FINAL REPORT



Contents: Volume 1 of 2

Text, Summary Tables, and Appendices A-F

Study Title: An Oral (Gavage) 14-Day and Pilot Prenatal

Developmental Toxicity Study of Naphthenic Acid in Rats

Study Number: WIL-402010

Study Director:

<u>Data Requirements</u>: Not Applicable

Study Initiation Date: 11 May 2010

Study Completion Date: 8 December 2010

<u>Performing Laboratory</u>: WIL Research Laboratories, LLC

1407 George Road

Ashland, OH 44805-8946

Sponsor: American Petroleum Institute

1220 L Street, NW

Washington, DC 20005

COMPLIANCE STATEMENT

This study, designated WIL-402010, was conducted in compliance with the United States EPA GLP Standards (40 CFR Part 792), 18 September 1989; the OECD Principles of GLP [C(97) 186/Final], 26 November 1997; WIL Research's SOPs; and the protocol as approved by the Sponsor.

Staff Toxicologist, Developmental and
Reproductive Toxicology
Study Director

Sponsor Representative

Applicant/Submitter

Date

Date

TABLE OF CONTENTS

| VOLUME 1 Page | | |
|---------------|--|----|
| | Compliance Statement | 2 |
| | Table of Contents | 3 |
| | Index of Tables | 6 |
| | Index of Appendices | 8 |
| 1. | Summary | 10 |
| 1.1. | Objective | 10 |
| 1.2. | Study Design | 10 |
| 1.3. | Results | 10 |
| 1.4. | Conclusion | 12 |
| 2. | Introduction | 14 |
| 2.1. | General Study Information | 14 |
| 2.2. | Key Study Dates | 14 |
| 2.3. | WIL Research Key Study Personnel | 15 |
| 3. | Study Design | 16 |
| 4. | Experimental Procedures - Materials and Methods | 17 |
| 4.1. | Test Substance and Vehicle | 17 |
| 4.1.1. | Test Substance | 17 |
| 4.1.2. | Vehicle | 17 |
| 4.1.3. | Preparation | 17 |
| 4.1.4. | Sampling and Analyses | 18 |
| 4.2. | Test System, Animal Receipt, and Acclimation | 19 |
| 4.3. | Animal Housing | 19 |
| 4.4. | Diet, Drinking Water, and Maintenance | 20 |
| 4.5. | Environmental Conditions | 20 |
| 4.6. | Assignment of Animals to Treatment Groups and Breeding Procedures | 21 |
| 4.7. | Organization of Test Groups, Dosage Levels, and Treatment Regimen | 22 |
| 5 | Parameters Evaluated | 23 |

| VOLU | JME 1 (continued) | <u>Page</u> |
|----------|--|-------------|
| 5.1. | Clinical Observations and Survival | 23 |
| 5.2. | Body Weights | 23 |
| 5.3. | Food Consumption | 23 |
| 5.4. | Macroscopic Examinations (Males) | 24 |
| 5.5. | Gestation Day 20 Laparohysterectomy | 24 |
| 5.6. | Organ Weights | 25 |
| 5.7. | External Fetal Morphological Examination | 25 |
| 5.8. | Data Acquisition and Analysis | 26 |
| 5.8.1. | Acquisition and Reporting | 26 |
| 5.8.2. | Statistical Analyses | 27 |
| 6. | Results and Discussion | 28 |
| 6.1. | Analytical Chemistry | 28 |
| 6.2. | Males | 29 |
| 6.2.1. | Clinical Observations and Survival | 29 |
| 6.2.2. | Body Weights | 29 |
| 6.2.3. | Food Consumption | 30 |
| 6.2.4. | Macroscopic Examinations | 30 |
| 6.2.4.1. | Organ Weights | 31 |
| 6.3. | Females | 31 |
| 6.3.1. | Clinical Observations and Survival | 31 |
| 6.3.2. | Body Weights | 32 |
| 6.3.3. | Food Consumption | 33 |
| 6.3.4. | Macroscopic Examinations | 34 |
| 6.3.5. | Organ Weights | 34 |
| 6.3.6. | Gestation Day 20 Laparohysterectomy | 34 |
| 6.3.7. | External Fetal Morphological Data | 36 |
| 7. | Conclusions | 38 |
| 8. | Report Review and Approval | 39 |
| Q | Quality Assurance Unit Statement | 40 |

| VOLUME 1 (continued) | | <u>Page</u> |
|----------------------|------------------|-------------|
| 9.1. | Phases Inspected | 40 |
| 9.2. | Approval | 42 |
| 10. | References | 43 |
| 11. | Data Retention | 44 |
| 12. | Abbreviations | 45 |

INDEX OF TABLES

Note: Some table titles have been shortened due to software spacing constraints.

| VOLUME 1 (continued) | <u>Page</u> |
|--|-------------|
| SUMMARY DATA - MALES | |
| S1. Summary of Clinical Findings (Males - Daily Examinations) | 47 |
| S2. Summary of Post-Dose Findings (Males) | 48 |
| S3. Summary of Body Weights [g] (Males) | 49 |
| S4. Summary of Body Weight Changes [g] (Males) | 50 |
| S5. Summary of Food Consumption [g/animal/day] (Males) | 51 |
| S6. Summary of Food Consumption [g/kg/day] (Males) | 52 |
| S7. Summary of Macroscopic Findings (Males) | 53 |
| S8. Summary of Organ Weights and Organ Wts. Relative to Body Wts | (Males)54 |
| SUMMARY DATA - FEMALES | |
| S9. Summary of Maternal Survival and Pregnancy Status | 57 |
| S10. Summary of Clinical Findings (Females - Daily Examinations) | 58 |
| S11. Summary of Post-Dose Findings (Females) | 61 |
| S12. Summary of Body Weights during Gestation [g] | 63 |
| S13. Summary of Body Weight Changes during Gestation [g] | 65 |
| S14. Summary of Gravid Uterine Wts. and Net Body Wt. Changes [g] | 67 |
| S15. Summary of Food Consumption during Gestation [g/animal/day] | 68 |
| S16. Summary of Food Consumption during Gestation [g/kg/day] | 70 |
| S17. Summary of Maternal Macroscopic Findings | 72 |
| S18 Summary of Organ Weights and Organ Wts Relative to Body Wt | ts 73 |

| WIL-402010 |
|------------------------------|
| American Petroleum Institute |

Naphthenic Acid

| | VOLUME 1 (continued) | <u>Page</u> |
|------|--|-------------|
| S19. | Summary of Fetal Data at Scheduled Necropsy | 76 |
| S20. | Summary of Fetal Data at Scheduled Necropsy [% per litter] | 77 |
| S21. | Summary of Fetuses and Litters with Malformations [absolute no.] | 80 |
| S22. | Summary of Fetuses and Litters with Variations [absolute no.] | 81 |

INDEX OF APPENDICES

| | VOLUME 1 (continued) | <u>Page</u> |
|----|--|-------------|
| A. | Study Protocol | 82 |
| B. | Test Substance Characterization (WIL Research Laboratories, LLC) | 101 |
| C. | Material Safety Data Sheet (Sponsor-Provided Data) | 116 |
| D. | Analyses of Dosing Formulations (WIL Research Laboratories, LLC) | 124 |
| E. | Animal Room Environmental Conditions | 158 |
| F. | WIL Developmental Historical Control Data Version 3.9 [Crl:CD(SD) Rats] | 164 |
| | VOLUME 2 | |
| G. | Individual Animal Data | 179 |
| | MALES | |
| | I1. Individual Clinical Observations (Males - Daily Examinations) | 180 |
| | I2. Individual Clinical Observations (Males - At Time of Dosing) | 195 |
| | I3. Individual Clinical Observations (Males - 1 Hour Post-Dosing) | 196 |
| | I4. Individual Body Weights [g] (Males) | 210 |
| | I5. Individual Body Weight Changes [g] (Males) | 215 |
| | I6. Individual Food Consumption [g/animal/day] (Males) | 220 |
| | I7. Individual Food Consumption [g/kg/day] (Males) | 225 |
| | I8. Individual Macroscopic Findings (Males) | 230 |
| | I9. Individual Organ Weights and Final Body Weights [g] (Males) | 255 |
| | I10. Individual Organ Wts. Relative to Final Body Wts. [g/100 g] (Males) | 260 |
| | FEMALES | |
| | II1. Individual Clinical Observations (Females - Daily Examinations) | 265 |

| VO | LUME 2 (continued) | Page |
|------|---|------|
| I12. | Individual Clinical Observations (Females - At Time of Dosing) | 272 |
| I13. | Individual Clinical Observations (Females - 1 Hour Post-Dosing) | 273 |
| I14. | Individual Body Weights during Gestation [g] | 292 |
| I15. | Individual Body Weight Changes during Gestation [g] | 297 |
| I16. | Individual Gravid Uterine Wts. and Net Body Wt. Changes [g] | 302 |
| I17. | Individual Food Consumption during Gestation [g/animal/day] | 307 |
| I18. | Individual Food Consumption during Gestation [g/kg/day] | 312 |
| I19. | Individual Maternal Macroscopic Findings | 317 |
| I20. | Individual Organ Weights and Final Body Weights [g] | 322 |
| I21. | Individual Organ Wts. Relative to Final Body Wts. [g/100 g] | 327 |
| I22. | Individual Fetal Data at Scheduled Necropsy | 332 |
| I23. | Individual Fetal Data at Scheduled Necropsy [% per litter] | 337 |
| I24. | Individual Fetal Weights [g] | 342 |
| I25. | Individual Fetal External Findings | 347 |

1. SUMMARY

1.1. OBJECTIVE

The study was designed to determine the dosage levels of the test substance to be evaluated in a potential combined repeated dose toxicity study with reproduction/developmental toxicity screening test (OECD 422) in rats.

1.2. STUDY DESIGN

The test substance, naphthenic acid, in the vehicle, corn oil, was administered orally by gavage once daily to 4 groups of Crl:CD(SD) rats, each group consisting of 5 males and 5 bred females. Dosage levels were 30, 100, 300, and 1000 mg/kg/day administered at a dosage volume of 10 mL/kg. A concurrent control group of 5 rats/sex/group received the vehicle (corn oil) on a comparable regimen. Males and females were approximately 9 and 13 weeks of age, respectively, at the beginning of test substance administration. Males were dosed daily for 14 consecutive days (study days 0-13). Following mating with untreated resident males, females were dosed during gestation days 1-20. All animals were observed twice daily for mortality and moribundity. Clinical observations, body weights, and food consumption were recorded at appropriate intervals. Gross necropsies were conducted on all males on study day 14. On gestation day 20, a laparohysterectomy was performed on each female. The uteri, placentae, and ovaries were examined, and the numbers of fetuses, early and late resorptions, total implantations, and corpora lutea were recorded. Gravid uterine weights were recorded, and net body weights and net body weight changes were calculated. The fetuses were weighed, sexed, and examined for external malformations and developmental variations. Selected organs from all animals at the scheduled necropsies were weighed.

1.3. RESULTS

All males and females survived to the scheduled necropsies. At approximately 1 hour following dose administration, clinical findings noted in the 1000 mg/kg/day group males and females included salivation and clear and/or red material around the nose and mouth and on the forelimbs and/or neck. Clear material around the nose and mouth were also

observed in the 100 and 300 mg/kg/day group females approximately 1 hour following dose administration. These findings generally did not persist to the daily examinations. In addition, behavior-related clinical findings consisting of hunched posture, rocking, lurching, or swaying while ambulating, and prostration were noted in 2 females in the 1000 mg/kg/day group at the daily examinations and/or approximately 1 hour following dose administration. At the daily examinations, an increase in the incidence of yellow material on the urogenital area was noted in the 1000 mg/kg/day group males and females.

Lower mean body weight gains and corresponding reduced food consumption were noted in the 1000 mg/kg/day group males throughout the treatment period (study days 0-14). As a result, mean body weight in this group was 9.1 % lower than the control group on study day 14. Mean body weights, body weight gains, and food consumption in the 30, 100, and 300 mg/kg/day male groups were unaffected by test substance administration.

A mean body weight loss during gestation days 0-6 with corresponding reduced food consumption was noted in the 1000 mg/kg/day group females when compared to the control group. This loss was primarily due to 2 females that lost 20 g to 27 g of body weight and were noted with behavioral-related clinical findings at the daily examinations and/or approximately 1 hour following dose administration during this period. Mean body weight gain in this group was similar to the control group during gestation days 6-14, and mean food consumption was similar to the control group throughout the remainder of the gestation period. However, lower mean body weight gains were noted in the 1000 mg/kg/day group during the last week of treatment (gestation days 14-20) and was attributed to the 3 females in this group; 2 females with completely resorbed litters and 1 female with a litter composed of 2 dead fetuses and 1 early resorption. Consequently, mean maternal body weight in the 1000 mg/kg/day group on gestation day 20 was 15.6% lower than the control group and mean terminal body weight, gravid uterine weights, hody weight gains, food consumption, gravid uterine weights, net

WIL-402010 Naphthenic Acid American Petroleum Institute

body weights, and net body weight changes in the 30, 100, and 300 mg/kg/day female groups were unaffected by test substance administration.

At the scheduled necropsies, no remarkable macroscopic findings were noted in the males and females at any dosage level. Increased mean absolute and relative (to final body weight) kidney and liver weights and decreased mean absolute and/or relative (to final body weight) thymus weights were noted in the 1000 mg/kg/day group males and females. Lower mean absolute ovarian weights were also noted in the 1000 mg/kg/day group females. In addition, lower mean final body weights were noted in the males and females in this group compared to the control group. In the 300 mg/kg/day group males, increased mean absolute and relative (to final body weight) liver weights were noted compared to the control group. No other effects on organ weights were noted when the test substance-treated groups were compared to the control group.

An increase in the mean litter proportion of pre-implantation loss and a corresponding lower mean number of implantation sites were noted in the 1000 mg/kg/day group compared with the control group. In addition, an increase in the mean litter proportion of postimplantation loss (primarily early resorptions) with a corresponding decrease in the number and proportion of viable fetuses and an increase in the mean litter proportion of dead fetuses were noted in the 1000 mg/kg/day group compared to the concurrent control group. There were no effects on mean fetal body weight or fetal sex ratio noted in the 1000 mg/kg/day group. Intrauterine growth and survival in the 30, 100, and 300 mg/kg/day groups were unaffected by test substance administration.

1.4. Conclusion

Clinical signs were noted in the 1000 mg/kg/day group females and consisted of hunched posture, rocking, lurching, or swaying while ambulating and prostration. Test substance-related lower mean body weight and reduced food consumption were noted in the 1000 mg/kg/day group males and females. Increased absolute and relative (to final body weight) kidney and liver weights and decreased mean absolute and/or relative (to final body weight) thymus weights were noted in the 1000 mg/kg/day group males and

WIL-402010 Naphthenic Acid
American Petroleum Institute

females. In addition, lower mean absolute ovarian weights were noted in the 1000 mg/kg/day group females. In the 300 mg/kg/day group males, a higher mean absolute and relative (to final body weight) liver weight was also observed. Two females in the 1000 mg/kg/day group had completely resorbed litters and 1 female had a litter composed of 2 dead fetuses and 1 early resorption. An increase in the mean litter proportion of pre-implantation and postimplantation loss were noted in the 1000 mg/kg/day group which corresponded to a decrease in the mean number and litter proportion of viable fetuses. No treatment-related effects on external fetal malformations and no developmental variations were noted in this study at any dosage level evaluated. Based on these results, dosage levels of 100, 300, and 900 mg/kg/day were selected for a combined repeated dose toxicity study with reproduction/developmental toxicity screening test (OECD 422) in Crl:CD(SD) rats administered naphthenic acid orally by gavage.

2. Introduction

2.1. GENERAL STUDY INFORMATION

This report presents the data from "An Oral (Gavage) 14-Day and Pilot Prenatal Developmental Toxicity Study of Naphthenic Acid in Rats." Due to software spacing constraints, the study title is presented as "Oral 14-Day/Pilot Dev Tox Study of Naphthenic Acid in Rats" on the report tables.

The study protocol is presented in Appendix A. A list of abbreviations potentially used in this report is presented in Section 12. (Abbreviations).

For the data collection process, each phase of the study was separated into what were termed WIL Research computer protocols. The computer protocol reference numbers and types of data collected were identified as follows:

| Computer Protocol | Type of Data Collected |
|--------------------------|------------------------|
| WIL-402010M | Male study data |
| WIL-402010F | Female study data |

2.2. KEY STUDY DATES¹

| <u>Date(s)</u> | Event(s) |
|----------------|---|
| 17 May 2010 | |
| | characterization of the test substance) |
| 2 June 2010 | Experimental start date (first test substance |
| | administration) |
| 2-15 June 2010 | Test substance administration period |
| | (males) |
| 2-24 June 2010 | Test substance administration period |
| | (females) |
| 16 June 2010 | Male necropsy |
| 24 June 2010 | Last female laparohysterectomy |
| 6 August 2010 | Experimental termination/completion date |
| J | (last characterization analysis) |
| | |

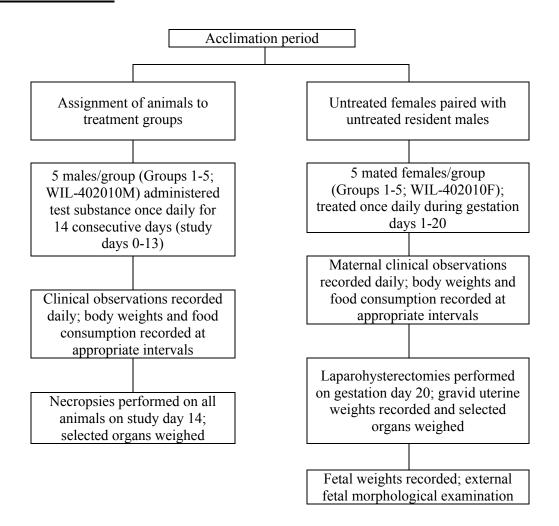
¹ Date ranges represent the first and last dates of evaluation.

2.3. WIL RESEARCH KEY STUDY PERSONNEL



Assistant Director, Analytical Chemistry
Senior Operations Manager, Vivarium
Manager, Gross Pathology and
Developmental Toxicology Laboratory
Group Manager, Formulations Laboratory
Director, Operations
Operations Manager, Reporting and
Regulatory Technical Services

3. STUDY DESIGN



American Petroleum Institute

4. Experimental Procedures - Materials and Methods

4.1. TEST SUBSTANCE AND VEHICLE

4.1.1. TEST SUBSTANCE

The test substance, naphthenic acid, was received from EPL Archives, Inc., Sterling, VA, on behalf of the Sponsor, on 1 April 2010, as follows:

| Identification | Physical Description |
|---|---------------------------|
| Naphthenic Acid Lot no. CP006002 CAS no.: 1338-24-5 [WIL log no. 10006B] | Clear, dark yellow liquid |

Characterization of the test substance was performed at WIL Research; details about the methodology and results of these analyses are presented in Appendix B. The purity of the test substance was 100%. The test substance was stored at room temperature, and according to the Material Safety Data Sheet was considered stable under this condition (see Appendix C). A reserve sample of the test substance was collected and stored in the Archives at WIL Research.

4.1.2. VEHICLE

The vehicle used in preparation of the test substance formulations and for administration to the control group was corn oil (lot no. YF0793, exp. date: 22 May 2011, received from Spectrum Chemical Manufacturing Corporation, New Brunswick, NJ).

4.1.3. PREPARATION

For the control group (Group 1), a sufficient amount of vehicle was transferred into a glass container. The vehicle was divided into aliquots for daily dispensation and stored at room temperature. The vehicle was mixed throughout preparation, sampling, and dose administration procedures.

American Petroleum Institute

Dosing formulations were prepared at the test substance concentrations indicated in the following table:

| Group Number | Treatment | Dosage Level (mg/kg/day) | Test substance Concentration ^a (mg/mL) |
|-----------------|-----------------|-----------------------------|---|
| 1 | Waliala Cantual | 0 | 0 |
| 1 | Vehicle Control | 0 | 0 |
| 2 | Naphthenic Acid | 30 | 3 |
| 3 | Naphthenic Acid | 100 | 10 |
| 4 | Naphthenic Acid | 300 | 30 |
| 5 | Naphthenic Acid | 1000 | 100 |

^a = The dosing formulations were not adjusted for purity.

The test substance formulations were prepared approximately weekly as single formulations for each dosage level, divided into aliquots for daily dispensation, and stored at room temperature. The test substance formulations were stirred continuously throughout the preparation, sampling, and dose administration procedures.

The first dosing formulations were visually inspected by the Study Director and were found to be visibly homogeneous and acceptable for administration.

4.1.4. Sampling and Analyses

Prior to the initiation of dose administration, samples for homogeneity determination were collected from the top, middle, and bottom strata of the 3 and 100 mg/mL dosing formulations. Samples for resuspension homogeneity were collected from the top and bottom strata of aliquots (similar in size needed for 1 day of dose administration) prepared from the same dosing formulations following 11-days of room temperature storage; aliquots were mixed for at least 30 minutes prior to sampling. Samples for concentration analysis were collected from the middle stratum of the dosing formulations (including the control group) prepared for the first and last weeks of dose administration. One set of samples from each collection was subjected to the appropriate analyses. The remaining set of samples was stored frozen (approximately -20°C) as back-up. All analyses were conducted by the Analytical Chemistry Department at WIL Research using

a validated gas chromatography method using mass spectrometric detection. Details about the methodology and results of these analyses are presented in Appendix D, and the

4.2. TEST SYSTEM, ANIMAL RECEIPT, AND ACCLIMATION

results are summarized in Section 6.1.

Sexually mature male and virgin female Sprague Dawley [Crl:CD(SD)] rats were used as the test system on this study. The animal model, the Crl:CD(SD) rat, is recognized as appropriate for reproduction studies and has been proven to be susceptible to the effects of reproductive toxicants. In addition, WIL Research has reproductive historical control data in the Crl:CD(SD) rat. The number of animals selected for this study (see Section 4.7.) was the minimum number of animals to allow for meaningful extrapolation to the definitive toxicity study.

