	0.75	26.4	23.7	43.9	4.46	55.1	39.9	106	1.0	108	105
27	25.8	1.09	26.6	22.5	44.4	4.34	53.1	39.2	106	0.8	107	105
28	26.3	0.73	26.7	24.1	46.5	1.86	49.4	43.5	105	0.8	106	104
29	25.7	0.76	26.1	23.4	50.1	2.28	55.1	45.3	106	0.8	107	105

Appendix B
Daily Exposure Chamber Environmental Conditions

Toxicity Females 100 ppm

Exposure	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Day	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	24.2	0.38	24.5	23.1	49.5	6.72	57.1	37.4	110	3.4	113	104
2	25.0	1.16	25.9	22.1	45.4	3.31	52.3	41.4	104	0.9	106	103
3	24.7	0.80	25.3	22.3	42.7	2.86	49.0	39.1	105	1.3	107	104
4	25.3	1.15	26.1	22.1	42.8	3.28	50.6	40.0	103	1.9	108	101
5	25.0	0.70	25.5	23.0	46.8	2.38	49.7	40.9	105	0.3	105	104
6	25.0	0.80	25.4	22.5	46.4	2.99	50.2	40.0	106	0.8	108	105
7	25.0	0.86	25.5	22.4	43.3	2.48	48.9	40.4	104	0.5	105	103
8	25.5	1.00	26.3	22.6	47.3	3.02	50.7	40.0	107	1.2	108	105
9	24.9	1.01	25.9	22.2	46.2	2.83	51.0	42.4	106	0.9	107	104
10	25.3	0.70	25.9	23.4	47.3	2.69	52.9	43.7	106	0.5	107	106
11	25.1	0.77	25.5	22.9	45.4	2.72	51.2	42.1	106	0.9	108	105
12	25.5	0.87	26.0	22.8	46.4	3.08	53.5	42.5	103	0.8	104	102
13	24.4	0.77	25.0	22.3	50.3	3.76	57.6	45.9	106	1.0	107	104
14	24.6	0.61	25.0	22.9	50.3	1.61	53.9	47.5	105	0.8	106	104
15	25.4	0.84	25.9	22.8	44.8	4.42	54.3	39.5	103	0.9	105	102
16	25.4	0.84	25.9	23.0	46.6	2.43	50.2	42.7	101	2.9	107	97
17	25.0	0.71	25.6	22.9	47.2	3.86	54.2	42.1	101	0.6	102	100
18	25.3	0.84	25.9	23.0	49.0	1.93	52.6	45.5	103	1.0	104	101
19	25.5	0.85	26.0	23.0	52.1	3.36	59.3	49.0	104	1.4	107	103
20	25.3	0.78	26.0	23.0	49.7	3.65	57.9	45.4	102	1.6	104	100
21	25.2	0.74	25.7	23.1	53.9	4.40	64.6	49.3	101	0.5	102	100
22	25.1	0.81	25.6	22.6	52.5	4.39	65.0	48.7	100	0.6	101	99
23	24.9	0.75	25.5	22.6	54.1	2.57	61.4	50.7	103	0.4	103	102
24	25.6	1.09	26.2	22.2	53.7	4.64	67.0	50.0	101	0.4	102	101
25	25.3	0.63	25.7	23.4	54.1	2.57	60.9	52.1	101	0.7	102	100
26	25.7	0.85	26.3	23.3	51.2	5.40	62.1	45.9	100	1.7	104	97
27	25.3	0.66	25.9	23.3	48.2	4.08	55.9	42.7	104	0.9	106	102
28	25.7	0.93	26.3	22.9	52.0	2.99	58.2	47.1	102	0.9	104	101
29	25.3	1.05	26.3	22.9	50.3	2.57	54.2	46.1	99	2.6	105	96

Appendix B
Daily Exposure Chamber Environmental Conditions

Toxicity Females 500 ppm

Exposure	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Day	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	26.1	1.35	27.0	22.3	37.8	5.42	43.7	26.3	107	1.1	108	104
2	26.2	1.20	26.8	22.5	36.6	2.59	41.8	32.5	108	0.0	108	108
3	25.9	1.24	26.6	22.0	34.1	2.56	39.6	30.7	105	1.4	107	103
4	26.2	1.42	27.0	21.9	33.2	2.49	38.1	30.7	108	0.7	108	106
5	26.4	1.24	27.0	22.6	35.7	2.18	37.7	30.2	105	1.0	106	104
6	26.1	1.11	26.7	22.6	37.7	3.04	41.4	31.3	109	0.7	110	108
7	26.5	1.28	27.1	22.5	34.2	2.87	41.2	31.2	104	0.3	104	103
8	26.5	1.08	27.2	23.3	37.6	3.03	40.3	29.6	109	2.7	111	103
9	26.0	1.25	26.8	22.3	36.7	2.61	41.5	33.3	106	1.5	108	103
10	26.3	1.09	27.1	23.5	37.2	2.97	41.2	31.6	107	3.5	113	103
11	26.3	0.87	26.8	23.8	36.4	2.28	40.8	33.3	108	2.0	110	104
12	26.0	0.92	26.6	23.3	37.7	2.02	41.3	35.2	107	1.8	108	103
13	25.9	0.77	26.4	23.7	38.4	2.56	44.8	35.9	106	1.3	109	104
14	26.1	0.81	26.6	23.8	40.9	2.49	45.0	37.0	106	1.2	107	104
15	26.1	1.01	26.7	23.0	36.0	3.32	42.0	32.3	107	0.4	107	106
16	26.3	1.06	26.9	23.2	37.0	2.74	41.9	32.9	106	0.7	107	104
17	26.1	0.73	26.7	24.0	37.8	2.78	43.0	34.5	108	0.6	109	107
18	26.1	1.11	26.8	22.8	39.9	2.65	42.6	33.1	108	1.0	109	106
19	26.3	0.89	26.8	23.7	43.0	2.13	48.7	41.2	107	1.4	110	104
20	26.1	0.81	26.7	23.7	39.5	2.82	44.5	36.1	106	1.9	108	103
21	26.0	0.86	26.6	23.5	43.8	3.18	51.3	39.4	104	2.2	109	103
22	26.2	1.10	26.8	22.8	42.8	3.36	53.1	40.8	107	0.8	108	106
23	26.3	1.31	27.2	22.3	48.0	4.07	60.5	45.1	107	0.7	108	106
24	26.2	1.31	26.9	22.2	44.5	3.85	56.7	42.7	104	0.6	106	104
25	26.4	0.99	26.9	23.4	43.0	1.94	49.0	41.6	109	0.7	110	108
26	26.6	0.78	27.1	24.3	40.6	3.72	49.8	36.4	106	1.2	108	104
27	26.1	0.86	26.8	23.5	39.6	2.78	45.5	34.5	105	1.5	108	103
28	26.5	0.90	27.0	23.8	43.3	3.07	47.1	36.6	107	0.9	108	106
29	26.3	0.85	27.0	23.9	45.8	2.17	51.5	43.5	99	1.7	105	98

Appendix B
Daily Exposure Chamber Environmental Conditions

Toxicity Females 2500 ppm

Exposure	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Day	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	25.9	0.99	26.7	23.3	41.0	3.30	44.6	34.5	107	1.7	110	104
2	26.1	1.08	26.7	22.8	39.1	1.84	42.0	35.9	106	0.8	108	105
3	25.1	1.02	25.8	22.0	38.7	3.24	43.0	33.6	107	1.3	109	106
4	25.7	1.12	26.4	22.1	38.0	3.81	46.1	33.8	106	1.3	108	103
5	25.3	1.02	25.9	22.2	41.8	2.41	44.0	36.7	107	1.2	108	105
6	25.0	0.88	25.6	22.3	43.6	2.52	47.1	38.5	108	1.4	110	106
7	25.4	1.05	26.0	22.3	40.7	3.11	47.9	36.8	105	0.8	106	104
8	25.5	1.33	26.4	21.7	43.3	3.47	50.6	36.3	112	2.5	114	106
9	25.6	1.37	26.6	21.8	40.4	3.77	49.5	35.9	106	1.8	109	103
10	25.7	1.04	26.7	23.3	42.4	3.01	49.0	38.2	106	0.8	108	105
11	25.7	0.95	26.6	23.4	41.1	3.92	49.2	37.1	106	1.0	108	105
12	25.4	0.94	26.0	22.8	42.9	3.39	49.6	38.5	108	1.3	109	105
13	25.1	0.79	25.9	23.2	44.7	3.62	54.5	40.7	108	1.0	109	106
14	25.3	0.73	25.8	23.4	47.9	4.53	56.3	42.5	106	1.4	109	104
15	25.2	1.09	25.8	22.0	42.3	5.23	54.4	37.5	107	1.1	108	105
16	25.7	1.00	26.4	23.1	43.2	3.64	49.7	37.7	108	0.8	109	106
17	25.8	0.69	26.5	24.0	41.3	3.76	48.6	36.3	107	1.8	110	105
18	25.3	0.97	26.1	22.9	45.0	2.63	48.3	40.9	108	1.5	110	106
19	26.4	0.85	27.1	24.2	46.2	3.23	52.9	43.7	108	0.9	109	106
20	25.5	0.91	26.4	23.2	45.2	4.96	54.0	39.6	108	2.0	113	106
21	25.9	0.93	26.5	23.5	49.6	5.27	61.6	45.1	109	1.0	112	108
22	25.7	1.04	26.5	22.6	48.2	5.40	62.6	43.2	105	1.8	109	102
23	26.0	1.20	26.9	22.6	49.4	4.50	59.9	45.8	108	1.6	109	105
24	25.5	1.09	26.1	22.2	50.1	4.45	62.4	46.7	109	2.1	112	105
25	25.7	0.78	26.4	23.7	49.2	3.29	57.1	45.6	106	1.7	109	105
26	26.4	0.77	27.1	24.6	45.1	5.74	56.8	40.5	110	0.8	111	109
27	26.1	0.96	26.7	23.3	42.7	3.97	52.2	37.8	111	1.5	113	109
28	26.0	0.93	26.7	23.4	47.9	3.75	54.6	41.5	108	1.3	110	106
29	26.4	1.13	27.5	23.8	47.5	3.21	53.8	42.0	102	2.4	107	97

Appendix B
Daily Exposure Chamber Environmental Conditions

Reproductive Females Control

Exposure	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Day	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	25.2	1.11	25.9	22.0	42.0	4.74	48.7	32.3	107	1.0	108	104
2	25.4	0.92	26.1	22.8	39.0	2.07	44.1	37.0	107	0.8	108	106
3	25.8	0.95	26.4	22.9	37.2	4.28	45.8	32.6	106	1.1	108	104
4	25.7	1.16	26.4	22.3	35.9	2.44	41.2	34.2	108	0.8	110	107
5	25.6	1.00	26.2	22.6	40.3	2.46	45.5	34.7	103	0.5	103	102
6	25.5	0.93	26.0	22.6	41.3	2.66	43.8	35.1	106	0.9	108	105
7	25.5	0.97	26.0	22.5	38.9	2.55	44.6	36.4	108	0.5	109	108
8	25.7	0.94	26.4	22.9	41.1	3.16	46.0	33.7	108	0.6	109	107
9	25.5	1.13	26.3	22.2	39.9	3.29	46.5	35.2	107	1.1	108	105
10	25.6	0.90	26.2	23.2	42.9	3.19	51.1	39.6	106	0.9	107	105
11	25.8	0.92	26.4	23.2	39.9	2.57	44.2	35.9	107	0.7	108	106
12	25.9	0.91	26.4	23.2	41.0	3.09	48.7	37.8	104	0.5	105	103
13	25.5	0.86	26.2	23.1	41.6	3.73	50.0	37.6	107	0.9	108	105
14	25.9	0.88	26.5	23.3	43.6	3.41	50.6	39.6	106	0.6	107	105
15	25.9	0.98	26.4	22.8	39.0	4.52	50.1	33.6	106	0.6	107	105
16	26.2	0.97	26.8	23.3	39.5	2.64	44.4	35.9	105	0.5	106	105
17	26.0	0.84	26.7	23.6	41.0	4.51	47.2	35.2	106	0.7	107	105
18	25.9	1.05	26.5	22.9	42.0	2.51	46.8	35.9	107	0.8	107	105
19	25.9	0.94	26.4	23.2	47.2	4.48	58.1	42.9	108	1.5	110	106
20	26.0	0.76	26.5	23.7	43.6	3.27	50.4	40.0	105	1.3	107	103
21	25.7	0.94	26.3	22.9	48.1	4.58	60.8	44.6	104	0.0	104	104
22	26.1	1.00	26.6	23.0	45.0	4.04	55.6	42.0	104	0.5	105	104
23	26.0	1.11	26.9	22.7	49.6	4.31	62.1	46.0	108	0.7	108	106
24	25.6	1.06	26.1	22.2	47.1	4.28	60.0	44.3	104	0.0	104	104
25	25.9	0.89	26.4	23.2	46.9	3.81	56.9	43.7	107	0.5	107	106
26	25.9	0.75	26.4	23.7	43.9	4.46	55.1	39.9	106	1.0	108	105
27	25.8	1.09	26.6	22.5	44.4	4.34	53.1	39.2	106	0.8	107	105
28	26.3	0.73	26.7	24.1	46.5	1.86	49.4	43.5	105	0.8	106	104
29	25.7	0.76	26.1	23.4	50.1	2.28	55.1	45.3	106	0.8	107	105
30	24.4	0.24	24.9	24.1	48.4	3.25	55.5	40.9	105	0.5	106	104
31	24.7	0.52	25.1	23.5	53.2	4.88	58.9	38.4	104	0.5	105	104
32	25.8	0.55	26.1	24.0	51.1	2.85	57.6	48.6	108	0.4	108	107
33	25.2	0.67	25.7	23.1	49.2	4.29	56.1	40.0	106	0.5	107	106
34	25.4	0.76	25.8	23.1	53.1	3.78	58.4	42.1	106	0.6	108	106
35	24.9	0.52	25.1	23.3	51.1	3.24	54.5	41.0	107	0.9	108	106
36	24.7	0.51	25.1	23.3	45.5	8.01	52.7	33.2	106	0.5	106	105
37	24.2	0.35	24.7	23.6	48.0	4.30	51.2	34.2	108	0.0	108	108
38	23.8	0.42	24.3	23.2	45.1	4.55	49.3	31.7	106	0.6	107	105
39	24.1	0.53	24.6	22.9	42.0	1.74	44.6	39.7	105	0.5	106	105

Appendix B
Daily Exposure Chamber Environmental Conditions

Reproductive Females 100 ppm

Exposure	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Day	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	24.2	0.38	24.5	23.1	49.5	6.72	57.1	37.4	110	3.4	113	104
2	25.0	1.16	25.9	22.1	45.4	3.31	52.3	41.4	104	0.9	106	103
3	24.7	0.80	25.3	22.3	42.7	2.86	49.0	39.1	105	1.3	107	104
4	25.3	1.15	26.1	22.1	42.8	3.28	50.6	40.0	103	1.9	108	101
5	25.0	0.70	25.5	23.0	46.8	2.38	49.7	40.9	105	0.3	105	104
6	25.0	0.80	25.4	22.5	46.4	2.99	50.2	40.0	106	0.8	108	105
7	25.0	0.86	25.5	22.4	43.3	2.48	48.9	40.4	104	0.5	105	103
8	25.5	1.00	26.3	22.6	47.3	3.02	50.7	40.0	107	1.2	108	105
9	24.9	1.01	25.9	22.2	46.2	2.83	51.0	42.4	106	0.9	107	104
10	25.3	0.70	25.9	23.4	47.3	2.69	52.9	43.7	106	0.5	107	106
11	25.1	0.77	25.5	22.9	45.4	2.72	51.2	42.1	106	0.9	108	105
12	25.5	0.87	26.0	22.8	46.4	3.08	53.5	42.5	103	0.8	104	102
13	24.4	0.77	25.0	22.3	50.3	3.76	57.6	45.9	106	1.0	107	104
14	24.6	0.61	25.0	22.9	50.3	1.61	53.9	47.5	105	0.8	106	104
15	25.4	0.84	25.9	22.8	44.8	4.42	54.3	39.5	103	0.9	105	102
16	25.4	0.84	25.9	23.0	46.6	2.43	50.2	42.7	101	2.9	107	97
17	25.0	0.71	25.6	22.9	47.2	3.86	54.2	42.1	101	0.6	102	100
18	25.3	0.84	25.9	23.0	49.0	1.93	52.6	45.5	103	1.0	104	101
19	25.5	0.85	26.0	23.0	52.1	3.36	59.3	49.0	104	1.4	107	103
20	25.3	0.78	26.0	23.0	49.7	3.65	57.9	45.4	102	1.6	104	100
21	25.2	0.74	25.7	23.1	53.9	4.40	64.6	49.3	101	0.5	102	100
22	25.1	0.81	25.6	22.6	52.5	4.39	65.0	48.7	100	0.6	101	99
23	24.9	0.75	25.5	22.6	54.1	2.57	61.4	50.7	103	0.4	103	102
24	25.6	1.09	26.2	22.2	53.7	4.64	67.0	50.0	101	0.4	102	101
25	25.3	0.63	25.7	23.4	54.1	2.57	60.9	52.1	101	0.7	102	100
26	25.7	0.85	26.3	23.3	51.2	5.40	62.1	45.9	100	1.7	104	97
27	25.3	0.66	25.9	23.3	48.2	4.08	55.9	42.7	104	0.9	106	102
28	25.7	0.93	26.3	22.9	52.0	2.99	58.2	47.1	102	0.9	104	101
29	25.3	1.05	26.3	22.9	50.3	2.57	54.2	46.1	99	2.6	105	96
30	25.1	0.42	25.5	23.8	45.1	2.88	52.1	42.1	99	0.7	100	98
31	25.1	0.98	25.9	22.9	50.7	4.44	56.6	37.8	101	1.4	102	98
32	25.9	0.96	26.6	23.2	46.8	4.53	57.7	43.4	99	0.9	101	98
33	25.5	0.76	26.2	23.6	46.5	3.82	54.5	41.9	99	1.5	102	98
34	25.9	0.89	26.4	23.3	49.3	3.16	55.9	42.9	98	0.4	99	98
35	24.5	0.65	24.9	22.7	47.7	2.51	51.3	41.8	100	0.8	101	99
36	24.6	0.46	25.0	23.4	42.1	8.04	49.8	29.8	101	0.5	101	100
37	24.4	0.49	25.0	23.6	44.2	3.74	46.5	32.2	99	0.5	99	98

Appendix B
Daily Exposure Chamber Environmental Conditions

Reproductive Females 500 ppm

Exposure Day	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	26.1	1.35	27.0	22.3	37.8	5.42	43.7	26.3	107	1.1	108	104
2	26.2	1.20	26.8	22.5	36.6	2.59	41.8	32.5	108	0.0	108	108
3	25.9	1.24	26.6	22.0	34.1	2.56	39.6	30.7	105	1.4	107	103
4	26.2	1.42	27.0	21.9	33.2	2.49	38.1	30.7	108	0.7	108	106
5	26.4	1.24	27.0	22.6	35.7	2.18	37.7	30.2	105	1.0	106	104
6	26.1	1.11	26.7	22.6	37.7	3.04	41.4	31.3	109	0.7	110	108
7	26.5	1.28	27.1	22.5	34.2	2.87	41.2	31.2	104	0.3	104	103
8	26.5	1.08	27.2	23.3	37.6	3.03	40.3	29.6	109	2.7	111	103
9	26.0	1.25	26.8	22.3	36.7	2.61	41.5	33.3	106	1.5	108	103
10	26.3	1.09	27.1	23.5	37.2	2.97	41.2	31.6	107	3.5	113	103
11	26.3	0.87	26.8	23.8	36.4	2.28	40.8	33.3	108	2.0	110	104
12	26.0	0.92	26.6	23.3	37.7	2.02	41.3	35.2	107	1.8	108	103
13	25.9	0.77	26.4	23.7	38.4	2.56	44.8	35.9	106	1.3	109	104
14	26.1	0.81	26.6	23.8	40.9	2.49	45.0	37.0	106	1.2	107	104
15	26.1	1.01	26.7	23.0	36.0	3.32	42.0	32.3	107	0.4	107	106
16	26.3	1.06	26.9	23.2	37.0	2.74	41.9	32.9	106	0.7	107	104
17	26.1	0.73	26.7	24.0	37.8	2.78	43.0	34.5	108	0.6	109	107
18	26.1	1.11	26.8	22.8	39.9	2.65	42.6	33.1	108	1.0	109	106
19	26.3	0.89	26.8	23.7	43.0	2.13	48.7	41.2	107	1.4	110	104
20	26.1	0.81	26.7	23.7	39.5	2.82	44.5	36.1	106	1.9	108	103
21	26.0	0.86	26.6	23.5	43.8	3.18	51.3	39.4	104	2.2	109	103
22	26.2	1.10	26.8	22.8	42.8	3.36	53.1	40.8	107	0.8	108	106
23	26.3	1.31	27.2	22.3	48.0	4.07	60.5	45.1	107	0.7	108	106
24	26.2	1.31	26.9	22.2	44.5	3.85	56.7	42.7	104	0.6	106	104
25	26.4	0.99	26.9	23.4	43.0	1.94	49.0	41.6	109	0.7	110	108
26	26.6	0.78	27.1	24.3	40.6	3.72	49.8	36.4	106	1.2	108	104
27	26.1	0.86	26.8	23.5	39.6	2.78	45.5	34.5	105	1.5	108	103
28	26.5	0.90	27.0	23.8	43.3	3.07	47.1	36.6	107	0.9	108	106
29	26.3	0.85	27.0	23.9	45.8	2.17	51.5	43.5	99	1.7	105	98
30	25.7	0.47	26.1	24.3	43.5	2.86	48.7	38.7	100	0.4	101	100
31	25.3	1.05	26.1	22.9	49.8	3.66	55.7	39.9	101	1.2	104	100
32	26.3	0.70	26.9	24.1	46.2	3.29	52.7	41.5	99	0.5	100	99
33	25.7	0.86	26.3	23.1	46.0	3.23	52.1	41.3	99	0.8	101	99
34	26.0	0.92	26.5	23.3	47.5	3.35	52.6	40.5	98	0.7	100	98
35	25.2	0.68	25.7	23.4	45.7	2.51	50.3	39.9	99	0.5	100	99
36	25.4	0.63	25.9	23.7	40.4	6.74	47.1	28.3	99	0.4	100	99
37	24.7	0.55	25.3	23.6	44.2	3.68	47.9	32.9	100	0.5	101	100

Appendix B
Daily Exposure Chamber Environmental Conditions

Reproductive Females 2500 ppm

	Temperature (°C)				Humidity (%RH)				Air Flow (LPM)			
Exposure Day	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min	Mean	Std Dev	Max	Min
1	25.9	0.99	26.7	23.3	41.0	3.30	44.6	34.5	107	1.7	110	104
2	26.1	1.08	26.7	22.8	39.1	1.84	42.0	35.9	106	0.8	108	105
3	25.1	1.02	25.8	22.0	38.7	3.24	43.0	33.6	107	1.3	109	106
4	25.7	1.12	26.4	22.1	38.0	3.81	46.1	33.8	106	1.3	108	103
5	25.3	1.02	25.9	22.2	41.8	2.41	44.0	36.7	107	1.2	108	105
6	25.0	0.88	25.6	22.3	43.6	2.52	47.1	38.5	108	1.4	110	106
7	25.4	1.05	26.0	22.3	40.7	3.11	47.9	36.8	105	0.8	106	104
8	25.5	1.33	26.4	21.7	43.3	3.47	50.6	36.3	112	2.5	114	106
9	25.6	1.37	26.6	21.8	40.4	3.77	49.5	35.9	106	1.8	109	103
10	25.7	1.04	26.7	23.3	42.4	3.01	49.0	38.2	106	0.8	108	105
11	25.7	0.95	26.6	23.4	41.1	3.92	49.2	37.1	106	1.0	108	105
12	25.4	0.94	26.0	22.8	42.9	3.39	49.6	38.5	108	1.3	109	105
13	25.1	0.79	25.9	23.2	44.7	3.62	54.5	40.7	108	1.0	109	106
14	25.3	0.73	25.8	23.4	47.9	4.53	56.3	42.5	106	1.4	109	104
15	25.2	1.09	25.8	22.0	42.3	5.23	54.4	37.5	107	1.1	108	105
16	25.7	1.00	26.4	23.1	43.2	3.64	49.7	37.7	108	0.8	109	106
17	25.8	0.69	26.5	24.0	41.3	3.76	48.6	36.3	107	1.8	110	105
18	25.3	0.97	26.1	22.9	45.0	2.63	48.3	40.9	108	1.5	110	106
19	26.4	0.85	27.1	24.2	46.2	3.23	52.9	43.7	108	0.9	109	106
20	25.5	0.91	26.4	23.2	45.2	4.96	54.0	39.6	108	2.0	113	106
21	25.9	0.93	26.5	23.5	49.6	5.27	61.6	45.1	109	1.0	112	108
22	25.7	1.04	26.5	22.6	48.2	5.40	62.6	43.2	105	1.8	109	102
23	26.0	1.20	26.9	22.6	49.4	4.50	59.9	45.8	108	1.6	109	105
24	25.5	1.09	26.1	22.2	50.1	4.45	62.4	46.7	109	2.1	112	105
25	25.7	0.78	26.4	23.7	49.2	3.29	57.1	45.6	106	1.7	109	105
26	26.4	0.77	27.1	24.6	45.1	5.74	56.8	40.5	110	0.8	111	109
27	26.1	0.96	26.7	23.3	42.7	3.97	52.2	37.8	111	1.5	113	109
28	26.0	0.93	26.7	23.4	47.9	3.75	54.6	41.5	108	1.3	110	106
29	26.4	1.13	27.5	23.8	47.5	3.21	53.8	42.0	102	2.4	107	97
30	25.7	0.51	26.4	24.4	44.1	3.03	52.7	40.7	104	0.0	104	104
31	25.5	1.13	26.4	23.1	46.5	3.90	52.9	37.2	102	1.4	104	100
32	26.4	0.73	27.4	24.5	48.3	4.75	57.5	43.3	101	0.9	102	99
33	26.2	0.83	26.7	23.9	48.8	4.90	57.7	43.4	103	1.3	106	102
34	26.1	0.97	26.8	23.5	52.8	4.50	59.1	42.9	100	0.6	102	100
35	25.9	0.92	26.6	23.5	45.9	3.53	54.7	42.8	104	0.6	104	102
36	24.9	0.56	25.4	23.5	40.5	7.08	46.6	29.4	102	0.0	102	102
37	24.5	0.64	25.3	23.4	43.5	3.68	46.5	32.1	102	0.5	103	102

Appendix C

Individual Clinical Observations and Detailed Physical Exam Data

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Males - Controls

Animal ID	Clinical Sign	DPE	Study Day 1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
Group 1 - Controls																
D0041	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0042	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0043	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0044	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0045	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0046	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0047	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0048	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0049	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0050	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Animal ID	Clinical Sign	Study Day 15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29
Group 1 - Controls																
D0041	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0042	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0043	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0044	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0045	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0046	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0047	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0048	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0049	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0050	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Males - 100 ppm

Animal ID	Clinical Sign	DPE	Study Day 1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
Group 2 - 100 ppm																
D0051	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0052	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0053	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0054	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0055	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0056	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0057	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0058	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0059	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0060	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Animal ID	Clinical Sign	Study Day 15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29
Group 2 - 100 ppm																
D0051	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0052	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0053	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0054	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0055	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0056	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0057	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0058	Normal	P	P	P	P	N	P	P	P	P	P	P	P	P	P	P
D0058	Probable Porphyrin Staining, head	N	N	N	N	P	N	N	N	N	N	N	N	N	N	N
D0058	Salivation	N	N	N	N	P	N	N	N	N	N	N	N	N	N	N
D0059	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0060	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Males - 500 ppm

Animal ID	Clinical Sign	DPE	1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
		Study Day														
Group 3 - 500 ppm																
D0061	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0062	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0063	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0064	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0065	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0066	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0067	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0068	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0069	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0070	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Animal ID	Clinical Sign	15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29
		Study Day														
Group 3 - 500 ppm																
D0061	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0062	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0063	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0064	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0065	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0066	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0067	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0068	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0069	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0070	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Males - 2500 ppm

Animal ID	Clinical Sign	DPE	Study Day 1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
Group 4 - 2500 ppm																
D0071	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0072	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0073	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0074	Normal	P	P	P	P	P	P	P	P	P	P	N	P	P	P	P
D0074	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	N	N	P	N	N	N	N
D0075	Normal	P	P	P	P	N	N	N	N	N	N	N	N	N	N	N
D0075	Anogenital Staining	N	N	N	N	P	P	P	P	P	P	P	P	P	P	P
D0076	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0077	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0078	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0079	Normal	P	P	N	P	P	P	P	P	P	P	P	P	P	P	P
D0079	Anogenital Staining	N	N	P	N	N	N	N	N	N	N	N	N	N	N	N
D0080	Normal	P	N	N	P	P	N	P	P	P	P	P	P	P	P	P
D0080	Anogenital Staining	N	P	P	N	N	P	N	N	N	N	N	N	N	N	N

DPE = Detailed physical exam
P = Present
N = Not Present

Animal ID	Clinical Sign	Study Day 15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29
Group 4 - 2500 ppm																
D0071	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0072	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0073	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0074	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0075	Normal	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
D0075	Anogenital Staining	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0076	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0077	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0078	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0079	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0080	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Toxicity Females - Controls

Animal ID	Clinical Sign	DPE	1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
			Study Day													
Group 1 - Controls																
D0081	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0082	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0083	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0084	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0085	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0086	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0087	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0088	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0089	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0090	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Animal ID	Clinical Sign	15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29	30
		Study Day															
Group 1 - Controls																	
D0081	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0082	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0083	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0084	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0085	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0086	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0087	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0088	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0089	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0090	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Toxicity Females - 100 ppm

Animal ID	Clinical Sign	DPE	Study Day 1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
Group 2 - 100 ppm																
D0091	Normal	P	P	P	N	P	P	P	P	P	P	P	P	P	P	P
D0091	Bloody Knuckle	N	N	N	P	N	N	N	N	N	N	N	N	N	N	N
D0092	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0093	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0094	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0095	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0096	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0097	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0098	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0099	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0100	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Animal ID	Clinical Sign	Study Day 15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29	30
Group 2 - 100 ppm																	
D0091	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0092	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0093	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0094	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0095	Normal	P	P	N	P	P	P	P	P	P	P	P	P	P	P	P	P
D0095	Probable Porphyrin Staining, head	N	N	P	N	N	N	N	N	N	N	N	N	N	N	N	N
D0096	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0097	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0098	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0099	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0100	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Toxicity Females - 500 ppm

Animal ID	Clinical Sign	DPE	Study Day 1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
Group 3 - 500 ppm																
D0101	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0102	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0103	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0104	Normal	P	N	P	P	P	P	P	P	P	P	P	P	P	P	P
D0104	Toenail Bleeding	N	P	N	N	N	N	N	N	N	N	N	N	N	N	N
D0105	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0106	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0107	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0108	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0109	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0110	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Animal ID	Clinical Sign	Study Day 15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29	30
Group 3 - 500 ppm																	
D0101	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0102	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0103	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0104	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0105	Normal	P	P	P	N	P	P	P	P	P	P	P	P	P	P	P	P
D0105	Probable Porphyrin Staining, head	N	N	N	P	N	N	N	N	N	N	N	N	N	N	N	N
D0106	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0107	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0108	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0109	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0110	Normal	N	P	P	N	N	N	P	P	P	N	P	P	P	P	P	P
D0110	Probable Porphyrin Staining, head	P	N	N	P	P	P	N	N	N	P	N	N	N	N	N	N