Crl:CD(SD) rats (30 males and 32 females) were received in good health from Charles River Laboratories, Inc., Raleigh, NC on 18 May 2010. The males and females were approximately 46 and 79 days old, respectively, upon receipt. Each animal was examined by a qualified biologist on the day of receipt. The day following receipt, all animals were weighed and clinical observations were recorded. Each rat was uniquely identified by a Monel® metal ear tag displaying the animal number. The males were housed for an acclimation period of 15 days prior to the first day of treatment. The females were housed for an acclimation period of 14 days prior to breeding. During the acclimation period, the animals were observed twice daily for mortality and general changes in appearance and behavior. Female body weights were recorded prior to the initiation of breeding.

4.3. Animal Housing

Upon arrival (males and females) and until cohabitation (females), all rats were individually housed in clean, stainless steel, wire-mesh cages suspended above cage-board. The cage-board was changed at least 3 times per week. A separate group of untreated resident males were used for mating with the test substance-treated females. The female rats were paired for mating in the home cage of the untreated resident male.

Following positive evidence of mating, the females were returned to individual cages. Nesting material was not required, as euthanasia was scheduled prior to the date of expected parturition. Animals were maintained in accordance with the *Guide for the Care and Use of Laboratory Animals* (National Research Council, 1996). The animal facilities at WIL Research are fully accredited by AAALAC International.

4.4. <u>DIET, DRINKING WATER, AND MAINTENANCE</u>

The basal diet used in this study, PMI Nutrition International, LLC Certified Rodent LabDiet[®] 5002, was a certified feed with appropriate analyses performed by the manufacturer and provided to WIL Research. Feed lots used during the study were documented in the study records. Feeders were changed and sanitized once per week. Municipal water supplying the facility was sampled for contaminants according to WIL Research's SOPs. The results of the diet and water analyses are maintained at WIL Research. No contaminants were present in animal feed or water at concentrations sufficient to interfere with the objectives of this study. Reverse osmosis-purified (on-site) drinking water, delivered by an automatic watering system, and the basal diet were provided *ad libitum* throughout the acclimation period and during the study.

4.5. Environmental Conditions

All rats were housed throughout the acclimation period and during the study in an environmentally controlled room. The room temperature and humidity controls were set to maintain environmental conditions of $71^{\circ}F \pm 5^{\circ}F$ ($22^{\circ}C \pm 3^{\circ}C$) and $50\% \pm 20\%$ relative humidity. Room temperature and relative humidity data were recorded approximately hourly and are summarized in Appendix E. Actual mean daily temperature ranged from $70.6^{\circ}F$ to $73.6^{\circ}F$ ($21.4^{\circ}C$ to $23.1^{\circ}C$) and mean daily relative humidity ranged from 42.3% to 57.7% during the study. Fluorescent lighting provided illumination for a 12-hour light (0600 hours to 1800 hours)/12-hour dark photoperiod. Air handling units were set to provide a minimum of 10 fresh air changes per hour.

4.6. <u>Assignment of Animals to Treatment Groups and Breeding</u> Procedures

At the conclusion of the male acclimation period, all available males were weighed and examined in detail for physical abnormalities. At the discretion of the Study Director, each animal judged to be in good health and meeting acceptable body weight requirements was selected for use in the computerized randomization procedure. At that time, the individual body weights and corresponding animal identification numbers were entered into WTDMSTM. A printout containing the animal numbers, corresponding body weights, and individual group assignments was generated based on body weight stratification randomized in a block design. The animals then were arranged into groups according to the printout. These males were not used for mating.

At the conclusion of the female acclimation period, all available females were weighed and examined in detail for physical abnormalities. At the discretion of the Study Director, each animal judged to be in good health and meeting acceptable body weight requirements was placed in a suspended wire-mesh cage with a resident male from the same strain and source for breeding. Resident males were untreated, sexually mature rats utilized exclusively for breeding. These rats were maintained under similar laboratory conditions as the females. A breeding record containing the male and female identification numbers and the dates of cohabitation was maintained. Positive evidence of mating was confirmed by the presence of a vaginal copulatory plug or the presence of sperm in a vaginal lavage and verified by a second biologist. Each mating pair was examined daily. The day on which evidence of mating was identified was termed gestation day 0 and the animals were separated. The selected females were approximately 13 weeks old when paired for breeding.

The experimental design, consisted of 4 test substance-treated groups and 1 control group composed of 5 rats/sex/group. At the initiation of dose administration (study day 0), the males were approximately 9 weeks old; body weights ranged from 281 g to 352 g. The bred females were assigned to groups using a WTDMSTM computer program which

randomized the animals based on stratification of the gestation day 0 body weights in a block design. Body weight values of the selected females ranged from 225 g to 290 g on gestation day 0 (the day prior to the first day of dose administration). Animals not assigned to study were transferred to the WIL Research stock colony.

4.7. ORGANIZATION OF TEST GROUPS, DOSAGE LEVELS, AND TREATMENT REGIMEN

The vehicle and test substance formulations were administered orally by gavage, via an appropriately sized flexible, Teflon[®]-shafted, stainless steel ball-tipped dosing cannula (Natume, Japan) once daily. The males were dosed once daily for 14 consecutive days (study days 0-13). The females were dosed once daily during gestation days 1-20. The dosage volume for all groups was 10 mL/kg. Individual dosages were based on the most recently recorded body weights to provide the correct mg/kg/day dose. All animals were dosed at approximately the same time each day.

The following table presents the study group assignment:

| Group Number | Treatment | Dosage Level (mg/kg/day) | Dosage Volume (mL/kg) | Number of Males | Number of Females |
|-----------------|-----------------|--------------------------------|-----------------------------|-----------------|-------------------|
| | | | | | |
| 1 | Vehicle | 0 | 10 | 5 | 5 |
| 2 | Naphthenic Acid | 30 | 10 | 5 | 5 |
| 3 | Naphthenic Acid | 100 | 10 | 5 | 5 |
| 4 | Naphthenic Acid | 300 | 10 | 5 | 5 |
| 5 | Naphthenic Acid | 1000 | 10 | 5 | 5 |

Dosage levels were selected based on results of previous studies and were provided by the Sponsor after consultation with the Study Director.

The selected route of administration for this study was oral (gavage) because this is a potential route of exposure for humans and historically this route has been used extensively for studies of this nature.

5. PARAMETERS EVALUATED

5.1. CLINICAL OBSERVATIONS AND SURVIVAL

All rats were observed twice daily, once in the morning and once in the afternoon, for moribundity and mortality. Individual clinical observations were recorded daily, including the day of necropsy (prior to test substance administration during the treatment period). Each male and female was also observed for signs of toxicity approximately 1 hour following dose administration. The absence or presence of findings was recorded for individual animals. In addition, the presence of findings at the time of dose administration was recorded for individual animals.

5.2. BODY WEIGHTS

Individual male body weights were recorded weekly throughout the study and prior to the scheduled euthanasia. Mean weekly body weights and body weight changes are presented for each interval. In addition, cumulative mean body weight changes are presented for the entire treatment period (study days 0-14). Female body weights were recorded on gestation days 0, 6, 7, 10, 14, 17, and 20. Mean gestation body weights and corresponding mean body weight changes are presented for these intervals and for the overall gestation interval (days 0-20).

Gravid uterine weight was collected and net body weight (the gestation day 20 body weight exclusive of the weight of the uterus and contents) and net body weight change (the gestation day 0-20 body weight change exclusive of the weight of the uterus and contents) were calculated and presented for each gravid female at the scheduled laparohysterectomy.

5.3. FOOD CONSUMPTION

Individual male food consumption was recorded weekly throughout the study. Individual female food consumption was recorded on gestation days 0, 6, 7, 10, 14, 17, and 20. Food consumption was reported as g/animal/day and g/kg/day for the corresponding body weight change intervals. Calculation of the comprehensive intervals excludes all erroneous values such as total food spillage.

WIL-402010 Naphthenic Acid American Petroleum Institute

When food consumption could not be determined for an animal during a given interval (due to a weighing error, food spillage, obvious erroneous value, *etc.*), group mean values were calculated for that interval using the available data. The time periods when food consumption values were unavailable for a given animal were designated as "NA" on the individual report tables.

5.4. Macroscopic Examinations (Males)

All males were euthanized by carbon dioxide inhalation on study day 14. A gross necropsy was performed on all animals and included examination of the external surfaces, all orifices, the cranial cavity, the external surfaces of the brain and the thoracic, abdominal, and pelvic cavities including viscera. Tissues were preserved in 10% neutral-buffered formalin for possible future histopathologic examination only as indicated by the gross findings. The carcasses were then discarded.

5.5. GESTATION DAY 20 LAPAROHYSTERECTOMY

The laparohysterectomies and macroscopic examinations were performed blind to treatment group. All females were euthanized on gestation day 20 by carbon dioxide inhalation. The thoracic, abdominal, and pelvic cavities were opened by a ventral mid-line incision, and the contents were examined. In all instances, the *postmortem* findings were correlated with the *antemortem* comments, and any abnormalities were recorded. The uterus and ovaries were then exposed and excised. The number of corpora lutea on each ovary was recorded. The trimmed uterus was weighed and opened, and the number and location of all fetuses, early and late resorptions and the total number of implantation sites were recorded. The placentae were also examined. The individual uterine distribution of implantation sites was documented using the following procedure. All implantation sites, including resorptions, were numbered in consecutive order beginning with the left distal to the left proximal uterine horn, noting the position of the cervix, and continuing from the right proximal to the right distal uterine horn. The carcass of each female was then discarded.

WIL-402010 Naphthenic Acid American Petroleum Institute

Uteri with no macroscopic evidence of implantation were opened and subsequently placed in 10% ammonium sulfide solution for detection of early implantation loss (Salewski, 1964).

Intrauterine data were summarized using 2 methods of calculation. An example of each method of calculation follows:

1. Group Mean Litter Basis:

2. <u>Proportional Litter Basis</u>:

Where:

Postimplantation Loss/Litter (%) =
$$\frac{\text{No. Dead Embryos,}}{\text{Resorptions (Early/Late)/Litter}} \times 100$$

5.6. ORGAN WEIGHTS

The following organs were weighed from all animals at the scheduled necropsy.

Kidneys Liver
Testes^a Ovaries (with oviducts)^a
Thymus

Except as noted, paired organs were weighed together. Organ-to-final body weight ratios were calculated.

5.7. EXTERNAL FETAL MORPHOLOGICAL EXAMINATION

Fetal examinations were performed blind to treatment group. A detailed external examination of each fetus was conducted to include, but was not limited to, an

^a = These paired organs were weighed separately.

examination of the eyes, palate and external orifices. Findings were recorded as either developmental variations (alterations in anatomic structure that are considered to have no significant biological effect on animal health or body conformity and/or occur at high incidence, representing slight deviations from normal) or malformations (those structural anomalies that alter general body conformity, disrupt or interfere with normal body function, or may be incompatible with life). Each fetus was weighed, sexed, euthanized by a subcutaneous injection of sodium pentobarbital in the scapular region, and discarded.

5.8. DATA ACQUISITION AND ANALYSIS

5.8.1. ACQUISITION AND REPORTING

| Program/System | Description |
|---|---|
| Archive Management System (AMS) | In-house developed application for storage, maintenance, and retrieval information for archived materials (<i>e.g.</i> , lab books, study data, wet tissues, slides, <i>etc.</i>) |
| Formulations Dose Dispensing Management System (FDDMS) | In-house developed system used to assign unique barcodes to formulation containers and individual containers used for dispensing dosing formulations. |
| InSight® Publisher | Electronic publishing system (output is Adobe Acrobat, PDF) |
| Master Schedule | Maintains the master schedule for the company. |
| Metasys DDC Electronic Environmental Control System | Controls and monitors animal room environmental conditions. |
| Microsoft® Office 2002 and 2007 | Used in conjunction with the publishing software to generate study reports. |
| Provantis Dispense TM | Comprehensive system (Instem LSS Limited) to manage test materials, including receipt, formulation instructions, and accountability. |
| WIL Formulations Dispense System (WFDS) | In-house developed system for use in conjunction with Provantis Dispense [™] to ensure proper storage and use of formulations. |
| WIL Metasys | In-house developed system used to record and report animal room environmental conditions. |

WIL-402010 American Petroleum Institute

| Program/System | Description | | | |
|---|--|--|--|--|
| WIL Toxicology Data Management System TM (WTDMS TM) | In-house developed system used for collection and reporting of in-life and <i>postmortem</i> data. | | | |
| Note: Version numbers of WTDMS TM programs used for the study are presented on the | | | | |

Note: Version numbers of WTDMSTM programs used for the study are presented on the report data tables (reporting programs); version numbers and release dates are otherwise maintained in the study records and/or facility records.

5.8.2. STATISTICAL ANALYSES

All statistical tests were performed using WTMDSTM unless otherwise noted. Analyses were conducted using two-tailed tests (except as noted otherwise) for minimum significance levels of 1% and 5%, comparing each test substance-treated group to the control group by sex. Each mean was presented with the standard deviation (S.D.), standard error (S.E.), and the number of animals (N) used to calculate the mean. Due to the different rounding conventions inherent in the types of software used, the means, standard deviations, and standard errors on the summary and individual tables may differ by ± 1 in the last significant figure. Where applicable, the litter was used as the experimental unit.

Mean male and female body weights, body weight changes, food consumption, and absolute and relative organ weights; female net body weight, net body weight changes, and gravid uterine weights, numbers of corpora lutea, implantation sites, and viable fetuses, fetal body weights (separately by sex and combined), were subjected to a parametric one-way ANOVA (Snedecor and Cochran, 1980) to determine intergroup differences between the control and test substance-treated groups. If the ANOVA revealed significant (p<0.05) intergroup variance, Dunnett's test (Dunnett, 1964) was used to compare the test substance-treated groups to the control group. Mean litter proportions (percent per litter) of prenatal data (viable and nonviable fetuses, early and late resorptions, total resorptions, pre- and postimplantation loss, and fetal sex distribution) were subjected to the Kruskal-Wallis nonparametric ANOVA test (Kruskal and Wallis, 1952) to determine intergroup differences between the control and test substance-treated groups. If the ANOVA revealed significant (p<0.05) intergroup variance, Dunn's test (Dunn, 1964) was used to compare the test substance-treated groups to the control group.

6. RESULTS AND DISCUSSION

6.1. ANALYTICAL CHEMISTRY

Analytical Chemistry Report: Appendix D

The analyzed dosing formulations were within WIL Research's SOP range for suspensions (85% to 115%), were homogeneous, and were stable at room temperature for 11 days. Based on these results, the protocol-specified dosages of test substance were administered to the animals. No quantifiable amount of test substance was detected in the vehicle formulation that was administered to the control group (Group 1).

Results of the analyses of dosing formulations are summarized below.

Text Table 1. Results of Homogeneity Analyses

| | Group 2 (3 mg/mL) | Group 5 (100 mg/mL) | | | | | |
|---|-------------------|---------------------|--|--|--|--|--|
| Homogeneity Assessment of the 27 May 2010 Formulations | | | | | | | |
| Mean Concentration (mg/mL) | 3.04 | 101 | | | | | |
| RSD (%) | 2.0 | 5.0 | | | | | |
| Mean % of Target | 101 | 101 | | | | | |
| 11-Day Room Temperature Resuspension Homogeneity Assessment of the 27 May 2010 Formulations | | | | | | | |
| Mean Concentration (mg/mL) | 2.94 | 101 | | | | | |
| RSD (%) | 4.7 | 6.0 | | | | | |
| Mean % of Target | 97.9 | 101 | | | | | |

Text Table 2. Results of Stability Analyses

| | Mean Concentration, mg/mL (% of Time Zero) | | | |
|-------------------------|--|--------------|--|--|
| | Group 2 Group 5 | | | |
| | (3 mg/mL) | (100 mg/mL) | | |
| 11-Day Room Temperature | | | | |
| Storage | 2.94 (96.6) | 101 (100) | | |

American Petroleum Institute

Text Table 3. Results of Concentration Analyses

| | Mean Concentration, mg/mL (% of Target) | | | | |
|--------------|---|------------|-------------|-------------|--------------|
| Date of | Group 1 | Group 2 | Group 3 | Group 4 | Group 5 |
| Preparation | (0 mg/mL) | (3 mg/mL) | (10 mg/mL) | (30 mg/mL) | (100 mg/mL) |
| 1 June 2010 | NQ | 3.34 (111) | 9.22 (92.2) | 28.7 (95.8) | 97.4 (97.4) |
| 16 June 2010 | NQ | 3.04 (101) | 9.87 (98.7) | 27.9 (92.9) | 89.7 (89.7) |

NQ = Not quantifiable

6.2. MALES

6.2.1. CLINICAL OBSERVATIONS AND SURVIVAL

Summary Data: Table S1, Table S2

Individual Data: Table I1, Table I2, Table I3

All males in the control, 30, 100, 300, and 1000 mg/kg/day groups survived to the scheduled necropsy on study day 14. Clinical findings noted in the 1000 mg/kg/day group at approximately 1 hour following dose administration included clear material on the neck and forelimbs and around the nose and mouth beginning as early as study day 2 and generally continuing through study day 14. In addition, salivation prior to dosing was noted in all males in the 1000 mg/kg/day group on 1-6 occassions each during study days 7-14. At the daily examinations, an increased incidence of yellow material on the urogenital area was observed in 3 males in the 1000 mg/kg/day group during study days 3-14. All of the above-mentioned clinical findings were considered to be test substance-related. All other clinical findings noted in the test substance-treated groups occurred in frequently and/or in a manner that was not dose-related.

6.2.2. BODY WEIGHTS

Summary Data: Table S3, Table S4 Individual Data: Table I4, Table I5

A significantly (p<0.01) lower mean body weight gain was noted in the 1000 mg/kg/day group when compared to the control group during the first week of treatment (study days 0-7) followed by a lower (not statistically significant) mean body weight gain during

American Petroleum Institute

study days 7-14. As a result, a significantly (p<0.01) lower mean body weight gain was

noted in this group compared to the control group when the overall treatment period

(study days 0-14) was evaluated. Consequently, mean body weight in the

1000 mg/kg/day group was 9.1% lower than the control group on study day 14; the

difference was not statistically significant.

Mean body weights and body weight gains in the 30, 100, and 300 mg/kg/day groups

were unaffected by test substance administration. Differences from the control group

were slight and not statistically significant.

6.2.3. FOOD CONSUMPTION

Summary Data: Table S5, Table S6

Individual Data: Table I6, Table I7

Lower mean food consumption, evaluated as g/animal/day and g/kg/day, was noted in the

1000 mg/kg/day group during study days 0-7, 7-14, and when the overall treatment

period (study days 0-14) was evaluated; the difference from the control group was

significant (p<0.05) during study days 0-7. These reductions in food consumption

corresponded to the lower mean body weight gains observed in this group during the

treatment period.

Mean food consumption in the 30, 100, and 300 mg/kg/day groups was unaffected by test

substance administration. Differences from the control group were slight and not

statistically significant.

6.2.4. MACROSCOPIC EXAMINATIONS

Summary Data: Table S7

Individual Data: Table I8

All males survived to the scheduled necropsy on study day 14. Macroscopic findings

observed in the test substance-treated groups occurred infrequently and in a manner that

was not dose-related.

Page 30 of 360

American Petroleum Institute

6.2.4.1. ORGAN WEIGHTS

Summary Data: Table S8

Individual Data: Table I9, Table I10

Increased mean absolute and relative (to final body weight) kidney and liver weights were noted in the 1000 mg/kg/day group; differences were generally significant (p<0.01) when compared to the control group. A decrease in the mean absolute and relative (to final body weight) thymus weight was noted in the 1000 mg/kg/day group when compared with the control group; the difference was not statistically significant. In addition, a lower (not statistically significant) mean final body weight was noted in the 1000 mg/kg/day group compared to the control group. An increase in the mean absolute and relative (to final body weight) liver weight was noted in the 300 mg/kg/day group males when compared to the control group; only the difference in the relative to final body weight value was significant (p<0.01). No other effects on organ weight data were noted when the test substance-treated groups were compared to the control group.

6.3. FEMALES

6.3.1. CLINICAL OBSERVATIONS AND SURVIVAL

Summary Data: Table S9, Table S10, Table S11 Individual Data: Table I11, Table I12, Table I13

All females in the control, 30, 100, 300, and 1000 mg/kg/day groups survived to the scheduled necropsy. In the 1000 mg/kg/day group, behavioral clinical findings consisting of hunched posture, rocking, lurching, or swaying while ambulating, and/or prostration were noted for 2 females at approximately 1 hour following dose administration during gestation days 2-4. Hunched posture was also noted for 1 of these females at the daily examinations on gestation days 3 and 4. Clear material on the forelimbs and around the nose was noted in the 1000 mg/kg/day group at approximately 1 hour following dose administration beginning on gestation days 10 and 11, respectively, and generally continuing through gestation day 20. Additionally, clear or red material around the mouth was noted in this group at approximately 1 hour following

American Petroleum Institute

dose administration beginning as early as gestation day 2 and generally continuing through gestation day 20. Salivation prior to dosing was also noted for 2 females in this group during gestation days 10-18. At the daily examinations, an increase in the incidence of yellow material on the urogenital area was observed in the 1000 mg/kg/day group during gestation days 3-12. Clear material around the mouth and nose was also observed in the 100 and 300 mg/kg/day groups at approximately 1 hour following dose administration beginning as early as gestation day 4 and 7, respectively, and these observations continued sporadically until gestation day 20. In the 30 and 100 mg/kg/day groups, an increase in the incidences of hair loss on the forelimbs was observed at the daily examinations. In the absence of a dose-response, the hair loss noted in the 30 and 100 mg/kg/day groups was not attributed to test substance administration. There were no remarkable clinical findings noted in the 30 mg/kg/day group at the daily examinations or approximately 1 hour following dose administration.

6.3.2. BODY WEIGHTS

Summary Data: Table S12, Table S13, Table S14 Individual Data: Table I14, Table I15, Table I16

A mean body weight loss (2 g) was noted in the 1000 mg/kg/day group during gestation days 0-6 compared to a body weight gain in the control group (18 g); the difference was not statistically significant. This loss was primarily due to 2 females that lost 20 g to 27 g of body weight and were noted with behavioral-related clinical findings at the daily examinations and/or approximately 1 hour following dose administration during this period (see Section 6.3.1.). Mean body weight gains in this group were similar to the control group during gestation days 6-14. Lower mean body weight gains were noted in the 1000 mg/kg/day group during gestation days 14-20; the difference from the control group was significant (p<0.01) during gestation days 17-20. When the overall gestation period (gestation days 0-20) was evaluated, a significantly (p<0.01) lower mean body weight gain was noted in the 1000 mg/kg/day group compared to the control group. As a result of the lower mean body weight gains, mean body weights in the 1000 mg/kg/day

American Petroleum Institute

group were 15.6% lower than the control group on gestation day 20; the difference was

significant (p<0.01). Mean gravid uterine weight in this group was significantly (p<0.01)

lower than the control group and was attributed to the 2 females in this group with

complete litter resorptions and 1 female with a litter composed of 2 dead fetuses and 1

early resorption (see Section 6.3.6.). Mean net body weight and net body weight gain in

this group were slightly lower than the control group; the differences were not

statistically significant.

Mean maternal body weights, body weight gains, net body weights, net body weight

gains, and gravid uterine weights in the 30, 100, and 300 mg/kg/day groups were

unaffected by test substance administration. Differences from the control group were

slight and not statistically significant.

6.3.3. FOOD CONSUMPTION

Summary Data: Table S15, Table S16

Individual Data: Table I17, Table I18

Significantly (p<0.05) lower mean food consumption, evaluated as g/animal/day and

g/kg/day, was noted in the 1000 mg/kg/day group when compared to the control group

during gestation days 0-6 and corresponded to a mean body weight loss during this same

interval. Mean food consumption in this group was similar to the control group

throughout the remainder of the gestation period.

Mean food consumption in the 30, 100, and 300 mg/kg/day groups was unaffected by test

substance administration. Differences from the control group were slight and not

statistically significant with the exception of significantly (p<0.05) higher mean food

consumption (g/kg/day value only) in the 30 mg/kg/day group when the overall gestation

period (gestation days 0-20) was evaluated.

Page 33 of 360

American Petroleum Institute

6.3.4. MACROSCOPIC EXAMINATIONS

Summary Data: Table S17

Individual Data: Table I19

At the scheduled necropsy on gestation day 20, no remarkable internal findings were

observed at dosage levels of 30, 100, 300, and 1000 mg/kg/day. All females in the 30,

100, 300, and 1000 mg/kg/day groups were determined to be gravid.

6.3.5. ORGAN WEIGHTS

Summary Data: Table S18

Individual Data: Table I20, Table I21

Increased mean absolute and relative (to final body weight) kidney and liver weights

were noted in the 1000 mg/kg/day group; differences were generally significant (p<0.01)

when compared to the control group. Decreases in mean absolute thymus and ovarian

weights were also noted in this group when compared to the control group; the

differences were not statistically significant and the relative to final body weight values

were similar to the control group. Furthermore, the decrease in ovarian weights

corresponded to the increase in the mean litter proportions of pre- and postimplantation

loss noted in this group. In addition, a significantly (p<0.01) lower mean final body

weight was noted in the 1000 mg/kg/day group compared to the control group. No other

effects on organ weight data were noted when the test substance-treated groups were

compared to the control group.

6.3.6. GESTATION DAY 20 LAPAROHYSTERECTOMY

Summary Data: Table S19, Table S20

Individual Data: Table I22, Table I23, Table I24

Historical Control Data: Appendix F

An increase (not statistically significant) in the mean litter proportion of pre-implantation

loss was observed in the 1000 mg/kg/day group when compared to the concurrent control

group and the value exceeded the maximum mean value in the WIL historical control

Page 34 of 360

data. Consequently, a significantly (p<0.05) lower mean number of implantation sites was noted in the 1000 mg/kg/day group when compared to the control group. This reduction was primarily due to 3 females that had a lower number of implantation sites (3-5 implantation sites). The mean litter proportion of postimplantation loss (primarily early resorptions) in the 1000 mg/kg/day group was higher than the control group. Although the difference was not statistically significant when compared to the concurrent control group, the value exceeded the maximum mean value in the WIL historical control group data for definitive studies. A corresponding lower mean litter proportion and significantly (p<0.01) lower mean number of viable fetuses were noted in this group when compared to the concurrent control group and the values were below the minimum mean values in the WIL historical control data. These differences were attributed to 3 females (nos. 73718, 73701, and 73707) in this group; female nos. 73707 and 73718 had completely resorbed litters consisting of early resorptions and female no. 73701 had a litter composed of 2 dead fetuses and 1 early resorption. Additionally, an increase in the mean litter proportion of dead fetuses was observed in the 1000 mg/kg/day group compared to the concurrent control group. Although the difference was not statistically significant compared to the concurrent control group, the value exceeded the maximum mean value in the WIL historical control data. Mean fetal body weight and fetal sex ratio in the 1000 mg/kg/day group were unaffected by maternal test substance administration.

American Petroleum Institute

The gestation day 20 laparohysterectomy data are presented in the table below.

Text Table 4. Results of Gestation Day 20 Laparohysterectomy Data

| | Dosage Level (mg/kg/day) | | | | WIL HC ^a | |
|---------------------------------------|--------------------------|------|------|------|---------------------|------------------|
| Parameter | 0 | 30 | 100 | 300 | 1000 | Mean (Range) |
| Mean no. corpora lutea | 16.4 | 16.4 | 17.4 | 15.0 | 12.0 | 17.3 (14.5-19.4) |
| Mean no. implantation sites | 14.0 | 15.2 | 16.0 | 14.2 | 7.8* | 15.9 (13.0-17.6) |
| Mean early resorptions (% per litter) | 4.6 | 3.9 | 7.0 | 3.2 | 46.7 | 4.7 (1.5-9.9) |
| Pre-implantation loss (% per litter) | 9.6 | 6.3 | 8.1 | 5.3 | 36.1 | 7.4 (1.5-15.7) |
| Postimplantation loss (% per litter) | 4.6 | 3.9 | 7.0 | 3.2 | 60.0 | 4.8 (2.0-9.9) |
| Mean no. viable fetuses | 13.4 | 14.6 | 14.8 | 13.8 | 5.2** | 15.2 (12.2-17.1) |
| Viable fetuses (% per litter) | 95.4 | 96.2 | 93.0 | 96.8 | 40.0 | 95.2 (90.1-98.0) |
| Dead fetuses (% per litter) | 0.0 | 0.0 | 0.0 | 0.0 | 13.3 | 0.0 (0.0-0.5) |

^a = WIL historical control data

Intrauterine growth and survival were unaffected by maternal test substance administration at dosage levels of 30, 100, and 300 mg/kg/day. Mean numbers of corpora lutea and implantation sites and the mean litter proportions of pre-implantation loss in these groups were similar to the control group. Differences from the control group were slight and not statistically significant.

6.3.7. EXTERNAL FETAL MORPHOLOGICAL DATA

Summary Data: Table S21, Table S22

Individual Data: Table I25

Historical Control Data: Appendix F

The numbers of fetuses (litters) available for morphological evaluation were 67(5), 73(5), 74(5), 69(5), and 26(2) in the control, 30, 100, 300, and 1000 mg/kg/day groups, respectively. External malformations observed in the test substance-treated groups consisted of open eyelids (bilateral) in dead fetuses (fetus nos. 73701-02 and 73701-03) in the 1000 mg/kg/day group and omphalocele (several loops of intestine protruded through an opening in the umbilicus with remnants of a membranous sac) in fetus no. 73697-07 in the 30 mg/kg/day group. Because these findings were noted in single fetuses or litters and/or did not occur in a dose-related manner, they were not considered

^{* =} Statistically significant at p<0.05 compared to the control group.

^{** =} Statistically significant at p<0.01 compared to the control group.

WIL-402010 Naphthenic Acid American Petroleum Institute

to be test substance-related. No other external malformations were noted for fetuses in this study.

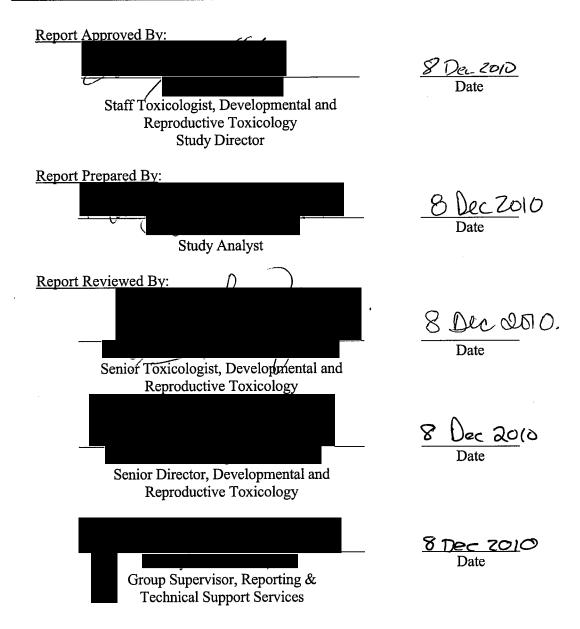
No external developmental variations were observed in fetuses in this study.

WIL-402010 Naphthenic Acid American Petroleum Institute

7. Conclusions

Clinical signs were noted in the 1000 mg/kg/day group females and consisted of hunched posture, rocking, lurching, or swaying while ambulating and prostration. substance-related lower mean body weight and reduced food consumption were noted in the 1000 mg/kg/day group males and females. Increased absolute and relative (to final body weight) kidney and liver weights and decreased mean absolute and/or relative (to final body weight) thymus weights were noted in the 1000 mg/kg/day group males and In addition, lower mean absolute ovarian weights were noted in the 1000 mg/kg/day group females. In the 300 mg/kg/day group males, a higher mean absolute and relative (to final body weight) liver weight was also observed. Two females in the 1000 mg/kg/day group had completely resorbed litters and 1 female had a litter composed of 2 dead fetuses and 1 early resorption. An increase in the mean litter proportion of pre-implantation and postimplantation loss were noted in the 1000 mg/kg/day group which corresponded to a decrease in the mean number and litter proportion of viable fetuses. No treatment-related effects on external fetal malformations and no developmental variations were noted in this study at any dosage level evaluated. Based on these results, dosage levels of 100, 300, and 900 mg/kg/day were selected for a combined repeated dose toxicity study with reproduction/developmental toxicity screening test (OECD 422) in Crl:CD(SD) rats administered naphthenic acid orally by gavage.

8. REPORT REVIEW AND APPROVAL



9. QUALITY ASSURANCE UNIT STATEMENT

9.1. PHASES INSPECTED

| Date(s) of Inspection(s) | Phase Inspected | Date(s) Findings Reported to Study Director | Date(s) Findings Reported to Management | Auditor(s) |
|---|--|---|---|------------|
| 2-Jun-2010 | Test Article Administration | 2-Jun-2010 | 21-Jul-2010 | |
| 11-Jun-2010 | Clinical Observations | 11-Jun-2010 | 21-Jul-2010 | |
| 22-Jun-2010 | Labparohysterectomy and Fetal Examination | 22-Jun-2010 | 21-Jul-2010 | |
| 13-Jul-2010 | Study Records (I-1) | 13-Jul-2010 | 20-Aug-2010 | |
| 13-Jul-2010 | Study Records (N-1, N-2) | 13-Jul-2010 | 20-Aug-2010 | |
| 14-Jul-2010 | Study Records (Rx-1) | 14-Jul-2010 | 20-Aug-2010 | |
| 24-Jul-2010, 28-Jul-2010 | Draft Report (Tables only) | 28-Jul-2010 | 20-Aug-2010 | |
| 27-Jul-2010, 28-Jul-2010, 29-Jul-2010 | Study Records (A-1, A-2, A-3) | 29-Jul-2010 | 20-Aug-2010 | |
| 10-Aug-2010 | Study Records (A-4) | 10-Aug-2010 | 23-Sep-2010 | |
| 11-Aug-2010 | Draft Report (Characterization Analyses) | 11-Aug-2010 | 23-Sep-2010 | |
| 11-Aug-2010 | Draft Report (Analytical Chemistry, Formulation Analysis) | 11-Aug-2010 | 23-Sep-2010 | |
| 13-Aug-2010 | Draft Report (Excluding Tables and Analytical Analysis Appendix) | 13-Aug-2010 | 23-Sep-2010 | |

This study was inspected in accordance with the United States EPA GLP Standards (40 CFR Part 792), the OECD Principles of GLP, WIL Research's SOPs, and the Sponsor's protocol. Quality Assurance findings, derived from the inspections during the conduct of the study and from the inspections of the raw data and draft report, are documented and have been reported to the Study Director. Review of the protocol as

WIL-402010 Naphthenic Acid American Petroleum Institute

well as a yearly internal facility inspection are conducted by the Quality Assurance Unit at WIL Research. A status report is submitted to management monthly.

This report accurately reflects the data generated during the study. The methods and procedures used in the study were those specified in the protocol and WIL Research's SOPs.

9.2. APPROVAL

This study was inspected according to the criteria discussed in Section 9.1.

Report Audited By:



Report Released By:



American Petroleum Institute

10. REFERENCES

Dunn, O.J. Multiple comparisons using rank sums. *Technometrics* **1964**, *6*(3), 241-252.

Dunnett, C.W. New tables for multiple comparisons with a control. *Biometrics* **1964**, *20*, 482-491.

Kruskal, W.H.; Wallis, W.A. Use of ranks in one-criterion variance analysis. *Journal of the American Statistical Association* **1952**, *47*, 583-621.

National Research Council. *Guide for the Care and Use of Laboratory Animals*, Institute of Laboratory Animal Resources, Commission on Life Sciences; National Academy Press: Washington, DC, **1996**.

Snedecor, G.W.; Cochran, W.G. One-Way Classifications; Analysis of Variance. In *Statistical Methods*, 7th ed.; The Iowa State University Press: Ames, IA, **1980**; pp 215-237.

WIL-402010 Naphthenic Acid American Petroleum Institute

11. DATA RETENTION

The Sponsor has title to all documentation records, raw data, specimens, or other work product generated during the performance of the study. Any remaining dosing formulation samples will be discarded upon issuance of the final report. All remaining work product generated by WIL Research, including raw paper data, pertinent electronic storage, and specimens, are retained in the Archives at WIL Research as specified in the study protocol.

Reserve samples of the test substance, pertinent electronic storage media, and the original final report are retained in the Archives at WIL Research in compliance with regulatory requirements.

WIL-402010 Naphthenic Acid American Petroleum Institute

12. ABBREVIATIONS

The following abbreviations may apply to this report:

AAALAC - Association for Assessment and Accreditation of Laboratory

Animal Care

ANOVA - analysis of variance

EPA - Environmental Protection Agency

g - gram

GLP - Good Laboratory Practices

hr - hour(s) kg - kilogram

L - liter

mg - milligram

mL - milliliter

NA - not applicable

OECD - Organisation for Economic Cooperation and Development

SOP - standard operating procedure

TSCA - Toxic Substance Control Act

WIL Research - WIL Research Laboratories, LLC

WTDMSTM - WIL Toxicology Data Management System

wt. - weight

wts. - weights

TABLES S1 - S22

Page 47 of 360

TABLE S1 (MALES - DAILY EXAMINATIONS) PROJECT NO.:WIL-402010M

ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SUMMARY OF CLINICAL FINDINGS: TOTAL OCCURRENCE/NO. OF ANIMALS SPONSOR: AMERICAN PETROLEUM

| | M A I | E | | | |
|--|-----------------------|--------------------|---------------------|--------|-------------|
| TABLE RANGE: | | 06-02-10 TO 06-16- | 10 | | |
| GROUP: | 1 | 2 | 3 | 4 | 5 |
| | | | | | |
| NORMAL OPERATOR OF THE OPERATOR | 84/5 | 84/5 | 84/5 | 75 / F | 48 / 5 |
| -NO SIGNIFICANT CLINICAL OBSERVATIONS | 74/ 5 | 74/ 5 | 74/ 5 | 75/ 5 | 47/ 5 |
| DISPOSITION | | | | | |
| -SCHEDULED EUTHANASIA | 5/ 5 | 5/ 5 | 5/ 5 | 5/ 5 | 5/ 5 |
| BODY/INTEGUMENT | | | | | |
| -WET YELLOW MATERIAL RIGHT INGUINAL AREA | 1/ 1 | 0/ 0 | 1/ 1 | 0/ 0 | 0/ 0 |
| -WET YELLOW MATERIAL LEFT INGUINAL AREA | 1/ 1 | 0/ 0 | 1/ 1 | 0/ 0 | 0/ 0 |
| -WET YELLOW MATERIAL UROGENITAL AREA | 1/ 1 | 0/ 0 | 0/ 0 | 0/ 0 | 15/ 3 |
| -WET YELLOW MATERIAL ANOGENITAL AREA | 0/ 0 | 1/ 1 | 0/ 0 | 0/ 0 | 0/0 |
| -HAIR LOSS VENTRAL ABDOMINAL AREA | 0/ 0 | 0/ 0 | 0/ 0 | 0/ 0 | 13/ 1 |
| -HAIR LOSS RIGHT HINDLIMB | 0/ 0 | 0/ 0 | 0/ 0 | 0/ 0 | 11/ 1 |
| -HAIR LOSS LEFT HINDLIMB | 0/ 0 | 0/ 0 | 0/ 0 | 0/ 0 | 11/ 1 |
| -HAIR LOSS LEFT LATERAL ABDOMINAL AREA | 0/ 0 | 0/ 0 | 0/ 0 | 0/ 0 | 1/ 1 |
| 1- 0 MG/KG/DAY 2- 30 MG/KG/DAY 3- 10 |) MG/KG/DAY 4- 30 | 0 MG/KG/DAY 5- | 1000 MG/KG/DAY | | |
| | 2 110, 110, 2111 1 30 | 0 110, 110, 2111 0 | 1000 110, 110, DIII | | DCCII++4 07 |

PCSUv4.07 07/15/2010

Page

48 of 360

TABLE S2 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM SUMMARY OF POST-DOSE FINDINGS: TOTAL OCCURRENCE/NO. OF ANIMALS

---- M A L E ----______ TABLE RANGE: 06-02-10 TO 06-15-10 GROUP: 1 2 3 4 5 NORMAL 1 HOUR POST-DOSING -NO SIGNIFICANT CLINICAL OBSERVATIONS 70/5 70/5 69/5 55/5 27/5 BODY/INTEGUMENT 1 HOUR POST-DOSING -WET CLEAR MATERIAL VENTRAL NECK 0/0 0/0 0/0 0/0 6/3 -WET CLEAR MATERIAL RIGHT FORELIMB 0/0 0/0 0/0 0/0 8/4 0/0 -WET CLEAR MATERIAL LEFT FORELIMB 0/0 0/0 0/0 9/4 CARDIO-PULMONARY TIME OF DOSE 0/0 0/0 0/0 0/0 1/1 -GASPING EYES/EARS/NOSE 1 HOUR POST-DOSING -WET CLEAR MATERIAL AROUND NOSE 0/0 0/0 0/0 0/0 11/5 -DRIED RED MATERIAL AROUND NOSE 0/0 0/0 0/0 0/0 1/1 ORAL/DENTAL TIME OF DOSE -SALIVATION PRIOR TO DOSING 0/0 0/0 2/1 0/0 14/5 1 HOUR POST-DOSING 1/1 -WET CLEAR MATERIAL AROUND MOUTH 0/0 0/0 11/2 36/5 -DRIED CLEAR MATERIAL AROUND MOUTH 0/0 0/0 0/0 4/1 6/3 -DRIED RED MATERIAL AROUND MOUTH 0/0 0/0 0/0 0/0 0/0 1/1 -SALIVATION 0/0 0/0 0/0 2/2 _______

1- 0 MG/KG/DAY 2- 30 MG/KG/DAY 3- 100 MG/KG/DAY 4- 300 MG/KG/DAY 5- 1000 MG/KG/DAY

PPDTSUv1.46 07/15/2010

PAGE 1

| | | MALI | ES | | |
|--------------|-------------|--------------|---------------|---------------|----------------|
| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
| DAY 0 | | | | | |
| MEAN | 315. | 317. | 317. | 316. | 318. |
| % DIFFERENCE | | 0.6 | 0.6 | 0.3 | 1.0 |
| S.D. | 11.9 | 11.8 | 13.9 | 19.0 | 26.2 |
| S.E. | 5.3 | 5.3 | 6.2 | 8.5 | 11.7 |
| N | 5 | 5 | 5 | 5 | 5 |
| DAY 7 | | | | | |
| MEAN | 359. | 359. | 354. | 360. | 341. |
| % DIFFERENCE | | 0.0 | -1.4 | 0.3 | -5.0 |
| S.D. | 15.3 | 14.2 | 17.6 | 27.1 | 23.1 |
| S.E. | 6.9 | 6.3 | 7.9 | 12.1 | 10.3 |
| N | 5 | 5 | 5 | 5 | 5 |
| DAY 14 | | | | | |
| MEAN | 406. | 398. | 396. | 399. | 369. |
| % DIFFERENCE | | -2.0 | -2.5 | -1.7 | -9.1 |
| S.D. | 20.6 | 17.5 | 22.8 | 37.3 | 35.8 |
| S.E. | 9.2 | 7.8 | 10.2 | 16.7 | 16.0 |
| N | 5 | 5 | 5 | 5 | 5 |

None significantly different from control group

PROJECT NO.:WIL-402010M

Page 49 of 360

SPONSOR: AMERICAN PETROLEUM

PJTBWSUv5.16 07/15/2010 TABLE S4 (MALES)

PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

SPONSOR:AMERICAN PETROLEUM SUMMARY OF BODY WEIGHT CHANGES [G]