DPE = Detailed physical exam
P = Present
N = Not Present

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Toxicity Females - 2500 ppm

Animal ID	Clinical Sign	DPE	Study Day 1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
Group 4 - 2500 ppm																
D0111	Normal	P	P	P	P	P	P	P	P	P	P	P	P	N	P	N
D0111	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	N	N	N	N	P	N	P
D0112	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	N
D0112	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	N	N	N	N	N	N	P
D0113	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0114	Normal	P	P	P	P	N	P	P	P	P	P	P	P	P	P	P
D0114	Anogenital Staining	N	N	N	N	P	N	N	N	N	N	N	N	N	N	N
D0115	Normal	P	P	P	P	P	N	P	P	P	P	P	P	P	P	P
D0115	Probable Porphyrin Staining, head	N	N	N	N	N	P	N	N	N	N	N	N	N	N	N
D0116	Normal	P	N	P	P	N	P	P	N	P	P	P	N	P	P	N
D0116	Anogenital Staining	N	P	N	N	P	N	N	N	N	N	N	P	N	N	N
D0116	Probable Porphyrin Staining, head	N	N	N	N	P	N	N	P	N	N	N	N	N	N	P
D0117	Normal	P	P	P	P	P	P	P	P	N	P	N	P	P	P	N
D0117	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	P	N	P	N	N	N	P
D0118	Normal	P	P	P	N	N	N	P	N	N	N	P	N	P	P	N
D0118	Anogenital Staining	N	N	N	P	P	P	N	N	N	N	N	N	N	N	N
D0118	Probable Porphyrin Staining, head	N	N	N	N	N	P	N	P	P	P	N	P	N	N	P
D0119	Normal	P	P	P	P	N	P	P	P	P	P	P	N	P	P	N
D0119	Probable Porphyrin Staining, head	N	N	N	N	P	N	N	N	N	N	N	P	N	N	P
D0040	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	N
D0040	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	N	N	N	N	N	N	P

DPE = Detailed physical exam
P = Present
N = Not Present

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Toxicity Females - 2500 ppm (continued)

Animal ID	Clinical Sign	Study Day 15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29	30
Group 4 - 2500 ppm																	
D0111	Normal	P	P	N	N	N	N	N	N	N	N	N	N	N	N	N	P
D0111	Probable Porphyrin Staining, head	N	N	P	P	P	P	P	P	P	P	P	P	P	P	P	N
D0112	Normal	P	P	P	N	P	P	N	N	N	P	P	N	P	P	P	P
D0112	Anogenital Staining	N	N	N	P	N	N	P	P	P	N	N	P	N	N	N	N
D0113	Normal	P	P	P	P	N	P	P	P	P	P	P	P	P	P	P	P
D0113	Probable Porphyrin Staining, head	N	N	N	N	P	N	N	N	N	N	N	N	N	N	N	N
D0114	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0115	Normal	P	P	P	P	P	N	P	P	P	N	N	P	P	P	P	P
D0115	Probable Porphyrin Staining, head	N	N	N	N	N	P	N	N	N	P	P	N	N	N	N	N
D0116	Normal	P	P	N	P	N	N	N	N	N	N	N	P	P	P	N	N
D0116	Anogenital Staining	N	N	P	N	P	P	P	P	P	P	P	N	N	N	P	N
D0116	Probable Porphyrin Staining, head	N	N	N	N	P	P	P	P	P	P	P	N	N	N	P	P
D0117	Normal	P	N	P	P	P	N	P	P	P	P	P	P	P	P	P	P
D0117	Probable Porphyrin Staining, head	N	P	N	N	N	P	N	N	N	N	N	N	N	N	N	N
D0118	Normal	P	P	P	P	P	P	N	N	N	P	P	N	N	N	P	P
D0118	Anogenital Staining	N	N	N	N	N	N	P	P	N	N	N	N	N	N	N	N
D0118	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	P	N	N	P	P	P	N	N
D0119	Normal	N	N	N	N	P	P	N	N	P	N	N	N	N	N	P	P
D0119	Probable Porphyrin Staining, head	P	P	P	P	N	N	P	P	N	P	P	P	P	P	N	N
D0040	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Reproductive Females - Controls

Animal ID	Clinical Sign	DPE	Study Day 1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
Group 1 - Controls																
D0001	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0002	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0003	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0004	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0005	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0006	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0007	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0008	Normal	P	N	P	P	P	N	P	P	P	P	P	P	P	P	P
D0008	Toenail Bleeding	N	P	N	N	N	P	N	N	N	N	N	N	N	N	N
D0009	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0010	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present
D = Deceased

Animal ID	Clinical Sign	Study Day 15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29
Group 1 - Controls																
D0001	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0002	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0003	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0004	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0005	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0006	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0007	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0008	Normal	P	P	P	P	P	P	N	P	P	P	P	P	P	P	P
D0008	Bleeding, ear	N	N	N	N	N	N	P	N	N	N	N	N	N	N	N
D0009	Normal	P	P	P	P	P	P	P	N	N	N	P	N	N	N	N
D0009	Generalized Hairloss, Anogenital region	N	N	N	N	N	N	N	P	P	P	N	P	P	P	P
D0010	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present
D = Deceased

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Reproductive Females - Controls (continued)

Animal ID	Clinical Sign	30	31	DPE 32	33	34	35	36	37	Study Day 38	DPE 39	40	41	42	43	44	45	DPE 46
Group 1 - Controls																		
D0001	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	D	D
D0002	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	N	N
D0003	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D	D
D0004	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D	D	D
D0005	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	D	D
D0006	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D	D
D0007	Normal	P	P	P	P	P	P	P	P	P	P	P	D	D	D	D	D	D
D0008	Normal	P	P	P	P	P	P	P	P	P	P	P	D	D	D	D	D	D
D0009	Normal	N	N	N	N	N	N	N	N	N	N	P	P	P	D	D	D	D
D0009	Generalized Hairloss, Anogenital Region	P	P	P	P	P	P	P	P	P	P	N	N	N	D	D	D	D
D0010	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D	D	D

DPE = Detailed physical exam
P = Present
N = Not Present
D = Deceased

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Reproductive Females - 100 ppm

Animal ID	Clinical Sign	DPE	Study Day 1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
Group 2 - 100 ppm		P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0011	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0012	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0013	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0014	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0015	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0016	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0017	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0018	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0019	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0020	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present
D = Deceased

Animal ID	Clinical Sign	Study Day 15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29
Group 2 - 100 ppm																
D0011	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0012	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0013	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0014	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0015	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0016	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0017	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0018	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0019	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0020	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present
D = Deceased

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Reproductive Females - 100 ppm (continued)

Animal ID	Clinical Sign	Study Day 30	31	DPE 32	33	34	35	36	37	38	DPE 39	40	41	42	43	44
Group 2 - 100 ppm																
D0011	Normal	P	P	P	P	P	P	P	P	P	P	P	D	D	D	D
D0012	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D
D0013	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D
D0014	Normal	P	P	N	P	P	P	P	P	P	P	P	P	D	D	D
D0014	Anogenital Staining	N	N	P	N	N	N	N	N	N	N	N	N	D	D	D
D0015	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	D
D0016	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D
D0017	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	D	D
D0018	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0019	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D
D0020	Normal	P	P	P	P	P	P	P	P	P	P	P	D	D	D	D

DPE = Detailed physical exam
P = Present
N = Not Present
D = Deceased

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Reproductive Females - 500 ppm

Animal ID	Clinical Sign	DPE	1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
		Study Day														
Group 3 - 500 ppm																
D0021	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	N	N
D0021	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	N	N	N	N	N	P	P
D0022	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0023	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0024	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0025	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0026	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0027	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0028	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0028	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
D0029	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0030	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present
D = Deceased

Animal ID	Clinical Sign	15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29
		Study Day														
Group 3 - 500 ppm																
D0021	Normal	N	N	P	N	N	N	N	N	N	P	P	P	P	P	P
D0021	Probable Porphyrin staining, head	P	P	N	P	P	P	P	P	P	N	N	N	N	N	N
D0022	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0023	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0024	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0025	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0026	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0027	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0027	Probable Porphyrin staining, head	N	N	N	N	N	N	N	N	N	N	N	N	N	N	N
D0028	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0029	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0030	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present
D = Deceased

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Reproductive Females - 500 ppm (continued)

Animal ID	Clinical Sign	Study Day 30	31	DPE 32	33	34	35	36	37	38	DPE 39	40	41	42	43	44
Group 3 - 500 ppm																
D0021	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D
D0022	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0023	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	D
D0024	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D
D0025	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0026	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D
D0027	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D
D0028	Normal	P	P	P	P	P	P	P	P	P	P	P	D	D	D	D
D0029	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	D	D
D0030	Normal	P	P	P	P	P	P	P	P	P	P	P	P	D	D	D

DPE = Detailed physical exam
P = Present
N = Not Present
D = Deceased

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Reproductive Females - 2500 ppm

Animal ID	Clinical Sign	DPE	Study Day 1	2	3	DPE 4	5	6	7	8	9	10	DPE 11	12	13	14
Group 4 - 2500 ppm																
D0031	Normal	P	P	P	P	P	P	P	N	N	P	N	N	N	P	N
D0031	Anogenital Staining	N	N	N	N	N	N	N	P	N	N	N	N	N	N	N
D0031	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	P	P	N	P	P	P	N	P
D0032	Normal	P	N	P	P	N	N	P	P	P	P	P	N	P	P	N
D0032	Anogenital Staining	N	P	N	N	P	N	N	N	N	N	N	N	N	N	N
D0032	Probable Porphyrin Staining, head	N	N	N	N	N	P	N	N	N	N	N	P	N	N	P
D0033	Normal	P	P	N	N	N	N	N	P	N	P	N	N	P	P	P
D0033	Anogenital Staining	N	N	P	P	P	P	P	N	P	N	P	P	N	N	N
D0033	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	N	N	N	P	N	N	N
D0034	Normal	P	P	P	P	N	P	P	P	P	N	P	P	P	P	P
D0034	Anogenital Staining	N	N	N	N	P	N	N	N	N	P	N	N	N	N	N
D0035	Normal	P	P	P	P	P	P	P	P	P	N	N	N	P	P	N
D0035	Anogenital Staining	N	N	N	N	N	N	N	N	N	P	P	P	N	N	N
D0035	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	N	N	P	N	N	N	P
D0035	Salivation	N	N	N	N	N	N	N	N	N	N	N	P	N	N	N
D0036	Normal	P	P	P	N	N	N	P	P	P	P	P	P	P	P	P
D0036	Anogenital Staining	N	N	N	P	P	P	N	N	N	N	N	N	N	N	N
D0037	Normal	P	P	N	P	N	P	P	P	P	P	P	P	P	P	P
D0037	Anogenital Staining	N	N	P	N	P	N	N	N	N	N	N	N	N	N	N
D0038	Normal	P	P	N	P	N	N	N	N	N	N	N	N	N	N	N
D0038	Anogenital Staining	N	N	P	N	P	N	N	N	P	N	N	N	N	N	N
D0038	Probable Porphyrin Staining, head	N	N	N	N	P	P	P	P	P	P	P	P	P	P	P
D0039	Normal	P	P	P	P	N	P	P	P	P	P	P	P	P	P	P
D0039	Anogenital Staining	N	N	N	N	P	N	N	N	N	N	N	N	N	N	N
D0039	Probable Porphyrin Staining, head	N	N	N	N	P	N	N	N	N	N	N	N	N	N	N
D0120	Normal	P	P	P	P	N	N	N	P	P	P	P	P	P	P	P
D0120	Anogenital Staining	N	N	N	N	P	P	P	N	N	N	N	N	N	N	N

DPE = Detailed physical exam
P = Present
N = Not Present
D = Deceased

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Reproductive Females - 2500 ppm (continued)

Animal ID	Clinical Sign	Study Day 15	16	17	DPE 18	19	20	21	22	23	24	DPE 25	26	27	28	29
Group 4 - 2500 ppm																
D0031	Normal	N	N	N	N	N	N	N	N	P	P	P	N	N	N	N
D0031	Probable Porphyrin Staining, head	P	P	P	P	P	P	P	P	N	N	N	P	P	P	P
D0032	Normal	P	P	N	N	P	P	P	P	P	P	P	P	P	P	P
D0032	Probable Porphyrin Staining, head	N	N	P	P	N	N	N	N	N	N	N	N	N	N	N
D0033	Normal	N	N	N	N	P	P	P	P	N	N	P	P	P	P	P
D0033	Anogenital Staining	P	P	P	P	N	N	N	N	P	P	N	N	N	N	N
D0033	Probable Porphyrin Staining, head	P	N	P	P	N	N	N	N	P	P	N	N	N	N	N
D0034	Normal	P	P	P	P	P	P	P	P	N	P	P	P	P	P	P
D0034	Probable Porphyrin Staining, head	N	N	N	N	N	N	N	N	P	N	N	N	N	N	N
D0035	Normal	P	P	P	N	P	P	N	N	N	N	P	N	N	P	P
D0035	Anogenital Staining	N	N	N	P	N	N	N	N	N	P	N	P	P	N	N
D0035	Probable Porphyrin Staining, head	N	N	N	N	N	N	P	P	P	N	N	P	P	N	N
D0036	Normal	P	N	N	P	N	N	N	N	P	P	N	N	N	P	N
D0036	Anogenital Staining	N	P	P	N	N	N	N	N	N	N	N	N	N	N	N
D0036	Probable Porphyrin Staining, head	N	N	N	N	P	P	P	P	N	N	P	P	P	N	P
D0037	Normal	N	P	P	P	P	P	N	N	P	P	P	P	P	P	P
D0037	Probable Porphyrin Staining, head	P	N	N	N	N	N	P	P	N	N	N	N	N	N	N
D0037	Toenail Bleeding	P	N	N	N	N	N	N	N	N	N	N	N	N	N	N
D0038	Normal	N	N	P	N	N	N	P	P	N	N	P	P	P	P	P
D0038	Probable Porphyrin Staining, head	P	P	N	P	P	P	N	N	P	P	N	N	N	N	N
D0039	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0120	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P

DPE = Detailed physical exam
P = Present
N = Not Present
D=Deceased

Appendix C
Individual Clinical Observations and Detailed Physical Exam Data

Reproductive Females - 2500 ppm (continued)

Animal ID	Clinical Sign	Study Day 30	31	DPE 32	33	34	35	36	37	38	DPE 39	40	41	42	43	44
Group 4 - 2500 ppm																
D0031	Normal	P	P	N	N	N	N	N	P	P	P	P	P	P	D	D
D0031	Probable Porphyrin Staining, head	N	N	P	P	P	P	P	N	N	N	N	N	N	D	D
D0032	Normal	P	N	P	N	N	P	P	P	P	P	P	P	D	D	D
D0032	Probable Porphyrin Staining, head	N	P	N	P	P	N	N	N	N	N	N	N	D	D	D
D0033	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0034	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	D	D
D0035	Normal	P	N	N	N	N	P	P	P	P	P	P	P	P	D	D
D0035	Anogenital Staining	N	P	P	N	P	N	N	N	N	N	N	N	N	D	D
D0035	Probable Porphyrin Staining, head	N	P	N	P	P	N	N	N	N	N	N	N	N	D	D
D0035	Salivation	N	P	N	N	N	N	N	N	N	N	N	N	N	D	D
D0036	Normal	N	P	P	P	N	N	N	P	P	P	P	P	P	D	D
D0036	Probable Porphyrin Staining, head	P	N	N	N	P	P	P	N	N	N	N	N	N	D	D
D0037	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
D0038	Normal	N	N	N	P	P	N	N	P	P	P	P	P	P	P	D
D0038	Probable Porphyrin Staining, head	P	P	P	N	N	P	P	N	N	N	N	N	N	N	D
D0039	Normal	P	P	P	P	P	P	P	P	P	P	P	P	P	D	D
D0120	Normal	P	P	N	P	N	P	P	P	P	P	P	P	D	D	D
D0120	Probable Porphyrin Staining, head	N	N	P	N	P	N	N	N	N	N	N	N	D	D	D

DPE = Detailed physical exam
P = Present
N = Not Present
D=Deceased

Appendix D

Individual Animal Body Weights

Appendix D
Individual Animal Body Weights
Males (grams)

Animal ID	Day 1 03/25/02	Day 8 04/01/02	Day 15 04/08/02	Day 22 04/15/02	Day 29 04/22/02
Group 1 - Controls					
D0041	249.8	270.0	281.5	298.7	293.7
D0042	286.8	328.6	346.1	372.7	368.7
D0043	293.9	332.7	367.8	393.7	384.7
D0044	284.2	316.2	337.6	368.3	360.3
D0045	280.4	316.6	334.1	350.0	307.0
D0046	294.5	325.0	342.9	389.8	379.7
D0047	281.9	316.3	327.5	367.5	362.4
D0048	283.2	312.9	325.9	367.4	354.0
D0049	312.0	359.1	381.4	418.0	406.0
D0050	313.0	338.6	362.1	396.0	394.3
Group 2 - 100 ppm					
D0051	300.4	343.3	376.0	413.0	406.3
D0052	281.2	302.0	303.5	331.9	322.7
D0053	285.8	307.6	319.6	357.9	351.1
D0054	276.7	310.2	318.3	346.6	342.9
D0055	288.7	300.7	309.9	338.4	338.7
D0056	292.3	323.6	345.3	374.7	358.8
D0057	293.6	337.4	369.1	396.1	389.3
D0058	306.8	345.8	375.1	407.5	395.8
D0059	293.0	326.1	339.4	373.5	364.3
D0060	313.4	353.5	358.2	398.7	396.8
Group 3 - 500 ppm					
D0061	276.9	305.8	334.0	361.5	354.4
D0062	288.5	329.4	324.2	372.3	362.0
D0063	293.0	326.1	353.4	380.9	372.4
D0064	282.9	320.8	341.2	379.6	364.8
D0065	293.7	334.3	344.8	378.1	366.7
D0066	280.4	326.8	358.7	375.0	358.7
D0067	300.1	328.0	348.4	372.9	366.3
D0068	289.9	324.6	334.2	365.2	349.9
D0069	300.5	338.4	358.8	395.0	382.3
D0070	310.6	343.3	359.5	389.0	391.4
Group 4 - 2500 ppm					
D0071	296.3	327.6	350.3	384.7	363.1
D0072	291.5	317.5	330.4	348.4	331.8
D0073	282.5	295.5	308.0	326.2	318.4
D0074	300.7	324.5	347.8	370.2	355.8
D0075	271.6	283.7	275.3	299.8	276.6
D0076	290.4	310.7	327.2	348.6	336.0
D0077	284.6	311.0	327.2	344.3	329.7
D0078	292.6	314.4	326.4	347.0	329.3
D0079	310.2	336.0	356.4	384.9	374.2
D0080	312.5	330.0	341.4	368.1	361.9

Appendix D
Individual Animal Body Weights
Toxicity Females (grams)

Animal ID	Day 1 03/25/02	Day 8 04/01/02	Day 15 04/08/02	Day 22 04/15/02	Day 30 04/23/02
Group 1 - Controls					
D0081	209.4	231.8	246.5	258.5	244.7
D0082	206.8	225.4	237.4	230.1	222.1
D0083	216.7	237.3	246.0	250.1	235.8
D0084	205.6	224.0	243.3	258.6	239.5
D0085	215.0	241.3	259.5	262.6	248.6
D0086	212.9	241.1	245.7	262.1	245.9
D0087	217.4	239.8	250.2	267.6	244.2
D0088	213.8	231.5	246.9	259.3	240.0
D0089	223.2	227.8	255.0	257.7	247.9
D0090	223.8	244.6	252.4	261.3	249.8
Group 2 - 100 ppm					
D0091	195.6	216.5	226.5	231.8	216.1
D0092	205.9	220.7	242.5	255.8	234.6
D0093	207.3	226.3	236.9	247.0	232.9
D0094	220.8	243.1	255.3	251.6	247.3
D0095	206.8	230.4	237.4	254.5	241.0
D0096	209.8	227.6	247.4	253.8	234.4
D0097	212.7	231.7	239.1	251.4	235.6
D0098	213.8	235.9	253.5	253.6	237.8
D0099	225.8	240.0	242.5	257.6	244.3
D0100	225.2	242.9	256.8	268.1	255.7
Group 3 - 500 ppm					
D0101	203.7	221.4	233.5	237.9	225.3
D0102	204.2	215.5	221.8	228.5	218.9
D0103	206.4	215.2	218.5	225.1	214.3
D0104	204.8	215.1	221.2	219.8	210.6
D0105	207.0	228.6	238.9	247.2	229.4
D0106	205.2	233.2	249.8	250.9	238.9
D0107	213.9	229.9	238.6	251.5	232.9
D0108	225.0	243.1	258.8	269.0	253.6
D0109	212.4	224.5	246.8	253.0	236.7
D0110	218.9	249.1	257.6	255.6	237.9
Group 4 - 2500 ppm					
D0111	186.5	202.6	207.3	229.1	210.6
D0112	212.5	234.0	242.0	250.8	241.9
D0113	212.2	228.7	246.0	258.2	234.2
D0114	205.4	218.6	234.2	244.5	223.8
D0115	195.4	207.2	218.2	223.7	207.8
D0116	209.8	221.9	228.0	237.3	219.6
D0117	203.8	219.3	222.0	242.5	227.7
D0118	210.1	238.6	274.4	270.1	244.0
D0119	214.9	225.5	252.2	257.4	238.3
D0040	213.6	217.1	229.2	231.7	222.4

Appendix D
Individual Animal Body Weights

Reproductive Females (grams)

Animal ID	Day 1 03/25/02	Day 8 04/01/02	Day 15 04/08/02	Gestation Day 0	Gestation Day 7	Gestation Day 14	Gestation Day 20	Post partum Day 0/1	Post Partum Day 4
Group 1 - Controls									
D0001	199.5	211.7	215.0	217.1	250.9	285.5	350.5	270.6	282.5
D0002	204.3	218.7	234.7	241.2	275.1	309.3	375.9	280.1	310.6
D0003	204.4	209.1	227.2	222.3	251.1	278.1	337.0	271.5	291.8
D0004	209.2	220.5	233.1	233.1	268.1	314.3	368.6	274.9	291.7
D0005	203.2	228.1	242.0	234.6	258.9	288.1	365.0	270.0	286.5
D0006	221.2	242.0	256.9	247.7	287.2	327.1	393.6	295.9	319.1
D0007	227.8	244.4	259.6	259.6	291.3	328.9	394.6	314.6	324.0
D0008	223.5	241.8	263.1	263.1	288.5	328.5	401.5	300.5	317.8
D0009	210.3	231.7	249.2	248.8	268.8	294.1	379.1	271.0	289.4
D0010	232.3	247.9	269.6	269.6	298.6	342.8	402.9	307.5	331.1
Group 2 - 100 ppm									
D0011	208.5	219.2	234.8	234.8	265.1	304.0	381.1	289.9	302.2
D0012	192.7	206.4	226.7	226.7	247.5	283.6	358.4	273.8	256.9
D0013	201.9	215.7	233.0	233.0	261.1	298.3	376.3	280.1	297.8
D0014	208.6	227.6	235.0	235.0	261.1	298.0	373.5	265.2	304.2
D0015	218.2	233.9	256.6	254.8	270.5	316.7	403.4	313.4	300.6
D0016	220.4	242.5	263.4	263.4	308.0	353.0	409.8	326.6	329.4
D0017	216.8	234.3	260.8	250.8	305.0	339.9	407.5	308.7	330.8
D0018	227.0	244.4	255.3	256.0	283.8	315.7	385.5	288.2	315.9
D0019	230.1	256.7	272.3	272.3	296.9	337.6	410.5	307.6	338.1
D0020	221.9	244.3	244.1	244.1	281.4	322.9	411.5	322.8	317.0

Appendix D
Individual Animal Body Weights

Reproductive Females (grams) - continued

Animal ID	Day 1 03/25/02	Day 8 04/01/02	Day 15 04/08/02	Gestation Day 0	Gestation Day 7	Gestation Day 14	Gestation Day 20	Post partum Day 0/1	Post Partum Day 4
Group 3 - 500 ppm									
D0021	197.7	205.3	218.5	218.5	263.6	305.6	376.4	294.3	315.7
D0022	201.0	215.8	223.8	224.8	245.2	273.7	355.9	266.4	272.5
D0023	196.7	210.4	212.2	212.8	227.6	263.0	320.3	250.2	260.4
D0024	217.5	235.8	249.6	249.6	263.8	295.2	369.1	280.7	301.0
D0025	221.0	230.3	232.2	245.8	270.4	303.1	397.1	277.4	320.7
D0026	210.8	228.7	232.6	232.6	257.7	286.8	363.4	261.4	290.4
D0027	216.1	231.3	240.7	240.7	270.8	298.1	383.9	269.0	299.3
D0028	221.7	236.9	257.1	257.1	278.9	324.7	400.9	315.7	321.9
D0029	203.0	225.3	241.8	229.3	267.5	298.4	337.0	289.2	297.8
D0030	238.4	261.3	280.7	280.7	302.0	329.6	408.8	295.6	325.8
Group 4 - 2500 ppm									
D0031	197.4	211.3	222.0	201.4	250.2	274.9	303.8	285.4	269.7
D0032	203.9	211.8	230.6	230.6	245.3	281.2	332.9	273.2	282.2
D0033	211.4	230.4	249.3	NA	NA	NA	NA	NA	NA
D0034	206.9	217.4	231.8	231.8	264.0	297.7	338.9	301.3	307.2
D0035	207.9	219.5	241.5	218.7	248.1	278.7	336.8	248.2	251.6
D0036	213.9	217.0	225.2	211.5	248.7	277.9	344.4	269.4	290.0
D0037	213.3	233.6	234.4	234.9	266.1	293.1	366.5	274.8	294.2
D0038	218.1	234.4	256.5	253.4	264.3	317.4	379.5	302.9	305.5
D0039	222.3	230.2	245.1	235.0	275.2	302.6	356.8	278.2	304.6
D0120	217.6	232.5	242.8	242.8	266.3	302.5	345.4	290.0	308.3

NA = Not Applicable (Animal not pregnant)

Appendix E

Individual Animal Body Weight Gains

Appendix E
Individual Animal Body Weight Gains

Males (grams)

Animal ID	Week 1	Week 2	Week 3	Week 4	Overall Weight Gain
Group 1 - Controls					
D0041	20.2	11.5	17.2	-5.0	43.9
D0042	41.8	17.5	26.6	-4.0	81.9
D0043	38.8	35.1	25.9	-9.0	90.8
D0044	32.0	21.4	30.7	-8.0	76.1
D0045	36.2	17.5	15.9	-43.0	26.6
D0046	30.5	17.9	46.9	-10.1	85.2
D0047	34.4	11.2	40.0	-5.1	80.5
D0048	29.7	13.0	41.5	-13.4	70.8
D0049	47.1	22.3	36.6	-12.0	94.0
D0050	25.6	23.5	33.9	-1.7	81.3
Group 2 - 100 ppm					
D0051	42.9	32.7	37.0	-6.7	105.9
D0052	20.8	1.5	28.4	-9.2	41.5
D0053	21.8	12.0	38.3	-6.8	65.3
D0054	33.5	8.1	28.3	-3.7	66.2
D0055	12.0	9.2	28.5	0.3	50.0
D0056	31.3	21.7	29.4	-15.9	66.5
D0057	43.8	31.7	27.0	-6.8	95.7
D0058	39.0	29.3	32.4	-11.7	89.0
D0059	33.1	13.3	34.1	-9.2	71.3
D0060	40.1	4.7	40.5	-1.9	83.4
Group 3 - 500 ppm					
D0061	28.9	28.2	27.5	-7.1	77.5
D0062	40.9	-5.2	48.1	-10.3	73.5
D0063	33.1	27.3	27.5	-8.5	79.4
D0064	37.9	20.4	38.4	-14.8	81.9
D0065	40.6	10.5	33.3	-11.4	73.0
D0066	46.4	31.9	16.3	-16.3	78.3
D0067	27.9	20.4	24.5	-6.6	66.2
D0068	34.7	9.6	31.0	-15.3	60.0
D0069	37.9	20.4	36.2	-12.7	81.8
D0070	32.7	16.2	29.5	2.4	80.8
Group 4 - 2500 ppm					
D0071	31.3	22.7	34.4	-21.6	66.8
D0072	26.0	12.9	18.0	-16.6	40.3
D0073	13.0	12.5	18.2	-7.8	35.9
D0074	23.8	23.3	22.4	-14.4	55.1
D0075	12.1	-8.4	24.5	-23.2	5.0
D0076	20.3	16.5	21.4	-12.6	45.6
D0077	26.4	16.2	17.1	-14.6	45.1
D0078	21.8	12.0	20.6	-17.7	36.7
D0079	25.8	20.4	28.5	-10.7	64.0
D0080	17.5	11.4	26.7	-6.2	49.4

Appendix E
Individual Animal Body Weight Gains

Toxicity Females (grams)

Animal ID	Week 1	Week 2	Week 3	Week 4	Overall Weight Gain
Group 1 - Controls					
D0081	22.4	14.7	12.0	-13.8	35.3
D0082	18.6	12.0	-7.3	-8.0	15.3
D0083	20.6	8.7	4.1	-14.3	19.1
D0084	18.4	19.3	15.3	-19.1	33.9
D0085	26.3	18.2	3.1	-14.0	33.6
D0086	28.2	4.6	16.4	-16.2	33.0
D0087	22.4	10.4	17.4	-23.4	26.8
D0088	17.7	15.4	12.4	-19.3	26.2
D0089	4.6	27.2	2.7	-9.8	24.7
D0090	20.8	7.8	8.9	-11.5	26.0
Group 2 - 100 ppm					
D0091	20.9	10.0	5.3	-15.7	20.5
D0092	14.8	21.8	13.3	-21.2	28.7
D0093	19.0	10.6	10.1	-14.1	25.6
D0094	22.3	12.2	-3.7	-4.3	26.5
D0095	23.6	7.0	17.1	-13.5	34.2
D0096	17.8	19.8	6.4	-19.4	24.6
D0097	19.0	7.4	12.3	-15.8	22.9
D0098	22.1	17.6	0.1	-15.8	24.0
D0099	14.2	2.5	15.1	-13.3	18.5
D0100	17.7	13.9	11.3	-12.4	30.5
Group 3 - 500 ppm					
D0101	17.7	12.1	4.4	-12.6	21.6
D0102	11.3	6.3	6.7	-9.6	14.7
D0103	8.8	3.3	6.6	-10.8	7.9
D0104	10.3	6.1	-1.4	-9.2	5.8
D0105	21.6	10.3	8.3	-17.8	22.4
D0106	28.0	16.6	1.1	-12.0	33.7
D0107	16.0	8.7	12.9	-18.6	19.0
D0108	18.1	15.7	10.2	-15.4	28.6
D0109	12.1	22.3	6.2	-16.3	24.3
D0110	30.2	8.5	-2.0	-17.7	19.0
Group 4 - 2500 ppm					
D0111	16.1	4.7	21.8	-18.5	24.1
D0112	21.5	8.0	8.8	-8.9	29.4
D0113	16.5	17.3	12.2	-24.0	22.0
D0114	13.2	15.6	10.3	-20.7	18.4
D0115	11.8	11.0	5.5	-15.9	12.4
D0116	12.1	6.1	9.3	-17.7	9.8
D0117	15.5	2.7	20.5	-14.8	23.9
D0118	28.5	35.8	-4.3	-26.1	33.9
D0119	10.6	26.7	5.2	-19.1	23.4
D0040	3.5	12.1	2.5	-9.3	8.8

Dow Corning Corporation
Study Number: 9658

Appendix E
Individual Animal Body Weight Gains

Reproductive Females (grams)