MALES

GROUP: 0 MG/KG/DAY 30 MG/KG/DAY 100 MG/KG/DAY 300 MG/KG/DAY 1000 MG/KG/DAY

| | GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
|-----|--------|-------------|--------------|---------------|---------------|----------------|
| DAY | 0- 7 | | | | | |
| | MEAN | 43. | 41. | 37. | 43. | 24.** |
| | S.D. | 6.5 | 3.9 | 5.3 | 11.6 | 11.5 |
| | S.E. | 2.9 | 1.7 | 2.4 | 5.2 | 5.1 |
| | N. | 5 | 5 | 5 | 5 | 5 |
| DAY | 7- 14 | | | | | |
| | MEAN | 47. | 40. | 41. | 40. | 28. |
| | S.D. | 5.6 | 6.1 | 6.5 | 10.6 | 17.9 |
| | S.E. | 2.5 | 2.7 | 2.9 | 4.7 | 8.0 |
| | N | 5 | 5 | 5 | 5 | 5 |
| DAY | 0- 14 | | | | | |
| | MEAN | 90. | 81. | 79. | 83. | 52.** |
| | S.D. | 11.6 | 8.2 | 11.1 | 21.3 | 20.9 |
| | S.E. | 5.2 | 3.7 | 4.9 | 9.5 | 9.3 |
| | N | 5 | 5 | 5 | 5 | 5 |

** = Significantly different from the control group at 0.01 using Dunnett's test MEAN DIFFERENCES CALCULATED FROM INDIVIDUAL DIFFERENCES

PJTBWSUv5.16 07/15/2010

TABLE S5 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM SUMMARY OF FOOD CONSUMPTION [G/ANIMAL/DAY]

______ MALES GROUP: 0 MG/KG/DAY 30 MG/KG/DAY 100 MG/KG/DAY 300 MG/KG/DAY 1000 MG/KG/DAY DAY 0- 7 20. 1.4 0.6 5 20. 1.7 0.7 20. 2.1 0.9 5 20. 2.4 1.1 17.* MEAN S.D. 2.0 S.E. 0.9 5 N 5 5 DAY 7- 14 20. 1.9 0.8 18. 3.1 19. 2.2 18. 2.0 MEAN 16. 2.4 S.D. 1.0 0.9 S.E. 1.4 1.1 5 5 5 5 N 5 DAY 0- 14 20. 1.7 MEAN 20. 19. 19. 16. 1.7 S.D. 2.1 2.7 2.0 0.8 S.E. 1.0 0.8 1.2 0.9 5 5 5

 \star = Significantly different from the control group at 0.05 using Dunnett's test

PJTFWSUv5.18 07/15/2010

Page 52 of 360

TABLE S6 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SUMMARY OF FOOD CONSUMPTION [G/KG/DAY]

______ MALES GROUP: 0 MG/KG/DAY 30 MG/KG/DAY 100 MG/KG/DAY 300 MG/KG/DAY 1000 MG/KG/DAY DAY 0- 7 60. 2.8 1.3 60. 3.0 1.3 59. 4.8 2.1 59. 4.6 2.1 5 50.* MEAN S.D. 6.8 S.E. 3.1 N 5 5 5 5 DAY 7- 14 52. 3.7 1.6 49. 3.1 1.4 5 50. 3.5 MEAN 48. 5.0 46. S.D. 4.4 S.E. 1.6 2.2 2.0 5 5 N 5 5 DAY 0- 14 56. 3.1 MEAN 55. 54. 53. 48. S.D. 4.0 2.8 4.9 4.9 S.E. 1.8 1.4 1.3 2.2 2.2 5 5 5

 \star = Significantly different from the control group at 0.05 using Dunnett's test

PJTFWSUv5.18 07/15/2010

Page 53 of 360

PROJECT NO.:WIL-402010M

TABLE S7 (MALES) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

| PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM SUMMARY OF MACROSCOPIC FINDINGS | | | | | | | |
|---|--------------------------------|-------------|--------------|--------|---|----------|------|
| | SCHEDULED NE | ECROPSY | | | | | |
| | GROUP: | 1 | M A I | E | 4 | 5 | |
| NUMBER OF ANIMALS EXAMINED | | 5 | 5 | 5 | 5 | 5 | |
| KIDNEYS -PALE | | 0 | 1 | 0 | 1 | 1 | |
| SKIN -HAIR LOSS | | 0 | 0 | 0 | 0 | 1 | |
| NO SIGNIFICANT CHANGES OBSERVED | - ALL EXAMINED TISSUES | 5 | 4 | 5 | 4 | 4 | |
| 1- 0 MG/KG/DAY 2- 30 MG/ | KG/DAY 3- 100 MG/KG/DAY 4- 300 |) MG/KG/DAY | 5- 1000 MG/F | KG/DAY | | PGRSI2v4 | 4.08 |

07/16/2010

TABLE S8 (MALES)

PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

SPONSOR:AMERICAN PETROLEUM SUMMARY OF ORGAN WEIGHTS AND ORGAN WTS. RELATIVE TO BODY WTS.

MALES GROUP: 0 MG/KG/DAY 30 MG/KG/DAY 100 MG/KG/DAY 300 MG/KG/DAY 1000 MG/KG/DAY FINAL BODY WT (G) MEAN 406. 398. 396. 400. 369. % DIFFERENCE -2.0 -2.5 -1.5 -9.1 S.D. 20.7 17.4 22.8 37.1 35.7 7.8 S.E. 9.3 10.2 16.6 16.0 N 5 5 5 5 5 KIDNEYS (G) MEAN 3.19 3.25 3.24 3.38 3.68 % DIFFERENCE 1.9 1.6 6.0 15.4 0.348 0.155 S.D. 0.365 0.319 0.266 0.485 0.142 0.119 S.E. 0.163 0.217 N 5 5 5 5 KIDNEYS (G/100 G FINAL BODY WEIGHT) MEAN 0.786 0.817 0.820 0.845 0.999** 0.0727 S.D. 0.0499 0.0768 0.0227 0.1265 0.0343 0.0325 0.0223 0.0101 S.E. 0.0566 5 5 5 5 LIVER (G) MEAN 18.40 18.10 17.97 20.78 23.87** % DIFFERENCE -2.3 12.9 29.7 -1.6 2.267 0.710 S.D. 1.170 2.886 2.575 1.014 S.E. 0.523 0.317 1.291 1.152 5 5

^{** =} Significantly different from the control group at 0.01 using Dunnett's test

TABLE S8 (MALES)
ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM SUMMARY OF ORGAN WEIGHTS AND ORGAN WTS. RELATIVE TO BODY WTS.

PAGE 2

| | | MALI | ES | | |
|---------------------------|------------------|--------------|---------------|---------------|----------------|
| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
| LIVER (G/100 G FINAL BOD) | Y WEIGHT) | | | | |
| MEAN | 4.525 | 4.545 | 4.548 | 5.182** | 6.465** |
| S.D. | 0.3517 | 0.2271 | 0.1694 | 0.2672 | 0.3077 |
| S.E. | 0.1573 | 0.1016 | 0.0757 | 0.1195 | 0.1376 |
| N | 5 | 5 | 5 | 5 | 5 |
| TESTIS, LEFT (G) | | | | | |
| MEAN | 1.71 | 1.80 | 1.75 | 1.69 | 1.72 |
| % DIFFERENCE | | 5.3 | 2.3 | -1.2 | 0.6 |
| S.D. | 0.147 | 0.136 | 0.210 | 0.183 | 0.222 |
| S.E. | 0.066 | 0.061 | 0.094 | 0.082 | 0.099 |
| N | 5 | 5 | 5 | 5 | 5 |
| TESTIS, LEFT (G/100 G FI | NAL BODY WEIGHT) | | | | |
| MEAN | 0.422 | 0.451 | 0.442 | 0.423 | 0.474 |
| S.D. | 0.0313 | 0.0332 | 0.0431 | 0.0298 | 0.0974 |
| S.E. | 0.0140 | 0.0149 | 0.0193 | 0.0133 | 0.0436 |
| N | 5 | 5 | 5 | 5 | 5 |
| TESTIS, RIGHT (G) | | | | | |
| MEAN | 1.74 | 1.83 | 1.71 | 1.71 | 1.75 |
| % DIFFERENCE | | 5.2 | -1.7 | -1.7 | 0.6 |
| S.D. | 0.162 | 0.131 | 0.217 | 0.154 | 0.201 |
| S.E. | 0.072 | 0.058 | 0.097 | 0.069 | 0.090 |
| N | 5 | 5 | 5 | 5 | 5 |

** = Significantly different from the control group at 0.01 using Dunnett's test

TABLE S8 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR: AMERICAN PETROLEUM SUMMARY OF ORGAN WEIGHTS AND ORGAN WTS. RELATIVE TO BODY WTS.

MALES GROUP: 0 MG/KG/DAY 30 MG/KG/DAY 100 MG/KG/DAY 300 MG/KG/DAY 1000 MG/KG/DAY TESTIS, RIGHT (G/100 G FINAL BODY WEIGHT) 0.429 0.429 MEAN 0.460 0.431 0.481 0.0456 0.0204 0.0341 S.D. 0.0401 0.0316 0.0886 0.0179 5 0.0153 0.0396 S.E. 0.0141 N 5 5 5 5 THYMUS (G) 0.6622 MEAN 0.5351 0.6467 0.6290 0.3974 % DIFFERENCE 17.5 20.9 23.8 -25.7 0.07683 0.03436 0.15073 0.13310 S.D. 0.18433 0.13331 0.06741 S.E. 0.03436 0.05952 0.08244 0.05962 5 5 N 5 5 5 THYMUS (G/100 G FINAL BODY WEIGHT) MEAN 0.131 0.163 0.159 0.164 0.108 S.D. 0.0320 0.0267 0.0368 0.0377 0.0386 0.0143 S.E. 0.0119 0.0165 0.0169 0.0173 N 5 5 5 5

None significantly different from control group

POFBSTv5.20 07/16/2010

TABLE S9
ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010F

PAGE 1 SUMMARY OF MATERNAL SURVIVAL AND PREGNANCY STATUS SPONSOR: AMERICAN PETROLEUM

| DOSE GROUP : | 1 | | 2 | | 3 | 3 | | 4 | į | 5 |
|--------------------------------|------------|---------|-------|------------|--------|-----------|------|-------|-----|---------------|
| | NO. | % | NO. | % | NO. | % | NO. | % | NO. | % |
| FEMALES ON STUDY | 5 | | 5 | | 5 | | 5 | | 5 | |
| FEMALES THAT ABORTED | | | | | | | | | | |
| OR DELIVERED | 0 0.0 | 0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 |
| FEMALES THAT DIED | 0 0.0 | 0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 |
| FEMALES THAT ABORTED | 0 0.0 | 0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 |
| NONGRAVID | 0 0.0 | 0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 |
| GRAVID | 0 0.0 | 0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 |
| FEMALES THAT WERE EUTHANIZED | 0 0.0 | 0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 |
| NONGRAVID | 0 0.0 | 0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 |
| GRAVID | 0 0.0 | 0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 | 0 | 0.0 |
| FEMALES EXAMINED AT | | | | | | | | | | |
| SCHEDULED NECROPSY | 5 100.0 | n | 5 1 | 00.0 | 5 - | .00.0 | 5 | 100.0 | 5 - | 100.0 |
| NONGRAVID | 0 0.0 | | | 0.0 | | 0.0 | | 0.0 | | 0.0 |
| GRAVID | 5 100.0 | | | 00.0 | | 100.0 | | 100.0 | | 100.0 |
| WITH RESORPTIONS ONLY | 0 0.0 | | | 0.0 | | 0.0 | | 0.0 | | 60.0 |
| WITH VIABLE FETUSES | 5 100.0 | | | 00.0 | | 100.0 | | 100.0 | | 40.0 |
| TOTAL FEMALES GRAVID | 5 100.0 | 0 | 5 1 | 00.0 | 5 3 | 100.0 | 5 | 100.0 | 5 3 | 100.0 |
| 1- 0 MG/KG/DAY 2- 30 MG/KG/DAY | 3- 100 MG, | /KG/DAY | 4- 30 | 0 MG/KG/DA | Y 5- 1 | 000 MG/KG | /DAY | | | -DC++4 01 |

PSPSv4.01 07/15/2010

TABLE S10 (FEMALES - DAILY EXAMINATIONS) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM SUMMARY OF CLINICAL FINDINGS: TOTAL OCCURRENCE/NO. OF ANIMALS

PAGE 1

---- F E M A L E ----______ 06-01-10 TO 06-24-10 2 TABLE RANGE: 1 GROUP: 3 5 -NO SIGNIFICANT CLINICAL OBSERVATIONS 99/5 57/5 53/3 80/5 35/5 DISPOSITION -SCHEDULED EUTHANASIA; GESTATION DAY 20 5/5 5/5 5/5 5/5 BEHAVIOR/CNS 0/0 0/0 0/0 0/0 2/1 -HUNCHED POSTURE BODY/INTEGUMENT 6/ 1 35/ 3 47/ 4 5/ 1 0/ 0 6/ 1 35/ 3 43/ 4 1/ 1 0/ 0 0/ 0 1/ 1 8/ 1 15/ 1 30/ 2 0/ 0 1/ 1 19/ 2 6/ 1 32/ 3 0/ 0 1/ 1 1/ 1 0/ 0 6/ 2 0/ 0 0/ 0 0/ 0 0/ 0 3/ 1 1/ 1 0/ 0 0/ 0 0/ 0 0/ 0 2/ 1 9/ 2 -HAIR LOSS RIGHT FORELIMB -HAIR LOSS LEFT FORELIMB -HAIR LOSS RIGHT HINDLIMB -HAIR LOSS VENTRAL ABDOMINAL AREA -WET YELLOW MATERIAL ANOGENITAL AREA 2/ 1 -HAIR LOSS VENTRAL NECK -WET YELLOW MATERIAL VENTRAL ABDOMINAL AREA -WET YELLOW MATERIAL UROGENITAL AREA 0/0 1/1 1/1 3/1 13/4
-WET YELLOW MATERIAL RIGHT INGUINAL AREA 0/0 0/0 1/1 0/0 5/1
-WET YELLOW MATERIAL LEFT INGUINAL AREA 0/0 0/0 1/1 0/0 5/2
-HAIR LOSS LEFT HINDLIMB 0/0 6/1 17/1 3/2 22/2

______ 1- 0 MG/KG/DAY 2- 30 MG/KG/DAY 3- 100 MG/KG/DAY 4- 300 MG/KG/DAY 5- 1000 MG/KG/DAY

59

EYES/EARS/NOSE

EXCRETA

-DRIED RED MATERIAL AROUND RIGHT EYE -DRIED RED MATERIAL AROUND LEFT EYE

-DRIED RED MATERIAL AROUND NOSE

-DECREASED DEFECATION

TABLE S10 (FEMALES - DAILY EXAMINATIONS) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM SUMMARY OF CLINICAL FINDINGS: TOTAL OCCURRENCE/NO. OF ANIMALS

---- F E M A L E ----

PAGE 2

06-01-10 TO 06-24-10 2 TABLE RANGE: 1 GROUP: 3 4 5 BODY/INTEGUMENT 5/ 1 3/ 1 21/ 2 3/ 1 9/ 1 -WET YELLOW MATERIAL RIGHT HINDLIMB -WET YELLOW MATERIAL LEFT HINDLIMB -HAIR LOSS RUMP -HAIR LOSS LEFT LATERAL ABDOMINAL AREA -DRIED RED MATERIAL VENTRAL ABDOMINAL AREA

 0/ 0
 0/ 0
 0/ 0

 0/ 0
 0/ 0
 0/ 0

 0/ 0
 0/ 0
 0/ 0

 0/ 0
 0/ 0
 0/ 0

 0/ 0
 0/ 0
 3/ 1
 1/ 1

 0/ 0
 0/ 0
 4/ 1
 0/ 0

 0/ 0
 0/ 0
 2/ 1
 0/ 0

 0/ 0
 0/ 0
 5/ 1
 0/ 0

 0/ 0
 0/ 0
 5/ 1
 5/ 1

 0/ 0
 0/ 0
 5/ 1
 0/ 0

 7/ 2 1/ 1 -DRIED RED MATERIAL UROGENITAL AREA -DRIED RED MATERIAL RIGHT INGUINAL AREA 7/ 1 3/ 2 -DRIED RED MATERIAL ANOGENITAL AREA -HAIR LOSS VENTRAL THORACIC AREA -SCABBING VENTRAL ABDOMINAL AREA 0/0 0/0 -SCABBING LEFT HINDLIMB 5/2 -HAIR LOSS UROGENITAL AREA -HAIR LOSS BASE OF TAIL -HAIR LOSS ANOGENITAL AREA

0/0 0/0 0/0 0/0 0/0 2/1 0/0 0/0 0/0 0/0 0/0 3/1 0/0 0/0 0/0 1/1 7/3

0/0 0/0 0/0 0/0 8/2

______ 1- 0 MG/KG/DAY 2- 30 MG/KG/DAY 3- 100 MG/KG/DAY 4- 300 MG/KG/DAY 5- 1000 MG/KG/DAY

| Page | |
|----------|--|
| 9 | |
| 0 | |
| of | |
| ω | |
| 9 | |
| 0 | |

-SALIVATION

TABLE S10 (FEMALES - DAILY EXAMINATIONS) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

1- 0 MG/KG/DAY 2- 30 MG/KG/DAY 3- 100 MG/KG/DAY 4- 300 MG/KG/DAY 5- 1000 MG/KG/DAY

SPONSOR: AMERICAN PETROLEUM SUMMARY OF CLINICAL FINDINGS: TOTAL OCCURRENCE/NO. OF ANIMALS ---- F E M A L E -----TABLE RANGE: 06-01-10 TO 06-24-10 GROUP: 1 2 TABLE RANGE: 3 5 4 BODY/INTEG II 7/ 1 0/ 0 3/ 1 5/ 1 3/2 -HAIR LOSS LEFT INGUINAL AREA -HAIR LOSS RIGHT INGUINAL AREA ORAL/DENTAL -DRIED RED MATERIAL AROUND MOUTH

> PCSUv4.07 07/28/2010

| | | | F E M A | L E | | | |
|---|--|-------------------|---------------------|-------------------|---------------------------------|--------------------------|--|
| | | : 06-02- : 1 | 10 TO 06-24-10 2 | 3 | 4 | 5 | |
| | NORMAL | | | | | | |
| | 1 HOUR POST-DOSING -NO SIGNIFICANT CLINICAL OBSERVATIONS | 100/5 | 99/5 | 88/5 | 87/5 | 53/5 | |
| | BEHAVIOR/CNS | | | | | | |
| | 1 HOUR POST-DOSING -HUNCHED POSTURE -ROCKS, LURCHES, OR SWAYS AS IT WALKS -PROSTRATE | 0/0 0/0 0/0 | 0/0 0/0 0/0 | 0/0 0/0 0/0 | 0/0 0/0 0/0 | 4/2 2/2 1/1 | |
| _ | BODY/INTEGUMENT | | | | | | |
| | 1 HOUR POST-DOSING -WET YELLOW MATERIAL ANOGENITAL AREA -WET CLEAR MATERIAL RIGHT FORELIMB -WET CLEAR MATERIAL LEFT FORELIMB | 0/0 0/0 0/0 | 0/0 0/0 0/0 | 0/0 1/1 1/1 | 1/1 0/0 0/0 | 0/0 6/3 7/4 | |
| 3 | EYES/EARS/NOSE | | | | | | |
| | -DRIED RED MATERIAL AROUND NOSE | 0/0 0/0 | 0/0 0/0 | 0/0 0/0 | 0/0 0/0 0/0 0/0 4/3 | 1/1 1/1 1/1 4/3 | |
| | BODY/INTEG II | | | | | | |
| | 1 HOUR POST-DOSING -WET CLEAR MATERIAL VENTRAL NECK | 0/0 | 0/0 | 1/1 | 2/2 | 2/1 | |

1- 0 MG/KG/DAY 2- 30 MG/KG/DAY 3- 100 MG/KG/DAY 4- 300 MG/KG/DAY 5- 1000 MG/KG/DAY

SUMMARY OF POST-DOSE FINDINGS: TOTAL OCCURRENCE/NO. OF ANIMALS

Page 61 of 360

TABLE S11 (FEMALES)

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM SUMMARY OF POST-DOSE FINDINGS: TOTAL OCCURRENCE/NO. OF ANIMALS

| TABLE RANGE: GROUP: | | O 06-24-10 2 | 3 | 4 | 5 | |
|------------------------------------|-----|---------------------|------|------|------|--|
| AL/DENTAL | | | | | | |
| TIME OF DOSE | | | | | | |
| -SALIVATION PRIOR TO DOSING | 0/0 | 0/0 | 0/0 | 0/0 | 6/2 | |
| 1 HOUR POST-DOSING | | | | | | |
| -WET CLEAR MATERIAL AROUND MOUTH | 0/0 | 0/0 | 12/4 | 11/3 | 26/5 | |
| -DRIED RED MATERIAL AROUND MOUTH | 0/0 | 1/1 | 0/0 | 1/1 | 9/4 | |
| -DRIED CLEAR MATERIAL AROUND MOUTH | 0/0 | 0/0 | 0/0 | 1/1 | 5/3 | |
| -WET RED MATERIAL AROUND MOUTH | 0/0 | 0/0 | 0/0 | 0/0 | 2/2 | |
| -SALIVATION | 0/0 | 0/0 | 0/0 | 0/0 | 2/1 | |

PPDTSUv1.46 07/15/2010

Page 63 of 360

| PAGE 1 |
|--------|
|--------|

| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
|--------------|-------------|--------------|---------------|---------------|----------------|
| DAY 0 | | | | | |
| MEAN | 253. | 248. | 262. | 256. | 260. |
| % DIFFERENCE | | -2.0 | 3.6 | 1.2 | 2.8 |
| S.D. | 9.8 | 13.4 | 19.0 | 21.1 | 15.5 |
| S.E. | 4.4 | 6.0 | 8.5 | 9.4 | 6.9 |
| N | 5 | 5 | 5 | 5 | 5 |
| DAY 6 | | | | | |
| MEAN | 272. | 267. | 281. | 268. | 257. |
| % DIFFERENCE | | -1.8 | 3.3 | -1.5 | -5.5 |
| S.D. | 13.2 | 14.6 | 22.4 | 23.4 | 23.5 |
| S.E. | 5.9 | 6.5 | 10.0 | 10.5 | 10.5 |
| N | 5 | 5 | 5 | 5 | 5 |
| DAY 7 | | | | | |
| MEAN | 275. | 269. | 284. | 272. | 264. |
| % DIFFERENCE | | -2.2 | 3.3 | -1.1 | -4.0 |
| S.D. | 10.8 | 13.6 | 15.8 | 19.6 | 23.0 |
| S.E. | 4.8 | 6.1 | 7.0 | 8.8 | 10.3 |
| N | 5 | 5 | 5 | 5 | 5 |
| DAY 10 | | | | | |
| MEAN | 287. | 285. | 298. | 290. | 278. |
| % DIFFERENCE | | -0.7 | 3.8 | 1.0 | -3.1 |
| S.D. | 9.6 | 15.7 | 18.2 | 16.7 | 18.3 |
| S.E. | 4.3 | 7.0 | 8.1 | 7.5 | 8.2 |
| N | 5 | 5 | 5 | 5 | 5 |
| DAY 14 | | | | | |
| MEAN | 303. | 303. | 314. | 304. | 290. |
| % DIFFERENCE | | 0.0 | 3.6 | 0.3 | -4.3 |
| S.D. | 13.3 | 15.6 | 17.2 | 21.0 | 29.1 |
| S.E. | 5.9 | 7.0 | 7.7 | 9.4 | 13.0 |
| N | 5 | 5 | 5 | 5 | 5 |

None significantly different from control group NONGRAVID WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN SPONSOR: AMERICAN PETROLEUM

TABLE S12
PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 2

SUMMARY OF BODY WEIGHTS DURING GESTATION [G]

| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
|--------------|-------------|--------------|---------------|---------------|----------------|
| DAY 17 | | | | | |
| MEAN | 338. | 339. | 347. | 335. | 304. |
| % DIFFERENCE | | 0.3 | 2.7 | -0.9 | -10.1 |
| S.D. | 16.9 | 21.5 | 24.9 | 23.9 | 19.4 |
| S.E. | 7.6 | 9.6 | 11.1 | 10.7 | 8.7 |
| N | 5 | 5 | 5 | 5 | 5 |
| DAY 20 | | | | | |
| MEAN | 379. | 385. | 392. | 373. | 320.** |
| % DIFFERENCE | | 1.6 | 3.4 | -1.6 | -15.6 |
| S.D. | 20.6 | 24.9 | 29.5 | 26.3 | 31.5 |
| S.E. | 9.2 | 11.1 | 13.2 | 11.7 | 14.1 |
| N | 5 | 5 | 5 | 5 | 5 |
| | | | | | |

** = Significantly different from the control group at 0.01 using Dunnett's test NONGRAVID WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

PGBWSUv5.10 07/15/2010 Page 65 of 360

PAGE 1

| | GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY | |
|-----|-------------------------------------|------------------------|-------------------------|-------------------------|-------------------------|-------------------------|--|
| DAY | 0- 6 MEAN S.D. S.E. N | 18. 4.7 2.1 5 | 19. 8.8 4.0 5 | 19. 17.2 7.7 5 | 12. 3.0 1.3 5 | -2. 20.0 8.9 5 | |
| DAY | 6- 7 MEAN S.D. S.E. N | 3. 4.4 2.0 5 | 2. 5.6 2.5 5 | 4. 8.2 3.7 5 | 4. 7.7 3.4 5 | 7. 10.1 4.5 5 | |
| DAY | 7- 10 MEAN S.D. S.E. N | 12. 8.3 3.7 5 | 17. 5.4 2.4 5 | 13. 7.1 3.2 5 | 18. 5.2 2.3 5 | 14. 7.1 3.2 5 | |
| DAY | 10- 14 MEAN S.D. S.E. N | 16. 7.5 3.4 5 | 17. 4.9 2.2 5 | 16. 5.9 2.7 5 | 15. 8.4 3.8 5 | 12. 12.0 5.4 5 | |
| DAY | 14- 17 MEAN S.D. S.E. N | 34. 6.1 2.7 5 | 37. 10.5 4.7 5 | 33. 9.3 4.2 5 | 31. 12.1 5.4 5 | 14. 19.4 8.7 5 | |

None significantly different from control group MEAN DIFFERENCES CALCULATED FROM INDIVIDUAL DIFFERENCES NONGRAVID WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

PROJECT NO.:WIL-402010F SPONSOR:AMERICAN PETROLEUM

| | GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY | |
|-----|-------------------------------------|--------------------------|--------------------------|---------------------------|--------------------------|----------------------------|--|
| DAY | 17- 20 MEAN S.D. S.E. N | 41. 6.0 2.7 5 | 46. 8.9 4.0 5 | 46. 9.2 4.1 5 | 38. 6.8 3.0 5 | 16.** 18.0 8.1 5 | |
| DAY | 0- 20 MEAN S.D. S.E. N | 126. 15.2 6.8 5 | 137. 17.3 7.7 5 | 131. 23.7 10.6 5 | 117. 11.4 5.1 5 | 60.** 36.2 16.2 5 | |

** = Significantly different from the control group at 0.01 using Dunnett's test MEAN DIFFERENCES CALCULATED FROM INDIVIDUAL DIFFERENCES
NONGRAVID WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

PGBWSUv5.10 07/15/2010 TABLE S14
PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 1

| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
|---------------------|-------------|--------------|---------------|---------------|----------------|
| INITIAL BODY WT. | | | | | |
| MEAN | 253. | 248. | 262. | 256. | 260. |
| S.D. | 9.8 | 13.4 | 19.0 | 21.1 | 15.5 |
| S.E. | 4.4 | 6.0 | 8.5 | 9.4 | 6.9 |
| N | 5 | 5 | 5 | 5 | 5 |
| TERMINAL BODY WT. | | | | | |
| MEAN | 379. | 385. | 392. | 373. | 320.** |
| S.D. | 20.6 | 24.9 | 29.5 | 26.3 | 31.5 |
| S.E. | 9.2 | 11.1 | 13.2 | 11.7 | 14.1 |
| N | 5 | 5 | 5 | 5 | 5 |
| GRAVID UTERINE WT. | | | | | |
| MEAN | 74.7 | 78.6 | 82.4 | 74.7 | 28.1** |
| S.D. | 7.61 | 8.55 | 11.00 | 9.23 | 35.71 |
| S.E. | 3.40 | 3.82 | 4.92 | 4.13 | 15.97 |
| N | 5 | 5 | 5 | 5 | 5 |
| JET BODY WT. | | | | | |
| MEAN | 304.1 | 306.6 | 310.0 | 298.5 | 291.5 |
| S.D. | 20.15 | 20.32 | 18.63 | 20.48 | 19.72 |
| S.E. | 9.01 | 9.09 | 8.33 | 9.16 | 8.82 |
| N | 5 | 5 | 5 | 5 | 5 |
| NET BODY WT. CHANGE | | | | | |
| MEAN | 50.9 | 58.2 | 48.4 | 42.1 | 31.9 |
| S.D. | 12.89 | 15.93 | 16.87 | 8.61 | 8.90 |
| S.E. | 5.77 | 7.13 | 7.54 | 3.85 | 3.98 |
| N | 5 | 5 | 5 | 5 | 5 |

** = Significantly different from the control group at 0.01 using Dunnett's test

PUTSUv5.07 07/15/2010 Page 68 of 360

| | | GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
|-----|-----|---------------------------------|------------------------|------------------------|------------------------|------------------------|------------------------|
| DAY | 0 - | 6 MEAN S.D. S.E. N | 14. 2.1 0.9 5 | 14. 2.2 1.0 5 | 14. 2.6 1.2 5 | 12. 1.9 0.9 5 | 9.* 3.2 1.4 5 |
| DAY | 6- | 7 MEAN S.D. S.E. N | 14. 1.8 0.8 5 | 14. 6.2 2.8 5 | 14. 2.3 1.0 5 | 14. 2.9 1.3 5 | 12. 3.9 1.7 5 |
| DAY | 7- | 10 MEAN S.D. S.E. N | 13. 1.5 0.7 5 | 16. 1.3 0.6 5 | 15. 1.9 0.9 5 | 15. 0.8 0.4 5 | 15. 2.9 1.3 5 |
| DAY | 10- | 14 MEAN S.D. S.E. N | 14. 2.9 1.3 5 | 17. 0.5 0.2 5 | 15. 1.9 0.8 5 | 14. 2.3 1.0 5 | 14. 2.9 1.5 |
| DAY | 14- | 17 MEAN S.D. S.E. N | 18. 2.2 1.0 5 | 18. 2.5 1.1 5 | 18. 2.4 1.1 5 | 17. 3.2 1.4 5 | 16. 1.5 0.7 5 |

 $^{^\}star$ = Significantly different from the control group at 0.05 using Dunnett's test NONGRAVID WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

TABLE S15

ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS
SUMMARY OF FOOD CONSUMPTION DURING GESTATION [G/ANIMAL/DAY]

| | GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY | |
|-----|-------------------------------------|-------------|------------------------|------------------------|------------------------|------------------------|--|
| DAY | 17- 20 MEAN S.D. S.E. N | 2.9 | 20. 1.5 0.7 5 | 18. 2.6 1.2 5 | 15. 1.8 0.8 5 | 16. 2.6 1.3 | |
| DAY | 0- 20 MEAN S.D. S.E. N | 1.5 | 17. 1.1 0.5 5 | 16. 1.8 0.8 5 | 14. 1.3 0.6 5 | 14. 1.0 0.5 4 | |

None significantly different from control group NONGRAVID WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

PGFWSUv5.16 07/16/2010

PAGE 2

Page 69 of 360

Page 70 of 360

| | GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
|-----|-------------------------------------|------------------------|--------------------------|------------------------|-------------------------|--------------------------|
| DAY | 0- 6 MEAN S.D. S.E. N | 51. 6.3 2.8 5 | 54. 7.3 3.3 5 | 53. 7.3 3.3 5 | 45. 4.2 1.9 5 | 35.* 11.0 4.9 5 |
| DAY | 6- 7 MEAN S.D. S.E. N | 50. 7.7 3.4 5 | 53. 22.3 10.0 5 | 51. 9.8 4.4 5 | 54. 12.1 5.4 5 | 45. 11.6 5.2 5 |
| DAY | 7- 10 MEAN S.D. S.E. N | 46. 5.4 2.4 5 | 56. 2.9 1.3 | 51. 6.5 2.9 5 | 53. 5.3 2.4 5 | 57. 12.3 5.5 5 |
| DAY | 10- 14 MEAN S.D. S.E. N | 49. 8.9 4.0 5 | 57. 3.9 1.7 5 | 49. 4.4 2.0 | 48. 6.7 3.0 5 | 50. 6.0 3.0 4 |
| DAY | 14- 17 MEAN S.D. S.E. N | 55. 4.8 2.1 5 | 57. 6.8 3.0 5 | 54. 4.3 1.9 5 | 53. 9.3 4.2 5 | 54. 9.4 4.2 5 |

 $^{^\}star$ = Significantly different from the control group at 0.05 using Dunnett's test NONGRAVID WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

| | GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY | |
|-----|-------------------------------------|------------------------|-------------------------|------------------------|------------------------|------------------------|--|
| DAY | 17- 20 MEAN S.D. S.E. N | 49. 6.2 2.8 5 | 56. 3.3 1.5 5 | 47. 5.0 2.2 5 | 43. 4.2 1.9 5 | 50. 5.9 2.9 4 | |
| DAY | 0- 20 MEAN S.D. S.E. N | 49. 3.6 1.6 5 | 55.* 3.2 1.4 5 | 50. 3.1 1.4 5 | 47. 3.5 1.6 5 | 48. 2.5 1.3 4 | |

 * = Significantly different from the control group at 0.05 using Dunnett's test NONGRAVID WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

PGFWSUv5.16 07/16/2010

| PROJECT NO.:WIL-402010F ORA SPONSOR:AMERICAN PETROLEUM | TABLE : AL 14-DAY/PILOT DEV TOX STUDY (SUMMARY OF MATERNAL MA | OF NAPHTHENI | | ATS | PAGE 1 |
|--|--|--------------|------------|-----------|--------------------------|
| | GROUP : | 1 2 | 3 4 | 5 | |
| NUMBER EXAMINED | | 5 5 | 5 5 | 5 | |
| NO SIGNIFICANT CHANGES OBSERVED GRAVID AMMONIUM SULFIDE POSITIV | 3 | 5 5 0 0 | 5 5 0 0 | 3 2 | |
| 1- 0 MG/KG/DAY 2- 30 MG/KG/DA | Y 3- 100 MG/KG/DAY 4- 30 |) MG/KG/DAY | 5- 1000 N | IG/KG/DAY | PMGSIv4.04 07/15/2010 |

SPONSOR: AMERICAN PETROLEUM

TABLE S18 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 1 SUMMARY OF ORGAN WEIGHTS AND ORGAN WTS. RELATIVE TO BODY WTS.

| | | FEMA: | LES | | |
|-----------------------|--------------|--------------|---------------|---------------|----------------|
| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
| INAL BODY WT (G) | | | | | |
| MEAN | 379. | 385. | 392. | 373. | 320.** |
| % DIFFERENCE | | 1.6 | 3.4 | -1.6 | -15.6 |
| S.D. | 20.6 | 24.9 | 29.5 | 26.3 | 31.5 |
| S.E. | 9.2 | 11.1 | 13.2 | 11.7 | 14.1 |
| N | 5 | 5 | 5 | 5 | 5 |
| IDNEYS (G) | | | | | |
| MEAN | 1.84 | 1.97 | 1.87 | 1.96 | 2.09 |
| % DIFFERENCE | | 7.1 | 1.6 | 6.5 | 13.6 |
| S.D. | 0.200 | 0.189 | 0.194 | 0.168 | 0.214 |
| S.E. | 0.090 | 0.085 | 0.087 | 0.075 | 0.096 |
| N | 5 | 5 | 5 | 5 | 5 |
| IDNEYS (G/100 G FINAL | BODY WEIGHT) | | | | |
| MEAN | 0.484 | 0.511 | 0.475 | 0.525 | 0.661** |
| S.D. | 0.0287 | 0.0319 | 0.0262 | 0.0216 | 0.1082 |
| S.E. | 0.0128 | 0.0143 | 0.0117 | 0.0097 | 0.0484 |
| N | 5 | 5 | 5 | 5 | 5 |
| IVER (G) | | | | | |
| MEAN | 14.82 | 15.49 | 16.18 | 16.01 | 20.11** |
| % DIFFERENCE | | 4.5 | 9.2 | 8.0 | 35.7 |
| S.D. | 1.446 | 1.206 | 1.266 | 1.111 | 1.039 |
| S.E. | 0.646 | 0.539 | 0.566 | 0.497 | 0.465 |
| N | 5 | 5 | 5 | 5 | 5 |

** = Significantly different from the control group at 0.01 using Dunnett's test NONGRAVID WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

SPONSOR: AMERICAN PETROLEUM

TABLE S18

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 2

SUMMARY OF ORGAN WEIGHTS AND ORGAN WTS. RELATIVE TO BODY WTS.

| FEMALES | | | | | | | | |
|-----------------------|----------------------|--------------|---------------|---------------|----------------|--|--|--|
| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY | | | |
| VER (G/100 G FINAL B | ODY WEIGHT) | | | | | | | |
| MEAN | 3.906 | 4.020 | 4.123 | 4.290 | 6.328** | | | |
| S.D. | 0.2509 | 0.1504 | 0.0887 | 0.1202 | 0.5218 | | | |
| S.E. | 0.1122 | 0.0673 | 0.0397 | 0.0537 | 0.2334 | | | |
| N | 5 | 5 | 5 | 5 | 5 | | | |
| ARY/OD, LEFT (G) | | | | | | | | |
| MEAN | 0.0688 | 0.0727 | 0.0745 | 0.0840 | 0.0625 | | | |
| % DIFFERENCE | | 5.7 | 8.3 | 22.1 | -9.2 | | | |
| S.D. | 0.01146 | 0.01345 | 0.00733 | 0.01814 | 0.01102 | | | |
| S.E. | 0.00512 | 0.00601 | 0.00328 | 0.00811 | 0.00493 | | | |
| N | 5 | 5 | 5 | 5 | 5 | | | |
| ARY/OD, LEFT (G/100 (| G FINAL BODY WEIGHT) | | | | | | | |
| MEAN | 0.018 | 0.019 | 0.019 | 0.022 | 0.020 | | | |
| S.D. | 0.0022 | 0.0030 | 0.0008 | 0.0037 | 0.0041 | | | |
| S.E. | 0.0010 | 0.0014 | 0.0004 | 0.0016 | 0.0018 | | | |
| N | 5 | 5 | 5 | 5 | 5 | | | |
| ARY/OD, RIGHT (G) | | | | | | | | |
| MEAN | 0.0839 | 0.0802 | 0.0891 | 0.0753 | 0.0619 | | | |
| % DIFFERENCE | | -4.4 | 6.2 | -10.3 | -26.2 | | | |
| S.D. | 0.01361 | 0.00701 | 0.01790 | 0.01514 | 0.01197 | | | |
| S.E. | 0.00609 | 0.00313 | 0.00800 | 0.00677 | 0.00535 | | | |
| N | 5 | 5 | 5 | 5 | 5 | | | |

** = Significantly different from the control group at 0.01 using Dunnett's test NONGRAVID WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

TABLE S18 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 3 SUMMARY OF ORGAN WEIGHTS AND ORGAN WTS. RELATIVE TO BODY WTS.

| FEMALES | | | | | | | | | | |
|-------------------------|----------------------|--------------|---------------|---------------|----------------|--|--|--|--|--|
| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY | | | | | |
| OVARY/OD, RIGHT (G/100 | G FINAL BODY WEIGHT) | | | | | | | | | |
| MEAN | 0.022 | 0.021 | 0.023 | 0.020 | 0.019 | | | | | |
| S.D. | 0.0031 | 0.0022 | 0.0034 | 0.0033 | 0.0029 | | | | | |
| S.E. | 0.0014 | 0.0010 | 0.0015 | 0.0015 | 0.0013 | | | | | |
| N | 5 | 5 | 5 | 5 | 5 | | | | | |
| THYMUS (G) | | | | | | | | | | |
| MEAN | 0.2481 | 0.2646 | 0.2890 | 0.2550 | 0.1944 | | | | | |
| % DIFFERENCE | | 6.7 | 16.5 | 2.8 | -21.6 | | | | | |
| S.D. | 0.04853 | 0.04155 | 0.06621 | 0.05381 | 0.08086 | | | | | |
| S.E. | 0.02170 | 0.01858 | 0.02961 | 0.02407 | 0.03616 | | | | | |
| N | 5 | 5 | 5 | 5 | 5 | | | | | |
| THYMUS (G/100 G FINAL B | ODY WEIGHT) | | | | | | | | | |
| MEAN | 0.066 | 0.068 | 0.074 | 0.068 | 0.063 | | | | | |
| S.D. | 0.0140 | 0.0077 | 0.0171 | 0.0108 | 0.0286 | | | | | |
| S.E. | 0.0062 | 0.0034 | 0.0076 | 0.0048 | 0.0128 | | | | | |
| N | 5 | 5 | 5 | 5 | 5 | | | | | |

None significantly different from control group NONGRAVID WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

SPONSOR: AMERICAN PETROLEUM

POFBSTv5.20 07/16/2010

TABLE S19 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 1 SPONSOR: AMERICAN PETROLEUM SUMMARY OF FETAL DATA AT SCHEDULED NECROPSY

| | | | | | | POST | | | PRE | FETAL | NO. OF |
|------|----------------|---------|---------|--------|------|------|--------------|-------|--------------|----------|---------|
| | SEX | VIABLE | DEAD | RESORF | | | IMPLANTATION | | IMPLANTATION | WEIGHTS | GRAVID |
| GROU | P M F | FETUSES | FETUSES | EARLY | LATE | LOSS | SITES | LUTEA | LOSS | IN GRAMS | FEMALES |
| | | | | | | | | | | | |
| 1 | TOTAL 39 28 | 67 | 0 | 3 | C | 3 | 70 | 82 | 12 | NA | 5 |
| | MEAN 7.8 5.6 | 13.4 | 0.0 | 0.6 | 0.0 | 0.6 | 14.0 | 16.4 | 2.4 | 3.8 | |
| | S.D. 1.64 1.67 | 2.07 | 0.00 | 0.89 | 0.00 | 0.89 | 1.41 | 5.98 | 4.83 | 0.18 | |
| | S.E. 0.73 0.75 | 0.93 | 0.00 | 0.40 | 0.00 | 0.40 | 0.63 | 2.68 | 2.16 | 0.08 | |
| 2 | TOTAL 30 43 | 73 | 0 | 3 | C | 3 | 76 | 82 | 6 | NA | 5 |
| | MEAN 6.0 8.6 | 14.6 | 0.0 | 0.6 | 0.0 | 0.6 | 15.2 | 16.4 | 1.2 | 3.6 | |
| | S.D. 2.24 2.30 | 2.19 | 0.00 | 0.55 | 0.00 | 0.55 | 2.39 | 3.65 | 1.30 | 0.27 | |
| | S.E. 1.00 1.03 | 0.98 | 0.00 | 0.24 | 0.00 | 0.24 | 1.07 | 1.63 | 0.58 | 0.12 | |
| 3 | TOTAL 37 37 | 74 | 0 | 6 | C | 6 | 80 | 87 | 7 | NA | 5 |
| | MEAN 7.4 7.4 | 14.8 | 0.0 | 1.2 | 0.0 | | 16.0 | 17.4 | 1.4 | 3.7 | |
| | S.D. 2.51 3.13 | 1.92 | 0.00 | 0.84 | 0.00 | | 2.55 | 2.30 | 1.34 | 0.27 | |
| | S.E. 1.12 1.40 | 0.86 | 0.00 | 0.37 | 0.00 | 0.37 | 1.14 | 1.03 | 0.60 | 0.12 | |
| 4 | TOTAL 40 29 | 69 | 0 | 2 | C | 2 | 71 | 75 | 4 | NA | 5 |
| | MEAN 8.0 5.8 | 13.8 | 0.0 | 0.4 | 0.0 | | 14.2 | 15.0 | 0.8 | 3.8 | |
| | S.D. 2.35 1.48 | 2.39 | 0.00 | 0.55 | 0.00 | | 1.92 | 1.87 | 0.84 | 0.31 | |
| | S.E. 1.05 0.66 | 1.07 | 0.00 | 0.24 | 0.00 | | 0.86 | 0.84 | 0.37 | 0.14 | |
| 5 | TOTAL 13 13 | 26 | 2 | 11 | C | 13 | 39 | 60 | 21 | NA | 5 |
| | MEAN 2.6 2.6 | 5.2** | 0.4 | 2.2 | 0.0 | | 7.8* | 12.0 | 4.2 | 3.5 | = |
| | S.D. 3.58 3.71 | 7.16 | 0.89 | 2.59 | 0.00 | | 4.87 | 3.39 | 4.44 | 0.21 | |
| | S.E. 1.60 1.66 | 3.20 | 0.40 | 1.16 | 0.00 | | 2.18 | 1.52 | 1.98 | 0.15 | |