Animal ID	Week 1 Pre Mating	Week 2 Pre Mating	Mating 0-5 days	Gestation Week 1	Gestation Week 2	Gestation Week 3	Post Delivery	4 days Post Partum	Overall Weight Gain
Group 1 - Controls									
D0001	12.2	3.3	2.1	33.8	34.6	65.0	-79.9	11.9	83.0
D0002	14.4	16.0	6.5	33.9	34.2	66.6	-95.8	30.5	106.3
D0003	4.7	18.1	-4.9	28.8	27.0	58.9	-65.5	20.3	87.4
D0004	11.3	12.6	0.0	35.0	46.2	54.3	-93.7	16.8	82.5
D0005	24.9	13.9	-7.4	24.3	29.2	76.9	-95.0	16.5	83.3
D0006	20.8	14.9	-9.2	39.5	39.9	66.5	-97.7	23.2	97.9
D0007	16.6	15.2	0.0	31.7	37.6	65.7	-80.0	9.4	96.2
D0008	18.3	21.3	0.0	25.4	40.0	73.0	-101.0	17.3	94.3
D0009	21.4	17.5	-0.4	20.0	25.3	85.0	-108.1	18.4	79.1
D0010	15.6	21.7	0.0	29.0	44.2	60.1	-95.4	23.6	98.8
Group 2 - 100 ppm									
D0011	10.7	15.6	0.0	30.3	38.9	77.1	-91.2	12.3	93.7
D0012	13.7	20.3	0.0	20.8	36.1	74.8	-84.6	-16.9	64.2
D0013	13.8	17.3	0.0	28.1	37.2	78.0	-96.2	17.7	95.9
D0014	19.0	7.4	0.0	26.1	36.9	75.5	-108.3	39.0	95.6
D0015	15.7	22.7	-1.8	15.7	46.2	86.7	-90.0	-12.8	82.4
D0016	22.1	20.9	0.0	44.6	45.0	56.8	-83.2	2.8	109.0
D0017	17.5	26.5	-10.0	54.2	34.9	67.6	-98.8	22.1	114.0
D0018	17.4	10.9	0.7	27.8	31.9	69.8	-97.3	27.7	88.9
D0019	26.6	15.6	0.0	24.6	40.7	72.9	-102.9	30.5	108.0
D0020	22.4	-0.2	0.0	37.3	41.5	88.6	-88.7	-5.8	95.1
Group 3 - 500 ppm									
D0021	7.6	13.2	0.0	45.1	42.0	70.8	-82.1	21.4	118.0
D0022	14.8	8.0	1.0	20.4	28.5	82.2	-89.5	6.1	71.5
D0023	13.7	1.8	0.6	14.8	35.4	57.3	-70.1	10.2	63.7
D0024	18.3	13.8	0.0	14.2	31.4	73.9	-88.4	20.3	83.5
D0025	9.3	1.9	13.6	24.6	32.7	94.0	-119.7	43.3	99.7
D0026	17.9	3.9	0.0	25.1	29.1	76.6	-102.0	29.0	79.6
D0027	15.2	9.4	0.0	30.1	27.3	85.8	-114.9	30.3	83.2
D0028	15.2	20.2	0.0	21.8	45.8	76.2	-85.2	6.2	100.2
D0029	22.3	16.5	-12.5	38.2	30.9	38.6	-47.8	8.6	94.8
D0030	22.9	19.4	0.0	21.3	27.6	79.2	-113.2	30.2	87.4
Group 4 - 2500 ppm									
D0031	13.9	10.7	-20.6	48.8	24.7	28.9	-18.4	-15.7	72.3
D0032	7.9	18.8	0.0	14.7	35.9	51.7	-59.7	9.0	78.3
D0033	19.0	18.9	NA	NA	NA	NA	NA	NA	NA
D0034	10.5	14.4	0.0	32.2	33.7	41.2	-37.6	5.9	100.3
D0035	11.6	22.0	-22.8	29.4	30.6	58.1	-88.6	3.4	43.7
D0036	3.1	8.2	-13.7	37.2	29.2	66.5	-75.0	20.6	76.1
D0037	20.3	0.8	0.5	31.2	27.0	73.4	-91.7	19.4	80.9
D0038	16.3	22.1	-3.1	10.9	53.1	62.1	-76.6	2.6	87.4
D0039	7.9	14.9	-10.1	40.2	27.4	54.2	-78.6	26.4	82.3
D0120	14.9	10.3	0.0	23.5	36.2	42.9	-55.4	18.3	90.7

NA = Not Applicaple (Animal not pregnant)

Appendix F

Individual Animal Food Consumption

Appendix F
Individual Animal Food Consumption

Males (grams)

Animal ID	Week 1	Week 2	Total
Group 1 - Controls			
D0041	145.8	123.2	269.0
D0042	204.5	169.3	373.8
D0043	170.6	151.8	322.4
D0044	159.7	137.5	297.2
D0045	170.0	139.8	309.8
D0046	163.5	145.4	308.9
D0047	172.0	152.3	324.3
D0048	168.4	151.7	320.1
D0049	197.9	166.5	364.4
D0050	178.2	160.0	338.2
Group 2 - 100 ppm			
D0051	178.6	162.8	341.4
D0052	143.1	116.5	259.6
D0053	155.9	137.3	293.2
D0054	157.0	136.3	293.3
D0055	164.0	139.5	303.5
D0056	156.7	141.9	298.6
D0057	176.9	154.6	331.5
D0058	192.2	168.2	360.4
D0059	166.3	152.8	319.1
D0060	199.1	163.6	362.7
Group 3 - 500 ppm			
D0061	156.9	140.0	296.9
D0062	183.3	160.1	343.4
D0063	174.5	147.7	322.2
D0064	159.9	142.9	302.8
D0065	178.9	158.6	337.5
D0066	175.5	149.6	325.1
D0067	184.3	152.5	336.8
D0068	165.4	145.8	311.2
D0069	187.0	164.5	351.5
D0070	172.7	150.3	323.0
Group 4 - 2500 ppm			
D0071	179.0	132.8	311.8
D0072	153.1	122.9	276.0
D0073	122.0	109.2	231.2
D0074	156.1	144.0	300.1
D0075	147.2	118.4	265.6
D0076	157.9	134.3	292.2
D0077	159.5	138.5	298.0
D0078	153.5	122.9	276.4
D0079	180.5	153.4	333.9
D0080	145.6	138.5	284.1

Appendix F
Individual Animal Food Consumption

Toxicity Females (grams)

Animal ID	Week 1	Week 2	Week 3	Week 4	Total
Group 1 - Controls					
D0081	138.9	119.5	187.1	148.5	594.0
D0082	125.2	101.8	141.9	122.9	491.8
D0083	152.6	128.5	166.6	147.5	595.2
D0084	136.9	132.9	187.6	134.9	592.3
D0085	154.4	129.0	175.6	150.4	609.4
D0086	146.7	124.6	172.1	139.5	582.9
D0087	145.0	127.2	171.2	138.8	582.2
D0088	129.3	110.5	153.5	128.6	521.9
D0089	134.0	134.0	170.6	147.7	586.3
D0090	150.5	128.9	164.9	152.2	596.5
Group 2 - 100 ppm					
D0091	128.3	106.3	149.6	128.3	512.5
D0092	142.0	121.0	174.4	139.0	576.4
D0093	137.4	103.6	151.3	124.3	516.6
D0094	143.8	114.7	173.4	154.0	585.9
D0095	129.4	122.1	160.8	120.7	533.0
D0096	129.3	112.5	159.5	129.5	530.8
D0097	138.2	119.8	162.6	139.5	560.1
D0098	121.0	100.8	138.6	115.5	475.9
D0099	135.4	116.4	165.2	138.5	555.5
D0100	142.0	126.3	181.8	143.6	593.7
Group 3 - 500 ppm					
D0101	118.4	101.7	148.7	121.1	489.9
D0102	126.9	97.6	137.7	121.9	484.1
D0103	138.7	114.7	140.4	129.4	523.2
D0104	123.0	105.9	161.0	137.4	527.3
D0105	128.1	113.4	156.6	127.0	525.1
D0106	139.7	114.1	157.2	137.8	548.8
D0107	126.4	110.1	157.6	120.4	514.5
D0108	144.0	124.4	165.6	135.4	569.4
D0109	132.7	123.9	164.5	143.9	565.0
D0110	161.1	116.1	167.2	139.1	583.5
Group 4 - 2500 ppm					
D0111	120.2	99.8	142.5	130.9	493.4
D0112	126.3	106.6	143.7	135.7	512.3
D0113	121.5	107.5	150.9	125.7	505.6
D0114	130.9	110.4	154.1	126.4	521.8
D0115	121.9	95.6	136.3	117.3	471.1
D0116	130.3	100.7	145.7	131.6	508.3
D0117	139.0	118.8	156.5	142.9	557.2
D0118	131.9	134.9	167.6	136.4	570.8
D0119	115.3	95.1	142.9	125.7	479.0
D0040	106.7	109.4	156.5	146.4	519.0

Appendix F
Individual Animal Food Consumption

Reproductive Females (grams)

Animal ID	Pre-mating Week 1	Pre-mating Week 2	Gestation Week 1	Gestation Week 2	Gestation Week 3	Gest. Day 20 to PP Day 0/1	Periparturiant Week	Total
Group 1 - Controls								
D0001	121.1	102.8	155.6	175.7	139.9	32.9	95.8	823.8
D0002	129.5	117.8	159.4	178.7	139.3	55.3	112.5	892.5
D0003	128.8	110.1	157.5	171.2	126.1	38.9	120.3	852.9
D0004	124.3	120.9	176.4	193.9	121.3	36.5	118.2	891.5
D0005	137.5	116.2	146.2	158.3	126.0	42.7	119.0	845.9
D0006	140.9	125.7	157.0	169.8	148.7	36.4	145.4	923.9
D0007	138.0	127.0	175.1	195.7	154.6	27.3	98.4	916.1
D0008	136.0	117.9	169.3	183.2	148.6	26.8	140.8	922.6
D0009	141.8	127.5	165.3	164.0	218.9	21.4	95.1	934.0
D0010	151.3	131.0	181.5	197.2	148.9	42.1	130.8	982.8
Group 2 - 100 ppm								
D0011	116.6	110.2	155.1	168.0	150.2	24.0	106.2	830.3
D0012	111.6	103.6	136.7	152.7	136.4	39.8	54.2	735.0
D0013	123.5	110.8	151.3	153.7	133.8	36.8	98.8	808.7
D0014	114.4	102.6	131.9	152.5	125.9	35.6	122.7	785.6
D0015	138.5	124.3	184.4	211.7	192.6	38.2	105.9	995.6
D0016	147.6	130.0	195.8	208.6	143.3	37.6	113.1	976.0
D0017	162.5	145.2	214.4	213.9	162.5	21.6	137.8	1057.9
D0018	143.2	113.1	159.3	180.3	127.9	37.7	128.1	889.6
D0019	156.5	127.6	163.7	178.0	129.2	40.1	180.8	975.9
D0020	139.7	120.3	168.3	191.0	155.8	30.6	113.3	919.0
Group 3 - 500 ppm								
D0021	110.8	104.4	169.4	186.6	135.3	46.8	116.3	869.6
D0022	114.0	90.3	125.5	151.5	119.5	41.9	101.0	743.7
D0023	109.9	88.7	125.1	143.1	115.0	37.2	92.6	711.6
D0024	141.6	116.0	150.7	159.8	143.9	41.2	140.4	893.6
D0025	135.5	123.2	171.6	178.5	149.6	46.4	158.6	963.4
D0026	129.4	117.0	146.6	159.0	143.3	33.6	140.9	869.8
D0027	131.5	119.7	160.4	162.4	142.4	35.9	127.0	879.3
D0028	131.0	117.8	143.5	182.2	168.1	33.2	126.5	902.3
D0029	129.2	113.2	164.3	169.5	145.9	46.1	88.4	856.6
D0030	165.8	136.8	186.4	182.7	143.8	40.1	119.2	974.8
Group 4 - 2500 ppm								
D0031	129.8	104.5	153.7	172.3	176.7	52.9	57.1	847.0
D0032	108.1	97.2	135.7	159.7	134.1	45.1	102.9	782.8
D0033	135.6	120.4	NP	NP	NP	NP	NP	NP
D0034	123.9	114.0	161.6	189.9	151.1	84.4	100.7	925.6
D0035	110.4	107.9	130.4	132.0	126.5	43.8	68.5	719.5
D0036	111.8	101.5	132.5	156.0	135.8	49.6	124.9	812.1
D0037	125.4	98.4	135.5	156.5	130.4	42.8	131.9	820.9
D0038	143.7	124.4	168.2	194.0	150.5	55.2	130.3	966.3
D0039	113.0	108.5	149.6	165.6	148.3	59.3	98.3	842.6
D0120	122.4	114.9	141.8	161.0	136.0	49.8	118.6	844.5

NP = Non pregnant (Values not use in calculations.)

Appendix G

Individual Animal FOB Ranked/Categorical Observations

Appendix G
Individual Animal FOB Ranked/Categorical Observations

Male Rats - Baseline

Animal ID	Resistance to Removal	Palpebral Closure	Pupil Reaction	Pupil Size	Lacrimation	Salivation	Muscle Tone	Extensor-Thrust	Reaction to Handling	Level of Activity	Response to Noise	Response to Touch	Response to Tail Pinch	Gait Evaluation	Urination	Defecation	Categoricals
Group 1 - Controls																	
D0041	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	none
D0042	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	none
D0043	3	3	3	3	3	3	3	3	3	3	3	3	3	1	2	1	none
D0044	2	3	3	3	3	3	3	3	3	3	3	3	3	1	4	1	none
D0045	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	2	none
D0046	2	3	3	3	3	3	3	3	2	3	2	3	3	1	2	1	none
D0047	2	3	3	3	3	3	3	3	2	3	3	3	3	1	1	1	none
D0048	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	2	none
D0049	3	3	3	3	3	3	3	3	3	3	3	3	3	1	2	3	none
D0050	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
Group 2 - 100 ppm																	
D0051	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
D0052	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	1	none
D0053	3	3	3	3	3	3	3	3	4	3	3	3	3	1	2	2	none
D0054	3	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	none
D0055	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	none
D0056	3	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
D0057	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
D0058	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	2	none
D0059	2	3	3	3	3	3	3	3	3	3	3	3	2	1	3	3	none
D0060	2	3	3	3	3	3	3	3	3	3	4	3	3	1	2	2	none

Appendix G
Individual Animal FOB Ranked/Categorical Observations

Male Rats - Baseline

Animal ID	Resistance to Removal	Palpebral Closure	Pupil Reaction	Pupil Size	Lacrimation	Salivation	Muscle Tone	Extensor-Thrust	Reaction to Handling	Level of Activity	Response to Noise	Response to Touch	Response to Tail Pinch	Gait Evaluation	Urination	Defecation	Categoricals
Group 3 - 500 ppm																	
D0061	3	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	Feces: soft
D0062	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	2	none
D0063	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	3	none
D0064	2	3	3	3	3	3	3	3	4	3	3	3	3	1	2	1	none
D0065	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
D0066	2	3	3	3	3	3	3	3	2	3	3	3	3	1	4	3	none
D0067	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	1	none
D0068	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	1	none
D0069	3	3	3	3	3	3	3	3	2	3	3	3	3	1	1	1	none
D0070	3	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	none
Group 4 - 2500 ppm																	
D0071	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	3	Feces: soft
D0072	2	3	3	3	3	3	3	3	2	3	3	3	3	1	1	2	none
D0073	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	1	none
D0074	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	1	none
D0075	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
D0076	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	2	none
D0077	2	3	3	3	3	3	3	3	3	3	2	3	3	1	2	2	none
D0078	2	3	3	3	3	3	3	3	2	3	3	3	2	1	3	1	none
D0079	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
D0080	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	2	none

Appendix G
Individual Animal FOB Ranked/Categorical Observations

Toxicity Group Female Rats - Baseline

Animal ID	Resistance to Removal	Palpebral Closure	Pupil Reaction	Pupil Size	Lacrimation	Salivation	Muscle Tone	Extensor-Thrust	Reaction to Handling	Level of Activity	Response to Noise	Response to Touch	Response to Tail Pinch	Gait Evaluation	Urination	Defecation	Categoricals
Group 1 - Controls																	
D0081	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0082	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
D0083	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	3	none
D0084	3	3	3	3	3	3	3	3	4	3	3	3	3	1	2	1	none
D0085	3	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0086	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	none
D0087	3	3	3	3	3	3	3	3	3	3	3	3	3	1	1	2	none
D0088	3	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0089	3	3	3	3	3	3	3	3	3	3	3	3	2	1	1	3	none
D0090	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	1	none
Group 2 - 100 ppm																	
D0091	2	3	4	3	3	3	3	3	3	3	3	3	3	1	2	2	none
D0092	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	2	none
D0093	3	3	3	3	3	3	3	3	3	3	3	3	3	1	3	1	none
D0094	3	3	3	3	3	3	3	3	3	3	3	3	2	1	2	2	none
D0095	2	3	3	3	3	3	3	3	3	3	2	3	3	1	2	3	none
D0096	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0097	3	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0098	2	3	3	3	3	3	3	3	2	3	3	3	2	1	2	3	none
D0099	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
D0100	3	3	3	3	3	3	3	3	3	3	3	3	3	1	1	3	none

Appendix G
Individual Animal FOB Ranked/Categorical Observations

Toxicity Group Female Rats - Baseline

Animal ID	Resistance to Removal	Palpebral Closure	Pupil Reaction	Pupil Size	Lacrimation	Salivation	Muscle Tone	Extensor-Thrust	Reaction to Handling	Level of Activity	Response to Noise	Response to Touch	Response to Tail Pinch	Gait Evaluation	Urination	Defecation	Categoricals
Group 3 - 500 ppm																	
D0101	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0102	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	1	none
D0103	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	1	none
D0104	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	none
D0105	2	3	3	3	3	3	3	3	4	3	3	3	3	1	2	2	none
D0106	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	1	none
D0107	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	1	none
D0108	2	3	3	3	3	3	3	3	2	3	3	3	3	1	1	1	none
D0109	3	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0110	3	3	3	3	3	3	3	3	4	3	3	3	3	1	2	2	none
Group 4 - 2500 ppm																	
D0111	3	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0112	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0113	3	3	3	3	3	3	3	3	3	3	2	3	3	1	1	1	none
D0114	3	3	3	3	3	3	3	3	3	3	3	3	3	1	1	2	none
D0115	2	3	3	3	3	3	3	3	2	3	3	3	3	1	1	1	none
D0116	2	3	4	3	3	3	3	3	3	3	3	3	3	1	2	2	none
D0117	3	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0118	3	3	3	3	3	3	3	3	3	3	3	3	3	1	1	2	none
D0119	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	1	none
D0120*	3	3	3	3	3	3	3	3	3	4	3	3	3	1	2	1	none

*Animal was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Appendix G
Individual Animal FOB Ranked/Categorical Observations

Male Rats - Treated

Animal ID	Resistance to Removal	Palpebral Closure	Pupil Reaction	Pupil Size	Lacrimation	Salivation	Muscle Tone	Extensor-Thrust	Reaction to Handling	Level of Activity	Response to Noise	Response to Touch	Response to Tail Pinch	Gait Evaluation	Urination	Defecation	Categoricals
Group 1 - Controls																	
D0041	2	3	3	3	3	3	3	3	3	3	3	3	3	1	4	3	none
D0042	2	3	3	3	3	3	3	3	3	3	4	3	3	1	2	2	none
D0043	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	4	none
D0044	2	3	3	3	3	3	3	3	3	3	2	3	3	1	3	5	none
D0045	3	3	3	3	3	3	3	3	2	3	3	3	3	1	4	2	none
D0046	2	3	3	3	3	3	3	3	2	2	3	3	3	1	2	2	none
D0047	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	2	none
D0048	2	3	3	3	3	3	3	3	2	3	4	3	3	1	3	2	none
D0049	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	3	none
D0050	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
Group 2 - 100 ppm																	
D0051	2	3	3	3	3	3	3	3	3	2	3	3	3	1	2	2	none
D0052	2	3	3	3	3	3	3	3	3	3	3	3	4	1	3	3	none
D0053	2	3	3	3	3	3	3	3	3	3	3	3	3	1	4	4	none
D0054	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	3	none
D0055	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	3	Soiling: perineal, urine
D0056	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	3	none
D0057	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	2	none
D0058	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	2	none
D0059	2	3	3	3	3	3	3	3	3	2	3	3	3	1	3	5	none
D0060	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	none

Appendix G
Individual Animal FOB Ranked/Categorical Observations

Male Rats - Treated

Animal ID	Resistance to Removal	Palpebral Closure	Pupil Reaction	Pupil Size	Lacrimation	Salivation	Muscle Tone	Extensor-Thrust	Reaction to Handling	Level of Activity	Response to Noise	Response to Touch	Response to Tail Pinch	Gait Evaluation	Urination	Defecation	Categoricals
Group 3 - 500 ppm																	
D0061	2	3	3	3	3	3	3	3	2	2	3	3	3	1	3	2	none
D0062	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	none
D0063	2	3	3	3	3	3	3	3	2	3	3	3	3	1	4	2	none
D0064	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
D0065	2	3	3	3	3	3	3	3	3	3	3	3	3	1	4	5	none
D0066	2	3	3	3	3	3	3	3	2	3	3	3	3	1	4	2	none
D0067	2	3	3	3	3	3	3	3	3	3	3	3	3	1	4	2	none
D0068	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	3	none
D0069	2	3	3	3	3	3	3	2	2	3	3	3	3	1	3	1	none
D0070	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
Group 4 - 2500 ppm																	
D0071	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	2	none
D0072	2	3	3	3	3	3	3	3	2	2	3	3	3	1	2	2	none
D0073	2	3	3	3	3	3	3	3	2	3	3	3	3	1	1	1	none
D0074	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	1	none
D0075	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	2	none
D0076	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	2	none
D0077	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	4	none
D0078	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	3	none
D0079	2	3	3	3	3	3	3	3	3	3	3	3	3	1	4	2	none
D0080	2	3	3	3	3	3	3	3	3	2	3	3	3	1	3	2	none

Appendix G
Individual Animal FOB Ranked/Categorical Observations

Toxicity Group Female Rats - Treated

Animal ID	Resistance to Removal	Palpebral Closure	Pupil Reaction	Pupil Size	Lacrimation	Salivation	Muscle Tone	Extensor-Thrust	Reaction to Handling	Level of Activity	Response to Noise	Response to Touch	Response to Tail Pinch	Gait Evaluation	Urination	Defecation	Categoricals
Group 1 - Controls																	
D0081	2	3	3	3	3	3	3	3	3	2	3	3	3	1	2	1	none
D0082	2	3	3	3	3	3	3	3	3	2	3	3	3	1	3	2	none
D0083	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	5	none
D0084	2	3	3	3	3	3	3	3	4	3	3	3	3	1	3	2	none
D0085	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	1	none
D0086	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	3	none
D0087	2	3	3	3	3	3	3	3	3	3	4	3	3	1	3	1	none
D0088	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	5	none
D0089	2	3	3	3	3	3	3	3	3	3	5	3	3	1	2	4	none
D0090	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	1	none
Group 2 - 100 ppm																	
D0091	2	3	3	3	3	3	3	3	3	2	3	3	3	1	2	4	none
D0092	2	3	3	3	3	3	3	3	3	2	3	3	4	1	3	5	none
D0093	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	1	none
D0094	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	1	none
D0095	2	3	3	3	3	3	3	3	3	3	1	3	3	1	2	4	none
D0096	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	1	none
D0097	2	3	3	3	3	3	3	3	3	3	3	3	3	1	4	5	none
D0098	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	1	none
D0099	2	3	3	3	3	3	3	3	3	3	2	3	3	1	2	2	none
D0100	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	5	none

Appendix G
Individual Animal FOB Ranked/Categorical Observations

Toxicity Group Female Rats - Treated

Animal ID	Resistance to Removal	Palpebral Closure	Pupil Reaction	Pupil Size	Lacrimation	Salivation	Muscle Tone	Extensor-Thrust	Reaction to Handling	Level of Activity	Response to Noise	Response to Touch	Response to Tail Pinch	Gait Evaluation	Urination	Defecation	Categoricals
Group 3 - 500 ppm																	
D0101	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	4	none
D0102	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	1	none
D0103	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	3	none
D0104	2	3	3	3	3	3	3	3	3	2	3	3	3	1	3	2	none
D0105	2	3	3	3	3	3	3	3	3	3	3	3	3	1	3	2	none
D0106	2	3	3	3	3	3	3	3	2	3	2	3	3	1	2	3	none
D0107	2	3	3	3	3	3	3	3	3	2	3	3	3	1	3	1	none
D0108	2	3	3	3	3	3	3	3	3	3	4	3	3	1	1	2	none
D0109	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	2	none
D0110	2	3	3	3	3	3	3	3	3	1	3	3	3	1	2	1	none
Group 4 - 2500 ppm																	
D0111	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	1	none
D0112	2	3	3	3	3	3	3	3	3	3	3	3	3	1	1	1	none
D0113	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	1	none
D0114	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	1	none
D0115	2	3	3	3	3	3	3	3	2	3	3	3	3	1	2	1	none
D0116	2	3	3	3	3	3	3	3	2	3	3	3	3	1	3	1	none
D0117	2	3	3	3	3	3	3	3	3	3	3	3	3	1	2	1	none
D0118	2	3	3	3	3	3	3	3	2	3	2	3	3	1	2	3	none
D0119	2	3	3	3	3	3	3	3	2	3	4	3	3	1	3	1	none
D0120*	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-

*Animal was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Appendix G
Individual Animal FOB Ranked/Categorical Observations

Resistance to Removal:
1 = no resistance to capture and removal
2 = minimally avoids capture but is not aggressive
3 = moderate; actively avoids capture but is not aggressive
4 = pronounced; actively avoids capture and/or is mildly aggressive
5 = exaggerated; very difficult to capture and is very aggressive (i.e. tries to bite)

Palpebral closure:
1 = closed
2 = 50% closed
3 = open
4 = protruding eyes

Pupil size (aided by pen light):
1 = decreased pupil size (completely constricted)
2 = decreased pupil size (slightly constricted)
3 = normal pupil size
4 = increased pupil size (slightly dilated)
5 = increased pupil size (completely dilated)

Pupil reactivity (aided by pen light):
1 = dilated, nonreactive pupil
2 = dilated, slowly reactive pupil
3 = dilated, briskly reactive pupil
4 = constricted, cannot assess

Muscle tone:
1 = scored as none but defined as limp
2 = slight; very relaxed, "soft"
3 = moderate; animal is neither very relaxed nor tense
4 = pronounced; animal feels slightly rigid or tense
5 = exaggerated; very rigid or stiff

Extensor-thrust response:
1 = no extensor thrust response
2 = slight; barely detectable response
3 = moderate; clearly detectable
4 = pronounced; pushes vigorously
5 = exaggerated; extreme thrust response, greater than number 4

Reactivity to handling:
1 = none, no struggling
2 = minimal struggling
3 = moderate struggling with little or no vocalization
4 = pronounced struggling, animal may vocalize
5 = exaggerated; aggressive escape behavior, tries to bite observer and usually vocalizes

Level of activity :
1 = none, rat doesn't leave the center circle (20 cm dia.)
2 = minimal; inactive more than 50% of the time
3 = moderate; inactive less than 50% of the time
4 = pronounced; leaves circle, occasional inactivity
5 = exaggerated; leaves circle, never inactive

Responsiveness to sharp noise:
1 = none, no response to click
2 = minimal; only the ears move
3 = moderate; clear movement of the head, neck and ears
4 = pronounced; clear movement of head, neck and ears, plus clear movement of body
5 = exaggerated; whole body response to stimulus, more intense than 4

Responsiveness to touch:
1 = no response to touch
2 = slight; does not spontaneously lift tail; but briefly holds tail in air when manually lifted
3 = moderate; spontaneously lifts tail when touched
4 = pronounced; spontaneously lifts tail and acts startled
5 = exaggerated; turns towards finger in an attack response

Appendix G
Individual Animal FOB Ranked/Categorical Observations

Responsiveness to tail pinch (nociception):

1 = none; no response to pinch
2 = minimal; animal flinches but does not vocalize or move rapidly
3 = moderate; animal flinches, moves rapidly and/or vocalizes
4 = pronounced; animal moves very quickly and bends towards the tail and vocalizes
5 = exaggerated response to pinch; same as number 4, but includes an attempt to attack the source of the stimulation

Gait evaluation:

1 = smooth and coordinated gait
2 = slight incoordination; slight knuckling of the paws and/or slight tendency to stumble
3 = moderate incoordination; clear knuckling and stumbling, appears to be incoordinated
4 = pronounced incoordination; to include but not limited to: poor coordination, falling, and/or clear and repetitive knuckling of paws, and/or dragging of one or more limbs
5 = exaggerated; animal is unable to walk

Urination:

1 = no drops of urine
2 = minimal; a few (1-4) small drops or a small pool (~1 cm)
3 = moderate; more than one small pool, or a large pool (> 1 cm)
4 = pronounced; clearly a large amount of urine (more than 3)
5 = exaggerated; excessive amount of urine

Lacrimation (clear wetness):

1 = very dry appearance of cornea
2 = slightly dry appearance of cornea
3 = glistening cornea
4 = slight periocular wetness
5 = extensive periocular wetness

The degree of salivation:

1 = extensive oral dryness
2 = slight oral dryness
3 = glistening oral cavity/lips, no wetness
4 = slight perioral wetness
5 = extensive perioral wetness or drop(s) of fluid in or around mouth

Defecation:

1 = no pellets voided
2 = minimal; I or 2 pellets voided
3 = moderate; 3 pellets voided
4 = pronounced; 4 pellets voided
5 = exaggerated; 5 or more pellets voided

Categoricals:

Positive Observations made at any time regarding the following categories:

Behavior
Skin/fur/mucous membranes
Respiration
Muscle movements
Eyes
Urine or Feces
Soiling
General abnormalities
Posture

Appendix H

Individual Animal Rectal Temperature and Grip Strength/Foot Splay Data

Appendix H
Individual Animal Rectal Temperature and Grip Strength/Foot Splay Data

Males - Baseline

Animal	ID	Rectal Temperature (°Celsius)	Hindlimb Grip Strength (gram-force)[1]	Forelimb Grip Strength (gram-force)[1]	Landing Foot Splay (centimeters)[1]
Group 1 - Controls					
	D0041	37.4	510	865	15.0
	D0042	37.6	818	1100	13.8
	D0043	37.5	653	1220	12.7
	D0044	37.1	848	870	10.4
	D0045	37.6	682	902	14.0
	D0046	37.5	967	832	8.9
	D0047	37.2	1048	1008	12.7
	D0048	38.0	853	825	10.3
	D0049	38.0	822	522	16.4
	D0050	38.1	870	1058	13.4
Group 2 - 100 ppm					
	D0051	37.8	737	888	13.2
	D0052	37.8	780	818	11.2
	D0053	38.0	540	665	13.0
	D0054	37.1	1033	1067	10.3
	D0055	37.5	980	1098	14.9
	D0056	36.8	895	1007	10.9
	D0057	36.8	708	952	12.2
	D0058	37.8	913	873	11.0
	D0059	36.9	795	697	14.0
	D0060	37.6	1007	728	11.4
Group 3 - 500 ppm					
	D0061	37.9	797	872	12.3
	D0062	37.1	892	848	13.3
	D0063	37.8	720	987	10.3
	D0064	37.3	1297	1085	15.1
	D0065	38.0	690	1025	12.6
	D0066	37.8	608	738	12.0
	D0067	37.6	698	807	10.1
	D0068	37.1	842	907	10.0
	D0069	37.4	647	782	11.2
	D0070	37.4	690	1108	10.2
Group 4 - 2500 ppm					
	D0071	38.1	667	983	12.5
	D0072	37.3	812	788	9.1
	D0073	37.5	653	993	12.6
	D0074	37.1	818	1000	11.3
	D0075	36.6	743	1020	14.3
	D0076	37.3	545	1058	11.1
	D0077	37.5	720	852	14.3
	D0078	37.6	1135	1007	13.8
	D0079	37.6	528	710	13.1
	D0080	37.1	807	918	12.6

[1]Represents the mean of three trials

Appendix H
Individual Animal Rectal Temperature and Grip Strength/Foot Splay Data

Toxicity Females - Baseline

Animal	ID	Rectal Temperature (°Celsius)	Hindlimb Grip Strength (gram-force)[1]	Forelimb Grip Strength (gram-force)[1]	Landing Foot Splay (centimeters)[1]
Group 1 - Controls					
	D0081	37.4	720	777	12.6
	D0082	37.2	673	863	11.4
	D0083	37.8	530	842	9.3
	D0084	37.9	700	832	11.1
	D0085	38.2	568	760	10.6
	D0086	37.6	768	655	11.9
	D0087	38.1	775	1178	12.8
	D0088	37.8	702	705	10.6
	D0089	37.5	550	983	12.0
	D0090	37.0	757	755	11.5
Group 2 - 100 ppm					
	D0091	37.4	835	677	10.2
	D0092	36.7	502	792	9.0
	D0093	37.4	777	752	13.0
	D0094	37.0	600	883	11.1
	D0095	37.9	588	775	13.6
	D0096	37.3	822	1172	12.8
	D0097	37.7	718	787	9.3
	D0098	37.5	835	965	8.6
	D0099	37.3	542	638	11.6
	D0100	37.7	817	1012	11.8
Group 3 - 500 ppm					
	D0101	37.7	843	987	12.0
	D0102	37.9	620	1037	11.2
	D0103	38.2	660	710	13.1
	D0104	37.7	837	763	11.8
	D0105	38.5	833	852	13.5
	D0106	37.6	732	902	9.4
	D0107	37.6	850	1057	13.7
	D0108	37.3	768	948	13.2
	D0109	37.8	658	1017	10.5
	D0110	38.0	630	702	10.7
Group 4 - 2500 ppm					
	D0111	38.1	600	895	9.1
	D0112	37.9	677	1073	10.0
	D0113	38.0	943	842	14.0
	D0114	38.0	677	863	8.6
	D0115	37.8	793	747	11.3
	D0116	37.7	733	1038	8.4
	D0117	37.3	775	797	11.4
	D0118	38.0	1073	808	11.0
	D0119	37.0	695	730	9.0
	D0120*	38.0	678	700	9.9