^{* =} Significantly different from the control group at 0.05 ** = Significantly different from the control group at 0.01

MEAN NUMBER OF VIABLE FETUSES, MEAN NUMBER OF IMPLANTATION SITES, MEAN NUMBER OF CORPORA LUTEA, FETAL WEIGHTS COMPARED USING DUNNETT'S TEST

1- 0 MG/KG/DAY 2- 30 MG/KG/DAY 3- 100 MG/KG/DAY 4- 300 MG/KG/DAY 5- 1000 MG/KG/DAY

PLSUv5.12 07/15/2010

NA = NOT APPLICABLE

SPONSOR: AMERICAN PETROLEUM

TABLE S20
PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 1

SUMMARY OF FETAL DATA AT SCHEDULED NECROPSY [% PER LITTER]

| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
|----------------------|-------------|--------------|---------------|---------------|----------------|
| ORPORA LUTEA | | | | | |
| MEAN | 16.4 | 16.4 | 17.4 | 15.0 | 12.0 |
| S.D. | 5.98 | 3.65 | 2.30 | 1.87 | 3.39 |
| S.E. | 2.68 | 1.63 | 1.03 | 0.84 | 1.52 |
| N | 5 | 5 | 5 | 5 | 5 |
| MPLANTATION SITES | | | | | |
| MEAN | 14.0 | 15.2 | 16.0 | 14.2 | 7.8* |
| S.D. | 1.41 | 2.39 | 2.55 | 1.92 | 4.87 |
| S.E. | 0.63 | 1.07 | 1.14 | 0.86 | 2.18 |
| N | 5 | 5 | 5 | 5 | 5 |
| TABLE FETUSES (%) | | | | | |
| MEAN | 95.4 | 96.2 | 93.0 | 96.8 | 40.0 |
| S.D. | 6.89 | 3.61 | 4.83 | 4.49 | 54.77 |
| S.E. | 3.08 | 1.61 | 2.16 | 2.01 | 24.49 |
| N | 5 | 5 | 5 | 5 | 5 |
| EAD FETUSES (%) | | | | | |
| MEAN | 0.0 | 0.0 | 0.0 | 0.0 | 13.3 |
| S.D. | 0.00 | 0.00 | 0.00 | 0.00 | 29.83 |
| S.E. | 0.00 | 0.00 | 0.00 | 0.00 | 13.34 |
| N | 5 | 5 | 5 | 5 | 5 |
| ARLY RESORPTIONS (%) | | | | | |
| MEAN | 4.6 | 3.9 | 7.0 | 3.2 | 46.7 |
| S.D. | 6.89 | 3.63 | 4.83 | 4.49 | 50.55 |
| S.E. | 3.08 | 1.62 | 2.16 | 2.01 | 22.61 |
| N | 5 | 5 | 5 | 5 | 5 |

PROPORTIONAL (%) DATA COMPARED USING DUNN'S TEST
CORPORA LUTEA AND IMPLANTATION SITES COMPARED USING DUNNETT'S TEST
MODIFIED STATISTICS USED. * INDICATES PARAMETRIC ANALYSIS AND + INDICATES NON-PARAMETRIC ANALYSIS.

* = Significantly different from the control group at 0.05

PROJECT NO.:WIL-402010F SPONSOR:AMERICAN PETROLEUM

TABLE S20 ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SUMMARY OF FETAL DATA AT SCHEDULED NECROPSY [% PER LITTER]

PAGE 2

| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
|----------------------|-------------|--------------|---------------|---------------|----------------|
| TE RESORPTIONS (%) | | | | | |
| MEAN | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 |
| S.D. | 0.00 | 0.00 | 0.00 | 0.00 | 0.00 |
| S.E. | 0.00 | 0.00 | 0.00 | 0.00 | 0.00 |
| N | 5 | 5 | 5 | 5 | 5 |
| TAL RESORPTIONS (%) | | | | | |
| MEAN | 4.6 | 3.9 | 7.0 | 3.2 | 46.7 |
| S.D. | 6.89 | 3.63 | 4.83 | 4.49 | 50.55 |
| S.E. | 3.08 | 1.62 | 2.16 | 2.01 | 22.61 |
| N | 5 | 5 | 5 | 5 | 5 |
| RE-IMPLANTATION LOSS | (%) | | | | |
| MEAN | 9.6 | 6.3 | 8.1 | 5.3 | 36.1 |
| S.D. | 17.68 | 6.24 | 8.42 | 5.42 | 32.08 |
| S.E. | 7.91 | 2.79 | 3.77 | 2.42 | 14.35 |
| N | 5 | 5 | 5 | 5 | 5 |
| ST-IMPLANTATION LOSS | (%) | | | | |
| MEAN | 4.6 | 3.9 | 7.0 | 3.2 | 60.0 |
| S.D. | 6.89 | 3.63 | 4.83 | 4.49 | 54.77 |
| S.E. | 3.08 | 1.62 | 2.16 | 2.01 | 24.49 |
| N | 5 | 5 | 5 | 5 | 5 |
| ALES (%) | | | | | |
| MEAN | 58.3 | 41.1 | 50.8 | 57.5 | 50.6 |
| S.D. | 10.32 | 13.34 | 18.62 | 11.00 | 10.94 |
| S.E. | 4.62 | 5.97 | 8.33 | 4.92 | 7.74 |
| N | 5 | 5 | 5 | 5 | 2 |

PROPORTIONAL (%) DATA COMPARED USING DUNN'S TEST
MODIFIED STATISTICS USED. * INDICATES PARAMETRIC ANALYSIS AND + INDICATES NON-PARAMETRIC ANALYSIS.
None significantly different from control group

PROJECT NO.:WIL-402010F SPONSOR:AMERICAN PETROLEUM

TABLE S20 ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SUMMARY OF FETAL DATA AT SCHEDULED NECROPSY [% PER LITTER]

| GROUP: | 0 MG/KG/DAY | 30 MG/KG/DAY | 100 MG/KG/DAY | 300 MG/KG/DAY | 1000 MG/KG/DAY |
|-----------------------|-------------|--------------|---------------|---------------|----------------|
| EMALES (%) | | | | | |
| MEAN | 41.7 | 58.9 | 49.2 | 42.5 | 49.4 |
| S.D. | 10.32 | 13.34 | 18.62 | 11.00 | 10.94 |
| S.E. | 4.62 | 5.97 | 8.33 | 4.92 | 7.74 |
| N | 5 | 5 | 5 | 5 | 2 |
| ALE FETAL WEIGHTS (g) | | | | | |
| MEAN | 3.9 | 3.7 | 3.8 | 3.9 | 3.6 |
| % DIFFERENCE | | -5.1 | -2.6 | 0.0 | -7.7 |
| S.D. | 0.24 | 0.27 | 0.26 | 0.33 | 0.14 |
| S.E. | 0.11 | 0.12 | 0.12 | 0.15 | 0.10 |
| N | 5 | 5 | 5 | 5 | 2 |
| EMALE FETAL WEIGHTS (| q) | | | | |
| MEAN | 3.6 | 3.6 | 3.6 | 3.6 | 3.3 |
| % DIFFERENCE | | 0.0 | 0.0 | 0.0 | -8.3 |
| S.D. | 0.15 | 0.29 | 0.33 | 0.28 | 0.14 |
| S.E. | 0.07 | 0.13 | 0.15 | 0.12 | 0.10 |
| N | 5 | 5 | 5 | 5 | 2 |
| OMBINED FETAL WEIGHTS | (q) | | | | |
| MEAN | 3.8 | 3.6 | 3.7 | 3.8 | 3.5 |
| % DIFFERENCE | | -5.3 | -2.6 | 0.0 | -7.9 |
| S.D. | 0.18 | 0.27 | 0.27 | 0.31 | 0.21 |
| S.E. | 0.08 | 0.12 | 0.12 | 0.14 | 0.15 |
| N | 5 | 5 | 5 | 5 | 2 |

PROPORTIONAL (%) DATA COMPARED USING DUNN'S TEST FETAL WEIGHTS COMPARED USING DUNNETT'S TEST

MODIFIED STATISTICS USED. * INDICATES PARAMETRIC ANALYSIS AND + INDICATES NON-PARAMETRIC ANALYSIS. None significantly different from control group

PLPSUv5.10 07/15/2010 R:07/15/2010

PAGE 3

TABLE S21
ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010F

| | | | | | | | | | | PAGE | 1 | |
|--|-------------------------|----|--------|---------|-----|------------|---------|---|------|---------|----|---------|
| SPONSOR: AMERICAN PETROLEUM SUMMARY OF FETUSES AND LITTERS WITH MALFORMATIONS [ABSOLUTE NO.] | | | | | | | | | | DAY | 20 | |
| | | | F | ETU | SES | | | L | ITTI | E R S | | |
| | DOSE GROUP: | 1 | 2 | 3 | 4 | 5 | 1 | 2 | 3 | 4 | 5 | |
| | | | | | | | | | | | | |
| NUMBER EXAMINED EXTERNALLY | | 67 | 73 | 74 | 69 | 26 | 5 | 5 | 5 | 5 | 2 | |
| OMPHALOCELE | | 0 | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | |
| TOTAL NUMBER WITH MALFORMATIONS | | | | | | | | | | | | |
| EXTERNAL : | | 0 | 1 | 0 | 0 | 0 | 0 | 1 | 0 | 0 | 0 | |
| 1- 0 MG/KG/DAY 2- 30 MG/ | KG/DAY 3- 100 MG/KG/DAY | 4- | 300 MG | G/KG/DA | Y 5 | 5- 1000 MG | /KG/DAY | | | | | |
| | | | | | | | | | | | | |

PMALv5.08 07/15/2010

| | | TABLE S22 |
|--|--|-----------|
| | | |

| PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE SPONSOR:AMERICAN PETROLEUM SUMMARY OF FETUSES AND LITTERS WITH VARIATIONS [ABSOLUTE NO.] | | | | | | | | PAGE | 1 | | | |
|--|---|----|--------|---------|------|-----------|--------|------|------|-------|---|--|
| SPONSOR:AMERICAN PETROLEUM | SUMMARY OF FEIUSES AND LITTERS WITH VARIATIONS [ABSOLUTE NO.] | | | | | | | | DAY | 20 | | |
| | | | F | ETU | SES | | | L | ITTI | E R S | | |
| | DOSE GROUP: | 1 | 2 | 3 | 4 | 5 | 1 | 2 | 3 | 4 | 5 | |
| | | | | | | | | | | | | |
| NUMBER EXAMINED EXTERNALLY | | 67 | 73 | 74 | 69 | 26 | 5 | 5 | 5 | 5 | 2 | |
| NUMBER WITH FINDINGS | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| 1- 0 MG/KG/DAY 2- 30 MC | G/KG/DAY 3- 100 MG/KG/DAY | 4- | 300 MG | G/KG/DA | .Y 5 | - 1000 MG | KG/DAY | | | | | |

PMALv5.08 07/15/2010

APPENDIX A

Study Protocol



PROTOCOL

AN ORAL (GAVAGE) 14-DAY AND PILOT PRENATAL DEVELOPMENTAL TOXICITY STUDY OF NAPHTHENIC ACID IN RATS

Submitted To:

American Petroleum Institute

1220 L Street, NW Washington, DC 20005

WIL Research Laboratories, LLC 1407 George Road Ashland, OH 44805-8946

1 OBJECTIVE:

To determine dose levels of the test substance to be evaluated in a potential combined repeated dose toxicity study with the reproduction/developmental toxicity screening test (OECD 422) in rats.

This study will be conducted in compliance with the EPA/TSCA (40 CFR 792) and OECD [C(97) 186/Final] Principles of Good Laboratory Practice.

2 PERSONNEL INVOLVED IN THE STUDY:

2.1 Sponsor Representative:

American Petroleum Institute 1220 L Street, NW Washington, DC 20005 Tel: (202) 682-8333 Email:

2.2 WIL Study Director:

Staff Toxicologist, Developmental and Reproductive Toxicology Tel: (419) 289-8700

Fax: (419) 289-3650

Email:

2.3 WIL Departmental Responsibilities:

Director, Developmental and Reproductive Toxicology **Emergency Contact**

Tel: (419) 289-8700 Fax: (419) 289-3650

Email:

President and Chief Operating Officer

Assistant Director, Developmental and Reproductive Toxicology



Director, Pathology

Assistant Director, Neurosciences

Director, Metabolism and Analytical Chemistry

Clinical Veterinarian, Head of Surgery and Experimental Medicine

Director, Informational Systems

Manager, Gross Pathology and Developmental Toxicology Laboratory

Manager, Quality Assurance

Operations Manager, Developmental and Reproductive Toxicology and the Formulations Laboratory

Manager, Reporting and Regulatory Technical Services



3 STUDY SCHEDULE:

Proposed Experimental Starting

(Animal Receipt) Date:

May 18, 2010

Proposed Experimental Start

(First Day of Dosing) Date:

June 2, 2010

Proposed Last Day of Dosing (Males)

June 15, 2010

Proposed Last Day of Dosing (Females)

June 25, 2010

Proposed Experimental

Completion/Termination Date:

June 25, 2010

Proposed Audited Report Date:

August 20, 2010

4 TEST SUBSTANCE DATA:

4.1 Test Substance Shipment:

Test substance and applicable documentation, including a Certificate of Analysis, will be shipped under Sponsor's responsibility to:

Formulations Laboratory (WIL-402010,

mulations Laboratory (WIL-402010,

Attn:

WIL Research Laboratories, LLC 1407 George Road

Ashland, Ohio 44805-8946

4.2 Identification:

Naphthenic Acid.

4.3 Lot Number:

CP006002

4.4 Expiration/Retest Date:

Test substance will be characterized prior to and following use for dose administration.



4.5 Purity:

100%

4.6 Storage Conditions:

Room temperature.

4.7 Stability:

The test substance is considered to be stable under the storage conditions provided by the Sponsor.

4.8 Physical Description:

To be documented by WIL Research Laboratories, LLC.

4.9 Reserve Samples:

Reserve samples of the test substance will be taken in accordance with WIL Standard Operating Procedures and stored in the Archives at WIL Research Laboratories, LLC indefinitely, unless otherwise specified.

4.10 Personnel Safety Data:

Included in the study records. It is the responsibility of the Sponsor to notify the testing facility of any special handling requirements for the test substance. A Material Safety Data Sheet (MSDS) should accompany the test substance upon arrival at the laboratory.

4.11 Test Substance Disposition:

With the exception of the reserve sample for each batch of test substance, all neat test substance remaining at study completion will be returned to the Sponsor. Alternatively, the test substance can be retained for subsequent studies.

5 TEST SYSTEM:

5.1 Species:

Rat

5.2 Breed/Strain:

Sprague-Dawley Crl:CD(SD)



5.3 Source:

Charles River Laboratories, Inc. (Facility to be included in study records)

5.4 Number of Animals:

25 Males and 25 Females (maximum of 30 Males and 32 Females purchased). Non-litter mates will be purchased. Animals not assigned to study will be transferred to the stock animal colony or will be euthanized by carbon dioxide inhalation and the carcasses discarded.

This is the minimum number of animals to allow for meaningful extrapolation to the definitive toxicity study.

5.5 Body Weight Range:

Males: 250-375 g at randomization.

Females: minimum of 220g at initiation of breeding

5.6 Approximate Age:

Males: 57-65 days at randomization

Females: 80 to 120 days at the initiation of breeding

5.7 Identification System:

The animals will be uniquely identified by a Monel® metal ear tag displaying the animal number. Individual cage cards will be affixed to each cage and will display the animal number, group number, study number, dosage level and sex of the animal.

5.8 Justification for Selection:

This species and strain of animal is recognized as appropriate for reproduction studies. WIL Research Laboratories, LLC has reproductive historical control data in the Crl:CD(SD) rat. This animal model has been proven to be susceptible to the effects of reproductive toxicants.

6 SPECIFIC MAINTENANCE SCHEDULE:

6.1 Animal Housing:

The animals will be individually housed (except during mating) in clean suspended wire-mesh cages in an environmentally controlled room during the



study. The cages will be elevated above cage-board or other suitable material, which will be changed at least three times each week. The cages will be subjected to routine cleaning at a frequency consistent with maintaining good animal health and WIL Standard Operating Procedures. The facilities at WIL Research Laboratories, LLC are accredited by the Association for Assessment and Accreditation of Laboratory Animal Care International (AAALAC International).

6.2 Environmental Conditions:

Controls will be set to maintain temperature at $71 \pm 5^{\circ}F$ ($22 \pm 3^{\circ}C$) and relative humidity at $50 \pm 20\%$. Temperature and relative humidity will be monitored continuously. Data for these two parameters will be scheduled for automatic collection on an hourly basis. Fluorescent lighting controlled by light timers will provide illumination for a 12-hour light/dark photoperiod. The ventilation rate will be set at a minimum of 10 room air changes per hour, 100% fresh air.

6.3 Drinking Water:

Reverse osmosis-purified water will be available ad libitum. Filters servicing the automatic watering system are changed regularly according to WIL Standard Operating Procedures. The municipal water supplying the laboratory is analyzed on a routine basis according to WIL Standard Operating Procedures to ensure that contaminants are not present in concentrations that would be expected to affect the outcome of the study.

6.4 Basal Diet:

PMI Nutrition International, LLC Certified Rodent LabDiet[®] 5002 will be offered *ad libitum* during the study. Periodic analyses of the certified feed are performed by the manufacturer to ensure that heavy metals and pesticides are not present at concentrations that would be expected to affect the outcome of the study. Results of the analyses are provided to WIL Research Laboratories, LLC by the manufacturer. Feeders will be changed and sanitized once per week.

7 EXPERIMENTAL DESIGN:

7.1 Animal Receipt and Quarantine:

Each animal will be inspected by a qualified technician upon receipt. Rats judged to be in good health and suitable as test animals will be immediately placed in quarantine for a minimum of 10 days. All rats will be initially weighed, permanently identified with a metal ear tag and receive a clinical observation. During the quarantine period, each rat will be observed twice daily



for changes in general appearance and behavior. Prior to the start of the in-life phase, those animals judged to be suitable test subjects will be identified.

7.2 Breeding Procedure:

At the conclusion of the quarantine period, female rats judged to be suitable test subjects and meeting acceptable body weight requirements will be cohabitated with untreated resident male rats (1:1) of the same strain and source in a suspended wire-mesh cage for mating. Detection of mating will be confirmed by evidence of a vaginal copulatory plug in the vagina or by a vaginal lavage for sperm. After confirmation of mating, the female will be returned to an individual suspended wire-mesh cage (assigned to a group), and the day will be designated as day 0 of gestation. Main study males will not be mated.

7.3 Randomization:

The mated females will be assigned to groups using a WIL Toxicology Data Management System (WTDMS computer program which assigns animals based on stratification of the gestation day 0 body weights into a block design to one control group and four test substance groups of five rats each.

At the conclusion of the male quarantine period, males judged to be suitable test subjects and meeting acceptable body weight requirements will be assigned to the study at random using a computer program. At that time, the animal numbers and corresponding body weights will be entered into the WIL Toxicology Data Management System (WTDMS[™]). A printout containing the animal numbers and individual group assignments will be generated based on body weight stratification into a block design. The animals will then be arranged into the groups according to the printout. The control group and four test substance groups will consist of 5 males each.

Any animal assigned to the study that is found dead, euthanized *in extremis* or exhibits abnormal clinical signs, reduced food consumption or body weight losses prior to the start of dosing may be replaced by an animal of appropriate age, when possible. Replacement animals will be arbitrarily assigned (not computer randomized) to the study based on comparable body weights (if possible) with respect to the animal that was replaced.

7.4 Route and Rationale of Test Substance Administration:

The route of administration will be oral (gavage) since this is a potential route of exposure for humans. Historically, this route has been used extensively for studies of this nature. Appropriately sized flexible, Teflon®-shafted, stainless steel ball-tipped dosing cannulae will be used for the oral administration by gavage.



7.5 Organization of Test Groups, Dosage Levels and Treatment Regimen:

7.5.1 Organization of Test Groups:

The dosage levels will be determined from results of previous studies and will be provided by the Sponsor Representative after consultation with the WIL Study Director.