[1]Represents the mean of three trials

*Animal was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Appendix H
Individual Animal Rectal Temperature and Grip Strength/Foot Splay Data

Males - Treated

Animal ID	Rectal Temperature (°Celsius)	Hindlimb Grip Strength (gram-force)[1]	Forelimb Grip Strength (gram-force)[1]	Landing Foot Splay (centimeters)[1]
Group 1 - Controls				
D0041	38.3	773	1330	14.6
D0042	38.4	913	1393	13.1
D0043	37.3	1072	1277	11.3
D0044	36.9	1488	1232	10.0
D0045	37.5	1207	1367	12.9
D0046	37.2	1553	1278	8.2
D0047	37.4	1385	1128	10.9
D0048	37.6	1178	1228	11.0
D0049	37.3	1745	1580	9.0
D0050	37.9	1167	1087	11.4
Group 2 - 100 ppm				
D0051	37.4	1303	1198	12.3
D0052	38.1	1374	1327	11.9
D0053	37.3	950	938	10.8
D0054	36.7	1347	1180	9.0
D0055	37.5	1368	1003	14.5
D0056	37.8	1192	1002	11.3
D0057	37.4	1052	992	12.1
D0058	38.6	1148	713	9.9
D0059	36.9	1193	948	11.6
D0060	37.8	1388	1057	12.1
Group 3 - 500 ppm				
D0061	36.9	1170	1158	7.6
D0062	37.5	1455	667	12.5
D0063	37.5	1228	1258	8.9
D0064	37.9	1473	1470	13.7
D0065	37.5	1445	862	13.0
D0066	37.5	1203	1080	11.3
D0067	37.6	1383	1418	12.5
D0068	38.0	1222	1040	9.3
D0069	37.4	1045	892	9.9
D0070	38.1	1127	1478	11.0
Group 4 - 2500 ppm				
D0071	37.0	1340	1520	11.5
D0072	37.5	1318	1237	9.4
D0073	36.6	1373	1240	11.8
D0074	36.5	1382	823	12.2
D0075	37.7	985	1142	13.9
D0076	36.9	843	913	11.0
D0077	37.5	1152	1412	12.9
D0078	38.4	1077	1012	10.3
D0079	38.4	1008	973	11.3
D0080	37.0	1182	1005	10.7

[1]Represents the mean of three trials

Appendix H
Individual Animal Rectal Temperature and Grip Strength/Foot Splay Data

Toxicity Females - Treated

Animal ID	Rectal Temperature (°Celsius)	Hindlimb Grip Strength (gram-force)[1]	Forelimb Grip Strength (gram-force)[1]	Landing Foot Splay (centimeters)[1]
Group 1 - Controls				
D0081	36.5	1088	1008	12.2
D0082	37.4	1278	1045	9.6
D0083	38.3	920	922	8.8
D0084	38.4	933	660	13.2
D0085	37.3	1102	982	10.6
D0086	37.7	1095	1115	11.1
D0087	39.0	1000	803	11.0
D0088	38.1	1122	1140	10.5
D0089	38.4	1183	1305	8.0
D0090	38.0	1252	1027	11.1
Group 2 - 100 ppm				
D0091	37.7	943	1005	7.9
D0092	37.3	1087	1020	7.9
D0093	38.2	1183	558	8.6
D0094	38.1	978	850	9.7
D0095	38.7	1092	960	11.1
D0096	38.1	1082	1058	11.7
D0097	37.3	1080	630	12.9
D0098	37.9	1223	1078	9.7
D0099	38.6	1053	903	9.6
D0100	37.9	1162	1043	10.5
Group 3 - 500 ppm				
D0101	37.6	1343	1378	9.0
D0102	37.8	965	1127	11.2
D0103	38.6	1118	952	10.2
D0104	37.1	1423	1367	9.1
D0105	38.0	1048	987	10.5
D0106	39.2	753	823	11.9
D0107	38.1	958	1072	12.9
D0108	38.3	1670	912	12.3
D0109	38.3	925	947	9.3
D0110	38.1	1078	615	9.6
Group 4 - 2500 ppm				
D0111	38.5	812	955	8.4
D0112	37.3	828	867	8.4
D0113	37.9	1275	1345	11.9
D0114	38.7	1042	1157	6.4
D0115	38.0	992	1115	11.3
D0116	37.8	897	968	8.1
D0117	38.0	1158	1073	9.5
D0118	38.5	993	977	10.6
D0119	37.6	748	877	10.3
D0120*	-	-	-	-

[1]Represents the mean of three trials

*Animal was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Appendix I

Individual Animal Motor Activity Data – Square Root Transformed

Appendix I
Individual Animal Motor Activity Data – Square Root Transformed[1]

Male Rats - Baseline

Animal ID	Intervals (8 minutes each) 1	2	3	4	5	6	7	8	9	10	Total
Group 1 - Controls											
D0041	12.81	11.00	8.72	7.68	7.81	3.74	0.00	1.00	2.00	0.00	54.76
D0042	14.49	11.92	6.24	9.49	3.74	0.00	0.00	0.00	0.00	1.73	47.61
D0043	17.26	14.39	12.29	11.18	6.86	7.42	8.94	0.00	0.00	0.00	78.33
D0044	16.85	13.30	11.53	8.83	6.16	13.78	2.45	0.00	0.00	0.00	72.92
D0045	16.91	9.33	3.46	1.73	7.42	3.16	3.46	1.00	11.09	11.14	68.70
D0046	12.88	13.82	9.59	7.62	7.55	1.00	0.00	0.00	0.00	2.24	54.70
D0047	14.39	9.38	2.24	0.00	1.73	0.00	2.00	0.00	0.00	1.41	31.15
D0048	15.59	13.00	11.27	10.15	2.00	3.00	0.00	11.00	11.14	5.66	82.80
D0049	13.60	8.77	6.00	8.66	7.87	5.66	4.12	0.00	2.00	1.41	58.10
D0050	16.16	13.82	11.36	11.27	7.35	2.45	1.00	0.00	2.65	2.45	68.50
Group 2 - 100 ppm											
D0051	13.64	14.00	13.67	12.88	11.18	11.40	12.65	8.89	12.41	10.82	121.54
D0052	14.80	5.83	0.00	0.00	0.00	0.00	9.90	10.00	4.58	1.00	46.11
D0053	15.84	13.75	13.19	11.75	11.66	11.66	5.83	11.62	2.83	0.00	98.13
D0054	18.08	13.75	10.34	8.49	1.00	0.00	0.00	0.00	2.65	0.00	54.31
D0055	13.49	11.05	2.65	0.00	7.07	8.83	5.92	1.00	0.00	0.00	50.00
D0056	14.25	12.33	13.04	8.31	12.61	12.49	7.55	1.73	0.00	0.00	82.30
D0057	12.65	11.14	9.95	8.66	9.38	9.43	4.12	0.00	0.00	0.00	65.33
D0058	17.00	13.27	10.72	12.08	11.79	10.39	8.66	11.27	7.55	2.83	105.56
D0059	14.14	11.53	8.83	6.78	7.42	4.00	0.00	0.00	0.00	0.00	52.70
D0060	17.26	13.00	6.08	1.73	2.00	11.36	3.61	0.00	0.00	0.00	55.04

[1] Square root transformation of raw data counts

Appendix I
Individual Animal Motor Activity Data – Square Root Transformed[1]

Male Rats - Baseline

Animal ID	Intervals (8 minutes each) 1	2	3	4	5	6	7	8	9	10	Total
Group 3 - 500 ppm											
D0061	12.81	12.49	6.16	0.00	1.00	0.00	0.00	0.00	0.00	0.00	32.46
D0062	10.82	5.20	2.83	8.43	8.66	0.00	2.45	0.00	0.00	0.00	38.38
D0063	15.20	13.60	10.05	11.14	8.54	0.00	8.25	8.94	13.71	13.19	102.62
D0064	14.70	12.81	8.37	8.31	7.68	0.00	1.00	1.41	0.00	0.00	54.27
D0065	14.28	7.21	6.93	9.06	3.74	0.00	0.00	8.43	10.77	8.49	68.90
D0066	13.08	10.20	8.60	8.37	9.49	3.61	8.54	11.22	9.80	11.70	94.61
D0067	16.79	13.93	7.48	7.94	6.24	0.00	0.00	0.00	0.00	0.00	52.39
D0068	15.91	12.77	11.22	10.58	9.85	8.94	6.78	10.49	11.09	2.24	99.87
D0069	17.38	13.67	7.94	11.75	14.56	1.00	7.21	9.06	3.87	0.00	86.44
D0070	13.96	12.65	13.08	8.12	8.06	1.73	0.00	0.00	0.00	8.72	66.33
Group 4 - 2500 ppm											
D0071	17.23	12.57	10.34	8.37	7.21	5.00	0.00	0.00	0.00	0.00	60.73
D0072	16.76	14.49	10.44	8.49	4.58	6.86	5.00	0.00	0.00	0.00	66.62
D0073	16.58	10.10	11.27	9.59	9.70	9.38	10.49	10.10	6.40	10.20	103.81
D0074	13.64	9.27	10.05	9.64	6.40	9.59	6.08	8.19	1.41	9.17	83.45
D0075	14.18	10.15	10.54	5.92	6.16	0.00	0.00	0.00	0.00	0.00	46.94
D0076	13.56	9.22	9.49	7.55	6.86	1.00	0.00	0.00	0.00	0.00	47.68
D0077	15.62	14.32	12.69	8.54	9.06	7.55	9.95	10.86	11.70	11.92	112.21
D0078	14.66	12.00	11.09	9.22	5.10	0.00	1.00	1.00	0.00	0.00	54.07
D0079	15.65	12.61	12.37	10.58	11.49	9.33	4.69	0.00	0.00	0.00	76.72
D0080	14.93	11.18	11.09	8.54	6.63	2.83	11.22	4.58	7.00	3.32	81.33

[1] Square root transformation of raw data counts

Appendix I
Individual Animal Motor Activity Data – Square Root Transformed[1]

Toxicity Group Female Rats - Baseline

Animal ID	Intervals (8 minutes each) 1	2	3	4	5	6	7	8	9	10	Total
Group 1 - Controls											
D0081	15.00	15.39	10.77	9.43	9.27	9.70	1.00	0.00	1.41	0.00	71.98
D0082	15.13	12.12	9.95	8.60	4.80	5.39	1.00	0.00	0.00	2.45	59.44
D0083	17.80	14.04	12.88	13.64	13.45	13.34	11.40	10.72	10.91	9.90	128.09
D0084	15.84	12.41	12.41	11.05	12.65	12.96	12.29	11.49	11.18	9.95	122.23
D0085	15.43	12.17	11.45	11.45	13.04	6.48	10.58	4.36	12.88	10.68	108.51
D0086	14.73	9.90	9.06	10.44	7.81	0.00	0.00	0.00	3.32	0.00	55.25
D0087	14.76	11.45	9.17	9.64	8.83	9.80	6.48	2.45	2.00	8.94	83.52
D0088	15.17	13.56	11.27	9.95	9.49	8.06	1.41	8.37	6.63	0.00	83.91
D0089	14.21	10.54	11.79	11.66	11.18	10.25	10.63	9.80	6.71	9.90	106.66
D0090	17.44	14.59	9.17	12.04	8.66	10.68	6.63	0.00	1.00	1.00	81.21
Group 2 - 100 ppm											
D0091	14.87	13.53	9.49	11.18	8.06	3.32	0.00	0.00	0.00	2.00	62.44
D0092	14.46	8.89	8.25	0.00	0.00	0.00	0.00	0.00	2.24	0.00	33.83
D0093	15.49	12.88	11.58	8.83	12.21	6.00	0.00	0.00	0.00	4.80	71.79
D0094	16.19	11.66	7.35	8.49	8.31	8.43	4.12	7.42	9.70	6.63	88.28
D0095	12.81	13.08	12.61	10.58	5.83	0.00	7.75	8.94	4.36	0.00	75.96
D0096	13.04	10.63	11.58	13.11	9.70	0.00	1.41	0.00	1.73	1.73	62.93
D0097	15.59	13.04	9.00	11.00	8.19	7.07	1.00	1.00	1.00	8.54	75.43
D0098	14.70	12.04	12.25	12.29	10.77	7.07	9.80	11.31	9.17	8.72	108.11
D0099	14.49	12.00	12.29	9.95	10.68	9.22	9.11	5.92	7.81	10.95	102.42
D0100	8.83	1.41	3.00	10.82	10.54	1.41	0.00	0.00	0.00	6.24	42.26

[1] Square root transformation of raw data counts

Appendix I
Individual Animal Motor Activity Data – Square Root Transformed[1]

Toxicity Group Female Rats - Baseline

Animal ID	Intervals (8 minutes each) 1	2	3	4	5	6	7	8	9	10	Total
Group 3 - 500 ppm											
D0101	16.12	13.30	10.20	0.00	3.46	9.54	0.00	0.00	8.19	4.58	65.40
D0102	15.36	14.56	12.37	9.27	5.10	4.80	0.00	0.00	0.00	0.00	61.46
D0103	16.43	12.65	13.34	10.05	11.96	7.55	0.00	0.00	0.00	0.00	71.98
D0104	13.53	11.58	9.90	9.75	8.00	11.18	9.80	1.73	1.00	0.00	76.46
D0105	15.46	14.11	11.79	10.91	11.14	9.22	10.34	12.00	10.86	12.69	118.52
D0106	13.75	9.95	11.09	7.42	7.94	7.94	9.22	0.00	0.00	2.45	69.75
D0107	14.42	10.82	4.47	11.70	6.86	0.00	1.73	0.00	0.00	3.00	53.00
D0108	15.49	12.57	10.86	11.40	4.12	9.85	8.49	12.45	11.09	8.12	104.45
D0109	18.57	13.82	13.78	14.35	12.08	7.42	10.91	4.58	4.90	6.78	107.20
D0110	16.52	11.49	11.83	10.54	11.00	10.49	4.90	3.00	0.00	0.00	79.77
Group 4 - 2500 ppm											
D0111	16.85	14.42	12.57	10.95	9.17	0.00	3.74	10.49	9.11	11.18	98.48
D0112	16.85	14.39	14.59	11.75	10.72	0.00	0.00	0.00	0.00	0.00	68.31
D0113	15.75	11.92	11.70	9.49	8.72	1.00	0.00	0.00	5.20	10.20	73.97
D0114	16.67	13.38	11.87	14.70	9.49	1.41	0.00	5.10	12.81	9.49	94.92
D0115	15.33	12.04	8.00	6.93	8.54	5.66	7.00	1.41	9.54	9.11	83.56
D0116	12.96	2.00	9.75	6.63	0.00	0.00	0.00	0.00	0.00	10.44	41.78
D0117	16.88	13.89	15.23	10.34	2.00	0.00	7.42	8.06	0.00	0.00	73.83
D0118	15.87	13.64	12.88	11.87	14.14	10.95	11.14	11.49	11.00	10.95	123.95
D0119	15.00	12.29	10.39	10.68	11.70	8.94	0.00	0.00	8.06	9.27	86.34
D0120*	15.36	12.61	9.22	11.83	8.00	3.46	3.61	0.00	0.00	0.00	64.09

[1] Square root transformation of raw data counts

*Animal was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Appendix I
Individual Animal Motor Activity Data – Square Root Transformed[1]

Male Rats - Treated

Animal ID	Intervals (8 minutes each) 1	2	3	4	5	6	7	8	9	10	Total
Group 1 - Controls											
D0041	14.21	8.66	7.62	8.72	7.94	1.41	1.00	0.00	2.00	9.49	61.04
D0042	12.85	9.90	9.06	6.78	8.77	5.20	1.00	7.94	8.77	2.00	72.27
D0043	15.26	11.49	3.87	0.00	7.75	6.78	1.41	0.00	7.87	3.32	57.76
D0044	13.89	11.96	9.33	5.20	7.68	3.87	0.00	2.00	8.66	8.89	71.48
D0045	19.77	12.61	10.49	3.87	1.41	0.00	0.00	0.00	1.41	0.00	49.57
D0046	14.21	9.11	6.56	3.16	9.80	9.95	1.41	4.58	0.00	0.00	58.79
D0047	13.23	10.34	4.36	4.80	8.66	9.70	4.12	4.47	4.24	2.24	66.16
D0048	15.26	11.05	9.38	9.95	3.46	1.00	1.73	1.41	2.45	2.00	57.70
D0049	13.30	8.94	9.38	7.00	5.20	1.41	6.71	3.32	6.32	8.37	69.96
D0050	15.78	12.45	9.64	10.15	9.06	6.56	8.72	7.14	6.32	1.41	87.23
Group 2 - 100 ppm											
D0051	15.59	13.11	9.64	5.20	10.91	8.25	8.77	9.06	4.80	7.68	93.01
D0052	14.18	12.17	11.49	10.30	11.27	5.29	1.73	9.54	1.73	0.00	77.69
D0053	15.56	12.33	8.60	7.68	12.65	8.43	0.00	0.00	5.66	7.87	78.77
D0054	15.07	12.25	10.91	8.06	8.89	4.24	0.00	0.00	0.00	0.00	59.42
D0055	12.53	8.12	6.78	6.78	5.39	1.00	4.47	1.00	0.00	4.12	50.20
D0056	15.87	13.71	11.53	8.72	7.75	5.57	11.05	11.18	7.48	9.22	102.08
D0057	10.91	10.30	8.37	8.72	7.81	9.70	6.24	4.80	2.65	0.00	69.48
D0058	15.62	11.09	9.43	10.30	10.58	6.48	7.81	9.11	10.77	8.19	99.38
D0059	11.92	8.83	6.00	5.20	8.00	1.00	4.47	6.56	7.28	5.74	65.00
D0060	13.86	10.68	11.14	7.68	8.37	8.43	5.39	9.00	7.87	1.00	83.40

[1] Square root transformation of raw data counts

Appendix I
Individual Animal Motor Activity Data – Square Root Transformed[1]

Male Rats - Treated

Animal ID	Intervals (8 minutes each) 1	2	3	4	5	6	7	8	9	10	Total
Group 3 - 500 ppm											
D0061	12.81	9.90	6.56	2.83	2.24	8.43	6.93	7.94	8.31	5.92	71.84
D0062	14.32	10.63	10.30	8.06	8.49	3.00	7.42	10.77	9.75	9.54	92.26
D0063	15.78	12.77	10.68	7.48	9.33	9.85	2.45	0.00	5.39	9.64	83.36
D0064	12.69	9.90	9.00	8.66	6.24	5.66	12.73	8.94	10.58	7.14	91.55
D0065	14.00	6.48	1.73	0.00	1.00	2.00	9.49	6.32	3.46	2.65	47.13
D0066	15.94	12.65	11.05	2.24	4.36	9.43	8.60	6.63	1.00	0.00	71.90
D0067	14.46	12.29	10.54	11.27	12.25	7.00	7.28	4.69	2.65	0.00	82.41
D0068	14.93	11.00	12.17	9.49	11.53	6.63	10.82	8.72	9.80	9.33	104.41
D0069	16.58	14.46	13.67	14.73	10.44	11.66	6.48	10.05	3.16	1.73	102.97
D0070	13.08	11.22	8.00	10.44	5.66	7.75	8.43	4.47	2.00	0.00	71.04
Group 4 - 2500 ppm											
D0071	12.12	10.10	7.48	7.81	5.92	4.00	0.00	1.73	1.00	1.41	51.58
D0072	12.85	9.00	7.42	5.29	2.45	0.00	2.83	0.00	0.00	1.73	41.56
D0073	12.61	7.14	7.68	2.65	0.00	1.41	0.00	0.00	0.00	0.00	31.49
D0074	9.49	7.87	5.39	2.00	0.00	2.00	1.41	0.00	5.48	0.00	33.64
D0075	11.40	11.14	7.07	4.47	1.41	4.24	3.32	0.00	0.00	0.00	43.05
D0076	10.95	10.39	6.78	3.32	1.41	0.00	0.00	0.00	0.00	0.00	32.86
D0077	13.23	8.77	6.71	3.16	5.29	1.00	3.46	1.00	0.00	5.39	48.01
D0078	13.15	9.59	9.00	6.93	4.58	2.83	1.00	2.83	5.83	5.20	60.94
D0079	11.83	7.48	2.00	2.24	2.00	2.24	2.24	3.87	3.46	0.00	37.36
D0080	11.66	10.49	8.83	5.20	4.00	3.16	1.41	0.00	0.00	2.00	46.75

[1] Square root transformation of raw data counts

Appendix I
Individual Animal Motor Activity Data – Square Root Transformed[1]

Toxicity Group Female Rats - Treated

	Intervals (8 minutes each)										
Animal ID	1	2	3	4	5	6	7	8	9	10	Total
Group 1 - Controls											
D0081	18.00	12.45	9.33	3.00	0.00	0.00	6.86	11.58	7.42	1.00	69.62
D0082	13.67	7.75	9.75	7.68	4.12	0.00	0.00	9.80	5.48	5.92	64.16
D0083	15.81	11.18	9.38	8.31	10.00	8.60	8.72	8.60	5.74	2.24	88.58
D0084	15.43	13.60	2.00	5.83	0.00	0.00	0.00	3.74	1.00	1.41	43.02
D0085	15.94	13.67	10.30	0.00	5.57	9.27	10.68	12.37	4.12	6.56	88.48
D0086	12.92	10.49	7.75	3.61	8.54	6.40	2.83	7.55	8.60	7.07	75.76
D0087	14.70	10.72	7.87	7.87	2.65	7.21	10.05	1.00	8.94	2.00	73.02
D0088	15.36	11.87	7.48	5.83	9.27	3.46	5.74	11.49	10.39	5.10	86.01
D0089	14.46	12.08	10.34	3.32	2.83	9.17	4.47	0.00	9.49	10.20	76.35
D0090	16.40	14.00	11.40	9.54	8.60	12.08	10.86	6.71	3.46	2.00	95.06
Group 2 - 100 ppm											
D0091	14.35	9.49	6.71	7.81	9.00	9.90	5.66	7.00	0.00	1.00	70.91
D0092	12.81	8.43	9.43	9.90	1.00	1.41	3.16	9.33	6.63	1.73	63.84
D0093	14.32	9.54	8.72	4.58	2.65	2.65	0.00	4.47	10.20	8.83	65.95
D0094	15.46	10.15	10.30	10.58	7.48	2.65	2.00	5.00	9.17	6.32	79.11
D0095	13.34	5.29	1.00	0.00	0.00	5.39	5.10	1.41	0.00	0.00	31.53
D0096	13.93	10.86	8.60	10.25	9.43	7.68	5.83	11.40	9.49	4.90	92.37
D0097	15.26	12.21	7.94	9.70	5.20	7.94	9.43	4.69	9.64	4.12	86.13
D0098	14.49	11.45	8.06	5.20	6.56	6.24	6.71	5.39	2.00	6.86	72.95
D0099	12.69	11.75	10.95	10.25	10.63	6.16	7.48	9.27	8.00	6.16	93.35
D0100	12.25	7.94	3.74	2.45	5.39	6.93	7.00	4.58	0.00	2.00	52.27

[1] Square root transformation of raw data counts

Appendix I
Individual Animal Motor Activity Data – Square Root Transformed[1]

Toxicity Group Female Rats - Treated

	Intervals (8 minutes each)										
Animal ID	1	2	3	4	5	6	7	8	9	10	Total
Group 3 - 500 ppm											
D0101	14.80	11.53	9.27	3.32	7.94	7.35	9.49	9.90	5.29	9.75	88.63
D0102	15.65	14.18	6.56	2.00	10.10	12.65	0.00	7.68	0.00	0.00	68.82
D0103	13.78	10.54	6.86	5.48	0.00	0.00	7.75	3.46	7.42	3.74	59.02
D0104	16.16	13.42	12.12	14.14	7.28	5.66	8.43	15.39	12.04	12.29	116.93
D0105	13.89	10.72	6.40	6.24	10.44	8.43	7.62	1.00	2.65	4.69	72.08
D0106	11.92	7.35	6.40	0.00	1.41	0.00	6.48	9.64	6.48	1.00	50.69
D0107	15.26	11.36	8.06	5.10	6.78	10.72	5.48	0.00	3.00	4.12	69.89
D0108	15.17	12.29	11.00	9.75	3.00	9.59	8.54	8.00	10.34	5.57	93.25
D0109	14.07	10.39	2.00	7.14	3.32	1.73	12.29	6.16	11.00	9.43	77.54
D0110	13.89	9.64	7.81	3.74	0.00	0.00	4.12	5.83	2.00	0.00	47.04
Group 4 - 2500 ppm											
D0111	13.15	9.59	8.49	8.12	3.16	6.16	0.00	0.00	0.00	10.34	59.02
D0112	15.52	14.07	10.58	12.81	10.82	4.90	1.00	1.00	0.00	0.00	70.70
D0113	13.04	10.77	6.32	6.86	7.75	6.08	5.57	0.00	5.66	7.21	69.25
D0114	12.17	8.37	1.00	3.46	9.27	5.39	0.00	1.00	4.12	9.70	54.47
D0115	11.09	7.75	6.40	5.57	7.28	4.24	3.16	1.41	8.00	7.00	61.91
D0116	12.21	7.94	4.80	1.41	3.46	7.75	1.41	3.61	0.00	0.00	42.58
D0117	14.07	12.61	10.82	9.54	5.83	3.16	0.00	0.00	5.10	8.25	69.38
D0118	13.27	10.20	9.38	6.00	0.00	0.00	1.00	9.85	7.87	10.58	68.15
D0119	11.70	7.14	5.83	1.00	1.73	2.83	4.24	6.93	5.39	6.00	52.79
D0120*	-	-	-	-	-	-	-	-	-	-	-

[1] Square root transformation of raw data counts

*Animal was removed from the Toxicity group and placed into the Reproductive group after baseline testing.

Appendix J

Individual Animal Hematology Data

Appendix J
Individual Animal Hematology Data

Males

Animal ID	White Blood Cells (X 10^3/cu mm)	Red Blood Cells (X 10^6/cu mm)	Hemoglobin (g/dL)	Hematocrit (%)	Mean Cell Volume (cubic microns)	Mean Cell Hemoglobin (picograms)	Mean Cell Hemaglobin Concentration (%)	Platlets (X 10^3/cu mm)
Group 1 - Controls								
D0041[a]								
D0042	13.7	8.70	17.5	47.8	54.9	20.1	36.6	1077
D0043	10.4	7.48	16.3	42.6	56.9	21.8	38.3	1040
D0044	11.8	8.33	16.9	46.0	55.2	20.3	36.7	1046
D0045	4.4	7.85	16.8	43.5	55.4	21.4	38.6	983
D0046	19.6	7.64	16.1	42.4	55.5	21.1	38.0	1069
D0047	13.2	7.49	16.2	41.5	55.4	21.6	39.0	1146
D0048	14.2	7.56	17.2	44.1	58.3	22.8	39.0	1210
D0049	10.8	8.42	17.2	44.8	53.2	20.4	38.4	1108
D0050	12.0	8.56	17.6	47.2	55.1	20.6	37.3	1145
Group 2 - 100 ppm								
D0051	7.0	7.46	16.0	41.4	55.5	21.4	38.6	998
D0052	11.2	7.64	15.6	41.1	53.8	20.4	38.0	1121
D0053	18.2	7.86	16.3	43.1	54.8	20.7	37.8	1078
D0054	20.0	8.54	17.2	46.4	54.3	20.1	37.1	1112
D0055	12.1	7.87	16.9	44.5	56.6	21.5	38.0	1037
D0056	13.7	8.35	17.0	44.6	53.4	20.4	38.1	1193
D0057	20.6	8.10	16.5	42.9	53.0	20.4	38.5	1143
D0058	8.3	8.63	17.0	46.0	53.3	19.7	37.0	1163
D0059	11.9	7.61	16.6	43.2	56.8	21.8	38.4	1183
D0060	7.3	7.68	17.0	43.4	56.5	22.1	39.2	1198
Group 3 - 500 ppm								
D0061	9.0	8.08	16.7	44.4	54.9	20.7	37.6	1169
D0062	9.4	7.60	16.2	42.6	56.1	21.3	38.0	1107
D0063	18.3	8.44	16.9	46.0	54.5	20.0	36.7	1002
D0064	16.7	7.88	16.3	43.3	55.0	20.7	37.6	1160
D0065	9.5	8.03	17.2	44.7	55.7	21.4	38.5	1078
D0066	11.8	8.15	16.7	43.8	53.7	20.5	38.1	1203
D0067	14.2	7.60	16.6	42.4	55.8	21.8	39.2	1221
D0068	12.7	7.53	15.9	41.0	54.5	21.1	38.8	1226
D0069	12.4	6.90	15.5	39.9	57.8	22.5	38.8	1081
D0070	7.4	7.48	15.8	42.2	56.4	21.1	37.4	1003
Group 4 - 2500 ppm								
D0071	12.9	7.99	16.2	43.1	54.0	20.3	37.6	1329
D0072	11.2	7.46	15.7	41.6	55.7	21.0	37.7	1125
D0073	11.9	7.71	15.4	40.6	52.6	20.0	37.9	1387
D0074	13.6	7.41	15.7	42.2	56.9	21.2	37.2	1254
D0075	13.1	8.52	15.9	43.8	51.4	18.7	36.3	1139
D0076	11.4	7.56	16.6	42.3	55.9	22.0	39.2	1430
D0077	8.3	8.01	15.8	42.1	52.6	19.7	37.5	1118
D0078	13.5	8.14	16.5	44.1	54.2	20.3	37.4	1453
D0079	7.4	7.66	15.4	41.1	53.6	20.1	37.5	1259
D0080	9.3	8.57	16.3	44.6	52.1	19.0	36.5	1300

[a] sample clotted

Appendix J
Individual Animal Hematology Data

Toxicity Females

Animal ID	White Blood Cells (X 10^3/cu mm)	Red Blood Cells (X 10^6/cu mm)	Hemoglobin (g/dL)	Hematocrit (%)	Mean Cell Volume (cubic microns)	Mean Cell Hemoglobin (picograms)	Mean Cell Hemaglobin Concentration (%)	Platlets (X 10^3/cu mm)
Group 1 - Controls								
D0081	8.2	7.54	16.0	42.4	56.2	21.2	37.7	1064
D0082	4.3	7.30	15.4	40.4	55.3	21.1	38.1	1144
D0083	14.4	7.52	13.4	35.3	46.9	17.8	38.0	1216
D0084	4.9	6.92	15.4	40.1	58.0	22.3	38.4	954
D0085	11.6	6.86	15.2	39.4	57.5	22.2	38.6	1151
D0086	6.4	6.89	14.9	38.0	55.1	21.6	39.2	1081
D0087	7.7	7.37	16.7	43.8	59.4	22.7	38.1	1111
D0088[a]								
D0089	7.4	7.13	15.3	39.9	55.9	21.5	38.3	1246
D0090	6.8	6.74	15.2	39.0	57.8	22.6	39.0	961
Group 2 - 100 ppm								
D0091	12.8	7.13	15.2	40.5	56.8	21.3	37.5	1153
D0092	7.8	7.39	15.8	42.3	57.2	21.4	37.4	1126
D0093	10.2	7.91	16.7	45.0	56.9	21.1	37.1	1219
D0094	4.4	6.65	14.7	37.5	56.4	22.1	39.2	1191
D0095[a]								
D0096	9.2	7.74	16.2	43.3	56.0	20.9	37.4	1005
D0097	6.4	6.89	16.0	41.4	60.1	23.2	38.6	890
D0098	4.9	7.33	15.9	42.5	58.0	21.7	37.4	1198
D0099	16.5	8.02	17.2	44.7	55.7	21.4	38.5	963
D0100	6.3	7.39	16.2	42.9	58.0	21.9	37.8	1243
Group 3 - 500 ppm								
D0101	7.1	6.69	14.5	38.1	56.9	21.7	38.1	1089
D0102	11.4	6.94	15.8	41.5	59.8	22.8	38.1	1093
D0103	7.4	7.37	15.5	41.2	55.9	21.0	37.6	1129
D0104	5.8	7.31	16.2	42.9	58.7	22.2	37.8	855
D0105	4.9	7.17	15.2	39.9	55.7	21.2	38.1	1010
D0106	5.3	6.91	15.4	41.0	59.4	22.3	37.6	1066
D0107	4.5	7.14	15.7	40.6	56.8	22.0	38.7	1258
D0108[a]								
D0109	9.0	7.54	16.0	42.0	55.7	21.2	38.1	1076
D0110	10.0	7.90	16.3	43.9	55.6	20.6	37.1	1175
Group 4 - 2500 ppm								
D0111	7.6	7.51	15.0	41.3	55.0	20.0	36.3	1237
D0112	9.0	7.18	15.7	41.4	57.7	21.9	37.9	1140
D0113	9.9	6.62	14.8	38.7	58.4	22.4	38.2	1237
D0114	8.6	6.39	14.6	38.3	59.9	22.8	38.1	1094
D0115	6.5	7.13	15.5	40.1	56.2	21.7	38.7	1066
D0116	6.1	7.11	15.7	42.8	60.2	22.1	36.7	1219
D0117	7.6	7.23	15.5	42.1	58.2	21.4	36.8	1200
D0118	7.3	6.92	15.4	40.7	58.8	22.3	37.8	969
D0119	7.0	6.77	15.0	40.3	59.5	22.2	37.2	1165
D0040	7.4	6.89	14.7	38.9	56.5	21.3	37.8	1038

[a] sample clotted

Appendix K

Individual Animal Hematology Differential Counts

Appendix K
Individual Animal Hematology Differential Counts

Males

Animal ID	Segmented Neutrophils	Band Neutrophils	Lymphocytes	Monocytes	Eosinophils	Basophils	Other	Polychromasia	Anisocytosis	Poikilocytosis	Platelets
Group 1 - Controls											
D0041[a]	.	.	.	.	.	.	.	.	.	.	.
D0042	8	0	92	0	0	0	0	+/-	-	-	N
D0043	7	0	93	0	0	0	0	+	-	-	N
D0044	11	0	88	1	0	0	0	+	-	-	N
D0045	24	1	75	0	0	0	0	+	-	+	N
D0046	11	0	87	2	0	0	0	+	-	-	N
D0047	18	0	79	2	1	0	0	+	+	+	N
D0048	16	1	82	1	0	0	0	+	-	-	N
D0049	16	0	82	2	0	0	0	+	-	-	N
D0050	10	0	90	0	0	0	0	+	-	-	N
Group 2 - 100 ppm											
D0051	10	0	90	0	0	0	0	+	-	-	N
D0052	16	0	84	0	0	0	0	+	-	-	N
D0053	13	0	87	0	0	0	0	+	-	-	N
D0054	12	0	85	3	0	0	0	+	-	-	N
D0055	7	0	93	0	0	0	0	+	-	+	N
D0056	7	0	90	3	0	0	0	+	-	-	N
D0057	13	0	87	0	0	0	0	+	-	+	N
D0058	7	0	93	0	0	0	0	+	-	-	N
D0059	11	0	89	0	0	0	0	+	-	-	N
D0060	12	0	85	3	0	0	0	+	-	-	N

[a] sample clotted

+ = slight or mild
++ = moderate
+++ = marked
++++ = severe

N= normal size and shape

Appendix K
Individual Animal Hematology Differential Counts

Males

Animal ID	Segmented Neutrophils	Band Neutrophils	Lymphocytes	Monocytes	Eosinophils	Basophils	Other	Polychromasia	Anisocytosis	Poikilocytosis	Platelets
Group 3 - 500 ppm											
D0061	22	0	77	1	0	0	0	+	-	-	N
D0062	19	0	81	0	0	0	0	+	-	-	N
D0063	9	0	91	0	0	0	0	+	+	+++	N
D0064	23	0	77	0	0	0	0	+	-	-	N
D0065	12	0	87	1	0	0	0	+	-	+	N
D0066	21	0	79	0	0	0	0	+	-	-	N
D0067	15	0	85	0	0	0	0	+	-	-	N
D0068	20	0	79	1	0	0	0	+	-	+	N
D0069	19	0	80	1	0	0	0	+	-	-	N
D0070	19	0	81	0	0	0	0	+	-	-	N
Group 4 - 2500 ppm											
D0071	10	1	87	2	0	0	0	+	-	-	N
D0072	15	0	84	1	0	0	0	+	-	++	N
D0073	10	0	88	2	0	0	0	+	-	++	N
D0074	15	0	85	0	0	0	0	+	-	+	N
D0075	7	0	93	0	0	0	0	+	-	-	N
D0076	14	0	85	1	0	0	0	+	-	-	N
D0077	12	0	88	0	0	0	0	+	-	-	N
D0078	19	0	80	1	0	0	0	+	-	-	N
D0079	13	0	87	0	0	0	0	+	-	+	N
D0080	15	0	85	0	0	0	0	+	-	-	N

+ = slight or mild
++ = moderate
+++ = marked
++++ = severe

N= normal size and shape

Appendix K
Individual Animal Hematology Differential Counts
Toxicity Females

Animal ID	Segmented Neutrophils	Band Neutrophils	Lymphocytes	Monocytes	Eosinophils	Basophils	Other	Polychromasia	Anisocytosis	Poikilocytosis	Platelets
Group 1 - Controls											
D0081	13	0	86	1	0	0	0	+	-	-	N
D0082	4	0	96	0	0	0	0	+	-	-	N
D0083	12	0	88	0	0	0	+ microcytosis + hypochromasia	+	-	-	N
D0084	17	0	83	0	0	0	0	+	-	-	N
D0085	5	0	95	0	0	0	0	+	-	-	N
D0086	7	0	93	0	0	0	0	+	-	-	N
D0087	10	0	90	0	0	0	0	+	-	-	N
D0088[a]	.	.	.	.	.	.	.	.	.	.	.
D0089	6	0	94	0	0	0	0	+	-	-	N
D0090	8	0	91	1	0	0	0	+	-	-	N
Group 2 - 100 ppm											
D0091	16	0	84	0	0	0	0	+	-	-	N
D0092	12	0	88	0	0	0	0	+	-	-	N
D0093	18	0	81	0	0	0	0	+	-	-	N
D0094	6	0	94	0	0	0	0	+	-	-	N
D0095[a]	.	.	.	.	.	.	.	.	.	.	.
D0096	21	0	79	0	0	0	0	+	-	-	N
D0097	13	0	85	2	0	0	0	+	-	-	N
D0098	4	0	95	1	0	0	0	+	-	-	N
D0099	15	0	85	0	0	0	0	+	-	-	N
D0100	9	0	91	0	0	0	0	+	-	-	N

[a] sample clotted

+ = slight or mild
++ = moderate
+++ = marked
++++ = severe

N= normal size and shape

Appendix K
Individual Animal Hematology Differential Counts

Toxicity Females

Animal ID	Segmented Neutrophils	Band Neutrophils	Lymphocytes	Monocytes	Eosinophils	Basophils	Other	Polychromasia	Anisocytosis	Poikilocytosis	Platelets
Group 3 - 500 ppm											
D0101	7	0	93	0	0	0	0	+	+	-	N
D0102	23	0	77	0	0	0	0	+	-	-	N
D0103	7	0	93	0	0	0	0	+	-	-	N
D0104	8	0	92	0	0	0	0	+	-	-	N
D0105	9	0	91	0	0	0	0	+	-	-	N
D0106	9	0	88	2	1	0	0	+	-	-	N
D0107	21	0	78	1	0	0	0	+	-	-	N
D0108[a]	.	.	.	.	.	.	.	.	.	.	.
D0109	23	0	77	0	0	0	0	+	-	-	N
D0110	25	0	75	0	0	0	0	+	-	-	N
Group 4 - 2500 ppm											
D0111	11	0	88	1	0	0	0	+	-	-	N
D0112	8	0	90	2	0	0	0	+	-	-	N
D0113	8	0	92	0	0	0	0	+	-	-	N
D0114	8	0	92	0	0	0	0	+	-	-	N
D0115	15	0	85	0	0	0	0	+	-	-	N
D0116	11	0	89	0	0	0	0	+	-	-	N
D0117	9	0	90	1	0	0	0	+	-	-	N
D0118	7	0	92	1	0	0	0	+	-	-	N
D0119	14	0	86	0	0	0	0	+	-	-	N
D0040	12	0	86	2	0	0	0	+	-	-	N

[a] sample clotted

+ = slight or mild
++ = moderate
+++ = marked
++++ = severe

N= normal size and shape

Appendix L

Individual Animal Prothrombin Times

Appendix L
Individual Animal Prothrombin Times

Males

Animal ID	Results (seconds)
Group 1 - Controls	
D0041[a]	.
D0042	15.9
D0043	15.6
D0044	16.5
D0045	18.5
D0046	17.0
D0047	16.0
D0048	18.0
D0049	16.5
D0050	15.0
Group 2 - 100 ppm	
D0051	16.4
D0052	15.4
D0053	18.0
D0054	15.0
D0055	16.0
D0056	16.0
D0057	15.5
D0058	17.0
D0059	16.0
D0060	15.5
Group 3 - 500 ppm	
D0061	18.0
D0062	17.5
D0063	16.0
D0064	19.0
D0065	21.0
D0066	17.0
D0067	16.0
D0068	18.0
D0069	16.5
D0070	15.5
Group 4 - 2500 ppm	
D0071	16.4
D0072	18.5
D0073	17.0
D0074	16.0
D0075	16.5
D0076	16.0
D0077	15.6
D0078	16.5
D0079	16.0
D0080	15.5

[a] specimen clotted

Appendix L
Individual Animal Prothrombin Times

Toxicity Females

Animal ID	Results (seconds)
Group 1 - Controls	
D0081	17.4
D0082	16.0
D0083	15.5
D0084	16.0
D0085	16.0
D0086	15.5
D0087	15.8
D0088[a]	.
D0089	16.0
D0090	15.0
Group 2 - 100 ppm	
D0091	17.5
D0092	16.0
D0093	16.0
D0094	15.5
D0095[a]	.
D0096	15.0
D0097[a]	.
D0098	15.5
D0099[a]	.
D0100	16.0
Group 3 - 500 ppm	
D0101	16.5
D0102	16.0
D0103	17.0
D0104[a]	.
D0105	16.5
D0106	16.0
D0107	16.0
D0108[a]	.
D0109	16.0
D0110	16.0
Group 4 - 2500 ppm	
D0111	16.0
D0112	17.5
D0113	17.5
D0114	17.0
D0115	15.5
D0116	17.0
D0117	16.0
D0118	16.5
D0119	16.0
D0040	16.0

[a] specimen clotted

Appendix M

Individual Animal Clinical Chemistry Data

Appendix M
Individual Animal Clinical Chemistry Data

Males

Animal ID	Albumin g/dL	Alkaline Phosphatase U/L	Alanine Aminotransferase U/L	Asparate Aminotransferase U/L	Total Bilirubin mg/dL	Calcium mg/dL	Creatinine mg/dL	Cholesterol mg/dL	Glucose mg/dL	Phosphorus mg/dL	Total Protein g/dL	Urea Nitrogen mg/dL	Sodium mmol/L	Potassium mmol/L	Chloride mmol/L
Group 1 - Controls															
D0041	2.4	178	28	97	0.2	9.8	0.8	29	101	9.1	6.0	10	147	5.8	107
D0042[a]	4.0	141	37	139	0.2	10.1	0.7	47	132	7.5	6.6	14	146	4.4	107
D0043[a]	3.9	129	32	113	0.2	10.1	0.6	41	117	7.5	6.4	17	148	4.7	109
D0044	3.6	174	38	106	0.1	10.2	0.7	41	113	9.0	5.5	16	146	5.6	110
D0045	3.8	110	28	99	0.0	9.6	0.8	43	136	6.3	6.1	24	152	3.6	113
D0046	3.6	77	21	98	0.2	9.8	0.7	40	132	7.9	5.8	16	145	4.5	109
D0047	3.7	168	18	77	0.1	10.1	0.5	35	156	8.3	5.8	10	148	4.3	111
D0048	3.8	118	18	86	0.2	9.8	0.7	37	123	8.0	6.1	14	146	4.7	111
D0049	4.0	109	23	87	0.2	10.6	0.6	38	112	7.7	6.6	13	144	4.2	107
D0050	4.0	150	25	84	0.2	10.2	0.6	47	119	7.6	6.2	15	145	4.2	107
Group 2 - 100 ppm															
D0051[a]	3.7	176	25	92	0.1	9.7	0.7	29	131	9.8	5.5	19	149	7.1	112
D0052	3.8	153	38	98	0.2	9.6	0.7	58	105	6.5	6.0	15	146	3.9	109
D0053[a]	4.1	157	31	95	0.1	9.4	0.7	44	128	8.2	6.0	17	146	5.0	108
D0054	3.9	145	28	85	0.1	10.6	0.7	73	161	7.0	6.0	18	145	3.9	108
D0055[a]	3.8	124	40	105	0.1	9.9	0.6	44	105	7.4	5.4	15	144	4.2	108
D0056	4.1	144	29	81	0.3	8.9	0.6	40	147	7.6	6.5	19	145	5.1	108
D0057	3.9	128	23	89	0.2	9.9	0.6	48	132	7.2	6.4	15	145	4.4	107
D0058b	3.8	112	21	84	0.2	10.5	0.6	20	105	11.7	6.1	15	145	9.2	109
D0059[a]	3.9	115	22	100	0.3	10.3	0.7	36	88	8.0	6.2	14	145	4.4	108
D0060	4.1	146	25	95	0.2	9.9	0.6	54	101	7.8	6.3	13	146	4.2	108

[a] slight hemolysis present
[b] potassium results verified

Appendix M
Individual Animal Clinical Chemistry Data

Males

Animal ID	Albumin g/dL	Alkaline Phosphatase U/L	Alanine Aminotransferase U/L	Asparate Aminotransferase U/L	Total Bilirubin mg/dL	Calcium mg/dL	Creatinine mg/dL	Cholesterol mg/dL	Glucose mg/dL	Phosphorus mg/dL	Total Protein g/dL	Urea Nitrogen mg/dL	Sodium mmol/L	Potassium mmol/L	Chloride mmol/L
Group 3 - 500 ppm															
D0061[c]	.	.	.	.	.	.	.	.	.	.	.	.	.	.	.
D0062[a]	4.0	182	32	94	0.1	9.6	0.7	46	120	7.0	6.2	18	144	4.2	108
D0063	3.9	120	43	102	0.1	9.5	0.7	45	117	6.7	6.2	18	145	4.0	107
D0064	3.6	129	34	103	0.0	9.6	0.6	32	98	8.2	5.6	15	149	4.6	113
D0065	3.8	159	32	98	0.1	9.7	0.7	35	100	7.9	6.0	16	145	4.6	109
D0066	4.0	125	24	74	0.2	10.1	0.6	46	133	7.7	6.4	16	147	4.2	110
D0067	3.7	153	27	98	0.2	9.9	0.6	47	125	7.6	5.9	14	146	4.4	110
D0068[a]	3.9	116	25	93	0.2	10.0	0.5	49	112	7.6	6.2	17	145	4.7	109
D0069	3.7	111	29	95	0.2	10.1	0.6	40	102	7.8	6.0	16	145	4.1	111
D0070	3.6	120	28	91	0.2	9.4	0.7	40	91	7.9	5.6	13	146	4.9	110
Group 4 - 2500 ppm															
D0071[a]	4.1	112	36	88	0.1	9.5	0.6	40	168	6.9	6.7	20	143	4.3	107
D0072[a]	3.9	93	38	88	0.1	10.4	0.7	39	195	9.4	6.7	21	144	6.3	105
D0073	3.9	94	30	74	0.1	10.2	0.7	43	126	8.0	6.0	19	144	4.4	108
D0074[a]	3.6	95	27	89	0.1	9.7	0.6	42	100	8.0	5.7	15	144	4.7	106
D0075	3.9	93	55	86	0.2	9.8	0.8	41	161	7.0	6.0	22	142	4.5	105
D0076	3.8	84	61	85	0.1	10.0	0.6	44	111	7.6	6.2	24	146	4.6	109
D0077[a]	3.6	107	27	87	0.2	9.5	0.7	40	90	7.7	6.0	23	144	5.3	109
D0078	3.8	115	26	89	0.2	9.9	0.5	53	114	7.4	6.3	13	144	4.7	108
D0079	3.6	90	36	106	0.2	9.4	0.6	44	85	8.6	5.7	15	146	5.1	111
D0080	3.7	102	30	78	0.1	9.9	0.5	32	125	7.8	6.4	19	147	4.2	110

[a] slight hemolysis present
[c] blood not collected

Appendix M
Individual Animal Clinical Chemistry Data

Toxicity Females

Animal ID	Albumin g/dL	Alkaline Phosphatase U/L	Alanine Aminotransferase U/L	Asparate Aminotransferase U/L	Total Bilirubin mg/dL	Calcium mg/dL	Creatinine mg/dL	Cholesterol mg/dL	Glucose mg/dL	Phosphorus mg/dL	Total Protein g/dL	Urea Nitrogen mg/dL	Sodium mmol/L	Potassium mmol/L	Chloride mmol/L
Group 1 - Controls															
D0081	4.3	62	19	88	0.2	10.3	0.6	66	148	5.6	6.7	13	144	3.9	110
D0082	4.6	74	26	96	0.2	10.2	1	54	155	4.9	7.0	16	144	3.7	110
D0083	4.3	58	27	89	0.2	10.0	0.7	46	152	5.2	6.5	13	142	3.4	110
D0084[a]	4.0	60	22	85	0.2	9.9	0.7	43	132	6.5	5.9	13	145	4.0	112
D0085	4.4	59	22	88	0.2	10.1	0.8	72	165	6.6	6.6	17	143	3.7	108
D0086	4.2	48	27	87	0.2	10.1	0.6	57	149	7.1	6.0	11	144	4.2	109
D0087	4.1	58	19	82	0.2	10.0	0.7	37	103	7.3	6.6	15	146	4.3	109
D0088	5.0	37	17	75	0.2	10.3	0.6	36	136	5.9	7.6	17	144	3.8	110
D0089	4.4	46	41	91	0.3	10.4	0.6	50	113	7.9	6.5	13	142	4.6	107
D0090	4.3	67	19	75	0.2	10.1	0.6	51	121	6.0	6.6	13	144	4.0	109
Group 2 - 100 ppm															
D0091	4.7	54	27	80	0.2	10.8	0.7	48	175	4.8	7.4	19	144	4.0	110
D0092	4.3	59	23	70	0.1	8.6	0.8	63	182	4.6	6.7	14	142	4.5	108
D0093[a,b]	4.6	51	26	85	0.2	10.8	0.9	63	109	7.3	7.2	15	145	7.5	112
D0094	4.9	92	23	96	0.2	9.8	0.7	55	150	6.3	5.4	12	143	3.8	109
D0095	4.3	56	20	77	0.2	9.7	0.7	52	150	5.6	6.3	17	143	4.3	110
D0096	4.6	72	29	104	0.2	10.2	0.7	58	111	5.8	7.3	19	144	4.3	109
D0097	4.5	78	24	86	0.2	10.3	0.6	28	124	6.2	7.1	15	146	3.7	112
D0098	4.1	63	19	78	0.2	8.5	0.7	60	100	8.3	6.5	16	145	5.7	110
D0099c	.	.	.	.	.	.	.	.	.	.	.	.	.	.	.
D0100	4.5	58	27	82	0.2	10.2	0.6	49	144	6.4	6.6	16	145	3.9	110

[a] slight hemolysis present
[b] potassium results verified
[c] blood not collected

Appendix M
Individual Animal Clinical Chemistry Data

Toxicity Females

Animal ID	Albumin g/dL	Alkaline Phosphatase U/L	Alanine Aminotransferase U/L	Asparate Aminotransferase U/L	Total Bilirubin mg/dL	Calcium mg/dL	Creatinine mg/dL	Cholesterol mg/dL	Glucose mg/dL	Phosphorus mg/dL	Total Protein g/dL	Urea Nitrogen mg/dL	Sodium mmol/L	Potassium mmol/L	Chloride mmol/L
Group 3 - 500 ppm															
D0101	4.2	68	23	77	0.2	9.8	0.8	34	156	5.0	6.4	14	143	4.1	110
D0102a	4.0	102	25	93	0.2	9.8	0.7	59	104	6.4	6.4	16	143	4.0	109
D0103[a]	4.3	75	26	112	0.2	9.7	0.9	42	108	6.0	6.5	20	145	3.9	111
D0104	4.3	61	31	122	0.2	9.5	0.8	54	92	7.2	6.3	18	144	3.5	110
D0105	4.1	50	21	78	0.2	9.6	0.6	44	117	8.5	6.2	13	147	4.3	112
D0106	4.2	55	21	103	0.2	9.7	0.8	49	103	6.3	6.5	17	146	4.5	111
D0107	4.5	67	21	74	0.2	10.5	0.7	69	122	5.8	7.2	16	146	4.2	113
D0108[a]	4.2	54	22	83	0.2	9.3	0.7	60	106	6.4	6.6	14	144	4.5	110
D0109	4.5	64	29	78	0.2	10.0	0.6	58	116	8.1	6.9	15	148	4.5	114
D0110	3.9	58	23	81	0.2	9.4	0.6	61	104	7.7	6.4	13	143	4.6	107
Group 4 - 2500 ppm															
D0111a	4.2	42	20	70	0.1	9.5	0.8	56	119	5.8	6.3	12	143	3.9	111
D0112	4.3	60	24	97	0.2	10.0	0.7	82	140	5.7	6.8	21	143	4.2	108
D0113	4.2	52	28	81	0.2	9.9	0.7	85	103	5.8	6.9	18	145	4.3	109
D0114	4.1	73	20	80	0.1	9.6	0.7	61	103	5.5	6.7	16	143	4.2	109
D0115	4.1	46	28	68	0.1	9.5	0.5	71	101	6.2	6.5	10	143	4.3	110
D0116	4.2	69	18	79	0.2	10.0	0.6	70	101	7.1	6.8	15	147	4.4	112
D0117	4.3	43	18	76	0.1	9.9	0.6	57	99	6.2	6.8	13	146	4.0	110
D0118	3.3	44	27	63	0.2	9.6	0.6	60	102	6.6	5.4	14	144	4.1	109
D0119	3.9	41	22	75	0.2	9.7	0.7	80	99	6.2	6.2	16	144	4.2	110
D0040	3.7	58	16	69	0.2	9.8	0.5	43	140	7.1	5.8	11	144	3.9	110

[a] slight hemolysis present

Appendix N

Individual Animal Reports

Appendix N
Individual Animal Reports

PATHOLOGY REPORT PAGE : 1/ 140
Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

EXPLANATION OF CODES AND SYMBOLS

CODES AND SYMBOLS USED AT ANIMAL LEVEL:

M = Male animal
F = Female animal
K0 = Terminal sacrifice group
K1...K9 = Interim sacrifice groups 1...9

CODES AND SYMBOLS USED AT FINDING LEVEL:

GRADE 1 = Minimal / very few / very small
GRADE 2 = Slight / few / small
GRADE 3 = Moderate / moderate number / moderate size
P = Finding present, severity not scored

Appendix N
Individual Animal Reports

PATHOLOGY REPORT PAGE : 2/ 140
Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

Additional codes used:

Codes used at group level:
T = Toxicity group females
R = Reproductive group females
Low = 100 ppm
Mid. = 500 ppm
High = 2500 ppm

Appendix N
Individual Animal Reports

PATHOLOGY REPORT PAGE : 3/ 140
INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

ANIMAL HEADING DATA
DOSE GROUP : R1, Control Repro.

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0001	F	K1	K1	44	25-MAR-02	07-MAY-02	07-MAY-02
D0002	F	K1	K1	46	25-MAR-02	09-MAY-02	09-MAY-02
D0003	F	K1	K1	42	25-MAR-02	05-MAY-02	05-MAY-02
D0004	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0005	F	K1	K1	44	25-MAR-02	07-MAY-02	07-MAY-02
D0006	F	K1	K1	42	25-MAR-02	05-MAY-02	05-MAY-02
D0007	F	K1	K1	40	25-MAR-02	03-MAY-02	03-MAY-02
D0008	F	K1	K1	40	25-MAR-02	03-MAY-02	03-MAY-02
D0009	F	K1	K1	42	25-MAR-02	05-MAY-02	05-MAY-02
D0010	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02

Appendix N
Individual Animal Reports

PATHOLOGY REPORT PAGE : 4/ 140
INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R1, Control Repro. FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 44 * ANIMAL NO. : D0001

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 46 * ANIMAL NO. : D0002

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

Appendix N
Individual Animal Reports

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

PAGE : 5/ 140
Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R1, Control Repro. FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 42 * ANIMAL NO. : D0003

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 41 * ANIMAL NO. : D0004

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

Appendix N
Individual Animal Reports

PATHOLOGY REPORT PAGE : 6/ 140
INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R1, Control Repro. FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 44 * ANIMAL NO. : D0005

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 42 * ANIMAL NO. : D0006

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

Appendix N
Individual Animal Reports

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

PAGE : 7/ 140
Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R1, Control Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 40 * ANIMAL NO. : D0007

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 40 * ANIMAL NO. : D0008

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

Appendix N
Individual Animal Reports

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

PAGE : 8/ 140
Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R1, Control Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 42
* ANIMAL NO. : D0009

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 41
* ANIMAL NO. : D0010

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

Appendix N
Individual Animal Reports

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

ANIMAL HEADING DATA
DOSE GROUP : M1, Control Male

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0041	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0042	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0043	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0044	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0045	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0046	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0047	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0048	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0049	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0050	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02

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Individual Animal Reports

PATHOLOGY REPORT PAGE : 10/ 140
INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M1, Control Male MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0041

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LIVER:
-Lymphoid cell infiltration, grade 2
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0042

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

URINARY BLADDER:
-Seminal Coagulum, grade 2
PROSTATE GLAND:
-Lymphoid cell infiltration, grade 3
ADRENAL GLANDS:
-hypertrophy, cortex, focal/multifocal, bilateral, grade 2

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M1, Control Male MALE

CONT./FF. ANIMAL NO. : D0042

ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0043

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
-Lymphoid cell infiltration, bilateral, grade 1
EPIDIDYMIDES:
-Intratubular cellular debris, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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Individual Animal Reports

PATHOLOGY REPORT PAGE : 12/ 140
INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M1, Control Male MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0044

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
EPIDIDYMIDES:
-Intratubular cellular debris, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0045

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
-Lymphoid cell infiltration, bilateral, grade 1

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PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M1, Control Male

MALE

CONT./FF. ANIMAL NO. : D0045

EPIDIDYMIDES:
-Intratubular cellular debris, bilateral, grade 1
PROSTATE GLAND:
-Lymphoid cell infiltration, grade 1
ADRENAL GLANDS:
Recuts requested
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0046

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 2
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
PROSTATE GLAND:
-Lymphoid cell infiltration, grade 2
SPLEEN:
-Lymphoid hyperplasia, grade 3
MESENTERIC LYMPH NODE:
-Lymphoid hyperplasia, grade 3
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M1, Control Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0047

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

PROSTATE GLAND:
-Lymphoid cell infiltration, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0048

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LIVER:
-Focal necrosis, grade 1
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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Individual Animal Reports

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M1, Control Male MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0049

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO MICROSCOPIC FINDINGS NOTED.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0050

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Inflammation, subacute, focal/multifocal, grade 2
-Alveolar histiocytosis, focal/multifocal, grade 1
-Granuloma, grade 2
PROSTATE GLAND:
-Lymphoid cell infiltration, grade 3
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

ANIMAL HEADING DATA
DOSE GROUP : T1, Control Tox.

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0081	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0082	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0083	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0084	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0085	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0086	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0087	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0088	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0089	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0090	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02

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PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T1, Control Tox.

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0081

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

OVARIES:
-Cyst(s), follicular, bilateral
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0082

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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Individual Animal Reports

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T1, Control Tox. FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0083

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

STOMACH:
-Inflammation, grade 2
-Chief cell hyperplasia, grade 2
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0084

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO MICROSCOPIC FINDINGS NOTED.

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T1, Control Tox. FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0085

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

SPLEEN:
-Lymphoid hyperplasia, grade 2
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0086

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T1, Control Tox. FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0087

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0088

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO MICROSCOPIC FINDINGS NOTED.

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Individual Animal Reports

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T1, Control Tox. FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0089

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0090

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
OVARIES:
-Cyst(s), follicular, bilateral

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Individual Animal Reports

PATHOLOGY REPORT PAGE : 22/ 140
INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T1, Control Tox. FEMALE

CONT./FF. ANIMAL NO. : D0090

UTERUS:
-Dilation, grade 3
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

ANIMAL HEADING DATA
DOSE GROUP : R2, Low Repro.

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0011	F	K1	K1	40	25-MAR-02	03-MAY-02	03-MAY-02
D0012	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0013	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0014	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0015	F	K1	K1	43	25-MAR-02	06-MAY-02	06-MAY-02
D0016	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0017	F	K1	K1	42	25-MAR-02	05-MAY-02	05-MAY-02
D0018	F	K1	K1	44	25-MAR-02	07-MAY-02	07-MAY-02
D0019	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0020	F	K1	K1	40	25-MAR-02	03-MAY-02	03-MAY-02

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R2, Low Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 40

* ANIMAL NO. : D0011

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 41

* ANIMAL NO. : D0012

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R2, Low Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 41

* ANIMAL NO. : D0013

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 41

* ANIMAL NO. : D0014

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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INDIVIDUAL ANIMAL DATA

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R2, Low Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 43

* ANIMAL NO. : D0015

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 41

* ANIMAL NO. : D0016

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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INDIVIDUAL ANIMAL DATA

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R2, Low Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 42

* ANIMAL NO. : D0017

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

* STATE AT NECROPSY: K1
DAYS ON TEST : 44

* ANIMAL NO. : D0018

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R2, Low Repro. FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 41 * ANIMAL NO. : D0019

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 40 * ANIMAL NO. : D0020

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

ANIMAL HEADING DATA
DOSE GROUP : M2, Low Male

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0051	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0052	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0053	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0054	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0055	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0056	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0057	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0058	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0059	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0060	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0051

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male MALE

CONT./FF. ANIMAL NO. : D0051

RECTUM:
Organ not examined
LIVER:
-Lymphoid cell infiltration, grade 1
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

Appendix N
Individual Animal Reports

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0052

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

Appendix N
Individual Animal Reports

PATHOLOGY REPORT PAGE : 33/ 140
INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male MALE

CONT./FF. ANIMAL NO. : D0052

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

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Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0053

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* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

Appendix N
Individual Animal Reports

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INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male MALE

CONT./FF. ANIMAL NO. : D0053

RECTUM:
Organ not examined
LIVER:
-Focal necrosis, grade 1
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
-Lymphoid cell infiltration, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0054

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* NECROPSY FINDINGS

LUNG:
Right phrenic lobe: dark focus.
NO OTHER NECROPSY OBSERVATIONS NOTED

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined

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INDIVIDUAL ANIMAL DATA

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

CONT./FF. ANIMAL NO. : D0054

COLON:
Organ not examined
RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
-Lymphoid cell infiltration, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0055

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* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male MALE

CONT./FF. ANIMAL NO. : D0055

RECTUM:
Organ not examined
LIVER:
-Lymphoid cell infiltration, grade 1
KIDNEYS:
Organ examined, no pathologic findings noted
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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Individual Animal Reports

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0056

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male MALE

CONT./FF. ANIMAL NO. : D0056

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ examined, no pathologic findings noted
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

PATHOLOGY REPORT
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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0057

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* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

Appendix N
Individual Animal Reports

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male MALE

CONT./FF. ANIMAL NO. : D0057

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ examined, no pathologic findings noted
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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Individual Animal Reports

PATHOLOGY REPORT
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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0058

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* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

CONT./FF. ANIMAL NO. : D0058

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ examined, no pathologic findings noted
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

PATHOLOGY REPORT
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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0059

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* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

CONT./FF. ANIMAL NO. : D0059

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
-Lymphoid cell infiltration, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0060

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M2, Low Male

MALE

CONT./FF. ANIMAL NO. : D0060

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 1
-Lymphoid cell infiltration, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

Appendix N
Individual Animal Reports

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

ANIMAL HEADING DATA
DOSE GROUP : T2, Low Toxicity

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0091	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0092	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0093	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0094	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0095	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0096	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0097	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0098	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0099	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0100	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0091

.....