The following table presents the study group arrangement.

| Group | Test Substance | Dosage Level | Dosage Concentration | Dosage Volume | Number of Animals | | |
|--------|-----------------|-----------------|-------------------------|------------------|----------------------|--------|--|
| Number | | (mg/kg/day) | (mg/mL) | (mL/kg) | Male | Female | |
| 1 | Vehicle Control | 0 | 0 | 10 | 5 | 5 | |
| 2 | Naphthenic Acid | 30 | 3 | 10 | 5 | 5 | |
| 3 | Naphthenic Acid | 100 | 10 | 10 | 5 | 5 | |
| 4 | Naphthenic Acid | 300 | 30 | 10 | - 5 | 5 | |
| 5 | Naphthenic Acid | 1000 | 100 | 10 | 5 | 5 | |

7.5.2 Vehicle Control Substance:

Corn oil.

7.5.3 Treatment Regimen:

All rats will be dosed at approximately the same time each day.

7.5.3.1 Males:

The test and control substances will be administered as a single daily dose from study day 0 until study day 13, inclusively.

7.5.3.2 Females:

The test and control substances will be administered as a single daily dose from gestation day 1 through 20, inclusively.

7.5.4 Adjustment of Doses:

Individual dosages will be calculated based on the most recent body weight to provide the proper mg/kg/day dosage.



7.6 Test Substance Characterization:

A sample of the test substance will be collected prior to and following use for test substance administration and characterized by the WIL Research Analytical Chemistry department according the method provided by the sponsor.

7.7 Preparation and Analysis of Test Substance Formulations:

7.7.1 Method and Frequency of Preparation:

Based on the physical characteristics of the test substance, appropriate methods will be used to ensure the best possible formulations of the test substance in the vehicle. Dosing formulations will be stored at room temperature for a period not to exceed established stability. The Study Director or designee will visually inspect the formulations prior to initiation of dosing. This visual inspection will be performed to ensure that the formulations are visibly homogeneous and acceptable for dosing. Any special procedures required for formulation will be documented according to Good Laboratory Practices and presented in the final report of this study. Test substance formulations will normally be prepared approximately weekly and divided into aliquots for daily dispensation. The test substance and vehicle formulations will be stirred continuously during dosing. Frequency of formulation may be adjusted based on stability results.

7.7.2 Homogeneity and Stability of Test Substance Formulations:

Analyses to demonstrate the stability of the test substance formulations and, for suspensions, the homogeneity of the test substance formulations will be conducted before the initiation of dosing, if possible, at concentrations spanning the range of concentrations to be used on the study.

For stability assessments, test substance formulations will be prepared, stored at room temperature and sampled over a minimum of 10 days. The samples will be analyzed to evaluate stability. Additional aspects of stability (e.g., stability of frozen samples) may also be investigated.

For homogeneity assessments, test substance formulations of sufficient volume for dosing a group of animals for up to one week (length dependent on stability results) will be prepared. Samples will be collected from the top, middle and bottom of the formulations and the samples analyzed to assess the homogeneity of the test substance in the mixtures. If aliquots of the test substance formulations will be stored prior to use, a volume of formulation similar in size to the amount



needed for one day will be stored under suitable conditions for the maximum length of time acceptable. After remixing for a minimum of 30 minutes using a magnetic stirrer, samples will be collected from the top and bottom of the formulations and analyzed to assess the homogeneity after storage and resuspension. Homogeneity assessments will not be performed for test substance formulations that are solutions provided that the Sponsor provides pertinent solubility data. Any backup samples kept at WIL Research Laboratories, LLC will be discarded after the study director's approval of the analytical results.

7.7.3 Concentration Analysis:

Samples will be taken from the middle of the first and last dosing formulation prepared during the in-life phase of the study. These samples will be analyzed to verify the concentration of the test substance in the dosing formulations. Any back-up samples kept at WIL Research Laboratories, LLC will be discarded after the study director's approval of the analytical results.

7.8 General Observations During the Experimental Period:

7.8.1 Appearance and Behavior:

Each animal will be observed twice daily for moribundity and mortality, once in the morning and once in the afternoon, for the duration of the study. Clinical observations will be recorded daily. Mortality and all signs of overt toxicity will be recorded on the day observed. The observations shall include, but are not limited to, evaluation for changes in appearance of skin and fur, eyes, mucous membranes, respiratory and circulatory systems, autonomic and central nervous systems, somatomotor activity and behavior. All animals will also be observed on the day of necropsy and any findings will be recorded.

During the treatment period, each animal will be observed at approximately one hour following each dose administration for findings that are potentially related to treatment or that might change before the next scheduled observation. Additional post-dosing observation periods may be necessary and will be documented in the study records.

7.8.2 Body Weights and Food Consumption:

7.8.2.1 Males:

Individual body weights and food consumption will be recorded weekly.



7.8.2.2 Females:

Individual body weights and food consumption will be recorded on gestation days 0, 6, 7, 10, 14, 17, and 20.

7.8.3 Deaths and Animals Euthanized in Extremis:

Males not surviving until the scheduled euthanasia will be necropsied, and the cause of death will be recorded, if possible. Animals not expected to survive to the next observation period (moribund) will be euthanized by carbon dioxide inhalation. The cranial, thoracic, abdominal and pelvic cavities will be opened and the contents examined. Tissues will be preserved in 10% neutral-buffered formalin only as deemed necessary by the gross findings. Organ weights will not be obtained. All carcasses will be discarded.

Females not surviving until the scheduled euthanasia will be necropsied and the cause of death recorded, if possible. Rats not expected to survive to the next observation period (moribund) will be euthanized by carbon dioxide inhalation. The cranial, thoracic, abdominal and pelvic cavities will be opened and the contents examined. The number and location of implantation sites and corpora lutea will be recorded. Uteri which appear nongravid by macroscopic examination will be opened and placed in a 10% ammonium sulfide solution (Salewski, 1964) for detection of early implantation loss. Tissue with gross lesions will be preserved in 10% neutral-buffered formalin for possible future histopathologic examination. Carcasses from adult animals will be discarded. Viable fetuses will be euthanized by a subcutaneous injection of sodium pentobarbital in the scapular region. Recognizable fetuses will be examined externally and discarded.

7.8.4 Premature Deliveries:

Females that deliver prematurely will be euthanized by carbon dioxide inhalation that day. The thoracic, abdominal and pelvic cavities will be opened and the organs examined. The number and location of former implantation sites and viable fetuses will be recorded. Corpora lutea will also be counted and recorded. Gross lesions will be preserved in 10% neutral-buffered formalin for possible future histopathologic examinations. Carcasses from adult animals will be discarded. Viable fetuses or pups will be euthanized by a subcutaneous (scapular region) or intraperitoneal injection of sodium pentobarbital (as appropriate). Recognizable fetuses or pups will be examined externally and discarded.



7.9 Macroscopic Examination:

7.9.1 Female Scheduled Necropsy – Gestation Day 20:

7.9.1.1 Laparohysterectomy and Macroscopic Examination:

Laparohysterectomies and macroscopic examinations will be performed blind to treatment group. All surviving rats will be euthanized by carbon dioxide inhalation on gestation day 20. The thoracic, abdominal, and pelvic cavities will be opened and the organs examined. The uterus of each dam will be excised and its adnexa trimmed. The number of corpora lutea will be counted and recorded. Gravid uterine weights will be obtained and recorded. The uterus of each dam will be opened and the number of viable and nonviable fetuses, early and late resorptions and the total number of implantation sites will be recorded, and the placentae will be examined. The individual uterine distribution will be documented using the following procedure: all implantation sites, including early and late resorptions, will be numbered in consecutive fashion beginning with the left distal uterine horn, noting the position of the cervix and continuing from the proximal to the distal right uterine horn. Uteri which appear nongravid by macroscopic examination will be opened and placed in a 10% ammonium sulfide solution (Salewski, 1964) for detection of early implantation loss. Maternal tissues will be preserved in 10% neutral-buffered formalin only as deemed necessary by the gross findings. Representative sections of corresponding organs from a sufficient number of controls will be retained for comparison, if possible. The carcasses will be discarded.

7.9.1.2 Fetal Examination:

Each viable fetus will be examined externally, sexed, weighed, euthanized by a subcutaneous injection of sodium pentobarbital in the scapular region and discarded. If autolysis is either absent or minimal, non-viable fetuses will be examined, crown rump by length measured, weighed and individually sexed. The crown-rump length of late resorptions (advanced degree of autolysis) will be measured, the degree of autolysis recorded, a gross external examination performed (if possible) and the tissue will be discarded. The findings will be recorded as either developmental variations or malformations. Representative photographs of all malformations, as



appropriate, will be included in the study records. Corresponding low magnification photographs, depicting both the malformed fetus and a comparison control fetus, or normal littermate, will also be included in the study records as needed and as appropriate for comparison, when possible. Selected specimens may be preserved at the discretion of the Study Director.

7.9.2 Male Scheduled Necropsy:

A gross necropsy will be conducted on surviving males on study day 14 following euthanasia by carbon dioxide inhalation. This will include examination of the external surface, all orifices, the cranial cavity, the external surface of the brain and the thoracic, abdominal and pelvic cavities including viscera. Tissues will be preserved in 10% neutral-buffered formalin only as deemed necessary by the gross findings. All carcasses will be discarded.

7.9.3 Organ Weights:

The following organs will be weighed from all animals at the scheduled necropsy. Organ-to-final-body-weight ratios will be calculated.

Kidneys Testes* Thymus

Liver

Ovaries (with oviducts)*

* - These paired organs will be weighed separately.

8 DURATION OF STUDY:

The acclimation and in-life phases of this study will require approximately 2 months.

9 STATISTICAL METHODS:

All analyses will be two-tailed for minimum significance levels of 5% and 1%. All means will be presented with standard deviations. All statistical tests will be performed using a computer with appropriate programming as referenced below.

9.1 In-Life Data:

Continuous data variables, [mean maternal body weights (absolute and net for females), body weight gains (absolute and net for females) and food consumption of each interval] will be subjected to a parametric one-way



analysis of variance (ANOVA) (Snedecor, 1980) to determine intergroup difference. If the results of the ANOVA are significant (p<0.05), Dunnett's test (Dunnett, 1964) will be applied to the data to compare the treated groups to the control group.

9.2 Laparohysterectomy Data:

The group mean numbers of viable fetuses, corpora lutea, implantation sites, maternal gravid uterine weights and mean fetal weight (separately by sex, and combined) will be subjected to a parametric one-way analysis of variance (ANOVA) (Snedecor, 1980) and Dunnett's test (Dunnett, 1964) as described above. The mean litter proportions of prenatal data (% per litter of viable and nonviable fetuses, early and late resorptions, total resorptions, pre- and post-implantation loss and the fetal sex distribution) will be subjected to the Kruskal-Wallis nonparametric ANOVA test (Kruskal, 1952) to determine intergroup difference. If the results of the ANOVA are significant (p<0.05), the Dunn's Test (Dunn, 1964) will be applied to the data to compare the treated groups to the control group.

9.3 Organ Weight Data:

Organ weights (absolute and relative to body weights) will be subjected to a parametric ANOVA test (Snedecor, 1980) and Dunnett's test (Dunnett, 1964) as described above.

10 QUALITY ASSURANCE:

The study will be audited by the WIL Quality Assurance Unit while in progress to assure compliance with Good Laboratory Practice Regulations, adherence to the protocol and to WIL Standard Operating Procedures. The raw data and draft report will be audited by the WIL Quality Assurance Unit prior to submission to the Sponsor to assure that the final report accurately describes the conduct and the findings of the study. This study will be conducted in compliance with the EPA/TSCA (40 CFR 792) and OECD [C(97) 186/Final] Principles of Good Laboratory Practice.

This study will be included on the WIL regulated master schedule.

11 RECORDS TO BE MAINTAINED:

All original raw data records, as defined by WIL SOPs and the applicable GLPs, will be stored as described in Section 12 in the Archives at WIL Research Laboratories, LLC.



12 WORK PRODUCT:

The Sponsor will have title to all documentation records, raw data, specimens and other work product generated during the performance of the study. Any remaining formulation samples will be discarded after issuance of the Final Report. All work product, including raw paper data, pertinent electronic storage media and specimens, will be retained at no charge for six months following issuance of the final report in the Archives at WIL Research Laboratories, LLC. Thereafter, WIL Research Laboratories, LLC will charge a monthly archiving fee for retention of all work product. All work product will be stored in compliance with regulatory requirements.

Any work product, including documents, specimens, and samples, that are required by this protocol, its amendments, or other written instructions of the Sponsor, to be shipped by WIL Research Laboratories, LLC to another location will be appropriately packaged and labeled as defined by WIL's SOPs and delivered to a common carrier for shipment. WIL Research Laboratories, LLC will not be responsible for shipment following delivery to the common carrier.

13 REPORTS:

The final report will contain a summary, test substance data, methods and procedures, maternal and fetal data, WIL Historical Control Data, the analytical chemistry report and an interpretation and discussion of the study results. The final report will be comprehensive and shall define level(s) inducing toxic effects under the condition of this investigation.

WIL Research Laboratories, LLC will submit one copy of an audited draft report in a timely manner upon completion of data collection prior to issuance of the final report. One revision will be permitted as part of the cost of the study, from which the Sponsor's reasonable revisions and suggestions will be incorporated into the final report, as appropriate. Additional changes or revisions may be made, at extra cost. It is expected that the Sponsor will review the draft report and provide comments to WIL Research Laboratories, LLC within a two-month time frame following submission. WIL Research Laboratories, LLC will submit the final report within one month following receipt of comments. If the Sponsor's comments and/or authorization to finalize the report have not been received at WIL Research Laboratories, LLC within one year following submission of the draft report, WIL Research Laboratories, LLC may elect to finalize the report following appropriate written notification to the Sponsor. Two electronic copies (PDF) of the final report on CD-R will be provided. Requests for paper copies of the final report may result in additional charges.



14 ANIMAL WELFARE ACT COMPLIANCE:

This study will comply with all applicable sections of the Final Rules of the Animal Welfare Act (AWA) regulations (9 CFR Parts 1, 2 and 3). The Sponsor should make particular note of the following:

- The Sponsor Representative's signature on this protocol documents for the Study Director the Sponsor's assurance that the study described in this protocol does not unnecessarily duplicate previous experiments.
- Whenever possible, procedures used in this study have been designed to avoid or minimize discomfort, distress or pain to animals. All methods are described in this study protocol or in written laboratory Standard Operating Procedures.
- Animals that experience severe or chronic pain or distress that cannot be relieved
 will be painlessly euthanized as deemed appropriate by the veterinary staff and
 Study Director. The Sponsor will be advised by the Study Director of all
 circumstances which could lead to this action in as timely a manner as possible.
- Methods of euthanasia used during this study are in conformance with the abovereferenced regulation.
- The Sponsor/Study Director has considered alternatives to procedures that may cause more than momentary or slight pain or distress to the animals and has provided a written narrative description (AWA covered species only) of the methods and sources used to determine that alternatives are not available.

15 PROTOCOL MODIFICATION:

Modification of the protocol may be accomplished during the course of this investigation. However, no changes will be made in the study design without the verbal or written permission of the Sponsor. In the event that the Sponsor verbally requests or approves a change in the protocol, such changes will be made by appropriate documentation in the form of a protocol amendment. All alterations of the protocol and reasons for the modification(s) will be signed by the Study Director and the Sponsor Representative.

16 REFERENCES:

Dunn, O.J. Multiple comparisons using rank sums. Technometrics 1964, 6(3), 241-252.

Dunnett, C.W. New tables for multiple comparisons with a control. *Biometrics* **1964**, 20, pp. 482-491.



Kruskal, W.H.; Wallis, W.A. Use of ranks in one-criterion variance analysis. *Journal of the American Statistical Association* **1952**, *47*, 583-621.

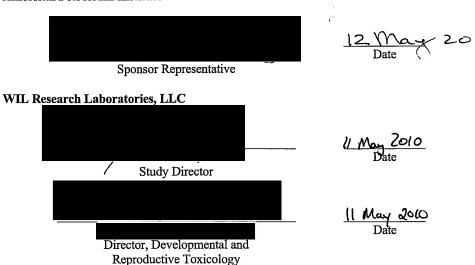
Salewski, E. Färbemethode zum makroskopischen Nachweis von Implantationsstellen am Uterus der Ratte. [Staining method for a macroscopic test for implantation sites in the uterus of the rat]. Naunyn - Schmiedebergs Archiv für Experimentelle Pathologie und Pharmakologie 1964, 247, 367.

Snedecor, G.W.; Cochran, W.G. One Way Classifications; Analysis of Variance. In *Statistical Methods*, 7th ed.; The Iowa State University Press: Ames, IA, **1980**; pp. 215-237.

17 PROTOCOL APPROVAL:

Sponsor approval received via $\underbrace{\mathcal{E}\mathcal{M}\omega'}_{\text{on}}$ on $\underbrace{\mathcal{S}\mathcal{M}\mathcal{H}\mathcal{T}\mathcal{D}\mathcal{P}}_{\text{Date}}$.

American Petroleum Institute





APPENDIX B

Test Substance Characterization (WIL Research Laboratories, LLC)

An Oral (Gavage) 14-Day and Pilot Prenatal Developmental Toxicity Study of Naphthenic Acid in Rats

Test Substance Characterization

Analytical Chemistry Department

WIL Research Laboratories, LLC



TABLE OF CONTENTS

| | | Page |
|--------|---|------|
| | Table of Contents | 2 |
| | Index of Figures | 2 |
| | Index of Tables | 3 |
| 1. | Introduction | 4 |
| 2. | Key Study Personnel | 4 |
| 3. | Experimental | 4 |
| 3.1. | Gas Chromatography | 4 |
| 3.2. | Mass Spectrometry | 5 |
| 3.3. | Sample Processing | 5 |
| 3.4. | Characterization | 5 |
| 4. | Results and Discussion | 6 |
| 4.1. | Test Article Characterization and Stability | 8 |
| 5. | Conclusion | 8 |
| 6. | Report Review and Approval | 9 |
| 7. | Abbreviations | 10 |
| | INDEX OF FIGURES | |
| Figure | e 1: Representative Distribution of Carbon Numbers and Z Families of Neat Naphthenic Acid Before Use for Dose Administration | 7 |
| Figure | 2: Representative Distribution of Carbon Numbers and Z Families of Neat Naphthenic Acid After Completion of Dose Administration | 8 |

INDEX OF TABLES

| | | <u>Page</u> | |
|----------|--|-------------|--|
| Table 1: | Before Use Characterization of Neat Test Substance | 13 | |
| Table 2: | After Use Characterization of Neat Test Substance | 14 | |

WIL-402010 American Petroleum Institute Naphthenic Acid

1. Introduction

This report provides a detailed description of a method using gas chromatography (GC) with mass spectrometric (MS) detection for naphthenic acid characterization. Test substance samples were collected and analyzed prior to use for dose administration and after completion of dose administration 12 weeks later.

A list of abbreviations that may have been used in this report is located in Section 7. (Abbreviations).

2. KEY STUDY PERSONNEL



Assistant Director, Analytical Chemistry
Data Coordinator, Reporting & Technical Support Services
Group Manager, Reporting & Technical Support Services

3. EXPERIMENTAL

3.1. Gas Chromatography

Instrument: Agilent 6890 gas chromatograph (GC) equipped with an

Agilent 5975 mass spectrometer (MS), an autosampler, and

MSD Security ChemStation data system, or equivalent

Column: Zebron ZB-5 column, $30 \text{ m} \times 0.32 \text{ mm ID}$,

0.25-um film-thickness.

Temperature Initial temperature 100°C, hold for 3 minutes.

Program: Ramp at 8°C/min to 300°C

Carrier Gas: Helium
Carrier Gas Pressure: 10.0 psi
Injector Temperature: 300°C

Injection Volume: 2.0 µL split

Split Ratio: 10:1

Run Time: 28 minutes

WIL-402010 Naphthenic Acid American Petroleum Institute

3.2. MASS SPECTROMETRY

Acquisition Parameters

Solvent Delay: 5.0 minutes

Ion Source: Electron Impact

Scan Range: 55-550 m/z

Transfer Line Temperature: 290°C Quadrupole Temperature: 150°C Source Temperature: 230°C

3.3. SAMPLE PROCESSING

Triplicate aliquots of the test substance (50.0 μL each, WIL ID no 10006B) were added to glass autosampler vials, and 450 μL of methylene chloride was added to each vial. The samples were mixed with vortex action. An aliquot (10.0 μL) of each diluted sample was removed and placed in another autosampler vial along with 100 μL of the derivatizing agent, N-methyl-N-(*tert*-butyldimethylsilyl) trifluoroacetamine (MTBSTFA) containing 1% *tert*-butyldimethylsilylchloride. The vials were capped and mixed with vortex action for approximately 20 minutes at approximately 60°C. The caps were removed and the solvent was evaporated under a stream of nitrogen until dry. The contents were reconstituted in 0.500 mL of methylene chloride.

3.4. CHARACTERIZATION

Single injections were made of each sample. The average ion intensity from approximately 10 minutes to 28 minutes was used to calculate the normalized response for carbon numbers (n) 5 through 33 and Z numbers -2 though -12 using the following equation:

$$C_nH_{2n} + {}_ZO_2$$

In total, there were 156 ions monitored for naphthenic acids. The following chart lists the ions monitored.

| n\z | 0 | -2 | -4 | -6 | -8 | -10 | -12 |
|-----|-----|-----|-----|-----|-----|-----|-----|
| 5 | 102 | | | | | | |
| 6 | 116 | | | | | | |
| 7 | 130 | 128 | | | | | |
| 8 | 144 | 142 | | | | | |
| 9 | 158 | 156 | | | | | |
| 10 | 172 | 170 | 168 | | | | |
| 11 | 186 | 184 | 182 | | | | |
| 12 | 200 | 198 | 196 | 194 | | | |
| 13 | 214 | 212 | 210 | 208 | | | |
| 14 | 228 | 226 | 224 | 222 | 220 | | |
| 15 | 242 | 240 | 238 | 236 | 234 | | |
| 16 | 256 | 254 | 252 | 250 | 248 | 246 | |
| 17 | 270 | 268 | 266 | 264 | 262 | 260 | |
| 18 | 284 | 282 | 280 | 278 | 276 | 274 | 272 |
| 19 | 298 | 296 | 294 | 292 | 290 | 288 | 286 |
| 20 | 312 | 310 | 308 | 306 | 304 | 302 | 300 |
| 21 | 326 | 324 | 322 | 320 | 318 | 316 | 314 |
| 22 | 340 | 338 | 336 | 334 | 332 | 330 | 328 |
| 23 | 354 | 352 | 350 | 348 | 346 | 344 | 342 |
| 24 | 368 | 366 | 364 | 362 | 360 | 358 | 356 |
| 25 | 382 | 380 | 378 | 376 | 374 | 372 | 370 |
| 26 | 396 | 394 | 392 | 390 | 388 | 386 | 384 |
| 27 | 410 | 408 | 406 | 404 | 402 | 400 | 398 |
| 28 | 424 | 422 | 420 | 418 | 416 | 414 | 412 |
| 29 | 438 | 436 | 434 | 432 | 430 | 428 | 426 |
| 30 | 452 | 450 | 448 | 446 | 444 | 442 | 440 |
| 31 | 466 | 464 | 462 | 460 | 458 | 456 | 454 |
| 32 | 480 | 478 | 476 | 474 | 472 | 470 | 468 |
| 33 | 494 | 492 | 490 | 488 | 486 | 484 | 482 |

4. RESULTS AND DISCUSSION

Figure 1 and Figure 2 are graphs displaying the representative distribution of carbon numbers and Z families for the test substance before use and after completion of dose administration. The sum of all the normalized intensities equals 100%.