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0091

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0092

.....

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

Appendix N
Individual Animal Reports

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

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Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0092

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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Individual Animal Reports

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

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Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0093

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0093

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

* STATE AT NECROPSY: KO
DAYS ON TEST : 30

* ANIMAL NO. : D0094

.....

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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Individual Animal Reports

PATHOLOGY REPORT PAGE : 58/ 140
INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity FEMALE

CONT./FF. ANIMAL NO. : D0094

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0095

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity FEMALE

CONT./FF. ANIMAL NO. : D0095

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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Individual Animal Reports

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0096

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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Individual Animal Reports

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0096

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0097

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity FEMALE

CONT./FF. ANIMAL NO. : D0097

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0098

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity FEMALE

CONT./FF. ANIMAL NO. : D0098

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0099

.....

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0099

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0100

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T2, Low Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0100

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

ANIMAL HEADING DATA
DOSE GROUP : R3, Mid. Repro.

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0021	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0022	F	K1	K1	44	25-MAR-02	07-MAY-02	07-MAY-02
D0023	F	K1	K1	43	25-MAR-02	06-MAY-02	06-MAY-02
D0024	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0025	F	K1	K1	44	25-MAR-02	07-MAY-02	07-MAY-02
D0026	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0027	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0028	F	K1	K1	40	25-MAR-02	03-MAY-02	03-MAY-02
D0029	F	K1	K1	42	25-MAR-02	05-MAY-02	05-MAY-02
D0030	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R3, Mid. Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 41

* ANIMAL NO. : D0021

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 44

* ANIMAL NO. : D0022

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R3, Mid. Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 43

* ANIMAL NO. : D0023

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 41

* ANIMAL NO. : D0024

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R3, Mid. Repro. FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 44 * ANIMAL NO. : D0025

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 41 * ANIMAL NO. : D0026

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R3, Mid. Repro. FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 41 * ANIMAL NO. : D0027

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

* STATE AT NECROPSY: K1
DAYS ON TEST : 40 * ANIMAL NO. : D0028

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R3, Mid. Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 42

* ANIMAL NO. : D0029

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 41

* ANIMAL NO. : D0030

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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Individual Animal Reports

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INDIVIDUAL ANIMAL DATA

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Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

ANIMAL HEADING DATA
DOSE GROUP : M3, Mid. Male

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0061	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0062	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0063	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0064	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0065	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0066	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0067	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0068	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0069	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0070	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0061

.....

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

CONT./FF. ANIMAL NO. : D0061

.....

RECTUM:
Organ not examined
LIVER:
-Lymphoid cell infiltration, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 1
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0062

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

CONT./FF. ANIMAL NO. : D0062

.....

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0063

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

CONT./FF. ANIMAL NO. : D0063

.....

RECTUM:
Organ not examined
LIVER:
-Lymphoid cell infiltration, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 1
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

PATHOLOGY REPORT
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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

* STATE AT NECROPSY: KO
DAYS ON TEST : 29

* ANIMAL NO. : D0064

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
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PATHOL. NO. : 00004 JWC
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PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

CONT./FF. ANIMAL NO. : D0064

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 1
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
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TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0065

.....

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

CONT./FF. ANIMAL NO. : D0065

RECTUM:
Organ not examined
LIVER:
-Focal necrosis, grade 1
KIDNEYS:
--Protein droplet nephropathy, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0066

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* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

CONT./FF. ANIMAL NO. : D0066

RECTUM:
Organ not examined
LIVER:
-Lymphoid cell infiltration, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0067

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
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SPONSOR : SEHSC

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DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male

MALE

CONT./FF. ANIMAL NO. : D0067

.....

RECTUM:
Organ not examined
LIVER:
-Focal necrosis, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 1
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
-Inflammation, perirenal, bilateral, grade 3
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
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SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0068

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
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SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male MALE

CONT./FF. ANIMAL NO. : D0068

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 1
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
-Lymphoid cell infiltration, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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Individual Animal Reports

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
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SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0069

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male MALE

CONT./FF. ANIMAL NO. : D0069

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ examined, no pathologic findings noted
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0070

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
SEMINAL VESICLES:
Organ examined, no pathologic findings noted
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
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SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M3, Mid. Male MALE

CONT./FF. ANIMAL NO. : D0070

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 1
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
URINARY BLADDER:
Organ not examined
TESTES:
Organ not examined
EPIDIDYMIDES:
Organ not examined
PROSTATE GLAND:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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Individual Animal Reports

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INDIVIDUAL ANIMAL DATA Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

ANIMAL HEADING DATA
DOSE GROUP : T3, Mid. Toxicity

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0101	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0102	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0103	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0104	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0105	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0106	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0107	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0108	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0109	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0110	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0101

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0101

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0102

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0102

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0103

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0103

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0104

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* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0104

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0105

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0105

.....

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

* STATE AT NECROPSY: KO
DAYS ON TEST : 30

* ANIMAL NO. : D0106

.....

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0106

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0107

.....

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0107

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0108

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0108

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

* STATE AT NECROPSY: KO
DAYS ON TEST : 30

* ANIMAL NO. : D0109

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined
RECTUM:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0109

.....

LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0110

.....

* NECROPSY FINDINGS

LUNG:
THE LEFT LUNG FAILED TO INFLATE WITH FORMALIN.
NO OTHER NECROPSY OBSERVATIONS NOTED

* MICROSCOPIC FINDINGS

LUNG:
Organ not examined
BRAIN:
Organ not examined
SPINAL CORD:
Organ not examined
SCIATIC NERVE (RIGHT):
Organ not examined
HEART:
Organ not examined
NOSE:
Organ not examined
TRACHEA:
Organ not examined
STOMACH:
Organ not examined
DUODENUM:
Organ not examined
JEJUNUM:
Organ not examined
ILEUM:
Organ not examined
CECUM:
Organ not examined
COLON:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T3, Mid. Toxicity

FEMALE

CONT./FF. ANIMAL NO. : D0110

RECTUM:
Organ not examined
LIVER:
Organ examined, no pathologic findings noted
KIDNEYS:
Organ not examined
URINARY BLADDER:
Organ not examined
OVARIES:
Organ not examined
UTERUS:
Organ not examined
THYROID GLAND (BOTH LOBES):
Organ not examined
ADRENAL GLANDS:
Organ not examined
SPLEEN:
Organ not examined
BONE MARROW (STERNUM):
Organ not examined
THYMUS:
Organ not examined
MEDIASTINAL LYMPH NODE:
Organ not examined
MESENTERIC LYMPH NODE:
Organ not examined
THORACIC CAVITY:
Organ not examined
ABDOMINAL CAVITY:
Organ not examined

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

ANIMAL HEADING DATA
DOSE GROUP : R4, High Repro.

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0031	F	K1	K1	42	25-MAR-02	05-MAY-02	05-MAY-02
D0032	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02
D0033	F	K1	K1	44	25-MAR-02	07-MAY-02	07-MAY-02
D0034	F	K1	K1	42	25-MAR-02	05-MAY-02	05-MAY-02
D0035	F	K1	K1	42	25-MAR-02	05-MAY-02	05-MAY-02
D0036	F	K1	K1	42	25-MAR-02	05-MAY-02	05-MAY-02
D0037	F	K1	K1	44	25-MAR-02	07-MAY-02	07-MAY-02
D0038	F	K1	K1	43	25-MAR-02	06-MAY-02	06-MAY-02
D0039	F	K1	K1	42	25-MAR-02	05-MAY-02	05-MAY-02
D0120	F	K1	K1	41	25-MAR-02	04-MAY-02	04-MAY-02

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INDIVIDUAL ANIMAL DATA

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Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R4, High Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 42

* ANIMAL NO. : D0031

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 41

* ANIMAL NO. : D0032

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R4, High Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 44

* ANIMAL NO. : D0033

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 42

* ANIMAL NO. : D0034

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R4, High Repro. FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 42 * ANIMAL NO. : D0035

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

* STATE AT NECROPSY: K1
DAYS ON TEST : 42 * ANIMAL NO. : D0036

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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Individual Animal Reports

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R4, High Repro. FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 44 * ANIMAL NO. : D0037

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 43 * ANIMAL NO. : D0038

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : R4, High Repro.

FEMALE

* STATE AT NECROPSY: K1
DAYS ON TEST : 42

* ANIMAL NO. : D0039

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

---

* STATE AT NECROPSY: K1
DAYS ON TEST : 41

* ANIMAL NO. : D0120

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

NO EXAMINATION PERFORMED.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

ANIMAL HEADING DATA
DOSE GROUP : M4, High Male

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0071	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0072	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0073	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0074	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0075	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0076	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0077	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0078	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0079	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02
D0080	M	K0	K0	29	25-MAR-02	22-APR-02	22-APR-02

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M4, High Male MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0071

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 3
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
-Lymphoid cell infiltration, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0072

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M4, High Male

MALE

CONT./FF. ANIMAL NO. : D0072

* MICROSCOPIC FINDINGS

LIVER:
-Centrilobular hepatocellular hypertrophy, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 2
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 2
EPIDIDYMIDES:
-Intratubular cellular debris, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0073

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 2
SEMINAL VESICLES:
-Atrophy, bilateral, grade 2
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 3
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
-Lymphoid cell infiltration, bilateral, grade 1

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M4, High Male

MALE

CONT./FF. ANIMAL NO. : D0073

ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0074

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 2
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 2
PROSTATE GLAND:
-Lymphoid cell infiltration, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M4, High Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0075

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* NECROPSY FINDINGS

SEMINAL VESICLES:
Diminished in size.
NO OTHER NECROPSY OBSERVATIONS NOTED

* MICROSCOPIC FINDINGS

SEMINAL VESICLES:
-Atrophy, bilateral, grade 2
BRAIN:
-Neuronal degeneration, grade 1
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 3
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
URINARY BLADDER:
-Lymphoid cell infiltration, grade 2
PROSTATE GLAND:
-Lymphoid cell infiltration, grade 3
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M4, High Male

MALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0076

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Inflammation, subacute, focal/multifocal, grade 1
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 3
-Lymphoid cell infiltration, bilateral, grade 1
URINARY BLADDER:
-Lymphoid cell infiltration, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 29 * ANIMAL NO. : D0077

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M4, High Male

MALE

CONT./FF. ANIMAL NO. : D0077

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
SEMINAL VESICLES:
-Atrophy, bilateral, grade 2
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 2
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 2
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0078

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
SEMINAL VESICLES:
-Atrophy, bilateral, grade 2
NOSE:
-hemorrhage, grade 2
There is unilateral hemorrhage in the nasal tissues associated with a fractured tooth.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M4, High Male

MALE

CONT./FF. ANIMAL NO. : D0078

STOMACH:
-Mucosal gland dilation, grade 2
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 2
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 2
ADRENAL GLANDS:
-hypertrophy, cortex, focal/multifocal, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0079

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 2
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
URINARY BLADDER:
-Seminal Coagulum, grade 2

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : M4, High Male

MALE

CONT./FF. ANIMAL NO. : D0079

TESTES:
-Multinucleated spermatid giant cells, focal/multifocal, bilateral, grade 1
EPIDIDYMIDES:
-Lymphoid cell infiltration, bilateral, grade 1
PROSTATE GLAND:
-Lymphoid cell infiltration, grade 2
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 29

* ANIMAL NO. : D0080

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 1
KIDNEYS:
-Protein droplet nephropathy, bilateral, grade 3
TESTES:
-Tubular atrophy, focal/multifocal, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

ANIMAL HEADING DATA
DOSE GROUP : T4, High Toxicity

ANIMAL NUMBER	SEX M/F	DEFINED AND FINAL STATE OF NECROPSY		TEST DAYS	FIRST AND LAST DAY UNDER TEST		DATE OF NECROPSY
D0040	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0111	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0112	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0113	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0114	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0115	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0116	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0117	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0118	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02
D0119	F	K0	K0	30	25-MAR-02	23-APR-02	23-APR-02

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T4, High Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0040

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 2
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 30

* ANIMAL NO. : D0111

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
LIVER:
-Tigroid basophilic focus, grade 1
-Centrilobular hepatocellular hypertrophy, grade 2

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T4, High Toxicity FEMALE

CONT./FF. ANIMAL NO. : D0111

ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0112

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

STOMACH:
-Inflammation, grade 1
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 2
KIDNEYS:
-Tubular degeneration/regeneration, focal/multifocal, bilateral, grade 1
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T4, High Toxicity FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0113

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LIVER:
-Centrilobular hepatocellular hypertrophy, grade 2
SPLEEN:
-Lymphoid hyperplasia, grade 2
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0114

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LIVER:
-Centrilobular hepatocellular hypertrophy, grade 2
ADRENAL GLANDS:
-ANOMALY, CORTEX, EXTRACAPSULAR NODULE, bilateral
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.:* 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T4, High Toxicity FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0115

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

STOMACH:
-Inflammation, grade 2
-Glandular hyperplasia, focal/multifocal, grade 1
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 2
KIDNEYS:
-Lymphoid cell infiltration, bilateral, grade 1
URINARY BLADDER:
-Lymphoid cell infiltration, grade 1
ADRENAL GLANDS:
Recuts requested
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0116

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

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TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T4, High Toxicity FEMALE

CONT./FF. ANIMAL NO. : D0116

* MICROSCOPIC FINDINGS

LIVER:
-Centrilobular hepatocellular hypertrophy, grade 2
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0117

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LUNG:
-Alveolar histiocytosis, focal/multifocal, grade 1
LIVER:
-Centrilobular hepatocellular hypertrophy, grade 2
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

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TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TEXT OF GROSS AND MICROSCOPIC FINDINGS
DOSE GROUP : T4, High Toxicity

FEMALE

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0118

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LIVER:
-Centrilobular hepatocellular hypertrophy, grade 2
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

---

* STATE AT NECROPSY: K0
DAYS ON TEST : 30 * ANIMAL NO. : D0119

* NECROPSY FINDINGS

NO NECROPSY OBSERVATIONS NOTED.

* MICROSCOPIC FINDINGS

LIVER:
-Centrilobular hepatocellular hypertrophy, grade 2
ALL OTHER PROTOCOL TISSUES WITHOUT PATHOLOGIC FINDINGS.

Appendix O

Individual Animal Organ Weights

Appendix O
Individual Animal Organ Weights

Males (grams)

Animal ID	Brain	Heart	Liver	Lungs	Kidneys	Adrenals	Thymus	Spleen	Testes	Epididymides	Seminal Vesicles	Prostate	Body Weight
Group 1 - Controls													
D0041	1.832	0.864	9.368	1.089	2.178	0.0413	0.263	0.518	2.659	0.905	0.955	0.560	293.7
D0042	1.996	1.104	10.412	1.228	2.615	0.0507	0.496	0.718	3.015	1.040	1.608	0.904	368.7
D0043	2.035	1.079	9.623	1.229	2.592	0.0541	0.337	0.740	3.004	0.903	1.305	1.065	384.7
D0044	2.107	1.369	8.905	1.314	2.464	0.0616	0.427	0.716	3.305	1.182	1.655	0.928	360.3
D0045	2.036	1.009	6.877	1.067	2.109	0.0489	0.232	0.443	3.050	1.021	1.404	0.678	307.0
D0046	2.155	1.268	10.029	1.443	2.681	0.0566	0.312	0.882	2.780	1.174	1.240	1.040	379.7
D0047	1.999	1.291	9.902	1.280	2.503	0.0514	0.377	not taken	3.098	1.199	1.558	0.883	362.4
D0048	1.860	1.351	9.062	1.268	2.451	0.0517	0.362	0.691	3.171	1.169	1.522	0.930	354.0
D0049	2.093	1.234	10.916	1.435	2.912	0.0754	0.442	0.836	3.306	1.293	1.198	1.034	406.0
D0050	2.246	1.242	12.287	1.433	2.924	0.0637	0.391	0.798	3.129	1.166	1.308	0.677	394.3
Group 2 - 100 ppm													
D0051	2.084	1.425	11.811	1.404	3.010	0.0763	0.326	0.778	3.176	1.172	2.131	0.905	406.3
D0052	1.705	0.946	7.699	1.253	2.143	0.0485	0.305	0.665	2.502	0.982	1.457	0.349	322.7
D0053	1.867	1.212	9.043	1.375	2.385	0.0425	0.394	0.713	2.929	1.156	1.160	1.134	351.1
D0054	2.017	0.918	8.827	1.247	2.326	0.0492	0.529	0.646	2.964	1.102	0.852	0.756	342.9
D0055	2.095	1.112	8.861	1.221	2.541	0.0558	0.202	0.591	3.603	1.180	1.287	0.968	338.7
D0056	2.001	1.045	9.351	1.333	2.813	0.0626	0.309	0.683	3.096	1.196	1.373	0.869	358.8
D0057	2.128	1.368	11.332	1.429	2.845	0.0463	0.520	0.916	2.655	0.934	0.928	0.756	389.3
D0058	1.999	1.207	11.273	1.479	3.002	0.0862	0.246	0.817	3.077	1.178	1.724	1.307	395.8
D0059	1.940	1.137	8.535	1.338	2.302	0.0489	0.368	0.593	3.020	1.036	1.501	0.992	364.3
D0060	2.090	1.228	9.965	1.388	2.748	0.0584	0.463	0.738	3.594	1.196	1.342	1.022	396.8

Appendix O
Individual Animal Organ Weights

Males (grams)

Animal ID	Brain	Heart	Liver	Lungs	Kidneys	Adrenals	Thymus	Spleen	Testes	Epididymides	Seminal Vesicles	Prostate	Body Weight
Group 3 - 500 ppm													
D0061	1.973	0.996	11.432	1.211	2.566	0.0472	0.365	0.714	2.896	1.077	1.842	0.711	354.4
D0062	2.133	1.090	10.137	1.174	2.558	0.0550	0.334	0.624	2.921	1.072	1.436	0.860	362.0
D0063	2.008	1.081	10.039	1.292	2.601	0.0509	0.603	0.781	2.952	0.999	1.216	0.857	372.4
D0064	2.006	1.040	8.716	1.425	2.379	0.0500	0.277	0.702	3.050	0.971	1.327	0.925	364.8
D0065	1.978	1.086	9.266	1.395	2.758	0.0761	0.469	0.685	3.332	1.103	1.790	1.028	366.7
D0066	1.962	0.938	9.203	1.270	2.543	0.0500	0.236	0.597	3.004	1.052	1.445	1.055	358.7
D0067	2.048	1.129	10.165	1.355	2.550	0.0486	0.327	0.656	3.484	1.154	1.491	0.843	366.3
D0068	1.953	1.244	9.975	1.187	2.565	0.0405	0.385	0.628	3.019	1.107	1.421	0.864	349.9
D0069	1.997	1.418	9.825	1.341	2.603	0.0745	0.302	0.761	3.022	1.082	1.509	1.064	382.3
D0070	1.988	1.156	10.347	1.540	2.570	0.0774	0.253	0.675	3.107	1.149	1.856	1.044	391.4
Group 4 - 2500 ppm													
D0071	2.071	1.180	14.181	1.281	2.804	0.0546	0.336	0.969	2.705	1.051	1.194	0.886	363.1
D0072	1.914	1.021	13.347	1.187	2.573	0.0402	0.150	0.487	2.854	0.840	1.304	0.628	331.8
D0073	1.958	1.192	10.407	1.357	2.279	0.0498	0.353	0.611	3.250	0.985	0.827	0.610	318.4
D0074	2.093	1.239	11.501	1.334	2.966	0.0786	0.268	0.630	3.106	1.061	0.931	0.699	355.8
D0075	1.752	1.113	11.330	1.125	2.159	0.0462	0.287	0.430	2.942	0.859	0.349	0.464	276.6
D0076	1.935	0.895	12.773	1.285	2.138	0.0668	0.250	0.653	3.323	1.086	1.082	0.687	336.0
D0077	2.143	1.026	10.671	1.283	2.637	0.0534	0.303	0.779	3.111	1.029	0.875	0.588	329.7
D0078	1.986	1.042	10.592	1.172	2.839	0.0482	0.282	0.583	2.885	1.016	1.023	0.809	329.3
D0079	2.083	1.085	10.858	1.308	3.342	0.0598	0.462	0.720	2.795	0.985	1.127	0.846	374.2
D0080	1.957	1.113	12.976	1.265	3.320	0.0664	0.344	0.797	2.994	0.976	0.852	0.633	361.9

Appendix O
Individual Animal Organ Weights

Toxicity Females (grams)

Animal ID	Brain	Heart	Liver	Lungs	Kidneys	Adrenals	Thymus	Spleen	Uterus	Ovaries	Body Weight
Group 1 - Control											
D0081	1.900	0.885	7.259	1.172	1.827	0.0798	0.375	0.577	0.614	0.1046	244.7
D0082	1.899	0.694	6.289	1.025	1.507	0.0662	0.264	0.415	0.497	0.1354	222.1
D0083	1.962	0.869	6.892	1.062	1.654	0.0765	0.270	0.574	0.515	0.1016	235.8
D0084	1.799	0.890	6.595	1.031	1.556	0.0817	0.315	0.457	0.808	0.1020	239.5
D0085	2.013	0.910	7.403	1.092	1.740	0.0868	0.482	0.518	0.572	0.0804	248.6
D0086	1.854	0.765	6.501	1.040	1.567	0.0635	0.293	0.548	0.921	0.0785	245.9
D0087	1.951	0.909	6.695	1.204	1.621	0.0706	0.254	0.570	0.464	0.1020	244.2
D0088	1.931	0.833	6.725	1.048	1.599	0.0810	0.271	0.519	1.144	0.0898	240.0
D0089	1.859	0.867	7.220	1.118	1.698	0.0774	0.247	0.605	0.544	0.0982	247.9
D0090	1.854	0.764	6.628	1.174	1.662	0.0671	0.332	0.578	1.188	0.1452	249.8
Group 2 - 100 ppm											
D0091	1.900	0.775	6.717	0.935	1.537	0.0671	0.382	0.506	0.462	0.0994	216.1
D0092	1.804	0.833	7.325	1.182	1.784	0.0723	0.333	0.508	0.433	0.1134	234.6
D0093	1.808	0.809	8.051	1.102	1.558	0.0679	0.385	0.482	0.368	0.1145	232.9
D0094	1.874	0.736	7.739	1.122	1.826	0.0672	0.267	0.449	0.627	0.0781	247.3
D0095	1.927	0.794	6.478	1.009	1.400	0.0602	0.355	0.382	0.950	0.1211	241.0
D0096	1.867	0.776	6.623	1.092	1.487	0.0572	0.399	0.577	0.392	0.0897	234.4
D0097	1.865	0.763	5.979	1.041	1.452	0.0603	0.287	0.406	0.897	0.1047	235.6
D0098	1.886	0.816	6.373	1.182	1.474	0.0585	0.312	0.470	0.402	0.1080	237.8
D0099	1.783	0.756	9.264	1.310	1.805	0.0658	0.305	0.556	1.147	0.1012	244.3
D0100	1.856	0.775	6.948	1.283	1.497	0.0707	0.394	0.543	1.171	0.0846	255.7

Appendix O
Individual Animal Organ Weights

Toxicity Females (grams)

Animal ID	Brain	Heart	Liver	Lungs	Kidneys	Adrenals	Thymus	Spleen	Uterus	Ovaries	Body Weight
Group 3 - 500 ppm											
D0101	1.908	0.794	6.696	1.060	1.495	0.0676	0.412	0.408	0.624	0.1182	225.3
D0102	1.920	0.687	6.394	1.030	1.476	0.0691	0.366	0.545	0.491	0.0766	218.9
D0103	1.900	0.701	6.273	1.111	1.557	0.0736	0.309	0.463	0.607	0.0821	214.3
D0104	1.916	0.731	6.403	1.084	1.492	0.0916	0.321	0.420	0.578	0.1834	210.6
D0105	1.809	0.796	6.716	1.130	1.661	0.0673	0.195	0.485	0.948	0.0856	229.4
D0106	2.048	0.742	6.959	1.119	1.795	0.0756	0.284	0.444	0.973	0.1505	238.9
D0107	1.905	0.741	7.063	1.111	1.739	0.0831	0.338	0.458	0.980	0.1206	232.9
D0108	2.000	0.752	7.295	1.139	1.547	0.0761	0.227	0.534	1.296	0.1185	253.6
D0109	1.959	0.808	7.262	1.171	1.756	0.0714	0.260	0.612	0.491	0.1061	236.7
D0110	1.901	0.845	7.446	1.229	1.747	0.0743	0.356	0.675	0.505	0.0946	237.9
Group 4 - 2500 ppm											
D0111	1.788	0.788	9.279	1.058	1.683	0.0858	0.216	0.471	0.610	0.0832	210.6
D0112	1.803	0.906	9.714	1.218	1.476	0.0626	0.325	0.675	1.161	0.1109	241.9
D0113	1.838	0.817	9.394	1.164	1.573	0.0703	0.361	0.512	0.442	0.0878	234.2
D0114	1.771	0.690	8.216	1.002	1.614	0.0800	0.242	0.535	0.546	0.1112	223.8
D0115	1.922	0.734	8.584	1.159	1.504	0.0738	0.214	0.459	0.407	0.0965	207.8
D0116	1.903	0.847	8.306	1.062	1.733	0.1031	0.264	0.536	0.555	0.0919	219.6
D0117	1.871	0.876	9.200	1.082	1.824	0.0982	0.343	0.410	0.727	0.0990	227.7
D0118	1.943	0.728	9.004	1.110	1.666	0.0815	0.395	0.601	0.489	0.1100	244.0
D0119	1.908	0.726	9.362	1.152	1.430	0.0734	0.252	0.632	0.479	0.1164	238.3
D0040	1.805	0.820	9.300	1.048	1.711	0.0852	0.224	0.546	0.526	0.1099	222.4

Appendix P

Individual Animal Organ to Body Weight Ratios

Appendix P
Individual Animal Organ to Body Weight Ratios

Males

Animal ID	Brains/ Body Weight Ratio	Heart/ Body Weight Ratio	Liver/ Body Weight Ratio	Lungs/ Body Weight Ratio	Kidneys/ Body Weight Ratio	Adrenals/ Body Weight Ratio	Thymus/ Body Weight Ratio	Spleen/ Body Weight Ratio	Testes/ Body Weight Ratio	Epididymides/ Body Weight Ratio	Seminal Vesicles/ Body Weight Ratio	Prostate/ Body Weight Ratio
Group 1 - Controls												
D0041	0.00624	0.00294	0.0319	0.00371	0.00742	0.000141	0.00090	0.00176	0.00905	0.00308	0.00325	0.00191
D0042	0.00541	0.00299	0.0282	0.00333	0.00709	0.000138	0.00135	0.00195	0.00818	0.00282	0.00436	0.00245
D0043	0.00529	0.00280	0.0250	0.00319	0.00674	0.000141	0.00088	0.00192	0.00781	0.00235	0.00339	0.00277
D0044	0.00585	0.00380	0.0247	0.00365	0.00684	0.000171	0.00119	0.00199	0.00917	0.00328	0.00459	0.00258
D0045	0.00663	0.00329	0.0224	0.00348	0.00687	0.000159	0.00076	0.00144	0.00993	0.00333	0.00457	0.00221
D0046	0.00568	0.00334	0.0264	0.00380	0.00706	0.000149	0.00082	0.00232	0.00732	0.00309	0.00327	0.00274
D0047	0.00552	0.00356	0.0273	0.00353	0.00691	0.000142	0.00104	NA	0.00855	0.00331	0.00430	0.00244
D0048	0.00525	0.00382	0.0256	0.00358	0.00692	0.000146	0.00102	0.00195	0.00896	0.00330	0.00430	0.00263
D0049	0.00516	0.00304	0.0269	0.00353	0.00717	0.000186	0.00109	0.00206	0.00814	0.00318	0.00295	0.00255
D0050	0.00570	0.00315	0.0312	0.00363	0.00742	0.000162	0.00099	0.00202	0.00794	0.00296	0.00332	0.00172
Group 2 - 100 ppm												
D0051	0.00513	0.00351	0.0291	0.00346	0.00741	0.000188	0.00080	0.00191	0.00782	0.00288	0.00524	0.00223
D0052	0.00528	0.00293	0.0239	0.00388	0.00664	0.000150	0.00095	0.00206	0.00775	0.00304	0.00452	0.00108
D0053	0.00532	0.00345	0.0258	0.00392	0.00679	0.000121	0.00112	0.00203	0.00834	0.00329	0.00330	0.00323
D0054	0.00588	0.00268	0.0257	0.00364	0.00678	0.000143	0.00154	0.00188	0.00864	0.00321	0.00248	0.00220
D0055	0.00619	0.00328	0.0262	0.00360	0.00750	0.000165	0.00060	0.00174	0.01064	0.00348	0.00380	0.00286
D0056	0.00558	0.00291	0.0261	0.00372	0.00784	0.000174	0.00086	0.00190	0.00863	0.00333	0.00383	0.00242
D0057	0.00547	0.00351	0.0291	0.00367	0.00731	0.000119	0.00134	0.00235	0.00682	0.00240	0.00238	0.00194
D0058	0.00505	0.00305	0.0285	0.00374	0.00758	0.000218	0.00062	0.00206	0.00777	0.00298	0.00436	0.00330
D0059	0.00533	0.00312	0.0234	0.00367	0.00632	0.000134	0.00101	0.00163	0.00829	0.00284	0.00412	0.00272
D0060	0.00527	0.00309	0.0251	0.00350	0.00693	0.000147	0.00117	0.00186	0.00906	0.00301	0.00338	0.00258

NA = Not Applicable

Appendix P
Individual Animal Organ to Body Weight Ratios

Males

Animal ID	Brains/ Body Weight Ratio	Heart/ Body Weight Ratio	Liver/ Body Weight Ratio	Lungs/ Body Weight Ratio	Kidneys/ Body Weight Ratio	Adrenals/ Body Weight Ratio	Thymus/ Body Weight Ratio	Spleen/ Body Weight Ratio	Testes/ Body Weight Ratio	Epididymides/ Body Weight Ratio	Seminal Vesicles/ Body Weight Ratio	Prostate/ Body Weight Ratio
Group 3 - 500 ppm												
D0061	0.00557	0.00281	0.0323	0.00342	0.00724	0.000133	0.00103	0.00201	0.00817	0.00304	0.00520	0.00201
D0062	0.00589	0.00301	0.0280	0.00324	0.00707	0.000152	0.00092	0.00172	0.00807	0.00296	0.00397	0.00238
D0063	0.00539	0.00290	0.0270	0.00347	0.00698	0.000137	0.00162	0.00210	0.00793	0.00268	0.00327	0.00230
D0064	0.00550	0.00285	0.0239	0.00391	0.00652	0.000137	0.00076	0.00192	0.00836	0.00266	0.00364	0.00254
D0065	0.00539	0.00296	0.0253	0.00380	0.00752	0.000208	0.00128	0.00187	0.00909	0.00301	0.00488	0.00280
D0066	0.00547	0.00261	0.0257	0.00354	0.00709	0.000139	0.00066	0.00166	0.00837	0.00293	0.00403	0.00294
D0067	0.00559	0.00308	0.0278	0.00370	0.00696	0.000133	0.00089	0.00179	0.00951	0.00315	0.00407	0.00230
D0068	0.00558	0.00356	0.0285	0.00339	0.00733	0.000116	0.00110	0.00179	0.00863	0.00316	0.00406	0.00247
D0069	0.00522	0.00371	0.0257	0.00351	0.00681	0.000195	0.00079	0.00199	0.00790	0.00283	0.00395	0.00278
D0070	0.00508	0.00295	0.0264	0.00393	0.00657	0.000198	0.00065	0.00172	0.00794	0.00294	0.00474	0.00267
Group 4 - 2500 ppm												
D0071	0.00570	0.00325	0.0391	0.00353	0.00772	0.000150	0.00093	0.00267	0.00745	0.00289	0.00329	0.00244
D0072	0.00577	0.00308	0.0402	0.00358	0.00775	0.000121	0.00045	0.00147	0.00860	0.00253	0.00393	0.00189
D0073	0.00615	0.00374	0.0327	0.00426	0.00716	0.000156	0.00111	0.00192	0.01021	0.00309	0.00260	0.00192
D0074	0.00588	0.00348	0.0323	0.00375	0.00834	0.000221	0.00075	0.00177	0.00873	0.00298	0.00262	0.00196
D0075	0.00633	0.00402	0.0410	0.00407	0.00781	0.000167	0.00104	0.00155	0.01064	0.00311	0.00126	0.00168
D0076	0.00576	0.00266	0.0380	0.00382	0.00636	0.000199	0.00074	0.00194	0.00989	0.00323	0.00322	0.00204
D0077	0.00650	0.00311	0.0324	0.00389	0.00800	0.000162	0.00092	0.00236	0.00944	0.00312	0.00265	0.00178
D0078	0.00603	0.00316	0.0322	0.00356	0.00862	0.000146	0.00086	0.00177	0.00876	0.00309	0.00311	0.00246
D0079	0.00557	0.00290	0.0290	0.00350	0.00893	0.000160	0.00123	0.00192	0.00747	0.00263	0.00301	0.00226
D0080	0.00541	0.00308	0.0359	0.00350	0.00917	0.000183	0.00095	0.00220	0.00827	0.00270	0.00235	0.00175

Appendix P
Individual Animal Organ to Body Weight Ratios

Toxicity Females

Animal ID	Brains/ Body Weight Ratio	Heart/ Body Weight Ratio	Liver/ Body Weight Ratio	Lungs/ Body Weight Ratio	Kidneys/ Body Weight Ratio	Adrenals/ Body Weight Ratio	Thymus/ Body Weight Ratio	Spleen/ Body Weight Ratio	Uterus/ Body Weight Ratio	Ovaries/ Body Weight Ratio
Group 1 - Controls										
D0081	0.00776	0.00362	0.0297	0.00479	0.00747	0.000326	0.00153	0.00236	0.00251	0.000427
D0082	0.00855	0.00312	0.0283	0.00462	0.00679	0.000298	0.00119	0.00187	0.00224	0.000610
D0083	0.00832	0.00369	0.0292	0.00450	0.00701	0.000324	0.00115	0.00243	0.00218	0.000431
D0084	0.00751	0.00372	0.0275	0.00430	0.00650	0.000341	0.00132	0.00191	0.00337	0.000426
D0085	0.00810	0.00366	0.0298	0.00439	0.00700	0.000349	0.00194	0.00208	0.00230	0.000323
D0086	0.00754	0.00311	0.0264	0.00423	0.00637	0.000258	0.00119	0.00223	0.00375	0.000319
D0087	0.00799	0.00372	0.0274	0.00493	0.00664	0.000289	0.00104	0.00233	0.00190	0.000418
D0088	0.00805	0.00347	0.0280	0.00437	0.00666	0.000338	0.00113	0.00216	0.00477	0.000374
D0089	0.00750	0.00350	0.0291	0.00451	0.00685	0.000312	0.00100	0.00244	0.00219	0.000396
D0090	0.00742	0.00306	0.0265	0.00470	0.00665	0.000269	0.00133	0.00231	0.00476	0.000581
Group 2 - 100 ppm										
D0091	0.00879	0.00359	0.0311	0.00433	0.00711	0.000311	0.00177	0.00234	0.00214	0.000460
D0092	0.00769	0.00355	0.0312	0.00504	0.00760	0.000308	0.00142	0.00217	0.00185	0.000483
D0093	0.00776	0.00347	0.0346	0.00473	0.00669	0.000292	0.00165	0.00207	0.00158	0.000492
D0094	0.00758	0.00298	0.0313	0.00454	0.00738	0.000272	0.00108	0.00182	0.00254	0.000316
D0095	0.00800	0.00329	0.0269	0.00419	0.00581	0.000250	0.00147	0.00159	0.00394	0.000502
D0096	0.00797	0.00331	0.0283	0.00466	0.00634	0.000244	0.00170	0.00246	0.00167	0.000383
D0097	0.00792	0.00324	0.0254	0.00442	0.00616	0.000256	0.00122	0.00172	0.00381	0.000444
D0098	0.00793	0.00343	0.0268	0.00497	0.00620	0.000246	0.00131	0.00198	0.00169	0.000454
D0099	0.00730	0.00309	0.0379	0.00536	0.00739	0.000269	0.00125	0.00228	0.00470	0.000414
D0100	0.00726	0.00303	0.0272	0.00502	0.00585	0.000276	0.00154	0.00212	0.00458	0.000331

Appendix P
Individual Animal Organ to Body Weight Ratios

Toxicity Females

Animal ID	Brains/ Body Weight Ratio	Heart/ Body Weight Ratio	Liver/ Body Weight Ratio	Lungs/ Body Weight Ratio	Kidneys/ Body Weight Ratio	Adrenals/ Body Weight Ratio	Thymus/ Body Weight Ratio	Spleen/ Body Weight Ratio	Uterus/ Body Weight Ratio	Ovaries/ Body Weight Ratio
Group 3 - 500 ppm										
D0101	0.00847	0.00352	0.0297	0.00470	0.00664	0.000300	0.00183	0.00181	0.00277	0.000525
D0102	0.00877	0.00314	0.0292	0.00471	0.00674	0.000316	0.00167	0.00249	0.00224	0.000350
D0103	0.00887	0.00327	0.0293	0.00518	0.00727	0.000343	0.00144	0.00216	0.00283	0.000383
D0104	0.00910	0.00347	0.0304	0.00515	0.00708	0.000435	0.00152	0.00199	0.00274	0.000871
D0105	0.00789	0.00347	0.0293	0.00493	0.00724	0.000293	0.00085	0.00211	0.00413	0.000373
D0106	0.00857	0.00311	0.0291	0.00468	0.00751	0.000316	0.00119	0.00186	0.00407	0.000630
D0107	0.00818	0.00318	0.0303	0.00477	0.00747	0.000357	0.00145	0.00197	0.00421	0.000518
D0108	0.00789	0.00297	0.0288	0.00449	0.00610	0.000300	0.00090	0.00211	0.00511	0.000467
D0109	0.00828	0.00341	0.0307	0.00495	0.00742	0.000302	0.00110	0.00259	0.00207	0.000448
D0110	0.00799	0.00355	0.0313	0.00517	0.00734	0.000312	0.00150	0.00284	0.00212	0.000398
Group 4 - 2500 ppm										
D0111	0.00849	0.00374	0.0441	0.00502	0.00799	0.000407	0.00103	0.00224	0.00290	0.000395
D0112	0.00745	0.00375	0.0402	0.00504	0.00610	0.000259	0.00134	0.00279	0.00480	0.000458
D0113	0.00785	0.00349	0.0401	0.00497	0.00672	0.000300	0.00154	0.00219	0.00189	0.000375
D0114	0.00791	0.00308	0.0367	0.00448	0.00721	0.000357	0.00108	0.00239	0.00244	0.000497
D0115	0.00925	0.00353	0.0413	0.00558	0.00724	0.000355	0.00103	0.00221	0.00196	0.000464
D0116	0.00867	0.00386	0.0378	0.00484	0.00789	0.000469	0.00120	0.00244	0.00253	0.000418
D0117	0.00822	0.00385	0.0404	0.00475	0.00801	0.000431	0.00151	0.00180	0.00319	0.000435
D0118	0.00796	0.00298	0.0369	0.00455	0.00683	0.000334	0.00162	0.00246	0.00200	0.000451
D0119	0.00801	0.00305	0.0393	0.00483	0.00600	0.000308	0.00106	0.00265	0.00201	0.000488
D0040	0.00812	0.00369	0.0418	0.00471	0.00769	0.000383	0.00101	0.00246	0.00237	0.000494

Appendix Q

Table of Individual Animal Microscopic Findings

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT

PAGE : 1/ 18
Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

EXPLANATION OF CODES AND SYMBOLS

CODES AND SYMBOLS USED AT ANIMAL LEVEL:

M = Male animal
F = Female animal
K0 = Terminal sacrifice group
K1...K9 = Interim sacrifice groups 1...9

CODES AND SYMBOLS USED AT ORGAN LEVEL:

R = Recut(s) requested
! = Gross observat.not checked off histologically
' = Histologic examination not required
+ = Organ examined, findings present
– = Organ examined, no pathologic findings noted (AOFT only)

CODES AND SYMBOLS USED AT FINDING LEVEL:

GRADE 1 = Minimal / very few / very small
GRADE 2 = Slight / few / small
GRADE 3 = Moderate / moderate number / moderate size
P = Finding present, severity not scored
* = Comment in text of individual animal data

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT PAGE : 2/ 18
Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane *PATHOL. NO.*: 00004 JWC
TEST SYSTEM : RAT, 28D, Inhalation *DATE* : 07-NOV-02
SPONSOR : SEHSC *PathData© System V6.1a*

Additional codes used:

Codes used at group level:
T = Toxicity group females
R = Reproductive group females
Low = 100 ppm
Mid. = 500 ppm
High = 2500 ppm

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : R1, Control Repro.

ANIMAL NUMBER :	D 0001 FK1	D 0002 FK1	D 0003 FK1	D 0004 FK1	D 0005 FK1	D 0006 FK1	D 0007 FK1	D 0008 FK1	D 0009 FK1	D 0010 FK1
LUNG	.	.	.	.	.	.	.	.	.	.
BRAIN	.	.	.	.	.	.	.	.	.	.
SPINAL CORD	.	.	.	.	.	.	.	.	.	.
SCIATIC NERVE,RIGHT	.	.	.	.	.	.	.	.	.	.
HEART	.	.	.	.	.	.	.	.	.	.
NOSE	.	.	.	.	.	.	.	.	.	.
TRACHEA	.	.	.	.	.	.	.	.	.	.
STOMACH	.	.	.	.	.	.	.	.	.	.
DUODENUM	.	.	.	.	.	.	.	.	.	.
JEJUNUM	.	.	.	.	.	.	.	.	.	.
ILEUM	.	.	.	.	.	.	.	.	.	.
CECUM	.	.	.	.	.	.	.	.	.	.
COLON	.	.	.	.	.	.	.	.	.	.
RECTUM	.	.	.	.	.	.	.	.	.	.
LIVER	.	.	.	.	.	.	.	.	.	.
KIDNEYS	.	.	.	.	.	.	.	.	.	.
URINARY BLADDER	.	.	.	.	.	.	.	.	.	.
OVARIES	.	.	.	.	.	.	.	.	.	.
UTERUS	.	.	.	.	.	.	.	.	.	.
THYROID GLAND	.	.	.	.	.	.	.	.	.	.
ADRENAL GLANDS	.	.	.	.	.	.	.	.	.	.
SPLEEN	.	.	.	.	.	.	.	.	.	.
BONE MARROW, STERNUM	.	.	.	.	.	.	.	.	.	.
THYMUS	.	.	.	.	.	.	.	.	.	.
MEDIAST. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
MESENT. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
THORACIC CAVITY	.	.	.	.	.	.	.	.	.	.
ABDOMINAL CAVITY	.	.	.	.	.	.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : M1, Control Male

ANIMAL NUMBER :	D 0041 MK0	D 0042 MK0	D 0043 MK0	D 0044 MK0	D 0045 MK0	D 0046 MK0	D 0047 MK0	D 0048 MK0	D 0049 MK0	D 0050 MK0
LUNG	-	-	-	+	-	+	-	-	-	+
- Inflammation	.	.	.	.	.	.	.	.	.	2.
- Alveolar histiocytosis	.	.	.	1.	.	2.	.	.	.	1.
- Granuloma	.	.	.	.	.	.	.	.	.	2.
SEMINAL VESICLES	-	-	-	-	-	-	-	-	-	-
BRAIN	-	-	-	-	-	-	-	-	-	-
SPINAL CORD	-	-	-	-	-	-	-	-	-	-
SCIATIC NERVE,RIGHT	-	-	-	-	-	-	-	-	-	-
HEART	-	-	-	-	-	-	-	-	-	-
NOSE	-	-	-	-	-	-	-	-	-	-
TRACHEA	-	-	-	-	-	-	-	-	-	-
STOMACH	-	-	-	-	-	-	-	-	-	-
DUODENUM	-	-	-	-	-	-	-	-	-	-
JEJUNUM	-	-	-	-	-	-	-	-	-	-
ILEUM	-	-	-	-	-	-	-	-	-	-
CECUM	-	-	-	-	-	-	-	-	-	-
COLON	-	-	-	-	-	-	-	-	-	-
LIVER	+	-	-	-	-	-	-	+	-	-
- Focal necrosis	.	.	.	.	.	.	.	1.	.	.
- Lymphoid cell infiltration	2.	.	.	.	.	.	.	.	.	.
KIDNEYS	+	-	+	+	+	+	-	+	-	-
- Tubular degeneration/regeneration	1.	.	1.	1.	1.	1.	.	1.	.	.
- Lymphoid cell infiltration	.	.	1.	.	1.	.	.	.	.	.
URINARY BLADDER	-	+	-	-	-	-	-	-	-	-
- Seminal Coagulum	.	2.	.	.	.	.	.	.	.	.
TESTES	-	-	-	-	-	-	-	-	-	-
EPIDIDYMIDES	-	-	+	+	+	-	-	-	-	-
- Intratubular cellular debris	.	.	1.	1.	1.	.	.	.	.	.
PROSTATE GLAND	-	+	-	-	+	+	+	-	-	+
- Lymphoid cell infiltration	.	3.	.	.	1.	2.	1.	.	.	3.
THYROID GLAND	-	-	-	-	-	-	-	-	-	-
ADRENAL GLANDS	-	+	-	-	-R	-	-	-	-	-
- hypertrophy	.	2.	.	.	.	.	.	.	.	.
SPLEEN	-	-	-	-	-	+	-	-	-	-
- Lymphoid hyperplasia	.	.	.	.	.	3.	.	.	.	.
BONE MARROW, STERNUM	-	-	-	-	-	-	-	-	-	-

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : M1, Control Male

ANIMAL NUMBER :	D 0041 MK0	D 0042 MK0	D 0043 MK0	D 0044 MK0	D 0045 MK0	D 0046 MK0	D 0047 MK0	D 0048 MK0	D 0049 MK0	D 0050 MK0
THYMUS	–	–	–	–	–	–	–	–	–	–
MEDIAST. LYMPH NODE	–	–	–	–	–	–	–	–	–	–
MESENT. LYMPH NODE	–	–	–	–	–	+	–	–	–	–
– Lymphoid hyperplasia	.	.	.	.	.	3.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : T1, Control Tox.

ANIMAL NUMBER :	D 0081 FK0	D 0082 FK0	D 0083 FK0	D 0084 FK0	D 0085 FK0	D 0086 FK0	D 0087 FK0	D 0088 FK0	D 0089 FK0	D 0090 FK0
LUNG	–	+	–	–	–	–	–	–	+	+
– Alveolar histiocytosis	.	1.	.	.	.	.	.	.	1.	1.
BRAIN	–	–	–	–	–	–	–	–	–	–
SPINAL CORD	–	–	–	–	–	–	–	–	–	–
SCIATIC NERVE,RIGHT	–	–	–	–	–	–	–	–	–	–
HEART	–	–	–	–	–	–	–	–	–	–
NOSE	–	–	–	–	–	–	–	–	–	–
TRACHEA	–	–	–	–	–	–	–	–	–	–
STOMACH	–	–	+	–	–	–	–	–	–	–
– Inflammation.	.	.	2.	.	.	.	.	.	.	.
– Chief cell hyperplasia	.	.	2.	.	.	.	.	.	.	.
DUODENUM	–	–	–	–	–	–	–	–	–	–
JEJUNUM	–	–	–	–	–	–	–	–	–	–
ILEUM	–	–	–	–	–	–	–	–	–	–
CECUM	–	–	–	–	–	–	–	–	–	–
COLON	–	–	–	–	–	–	–	–	–	–
LIVER	–	–	–	–	–	–	–	–	–	–
KIDNEYS	–	–	–	–	–	+	+	–	+	+
– Tubular degeneration/regeneration.	.	.	.	.	.	1.	1.	.	1.	1.
URINARY BLADDER	–	–	–	–	–	–	–	–	–	–
OVARIES	+	–	–	–	–	–	–	–	–	+
– Cyst(s)	P.	.	.	.	.	.	.	.	.	P.
UTERUS	–	–	–	–	–	–	–	–	–	+
– Dilation	.	.	.	.	.	.	.	.	.	3.
THYROID GLAND	–	–	–	–	–	–	–	–	–	–
ADRENAL GLANDS	–	–	–	–	–	–	–	–	–	–
SPLEEN	–	–	–	–	+	–	–	–	–	–
– Lymphoid hyperplasia	.	.	.	.	2.	.	.	.	.	.
BONE MARROW, STERNUM	–	–	–	–	–	–	–	–	–	–
THYMUS	–	–	–	–	–	–	–	–	–	–
MEDIAST. LYMPH NODE	–	–	–	–	–	–	–	–	–	–
MESENT. LYMPH NODE	–	–	–	–	–	–	–	–	–	–

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : R2, Low Repro.

ANIMAL NUMBER :	D 0011 FK1	D 0012 FK1	D 0013 FK1	D 0014 FK1	D 0015 FK1	D 0016 FK1	D 0017 FK1	D 0018 FK1	D 0019 FK1	D 0020 FK1
LUNG	.	.	.	.	.	.	.	.	.	.
GENERAL OBSERVATIONS	.	.	.	.	.	.	.	.	.	.
BRAIN	.	.	.	.	.	.	.	.	.	.
SPINAL CORD	.	.	.	.	.	.	.	.	.	.
SCIATIC NERVE,RIGHT	.	.	.	.	.	.	.	.	.	.
HEART	.	.	.	.	.	.	.	.	.	.
NOSE	.	.	.	.	.	.	.	.	.	.
TRACHEA	.	.	.	.	.	.	.	.	.	.
STOMACH	.	.	.	.	.	.	.	.	.	.
DUODENUM	.	.	.	.	.	.	.	.	.	.
JEJUNUM	.	.	.	.	.	.	.	.	.	.
ILEUM	.	.	.	.	.	.	.	.	.	.
CECUM	.	.	.	.	.	.	.	.	.	.
COLON	.	.	.	.	.	.	.	.	.	.
RECTUM	.	.	.	.	.	.	.	.	.	.
LIVER	.	.	.	.	.	.	.	.	.	.
KIDNEYS	.	.	.	.	.	.	.	.	.	.
URINARY BLADDER	.	.	.	.	.	.	.	.	.	.
OVARIES	.	.	.	.	.	.	.	.	.	.
UTERUS	.	.	.	.	.	.	.	.	.	.
THYROID GLAND	.	.	.	.	.	.	.	.	.	.
ADRENAL GLANDS	.	.	.	.	.	.	.	.	.	.
SPLEEN	.	.	.	.	.	.	.	.	.	.
BONE MARROW, STERNUM	.	.	.	.	.	.	.	.	.	.
THYMUS	.	.	.	.	.	.	.	.	.	.
MEDIAST. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
MESENT. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
THORACIC CAVITY	.	.	.	.	.	.	.	.	.	.
ABDOMINAL CAVITY	.	.	.	.	.	.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

PAGE : 8/ 18
Study No.9658

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : M2, Low Male

ANIMAL NUMBER :	D 0051 MK0	D 0052 MK0	D 0053 MK0	D 0054 MK0	D 0055 MK0	D 0056 MK0	D 0057 MK0	D 0058 MK0	D 0059 MK0	D 0060 MK0
LUNG	.	.	.	!	.	.	.	.	.	.
SEMINAL VESICLES	–	–	–	–	–	–	–	–	–	–
BRAIN	.	.	.	.	.	.	.	.	.	.
SPINAL CORD	.	.	.	.	.	.	.	.	.	.
SCIATIC NERVE,RIGHT	.	.	.	.	.	.	.	.	.	.
HEART	.	.	.	.	.	.	.	.	.	.
NOSE	.	.	.	.	.	.	.	.	.	.
TRACHEA	.	.	.	.	.	.	.	.	.	.
STOMACH	.	.	.	.	.	.	.	.	.	.
DUODENUM	.	.	.	.	.	.	.	.	.	.
JEJUNUM	.	.	.	.	.	.	.	.	.	.
ILEUM	.	.	.	.	.	.	.	.	.	.
CECUM	.	.	.	.	.	.	.	.	.	.
COLON	.	.	.	.	.	.	.	.	.	.
RECTUM	.	.	.	.	.	.	.	.	.	.
LIVER	+	–	+	–	+	–	–	–	–	–
- Focal necrosis	.	.	1.	.	.	.	.	.	.	.
- Lymphoid cell infiltration	1.	.	.	.	1.	.	.	.	.	.
KIDNEYS	+	+	+	+	–	–	–	–	+	+
- Protein droplet nephropathy	.	.	.	.	.	.	.	.	.	1.
- Tubular degeneration/regeneration	1.	1.	1.	1.	.	.	.	.	.	.
- Lymphoid cell infiltration	.	.	1.	1.	.	.	.	.	1.	1.
URINARY BLADDER	.	.	.	.	.	.	.	.	.	.
TESTES	.	.	.	.	.	.	.	.	.	.
EPIDIDYMIDES	.	.	.	.	.	.	.	.	.	.
PROSTATE GLAND	.	.	.	.	.	.	.	.	.	.
THYROID GLAND	.	.	.	.	.	.	.	.	.	.
ADRENAL GLANDS	.	.	.	.	.	.	.	.	.	.
SPLEEN	.	.	.	.	.	.	.	.	.	.
BONE MARROW, STERNUM	.	.	.	.	.	.	.	.	.	.
THYMUS	.	.	.	.	.	.	.	.	.	.
MEDIAST. LYMPH NODE	.	.	.	.	.	.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : M2, Low Male

ANIMAL NUMBER :	D 0051 MK0	D 0052 MK0	D 0053 MK0	D 0054 MK0	D 0055 MK0	D 0056 MK0	D 0057 MK0	D 0058 MK0	D 0059 MK0	D 0060 MK0
MESENT. LYMPH NODE	: .	.	.	.	.	.	.	.	.	.
THORACIC CAVITY	: .	.	.	.	.	.	.	.	.	.
ABDOMINAL CAVITY	: .	.	.	.	.	.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : T2, Low Toxicity

ANIMAL NUMBER :	D 0091 FK0	D 0092 FK0	D 0093 FK0	D 0094 FK0	D 0095 FK0	D 0096 FK0	D 0097 FK0	D 0098 FK0	D 0099 FK0	D 0100 FK0
LUNG	.	.	.	.	.	.	.	.	.	.
BRAIN	.	.	.	.	.	.	.	.	.	.
SPINAL CORD	.	.	.	.	.	.	.	.	.	.
SCIATIC NERVE,RIGHT	.	.	.	.	.	.	.	.	.	.
HEART	.	.	.	.	.	.	.	.	.	.
NOSE	.	.	.	.	.	.	.	.	.	.
TRACHEA	.	.	.	.	.	.	.	.	.	.
STOMACH	.	.	.	.	.	.	.	.	.	.
DUODENUM	.	.	.	.	.	.	.	.	.	.
JEJUNUM	.	.	.	.	.	.	.	.	.	.
ILEUM	.	.	.	.	.	.	.	.	.	.
CECUM	.	.	.	.	.	.	.	.	.	.
COLON	.	.	.	.	.	.	.	.	.	.
RECTUM	.	.	.	.	.	.	.	.	.	.
LIVER	-	-	-	-	-	-	-	-	-	-
KIDNEYS	.	.	.	.	.	.	.	.	.	.
URINARY BLADDER	.	.	.	.	.	.	.	.	.	.
OVARIES	.	.	.	.	.	.	.	.	.	.
UTERUS	.	.	.	.	.	.	.	.	.	.
THYROID GLAND	.	.	.	.	.	.	.	.	.	.
ADRENAL GLANDS	.	.	.	.	.	.	.	.	.	.
SPLEEN	.	.	.	.	.	.	.	.	.	.
BONE MARROW, STERNUM	.	.	.	.	.	.	.	.	.	.
THYMUS	.	.	.	.	.	.	.	.	.	.
MEDIAST. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
MESENT. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
THORACIC CAVITY	.	.	.	.	.	.	.	.	.	.
ABDOMINAL CAVITY	.	.	.	.	.	.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : R3, Mid. Repro.

ANIMAL NUMBER :	D 0021 FK1	D 0022 FK1	D 0023 FK1	D 0024 FK1	D 0025 FK1	D 0026 FK1	D 0027 FK1	D 0028 FK1	D 0029 FK1	D 0030 FK1
LUNG	.	.	.	.	.	.	.	.	.	.
BRAIN	.	.	.	.	.	.	.	.	.	.
SPINAL CORD	.	.	.	.	.	.	.	.	.	.
SCIATIC NERVE,RIGHT	.	.	.	.	.	.	.	.	.	.
HEART	.	.	.	.	.	.	.	.	.	.
NOSE	.	.	.	.	.	.	.	.	.	.
TRACHEA	.	.	.	.	.	.	.	.	.	.
STOMACH	.	.	.	.	.	.	.	.	.	.
DUODENUM	.	.	.	.	.	.	.	.	.	.
JEJUNUM	.	.	.	.	.	.	.	.	.	.
ILEUM	.	.	.	.	.	.	.	.	.	.
CECUM	.	.	.	.	.	.	.	.	.	.
COLON	.	.	.	.	.	.	.	.	.	.
RECTUM	.	.	.	.	.	.	.	.	.	.
LIVER	.	.	.	.	.	.	.	.	.	.
KIDNEYS	.	.	.	.	.	.	.	.	.	.
URINARY BLADDER	.	.	.	.	.	.	.	.	.	.
OVARIES	.	.	.	.	.	.	.	.	.	.
UTERUS	.	.	.	.	.	.	.	.	.	.
THYROID GLAND	.	.	.	.	.	.	.	.	.	.
ADRENAL GLANDS	.	.	.	.	.	.	.	.	.	.
SPLEEN	.	.	.	.	.	.	.	.	.	.
BONE MARROW, STERNUM	.	.	.	.	.	.	.	.	.	.
THYMUS	.	.	.	.	.	.	.	.	.	.
MEDIAST. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
MESENT. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
THORACIC CAVITY	.	.	.	.	.	.	.	.	.	.
ABDOMINAL CAVITY	.	.	.	.	.	.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : M3, Mid. Male

ANIMAL NUMBER :	D 0061 MK0	D 0062 MK0	D 0063 MK0	D 0064 MK0	D 0065 MK0	D 0066 MK0	D 0067 MK0	D 0068 MK0	D 0069 MK0	D 0070 MK0
LUNG	.	.	.	.	.	.	.	.	.	.
SEMINAL VESICLES	–	–	–	–	–	–	–	–	–	–
BRAIN	.	.	.	.	.	.	.	.	.	.
SPINAL CORD	.	.	.	.	.	.	.	.	.	.
SCIATIC NERVE,RIGHT	.	.	.	.	.	.	.	.	.	.
HEART	.	.	.	.	.	.	.	.	.	.
NOSE	.	.	.	.	.	.	.	.	.	.
TRACHEA	.	.	.	.	.	.	.	.	.	.
STOMACH	.	.	.	.	.	.	.	.	.	.
DUODENUM	.	.	.	.	.	.	.	.	.	.
JEJUNUM	.	.	.	.	.	.	.	.	.	.
ILEUM	.	.	.	.	.	.	.	.	.	.
CECUM	.	.	.	.	.	.	.	.	.	.
COLON	.	.	.	.	.	.	.	.	.	.
RECTUM	.	.	.	.	.	.	.	.	.	.
LIVER	+	–	+	–	+	+	+	–	–	–
- Focal necrosis	.	.	.	.	1.	.	1.	.	.	.
- Lymphoid cell infiltration	1.	.	1.	.	.	1.	.	.	.	.
KIDNEYS	+	+	+	+	+	+	+	+	–	+
- Protein droplet nephropathy	1.	1.	1.	1.	1.	1.	1.	1.	.	1.
- Tubular degeneration/regeneration	1.	.	1.	1.	.	.	1.	1.	.	1.
- Lymphoid cell infiltration	.	.	.	.	.	.	.	1.	.	.
- Inflammation	.	.	.	.	.	.	3.	.	.	.
URINARY BLADDER	.	.	.	.	.	.	.	.	.	.
TESTES	.	.	.	.	.	.	.	.	.	.
EPIDIDYMIDES	.	.	.	.	.	.	.	.	.	.
PROSTATE GLAND	.	.	.	.	.	.	.	.	.	.
THYROID GLAND	.	.	.	.	.	.	.	.	.	.
ADRENAL GLANDS	.	.	.	.	.	.	.	.	.	.
SPLEEN	.	.	.	.	.	.	.	.	.	.
BONE MARROW, STERNUM	.	.	.	.	.	.	.	.	.	.
THYMUS	.	.	.	.	.	.	.	.	.	.
MEDIAST. LYMPH NODE	.	.	.	.	.	.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : M3, Mid. Male

ANIMAL NUMBER :	D 0061 MK0	D 0062 MK0	D 0063 MK0	D 0064 MK0	D 0065 MK0	D 0066 MK0	D 0067 MK0	D 0068 MK0	D 0069 MK0	D 0070 MK0
MESENT. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
THORACIC CAVITY	.	.	.	.	.	.	.	.	.	.
ABDOMINAL CAVITY	.	.	.	.	.	.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : T3, Mid. Toxicity

ANIMAL NUMBER :	D 0101 FK0	D 0102 FK0	D 0103 FK0	D 0104 FK0	D 0105 FK0	D 0106 FK0	D 0107 FK0	D 0108 FK0	D 0109 FK0	D 0110 FK0
LUNG	.	.	.	.	.	.	.	.	.	!
BRAIN	.	.	.	.	.	.	.	.	.	.
SPINAL CORD	.	.	.	.	.	.	.	.	.	.
SCIATIC NERVE,RIGHT	.	.	.	.	.	.	.	.	.	.
HEART	.	.	.	.	.	.	.	.	.	.
NOSE	.	.	.	.	.	.	.	.	.	.
TRACHEA	.	.	.	.	.	.	.	.	.	.
STOMACH	.	.	.	.	.	.	.	.	.	.
DUODENUM	.	.	.	.	.	.	.	.	.	.
JEJUNUM	.	.	.	.	.	.	.	.	.	.
ILEUM	.	.	.	.	.	.	.	.	.	.
CECUM	.	.	.	.	.	.	.	.	.	.
COLON	.	.	.	.	.	.	.	.	.	.
RECTUM	.	.	.	.	.	.	.	.	.	.
LIVER	-	-	-	-	-	-	-	-	-	-
KIDNEYS	.	.	.	.	.	.	.	.	.	.
URINARY BLADDER	.	.	.	.	.	.	.	.	.	.
OVARIES	.	.	.	.	.	.	.	.	.	.
UTERUS	.	.	.	.	.	.	.	.	.	.
THYROID GLAND	.	.	.	.	.	.	.	.	.	.
ADRENAL GLANDS	.	.	.	.	.	.	.	.	.	.
SPLEEN	.	.	.	.	.	.	.	.	.	.
BONE MARROW, STERNUM	.	.	.	.	.	.	.	.	.	.
THYMUS	.	.	.	.	.	.	.	.	.	.
MEDIAST. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
MESENT. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
THORACIC CAVITY	.	.	.	.	.	.	.	.	.	.
ABDOMINAL CAVITY	.	.	.	.	.	.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : R4, High Repro.

ANIMAL NUMBER :	D 0031 FK1	D 0032 FK1	D 0033 FK1	D 0034 FK1	D 0035 FK1	D 0036 FK1	D 0037 FK1	D 0038 FK1	D 0039 FK1	D 0120 FK1
LUNG	.	.	.	.	.	.	.	.	.	.
BRAIN	.	.	.	.	.	.	.	.	.	.
SPINAL CORD	.	.	.	.	.	.	.	.	.	.
SCIATIC NERVE,RIGHT	.	.	.	.	.	.	.	.	.	.
HEART	.	.	.	.	.	.	.	.	.	.
NOSE	.	.	.	.	.	.	.	.	.	.
TRACHEA	.	.	.	.	.	.	.	.	.	.
STOMACH	.	.	.	.	.	.	.	.	.	.
DUODENUM	.	.	.	.	.	.	.	.	.	.
JEJUNUM	.	.	.	.	.	.	.	.	.	.
ILEUM	.	.	.	.	.	.	.	.	.	.
CECUM	.	.	.	.	.	.	.	.	.	.
COLON	.	.	.	.	.	.	.	.	.	.
RECTUM	.	.	.	.	.	.	.	.	.	.
LIVER	.	.	.	.	.	.	.	.	.	.
KIDNEYS	.	.	.	.	.	.	.	.	.	.
URINARY BLADDER	.	.	.	.	.	.	.	.	.	.
OVARIES	.	.	.	.	.	.	.	.	.	.
UTERUS	.	.	.	.	.	.	.	.	.	.
THYROID GLAND	.	.	.	.	.	.	.	.	.	.
ADRENAL GLANDS	.	.	.	.	.	.	.	.	.	.
SPLEEN	.	.	.	.	.	.	.	.	.	.
BONE MARROW, STERNUM	.	.	.	.	.	.	.	.	.	.
THYMUS	.	.	.	.	.	.	.	.	.	.
MEDIAST. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
MESENT. LYMPH NODE	.	.	.	.	.	.	.	.	.	.
THORACIC CAVITY	.	.	.	.	.	.	.	.	.	.
ABDOMINAL CAVITY	.	.	.	.	.	.	.	.	.	.

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO.: 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : M4, High Male

ANIMAL NUMBER :	D 0071 MK0	D 0072 MK0	D 0073 MK0	D 0074 MK0	D 0075 MK0	D 0076 MK0	D 0077 MK0	D 0078 MK0	D 0079 MK0	D 0080 MK0
LUNG	+	–	+	+	–	+	+	+	+	+
– Inflammation	.	.	.	.	.	1.	.	.	.	.
– Alveolar histiocytosis	1.	.	2.	1.	.	.	1.	1.	1.	1.
SEMINAL VESICLES	–	–	+	–	+!	–	+	+	–	–
– Atrophy	.	.	2.	.	2.	.	2.	2.	.	.
BRAIN	–	–	–	–	+	–	–	–	–	–
– Neuronal degeneration	.	.	.	.	1.	.	.	.	.	.
SPINAL CORD	–	–	–	–	–	–	–	–	–	–
SCIATIC NERVE,RIGHT	–	–	–	–	–	–	–	–	–	–
HEART	–	–	–	–	–	–	–	–	–	–
NOSE	–	–	–	–	–	–	–	+	–	–
– hemorrhage	.	.	.	.	.	.	.	2*	.	.
TRACHEA	–	–	–	–	–	–	–	–	–	–
STOMACH	–	–	–	–	–	–	–	+	–	–
– Mucosal gland dilation	.	.	.	.	.	.	.	2.	.	.
DUODENUM	–	–	–	–	–	–	–	–	–	–
JEJUNUM	–	–	–	–	–	–	–	–	–	–
ILEUM	–	–	–	–	–	–	–	–	–	–
CECUM	–	–	–	–	–	–	–	–	–	–
COLON	–	–	–	–	–	–	–	–	–	–
LIVER	+	+	+	+	+	+	+	+	+	+
– Centrilobular hepatocellular hypertrophy	1.	1.	1.	1.	1.	1.	1.	1.	1.	1.
KIDNEYS	+	+	+	+	+	+	+	+	+	+
– Protein droplet nephropathy	3.	2.	3.	2.	3.	3.	2.	2.	2.	3.
– Tubular degeneration/regeneration	1.	2.	1.	2.	1.	.	2.	2.	1.	.
– Lymphoid cell infiltration	1.	.	1.	.	.	1.	.	.	.	.
URINARY BLADDER	–	–	–	–	+	+	–	–	+	–
– Seminal Coagulum	.	.	.	.	.	.	.	.	2.	.
– Lymphoid cell infiltration	.	.	.	.	2.	1.	.	.	.	.
TESTES	–	–	–	–	–	–	–	–	+	+
– Tubular atrophy	.	.	.	.	.	.	.	.	.	1.
– Multinucleated spermatid giant cells	.	.	.	.	.	.	.	.	1.	.
EPIDIDYMIDES	–	+	–	–	–	–	–	–	+	–
– Lymphoid cell infiltration	.	.	.	.	.	.	.	.	1.	.
– Intratubular cellular debris	.	1.	.	.	.	.	.	.	.	.
PROSTATE GLAND	–	–	–	+	+	–	–	–	+	–
– Lymphoid cell infiltration	.	.	.	1.	3.	.	.	.	2.	.
THYROID GLAND	–	–	–	–	–	–	–	–	–	–

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
PathData© System V6.1a

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : M4, High Male

ANIMAL NUMBER :	D 0071 MK0	D 0072 MK0	D 0073 MK0	D 0074 MK0	D 0075 MK0	D 0076 MK0	D 0077 MK0	D 0078 MK0	D 0079 MK0	D 0080 MK0
ADRENAL GLANDS	-	-	-	-	-	-	-	+	-	-
- hypertrophy	.	.	.	.	.	.	.	1.	.	.
SPLEEN	-	-	-	-	-	-	-	-	-	-
BONE MARROW, STERNUM	-	-	-	-	-	-	-	-	-	-
THYMUS	-	-	-	-	-	-	-	-	-	-
MEDIAST. LYMPH NODE	-	-	-	-	-	-	-	-	-	-
MESENT. LYMPH NODE	-	-	-	-	-	-	-	-	-	-

Appendix Q
Table of Individual Animal Microscopic Findings

PATHOLOGY REPORT
INDIVIDUAL ANIMAL DATA

TEST ARTICLE : hexamethylcyclotrisiloxane
TEST SYSTEM : RAT, 28D, Inhalation
SPONSOR : SEHSC

PATHOL. NO. : 00004 JWC
DATE : 07-NOV-02
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : T4, High Toxicity

ANIMAL NUMBER :	D 0040 FK0	D 0111 FK0	D 0112 FK0	D 0113 FK0	D 0114 FK0	D 0115 FK0	D 0116 FK0	D 0117 FK0	D 0118 FK0	D 0119 FK0
LUNG	+	+	–	–	–	–	–	+	–	–
– Alveolar histiocytosis	1.	1.	.	.	.	.	.	1.	.	.
BRAIN	–	–	–	–	–	–	–	–	–	–
SPINAL CORD	–	–	–	–	–	–	–	–	–	–
SCIATIC NERVE,RIGHT	–	–	–	–	–	–	–	–	–	–
HEART	–	–	–	–	–	–	–	–	–	–
NOSE	–	–	–	–	–	–	–	–	–	–
TRACHEA	–	–	–	–	–	–	–	–	–	–
STOMACH	–	–	+	–	–	+	–	–	–	–
– Inflammation.	.	.	1.	.	.	2.	.	.	.	.
– Glandular hyperplasia.	.	.	.	.	.	1.	.	.	.	.
DUODENUM	–	–	–	–	–	–	–	–	–	–
JEJUNUM	–	–	–	–	–	–	–	–	–	–
ILEUM	–	–	–	–	–	–	–	–	–	–
CECUM	–	–	–	–	–	–	–	–	–	–
COLON	–	–	–	–	–	–	–	–	–	–
LIVER	+	+	+	+	+	+	+	+	+	+
– Tigroid basophilic focus.	.	1.	.	.	.	.	.	.	.	.
– Centrilobular hepatocellular hypertrophy	2.	2.	2.	2.	2.	2.	2.	2.	2.	2.
KIDNEYS	+	–	+	–	–	+	–	–	–	–
– Tubular degeneration/regeneration.	1.	.	1.	.	.	.	.	.	.	.
– Lymphoid cell infiltration	.	.	.	.	.	1.	.	.	.	.
URINARY BLADDER	–	–	–	–	–	+	–	–	–	–
– Lymphoid cell infiltration	.	.	.	.	.	1.	.	.	.	.
OVARIES	–	–	–	–	–	–	–	–	–	–
UTERUS	–	–	–	–	–	–	–	–	–	–
THYROID GLAND	–	–	–	–	–	–	–	–	–	–
ADRENAL GLANDS	–	–	–	–	+	–R	–	–	–	–
– ANOMALY	.	.	.	.	P.	.	.	.	.	.
SPLEEN	–	–	–	+	–	–	–	–	–	–
– Lymphoid hyperplasia	.	.	.	2.	.	.	.	.	.	.
BONE MARROW, STERNUM	–	–	–	–	–	–	–	–	–	–
THYMUS	–	–	–	–	–	–	–	–	–	–
MEDIAST. LYMPH NODE	–	–	–	–	–	–	–	–	–	–
MESENT. LYMPH NODE	–	–	–	–	–	–	–	–	–	–

Appendix R

Individual Animal Reproductive Parameters – Reproductive Group Female Rats

Appendix R
Individual Animal Reproductive Parameters – Reproductive Group Female Rats

		Litter							Initial		Final			
Animal ID	Days Gestation	Undetermined Sex	Male Pups	Female Pups	Males/ Females	Total Pups	Viable (Lived) Pups	Viable/Total	Litter Weight (Grams)	Average Pup Weight (Grams)	Litter Weight (Grams)	Average Pup Weight (Grams)	Total Implants	Corpora Lutea Counts
Group 1 - Controls														
D0001	22	0	6	6	1.0	12	12	1.0	73.1	6.1	115.1	9.6	16	19
D0002	22	0	6	9	0.7	15	14	0.9	105.1	7.0	139.5	10.0	16	21
D0003	22	0	8	4	2.0	12	12	1.0	77.7	6.5	125.4	10.5	13	14
D0004	22	0	6	9	0.7	15	15	1.0	94.8	6.3	146.8	9.8	15	16
D0005	22	0	11	4	2.8	15	15	1.0	95.0	6.3	139.5	9.3	18	18
D0006	22	0	10	6	1.7	16	16	1.0	103.2	6.5	174.1	10.9	17	18
D0007	21	0	7	8	0.9	15	15	1.0	79.8	5.3	128.1	8.5	17	21
D0008	21	0	10	8	1.3	18	16	0.9	99.1	5.5	140.6	8.8	18	23
D0009	21	1	6	11	0.5	18	14	0.8	89.8	5.0	115.6	8.3	19	21
D0010	22	0	5	12	0.4	17	17	1.0	109.8	6.5	164.6	9.7	16	14
Group 2 - 100 ppm														
D0011	21	0	8	6	1.3	14	14	1.0	80.7	5.8	124.7	8.9	15	17
D0012	22	0	11	2	5.5	13	0	0.0	81.4	6.3	NA	NA	15	19
D0013	22	0	9	8	1.1	17	17	1.0	98.5	5.8	140.9	8.3	17	16
D0014	22	1	8	9	0.9	18	17	0.9	97.4	5.4	133.2	7.8	18	17
D0015	22	0	7	10	0.7	17	17	1.0	103.6	6.1	148.1	8.7	17	19
D0016	22	0	5	10	0.5	15	15	1.0	95.8	6.4	156.7	10.4	16	16
D0017	22	0	9	5	1.8	14	14	1.0	88.9	6.4	159.1	11.4	15	15
D0018	22	0	3	12	0.3	15	15	1.0	96.6	6.4	146.1	9.7	15	17
D0019	22	0	8	9	0.9	17	15	0.9	103.0	6.1	155.1	10.3	17	19
D0020	21	0	9	5	1.8	14	14	1.0	85.7	6.1	138.2	9.9	14	15

NA = Not Applicable

Appendix R
Individual Animal Reproductive Parameters – Reproductive Group Female Rats

		Litter							Initial		Final			
Animal ID	Days Gestation	Undetermined Sex	Male Pups	Female Pups	Males/ Females	Total Pups	Viable (Lived) Pups	Viable/Total	Litter Weight (Grams)	Average Pup Weight (Grams)	Litter Weight (Grams)	Average Pup Weight (Grams)	Total Implants	Corpora Lutea Counts
Group 3 - 500 ppm														
D0021	22	0	6	9	0.7	15	14	0.9	92.0	6.1	141.1	10.1	16	16
D0022	22	0	8	8	1.0	16	13	0.8	99.2	6.2	129.2	9.9	16	15
D0023	22	0	8	5	1.6	13	13	1.0	81.9	6.3	118.2	9.1	13	19
D0024	22	0	8	6	1.3	14	14	1.0	98.8	7.1	144.9	10.4	14	16
D0025	22	0	8	12	0.7	20	20	1.0	119.0	6.0	180.8	9.0	20	21
D0026	22	0	5	10	0.5	15	14	0.9	98.1	6.5	147.2	10.5	16	16
D0027	22	0	8	8	1.0	16	15	0.9	95.0	5.9	135.1	9.0	16	16
D0028	21	0	8	9	0.9	17	16	0.9	91.4	5.4	139.1	8.7	17	27
D0029	22	0	2	4	0.5	6	6	1.0	47.6	7.9	82.3	13.7	6	15
D0030	22	0	8	10	0.8	18	17	0.9	108.4	6.0	145.0	8.5	18	16
Group 4 - 2500 ppm														
D0031	22	0	2	1	2.0	3	3	1.0	20.5	6.8	33.9	11.3	3	14
D0032	22	0	4	6	0.7	10	10	1.0	64.9	6.5	104.0	10.4	10	14
D0033	NP	NP	NP	NP	NP	NP	NP	NP	NP	NP	NP	NP	NP	NP
D0034	23	0	4	2	2.0	6	6	1.0	42.2	7.0	78.5	13.1	11	29
D0035	22	2	8	4	2.0	14	0	0.0	81.4	5.8	NA	NA	16	16
D0036	22	0	7	5	1.4	12	12	1.0	77.2	6.4	123.5	10.3	12	15
D0037	22	0	7	8	0.9	15	15	1.0	101.1	6.7	145.5	9.7	16	15
D0038	22	0	5	8	0.6	13	11	0.8	79.4	6.1	115.2	10.5	12	20
D0039	22	0	5	5	1.0	10	10	1.0	72.1	7.2	101.7	10.2	10	14
D0120	22	0	4	6	0.7	10	10	1.0	69.1	6.9	107.7	10.8	10	12

NA = Not Applicable
NP = Not Pregnant

Appendix S

Statistical Analysis – Individual Scientist Report

Appendix S
Statistical Analysis
Individual Scientist Report

Methods

All data analysis was carried out using SAS® version 8.2. In all comparisons, the family wise error rate was held at 5% ($\alpha = 0.05$). Prior to this analysis, an exploratory analysis was carried out on all variables tested. Bartlett's test was used to check for heterogeneity of variances (except data used in the Fisher's Exact Test) and a Kolmogorov-Smirnov test was used to test normality of the data (except data used in the Repeated Measures ANOVA and Fisher's Exact Test). Parametric data was analyzed using one-way Analysis of Variance (ANOVA); nonparametric data was analyzed using the Kruskal Wallis Test. A significant ANOVA was followed by a Dunnett's Test and a significant Kruskal Wallis Test was followed by a Wilcoxon Test. For variables that had multiple measurements across time, Repeated Measures ANOVA was used to analyze the data and if the Repeated Measures ANOVA was significant, Linear Contrasts were performed on the data to see which groups differed. Categorical data was tested for equal proportions using the Fisher's Exact Test. Analysis of Covariance (ANCOVA) was used to analyze data that was correlated.

Statistically significant probabilities are reported at either the $p \leq 0.05$ or $p \leq 0.01$ levels.

Motor Activity

Motor activity data was transformed (square root) prior to analysis to minimize problems of heterogeneity of variances and departures from normality. Repeated Measures ANOVA was used to analyze this data. Factors that were included were sex, epoch (interval), treatment and the repeated factor of time (baseline and treated).

Measured Functional Observational Battery (FOB) Data

Comparisons of the mean measured FOB data, including rectal temperatures, hindlimb grip strength, forelimb grip strength and landing foot splay were carried out using ANCOVA, including the factors of sex, treatment and the repeated factor of time. Baseline evaluations were used as the covariate.

Scored and categorical Functional Observational Battery (FOB) Data

The following scored FOB results were analyzed by comparing the proportion of observations within each group between the test and control groups, using Fisher's Exact Test: resistance to removal (males only), extensor thrust (males only), categoricals (male only), level of activity, response to handling, response to sharp noise, responsiveness to tail pinch, urination, and defecation. If scores were the same in all the groups then no test of proportion could be performed; the following FOB results were not evaluated using Fisher's

Exact Test: palpebral closure, lacrimation, pupil size, salivation, pupil reaction, muscle tone, response to touch, and gait evaluation.

Body Weights

The body weights for males and toxicity group females were analyzed using ANOVA and Kruskal Wallis.

The body weights for the reproductive group females were analyzed using ANOVA for the pre-mating, gestational time points, post partum, and terminal sacrifice time points.

Change in Body Weights

Changes in body weights for males and toxicity group females and reproductive group females were analyzed using ANOVA and Kruskal Wallis.

Food Consumption

Food consumption data was analyzed for males using ANOVA for the first and second week data.

Food consumption data was analyzed for the toxicity group females using ANOVA.

Food consumption data was analyzed for the reproductive group females using ANOVA and Kruskal Wallis.

Absolute Organ Weights

In males, ANOVA was used to analyze all the organ weight data, except kidneys and adrenals, which were analyzed using Kruskal Wallis.

In the females, all organ weight data, except liver and uterus, was analyzed using ANOVA. The Kruskal Wallis Test was used to analyze the liver and uterus weight data and a Wilcoxon test was performed on liver weights.

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Individual Scientist Report

Organ to Body Weight Ratios

In males, ANOVA was used to analyze all organs to body weight data, except kidneys and liver. The Kruskal Wallis Test was used to analyze the kidney and liver to body weight data. Individual Wilcoxon tests were performed on these data.

In the females, all organ/weight ratios, except liver, uterus and ovaries was analyzed using ANOVA. The Kruskal Wallis Test was used to analyze the liver, uterus and ovaries to weight ratios and individual Wilcoxon tests were performed on liver to body weight ratios.

Reproduction Parameters

For females, a comparison of the male/female ratio, initial litter weight, average initial pup weight, final litter weight and average final pup weight were analyzed using ANOVA. A comparison of mean gestation length, mean number of undetermined sex, mean male pups, female pups, litter size, live litter size, ratio live births/litter size, number of implants and number of corpora lutea were analyzed using the Kruskal Wallis Test. Wilcoxon tests were performed on total pups, viable pups and total implants.

Clinical Chemistry Values

For the males, comparisons of the mean aminotransferase, aspartate aminotransferase, calcium, glucose, total protein, urea nitrogen and chloride data were carried out using ANOVA. The Kruskal Wallis Test was used to compare mean albumin, alkaline phosphatase, alanine cholesterol, sodium, phosphorus, total bilirubin, creatinine, and potassium data. Wilcoxon tests were also performed on alkaline phosphatase and sodium.

For the toxicity group females, comparisons of the mean alkaline phosphatase, alanine aminotransferase, aspartate aminotransferase, cholesterol, glucose, phosphorus, total protein and urea nitrogen were carried out using ANOVA. A Dunnett's test was performed on the mean sodium and chloride data. The Kruskal Wallis Test was used to compare the mean albumin, total bilirubin, calcium, creatinine, sodium, chloride and potassium data. The Wilcoxon Test was performed on the mean calcium data.

Hematology Values

For males, all values were analyzed using ANOVA.

For females, all values were tested using ANOVA except for white blood cell counts and mean cell volume, which used the Kruskal Wallis Test.

Differentials

For males, all values were analyzed using Kruskal Wallis except for segmented neutrophils and lymphocyte data, which were analyzed using ANOVA.

For females, all values were tested using the Kruskal Wallis Test, except for lymphocytes, which was analyzed using ANOVA.

Prothrombin Times

Male and female prothrombin times were analyzed using ANOVA.

Clinical Observations

Fisher's Exact Test was performed to compare on the proportion of occurrences of anogenital staining and probable porphyrin staining in males, toxicity group females and reproductive group females.

Results

Motor Activity

The treatment*time*sex interaction was not significant; therefore, further analyses were carried out on combined male and female data. The time*treatment interaction was significant, ($p<0.01$). When linear contrasts were performed on the data, the 2500 ppm group had a significantly lower mean level of motor activity than the control ($p=0.0108$). This means that the mean levels of motor activity differed between the 2500 ppm and control groups over the two time periods (baseline and treated). Also, the time*treatment*epoch interaction was not significant, ($p=0.4398$), meaning that the mean motor activity did not differ amongst the test and control groups over the time*epoch interactions. When males and females were analyzed separately, the time*treatment interaction was not significant for females, ($p=0.3593$) but it was significant for males, ($p<0.01$). When linear contrasts were performed on the data, the 2500 ppm group had a significantly lower mean level of motor activity compared to the control, ($p<0.01$). The time*treatment*epoch interaction also was not significant for females, ($p=0.6937$) but it was significant for males, ($p<0.05$) with the 500 ppm group having a different time*epoch slope compared to the control group.

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Measured Functional Observational Battery (FOB) Data

The results from the ANCOVA were not significant meaning there were no significant differences in any of the mean FOB values of the test groups compared to the control group.

Scored and categorical Functional Observational Battery (FOB) Data

This analysis was carried out separately for males and females on the treated data only, (baseline data not evaluated); the only significant difference that was found was in the reaction to handling for females between the control and the 2500 ppm groups, ($p<0.01$).

Body Weights

There were no significant differences between mean weights for male rats.

For toxicity group females, there was a significantly lower mean body weight in the 2500 ppm group compared to the control, at days 8 and 30, ($p<0.05$).

For reproductive group females, there was a significantly lower mean body weight in the 2500 ppm group compared to the control, at gestation day 20, ($p<0.05$).

Change in Body Weights

The mean change in body weight in males was significantly smaller in the 2500 ppm group compared to the control group during week 1 and in the total weight change, ($p<0.05$).

There were no significant differences in the mean change in body weights for toxicity group female rats.

For reproductive group females, there were significantly smaller differences in the mean change in body weight in the 2500 ppm group compared to the control group during gestation week 3, using ANOVA ($p<0.05$).

Food Consumption

There were significantly lower mean levels of food consumption in the 2500 ppm group compared to the control group for males during weeks 1 and 2, ($p<0.05$).

For toxicity group females, there was a significantly lower mean level of food consumption in the 2500 ppm group compared to the control group during week 1, ($p<0.05$). The mean

levels of food consumption were also lower in both the 500 and 2500 ppm groups at weeks 2 and 3, ($p<0.05$).

For reproductive group female rats, there were no significant differences in the mean level of food consumption.

Absolute Organ Weights

For male rats, the mean liver weights were greater and the mean epididymides and seminal vesicle weights were smaller in the 2500 ppm group compared to the control, ($p<0.05$).

For female rats, there was a significantly lower mean heart weight in the 500 ppm group compared to the control, ($p<0.05$). There was also a significantly greater mean liver weight, ($p<0.01$) and lower mean overall body weight, ($p<0.05$) in the 2500 ppm group compared to the control. There was also a significantly lower mean adrenal weight in the 100 ppm group compared to the control group, ($p<0.05$). The ovaries data underwent a log10 transformation to make the variances more homogeneous, prior to analysis.

Organ to Body Weight

For male rats, the only organs, relative to body weight, which had a significantly ($p<0.05$) different mean weight when compared to the control group, were liver, kidneys and seminal vesicles for the 2500 ppm group. The mean liver and kidneys to body weight ratios were significantly greater ($p<0.05$) in the 2500 ppm group whereas the mean seminal vesicles to body weight ratio was smaller in the 2500 ppm group ($p<0.01$).

For female rats, the only organs, relative to body weight, which had a significantly ($p<0.05$) greater mean weight when compared to the control group, were brain, liver, lungs and adrenals. The mean brain, liver and lungs were greater in the 500 ppm group whereas the liver, lungs and adrenals were greater in the 2500 ppm group.

Reproduction Parameters

There were a significantly lower mean number of total pups born, viable pups, initial litter weight, final litter weight and total implants in the 2500 ppm group compared to the control, ($p<0.05$). The male/female ratio underwent a log10 transformation to normalize the data.

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Clinical Chemistry Values

Male rats showed significantly, ($p<0.05$) lower levels of alkaline phosphatase and sodium in the 2500 ppm groups compared to the controls and a significantly higher level of urea nitrogen, ($p<0.05$).

Toxicity group female rats showed a significantly ($p< 0.05$) lower level of calcium and glucose in 500 ppm and in the 2500 ppm groups. The mean cholesterol level was significantly higher ($p<0.05$) in the 2500 ppm group when compared to the control.

Hematology Values

For males, there were significantly lower ($p<0.05$) levels of mean hemoglobin and significantly higher levels of platelets ($p<0.05$) in the 2500 ppm group compared to the controls.

For females, there were no significant differences in mean hematology values in any treatment group compared to the controls.

Differentials

There were no significant differences in the mean differentials of any treatment group when compared to the control for males or females.

Prothrombin Times

There were no significant differences in the mean prothrombin times of any treatment groups when compared to the control for males or females.

Clinical Observations

In toxicity group females, the proportion of rats with probable porphyrin staining was significantly greater in the 2500 ppm group compared to the control group, ($p<0.01$).

In reproductive group females, the proportion of rats with anogenital staining was significantly greater in the 2500 ppm group compared to the control group, ($p<0.01$) and the proportion of rats with probable porphyrin staining was also significantly greater in the 2500 ppm group compared to the control group, ($p<0.01$).

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Documentation

A copy of the SAS® log and SAS® output for this analysis is maintained in the study file.

Trevor Newhook
Trevor W. Newhook
Senior Biostatistician

11/20/2002
Date

Appendix T

90-Day Study Abstract

Appendix T
90-Day Study Abstract

Thirteen-week Inhalation Toxicity Study (with Recovery) of Hexamethylcyclotrisiloxane in Rats

SUMMARY

This study was conducted to determine the subchronic toxic effects, and recovery therefrom, of Hexamethylcyclotrisiloxane following nose-only inhalation exposure. Three groups, each comprised of ten male and ten female Fischer 344 rats, were exposed to target concentration of 15, 150 and 600 ppm Hexamethylcyclotrisiloxane vapors. A fourth exposure group consisted of 20 rats per sex and was exposed to the test substance at a target vapor concentration of 2500 ppm. A fifth group, the Filtered Air Control group, was initially comprised of 25 rats per sex and was exposed to filtered air only. Five rats per sex from the Filtered Air Control group were designated as sentinel animals; after approximately six weeks, blood and feces samples were collected from these rats and stored frozen for possible future analysis. The sentinel rats were euthanized and discarded. Body weight and food consumption data for the sentinel animals are included as control animal data in this report.

All rats were exposed six hours plus T90 (the time required to reach 90% of target concentration) per day, five days per week for 13 weeks. After the 13 week exposure period, ten rats per sex per group were euthanized and necropsied. The remaining ten rats per sex in the control and high exposure level group (2500 ppm) were allowed a four-week, nonexposure recovery period, after which they were euthanized and necropsied. Endpoints evaluated included survival, clinical condition, body weight, food consumption, ophthalmic examination, hematology, clinical chemistry, selected organ weights, gross necropsy and microscopic examination of selected organs and tissues.

The study was conducted in accordance with OECD Guidelines for Testing of Chemicals, "Subchronic Inhalation Toxicity: 90-Day Study", No.413, May 12, 1981. Test atmospheres of Hexamethylcyclotrisiloxane were generated as vapors by controlled sublimation of the test substance. The average test atmosphere concentrations, as determined by gas chromatography, were 0, 14.9, 150, 601 and 2519 ppm for the control, low, mid, mid high and high concentration groups that were targeted for 0, 15, 150, 600 and 2500 ppm, respectively.

None of the rats died during the study. Test substance-related clinical signs were limited to the 600 and 2500 ppm groups (both sexes). There was an exposure-dependent increased incidence of wet ventral abdominal and dorsal rump fur in both of these groups. In addition, the incidences of yellow discolored inguinal fur, red material around the mouth and rough hair coat were increased in the 2500 ppm group. These signs were primarily observed immediately following daily exposure completion. No other test substance-related clinical signs were observed.

Body weight gain was reduced in the 2500 ppm males during the first three weeks of exposure. No other test substance-related effects on body weight were observed. The aforementioned reduced body weight gain resulted in 2500 ppm male mean body weights that were reduced relative to the control group throughout the study, however, body weight gain for the 2500 ppm males was similar to the control group for the remainder of the study, i.e., Weeks 4-17. Food consumption was unaffected by exposure to the test substance throughout the study. There were no test substance- related ophthalmic findings at the end of the 13-week exposure period.

Erythrocytes, hemoglobin (Hgb), hematocrit (Hct), mean corpuscular volume (MCV) and mean corpuscular hemoglobin (MCH) were decreased and platelets were increased for the 2500 ppm males at the end of the 13-week exposure period. Also at this time, exposure-dependent decrease's in erythrocytes, Hgb and Hct were present for the 600 ppm males. Similar changes were not present for the 600 or 2500 ppm females, or in other test substance-exposed groups. These same indices remained slightly different from control for the 2500 ppm males at the recovery evaluation, but to a lesser magnitude than observed at 13 weeks, thus indicating substantial partial recovery upon cessation of exposure to the test substance. In addition, at the end of the treatment period, the incidence of several

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atypical red cell types (anisocytes, poikilocytes, microcytes and hypochromic) was increased in the 2500 ppm males, and the incidence of microcytosis was somewhat increased for the 2500 ppm females. The incidence of anisocytes, microcytes and hypochronic cells remained elevated for the 2500 ppm males at recovery, but no increase in atypical cells was present for the 2500 ppm females at that time. There were no other apparent test substance-related changes in hematological parameters.

At the 13-week clinical chemistry evaluation, serum protein levels and gamma glutamyl transpeptidase (GGT) were increased and chloride was decreased for both sexes in the 2500 ppm group. Also in this group, potassium and calcium were increased for the males and triglycerides were increased for the females. Glucose was decreased for the 600 and 2500 ppm males. Exposure- related decreases in alkaline phosphatase (AP) were present in all male groups and in the 150, 600 and 2500 ppm females. At the recovery evaluation, AP remained somewhat decreased for the 2500 ppm males, but less so than at 13 weeks, and the 2500 ppm female AP level was comparable to control. All other clinical chemistry changes observed at 13 weeks had returned to levels comparable to control at the recovery evaluation; therefore, the effects seen at 13 weeks were reversible upon cessation of exposure. There were no other test substance-related changes in clinical chemistry parameters.

Liver weight was increased, generally in an exposure-dependent manner, in all treated groups at the end of exposure, although the differences were not uniformly statistically significant, and the increases for the 15 ppm females were slight. In the 2500 ppm group, thymus weight was decreased (both sexes) and adrenal weight was increased (females only). At the recovery necropsy, liver weight was comparable to control for the 2500 males (when body weight differences were taken into account) and only slightly increased for the 2500 ppm females in comparison to the end of exposure (≈10% versus ≈70%). Adrenal weight for the 2500 ppm females was also still somewhat increased, but, as for the liver, much less so than at 13 weeks (≈14% versus ≈ 35%). No effect on thymus weight was apparent at the recovery necropsy. All effects seen on organ weights at the end of exposure exhibited substantial partial to full recovery and were therefore reversible upon cessation of exposure. There were no other test substance-related changes or differences in organ weights.

There were no test substance-related gross necropsy findings. At the end of exposure, test substance-related histomorphologic changes were seen in the livers from all treated male groups and in all treated female groups except at 15 ppm. The change consisted of generalized (diffuse) or centrilobular hypertrophy of the hepatocytes that exhibited a exposure-response with respect to both severity and distribution. No hepatocytic hypertrophy was present in either males or females (2500 ppm) at the recovery examination. This change was fully reversible at the highest exposure level given (2500 ppm) within 28 days after cessation of exposure.

Adrenal cortical cell vacuolation was seen for essentially all rats on the study, including the controls, at both the end of exposure and recovery .The severity of this finding was higher in the 600 and 2500 ppm groups (both sexes) at the end of exposure, but similar between the control and 2500 ppm groups at recovery. This change was fully reversible at the highest exposure level given (2500 ppm) within 28 days after cessation of exposure.

Based on the results of this study, the No Observed Adverse Effect Level for Hexamethylcyclotrisiloxane vapors administered six hours per day, five days per week for thirteen weeks via nose-only exposure to male and female Fischer 344 rats was 150 ppm.