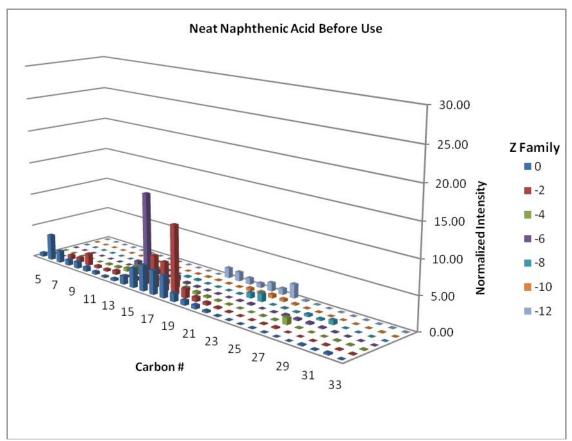


Figure 1: Representative Distribution of Carbon Numbers and Z Families of Neat Naphthenic Acid Before Use for Dose Administration

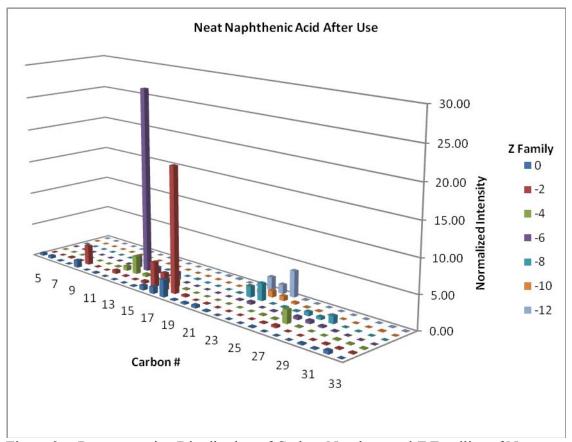


Figure 2: Representative Distribution of Carbon Numbers and Z Families of Neat Naphthenic Acid After Completion of Dose Administration

4.1. TEST ARTICLE CHARACTERIZATION AND STABILITY

The characterization of the test substance prior to use for dose administration is summarized in Table 1. Following 12 weeks of room temperature storage, the characterization assessment was repeated and is summarized in Table 2.

5. **CONCLUSION**

A GC/MS method was used to characterize neat naphthenic acid. Test substance samples were collected and analyzed prior to use for dose administration and after completion of dose administration 12 weeks later.

6. REPORT REVIEW AND APPROVAL

Report Approved by:

Staff Toxicologist, Developmental and Reproductive Toxicology Study Director 8 Dc-2010 Date

Report Prepared by:

Assistant Director, Analytical Chemistry

7. ABBREVIATIONS

The following abbreviations may apply to this report:

μ - micro

μL - microliter

ACN - acetonitrile

amu - atomic mass unit

AUC - area under the curve

cm - centimeter

conc. - concentration

DI - deionized

DMSO - dimethylsulfoxide

EPA - Environmental Protection Agency

ESI+ - positive electrospray ionization

FA - formic acid

FDA - Food and Drug Administration

FIFRA - Federal Insecticide, Fungicide and Rodenticide Act

g - gram

GLP - Good Laboratory Practices

GMP - Good Manufacturing Practices

HPLC - high performance liquid chromatography

hr - hour(s)

IS - internal standard

kg - kilogram

L - liter

LLOQ - lower limit of quantitation

M - molar

MC - methylcellulose

MeOH - methanol

mg - milligram

mL - milliliter

mm - millimeter

msec - milliseconds

MS - mass spectrometry

mM - millimolar

NA - not applicable

ND - not detected

ng - nanogram

nm - nanometer

ppm - parts per million

QC - quality control

Naphthenic Acid WIL-402010

American Petroleum Institute

%RE - percent relative error

RSD - relative standard deviation

SD - standard deviation

SOP - standard operating procedure

SPE - solid phase extraction

UV - ultraviolet

v - volume

w - weight

WIL - WIL Research Laboratories, LLC
WTDMSTM - WIL Toxicology Data Management System

TABLES 1 - 2

AN ORAL (GAVAGE) 14-DAY AND PILOT PRENATAL DEVELOPMENTAL TOXICITY STUDY OF NAPHTHENIC ACID IN RATS

Table 1: Before Use Characterization of Neat Test Substance (Analyzed 17 May 2010)

| | Pero | ent of Total Ion Intens | <u>stity</u> | |
|----------|-----------|-------------------------|--------------|-----------------|
| Carbon # | | (WIL ID no. 10006B) | | Mean % of Total |
| | Run # 002 | Run # 003 | Run # 004 | |
| 5 | 0.609 | 0.585 | 0.586 | 0.593 |
| 6 | 3.98 | 3.81 | 3.91 | 3.90 |
| 7 | 2.62 | 2.41 | 2.45 | 2.50 |
| 8 | 1.61 | 1.57 | 1.60 | 1.59 |
| 9 | 2.81 | 2.89 | 2.85 | 2.85 |
| 10 | 1.33 | 1.22 | 1.25 | 1.27 |
| 11 | 1.10 | 1.02 | 1.03 | 1.05 |
| 12 | 2.53 | 2.53 | 2.50 | 2.52 |
| 13 | 13.7 | 14.4 | 14.1 | 14.1 |
| 14 | 3.14 | 3.09 | 3.09 | 3.11 |
| 15 | 5.58 | 5.46 | 5.63 | 5.55 |
| 16 | 10.6 | 10.3 | 10.5 | 10.5 |
| 17 | 9.62 | 9.58 | 9.68 | 9.63 |
| 18 | 15.7 | 16.0 | 16.0 | 15.9 |
| 19 | 4.69 | 4.55 | 4.65 | 4.63 |
| 20 | 2.94 | 2.77 | 2.84 | 2.85 |
| 21 | 2.31 | 2.37 | 2.39 | 2.36 |
| 22 | 3.21 | 3.20 | 3.16 | 3.19 |
| 23 | 3.02 | 3.12 | 3.10 | 3.08 |
| 24 | 2.63 | 2.70 | 2.60 | 2.65 |
| 25 | 0.703 | 0.711 | 0.662 | 0.692 |
| 26 | 0.747 | 0.718 | 0.692 | 0.719 |
| 27 | 1.90 | 1.94 | 1.89 | 1.91 |
| 28 | 0.653 | 0.693 | 0.618 | 0.655 |
| 29 | 0.821 | 0.853 | 0.808 | 0.827 |
| 30 | 0.290 | 0.296 | 0.270 | 0.285 |
| 31 | 0.346 | 0.347 | 0.323 | 0.339 |
| 32 | 0.569 | 0.622 | 0.563 | 0.585 |
| 33 | 0.242 | 0.270 | 0.231 | 0.248 |

402010 Characterization data.xls Characterization Printed: 01Nov2010 8:48 AM

AN ORAL (GAVAGE) 14-DAY AND PILOT PRENATAL DEVELOPMENTAL TOXICITY STUDY OF NAPHTHENIC ACID IN RATS

Table 2: After Use Characterization of Neat Test Substance (Analyzed 6 August 2010)

| | Per | cent of Total Ion Inten | sity | |
|----------|-----------|-------------------------|-----------|-----------------|
| Carbon # | | (WIL ID no. 10006B) | | Mean % of Total |
| | Run # 010 | Run # 011 | Run # 012 | |
| 5 | 0.375 | 0.375 | 0.388 | 0.379 |
| 6 | 0.723 | 0.595 | 0.408 | 0.576 |
| 7 | 0.441 | 0.330 | 0.234 | 0.335 |
| 8 | 0.187 | 0.174 | 0.146 | 0.169 |
| 9 | 4.06 | 4.06 | 4.04 | 4.05 |
| 10 | 0.125 | 0.106 | 0.0768 | 0.103 |
| 11 | 0.268 | 0.250 | 0.230 | 0.250 |
| 12 | 2.97 | 3.00 | 2.95 | 2.97 |
| 13 | 28.7 | 29.3 | 30.5 | 29.5 |
| 14 | 1.62 | 1.53 | 1.39 | 1.51 |
| 15 | 1.15 | 1.02 | 0.796 | 0.988 |
| 16 | 6.73 | 6.61 | 6.39 | 6.58 |
| 17 | 5.62 | 5.39 | 5.21 | 5.41 |
| 18 | 20.7 | 21.0 | 21.5 | 21.1 |
| 19 | 0.738 | 0.618 | 0.525 | 0.627 |
| 20 | 0.312 | 0.273 | 0.234 | 0.273 |
| 21 | 1.49 | 1.50 | 1.48 | 1.49 |
| 22 | 4.45 | 4.49 | 4.46 | 4.47 |
| 23 | 5.19 | 5.19 | 5.20 | 5.19 |
| 24 | 4.51 | 4.54 | 4.58 | 4.54 |
| 25 | 0.738 | 0.720 | 0.663 | 0.707 |
| 26 | 1.06 | 1.15 | 0.966 | 1.06 |
| 27 | 3.40 | 3.45 | 3.44 | 3.43 |
| 28 | 1.01 | 1.01 | 0.966 | 0.993 |
| 29 | 1.39 | 1.37 | 1.35 | 1.37 |
| 30 | 0.364 | 0.360 | 0.307 | 0.344 |
| 31 | 0.430 | 0.421 | 0.392 | 0.414 |
| 32 | 0.940 | 0.917 | 0.877 | 0.911 |
| 33 | 0.297 | 0.292 | 0.291 | 0.293 |

402010 Characterization data.xls Characterization (3)

Printed: 01Nov2010 8:48 AM

APPENDIX C

Material Safety Data Sheet (Sponsor-Provided Data)

| REVI | SION 4: March 28, 2008 | MATERIAL SAFETY DATA SHEET | Page 1/6 |
|------|-----------------------------|--|----------|
| 1 | IDENTIFICATION OF TH | PRODUCT AND OF THE COMPANY | |
| 1.1 | Identification of the Produ | Naphthenic Acids (Carboxylic Acids, Fatty Acids) | |
| 1.2 | Product Code: | NAP ACID | |
| 1.3 | Company: | | |
| | | | |
| 1.4 | Transportation Emergency | USA 1-800-424-9300 (CHEMTREC) | |
| 1.5 | Product Information: | | |
| 1.6 | Intended Use : | For industrial use only. No other use is intended. | |
| 2 | HAZARDS IDENTIFICATI | 1 | |
| 2.1 | Classification: | ritant (XI). Harmful (Xn). | |
| 2.2 | Warning Statements: | auses eye and skin irritation. armful if swallowed – may enter lungs if swallowed or vomited. | |

| 2 | HAZARDS IDENTIFICA | TION | | |
|-----|--------------------------------------|---|---|--------------------------------|
| 2.1 | Classification: | Irritant (XI). Harmful (Xn). | | residente. |
| 2.2 | Warning Statements: | Causes eye and skin irritation. Harmful if swallowed – may enter lungs if swallow High vapor concentrations may cause drowsines tract. | ed or vomited. s and irritation of the e | yes or respiratory |
| 2.3 | Hazard Symbol(s): | | | |
| 2.4 | Risk Phrase(s): | R36/38 (Irritating to eyes and skin). R65 (Harmful: May cause lung damage if swallow | | |
| 2.3 | Potential Health Effects: | | | |
| | Eye Contact: | Causes eye irritation. Exposure may cause irritati | on, redness and tearing |). |
| | Skin Contact: | Causes skin irritation. Exposure may cause redne | ess, itching and inflamm | ation. |
| | Ingestion: | Expected to be a low ingestion hazard. Aspiratio and cause damage, | n hazard. If swallowed | , can enter lungs |
| | Inhalation: | High vapor concentrations may cause drowsine asthmatic breathing and breathlessness, | ss, respiratory tract irri | |
| | Chronic Effects: | No known deleterious effects. | N. S. DIGHES BADA | e bestructor |
| 2.4 | Other Hazards: | Target organs: Eyes, skin and central nervous sys | tem. | Same and the second |
| 3 | COMPOSITION/INFORM | MATION ON INGREDIENTS | | |
| | ince enic Acid eum Distillates | <u>CAS No.</u> 1338-24-5 8008-20-6 | EC No. 215-662-8 232-366-4 | % Present 70 - 99 1 - 30 |

| REVISION 4: March 28, 2008 | MATERIAL SAFETY DATA SHEET | Page 2/6 |
|----------------------------|--|----------|
| | Naphthenic Acids (Carboxylic Acids, Fatty Acids) | |

| 4 | FIRST-AI | D MEASURE | <u>ES</u> | |
|-------|-----------------|-----------------------|--------------|---|
| In ca | se of contact | with eyes: | Promptly f | lush eyes with running water for 15 minutes, including under eyelids. Seek medical all irritation develops. |
| In ca | se of contact | with skin: | Wash affe | cted area well with soap and water. Seek medical assistance if irritation develops. |
| In ca | se of ingestion | n: | poison cor | duce vorniting. Give 2-3 glasses of milk or water to dilute. Contact physician or ntrol center promptly for instructions. If vomiting occurs, keep head lower than hips event aspiration. Never give anything by mouth to an unconscious person. |
| In ca | se of inhalatio | n: | Remove to | o fresh air. Seek medical assistance if irritation develops. |
| 5 | FIRE-FIG | HTING MEA | SURES | |
| 5.1 | Suitable e | xtinguishing | media: | Water fog, fire fighting foam, dry chemical or carbon dioxide. |
| 5.2 | Unsuitable | extinguishir | ng media: | None |
| 5.3 | Specific h | azards: | | Combustion products are Carbon Oxides. |
| 5.4 | Personal | protective ed | quipment: | Wear Self Contained Breathing Apparatus and protective clothing appropriate for fire-fighting. |
| 5.5 | Other pred | cautions: | | Non-emergency personnel should be removed from the area immediately. Cool fire-exposed containers with water spray. Prevent water runoff from reaching drains, surface water and ground water. |
| 6 | ACCIDEN | TAL RELEA | SE MEASU | IRES |
| 6.1 | Personal r | orecautions: | | Avoid unnecessary exposure by wearing personal protective equipment specified in Section 8. Remove material from eyes, skin and clothing. |
| 6.2 | Spill clean | <u>up:</u> | | Suction up free liquids using non-sparking equipment: Liquid unable to be suctioned may be absorbed with a non-combustible material (vermiculite, sand, earth, etc.) and transferred to container(s) for later disposal. Remove contaminated sand, earth, etc. and transfer to container(s) for later disposal. |
| 6.3 | Environme | ental precauti | ions: | Keep away from drains, surface water and ground water. |
| 7 | HANDLING | G AND STOR | RAGE | |
| 7.1 | Handling: | Wear appr | opriate per | sonal protective equipment (see Section 8). |
| | | Avoid brea | thing vapo | rs, mists or spray. Use with adequate ventilation. |
| | | Avoid cont | act with ey | es, skin and clothing. |
| | | Wash thor | oughly afte | r handling. |
| | | Do not tas | te or swallo | w. |
| 7.2 | Storage: | Store in a and flame. | sealed con | stainer in a clean, dry, well-ventilated area away from oxidizers, strong bases, heat |
| | | Avoid use | of copper a | and brass alloys in storage and transfer equipment and process equipment. |
| | | | | |

| REVISION | 4: Mai | rch 28, | 2008 |
|----------|--------|---------|------|
|----------|--------|---------|------|

MATERIAL SAFETY DATA SHEET

Page 3/6

Naphthenic Acids (Carboxylic Acids, Fatty Acids)

| 8 | EXPOSURE CONTROLS/PI | ERSONAL PROTECTION |
|-----|-------------------------|--|
| 8.1 | Engineering controls: | Use local exhaust ventilation to control emissions at source. |
| 8.2 | Eye/face protection: | Wear safety glasses with side shields as minimum protection. Wear goggles or faceshield if a risk of splashing exists. |
| 8.3 | Skin protection: | Wear chemical resistant gloves. Heavy PVC, butyl rubber or Viton are recommended. |
| 8.4 | Respiratory protection: | An approved respirator must be worn if engineering controls do not maintain airborne concentrations below established exposure limits or, when limits have not been established, below irritant levels. Respirator selection must be based upon the airborne concentration. Consult a health and safety professional or manufacturer for specific recommendations: |
| 8.5 | Thermal hazards: | None |

8.6 Occupational Exposure Levels

| Chemical Name | Source | Туре | Exposure Limits | Notes |
|---|--------------------------|---------|-----------------------|------------------------------------|
| Cherosene (Non-Aerosol), Come Vapore totale Dell'Idrocarburo | Italy OEL's | TWA | 200 mg/m ³ | Skin Total Hydrocarbon Vapor |
| Kerosene | Poland MAC's | TWA | 100 mg/m ³ | |
| | Poland MAC's | STEL | 300 mg/m ³ | |
| Kerosine | Russian Federation MAC's | Ceiling | 300 mg/m ³ | As C |
| | Russian Federation MAC's | TWA | 600 mg/m ³ | As C |
| Kerosene (Non-Aerosol), As Total Hydrocarbon Vapor | ACGIH | TWA | 200 mg/m ³ | Irritation, CNS, Skin |
| Kerosene | NIOSH | REL | 100 mg/m ³ | |

| 9 PHYSICAL AND O | CHEMICAL PROPERTIES | | |
|-----------------------------|-------------------------------|---|-----------------------------|
| 9.1 Appearance: | Amber color | 9.2 Odour: | Hydrocarbon |
| 9.3 <u>pH</u> : | 5.2 (Saturated Solution) | 9.4 Boiling Pt./range: | 268°C (515°F) |
| 9.5 Freezing Pt./Range: | Not established | | |
| 9.6 Flash point: | >149°C (300°F) | | |
| 9.7 Flammability: | See 9.6 | 9.8 Autoflammability: | See 9.6 |
| 9.9 Explosive properties: | Not applicable | 9.10 Oxidizing properties: | Not an oxidizer |
| 9.11 Vapor pressure: | 0.005 mm Hg (37.8°C/100°F) | 9.12 Relative density (H ₂ O = 1): | 0.960 - 0.982 (15.6°C/60°F) |
| 0.13 Apparent density: | Not applicable | 9.14 Vapor density (Air = 1); | 6.5 |
| 0.14 Solubility: | Fat (type) - No | % by weight (15.6°C/60°F) at determined at determined | |
| 0.15 Partition coefficient: | Log Po/w (Octanol/water) | - Not determined | |
| 3.16 Other data: | Not Applicable | | |

| | Naphthenic Acids (Carboxylic Acids, Fatty Acids) | |
|----------------------------|--|----------|
| REVISION 4: March 28, 2008 | MATERIAL SAFETY DATA SHEET | Page 4/6 |

| 10 | STABILI' | TY AND REACTIVITY | | | | |
|------|---|--|---|--|--|--|
| 10.1 | Reactivit | A: | Not reactive under specified conditions of storage, shipment and use. | | | |
| 10.2 | Stability: | | Stable under specified conditions of storage, shipment and use. | | | |
| 0.3 | Condition | ns to avoid: | Heat and ignition sources. | | | |
| 0.4 | Incompat | tible materials: | Strong oxidizers and strong bases. | | | |
| 0.5 | Hazardou | us decomposition products: | Carbon Oxides. | | | |
| | | XICOLOGICAL INFORMATION | | | | |
| 1 | TOXICO | LOGICAL INFORMATION | | | | |
| | TOXICOI | | established as a respiratory tract irritant. May cause lung damage if aspirated | | | |
| | 200000000000000000000000000000000000000 | Eye and skin irritant. Not e | established as a respiratory tract irritant. May cause lung damage if aspirated | | | |
| | 200000000000000000000000000000000000000 | Eye and skin irritant. Not e | established as a respiratory tract irritant. May cause lung damage if aspirated Test Results Dermal LD ₅₀ (Rabbit): >2000 mg/kg | | | |
| | 200000000000000000000000000000000000000 | Eye and skin irritant. Not e Specified Substances Chemical Name | Test Results | | | |
| | 200000000000000000000000000000000000000 | Eye and skin irritant. Not e Specified Substances Chemical Name | Test Results Dermal LD ₅₀ (Rabbit): >2000 mg/kg | | | |
| | 200000000000000000000000000000000000000 | Eye and skin irritant. Not e Specified Substances Chemical Name | Test Results Dermal LD ₅₀ (Rabbit): >2000 mg/kg Oral LD ₅₀ (Rat): >5000 mg/kg | | | |
| | 200000000000000000000000000000000000000 | Eye and skin irritant. Not e Specified Substances Chemical Name Kerosene | Test Results Dermal LD ₅₀ (Rabbit): >2000 mg/kg Oral LD ₅₀ (Rat): >5000 mg/kg Inhalation LC ₅₀ : >5000 mg/m³, 4 H | | | |
| | 200000000000000000000000000000000000000 | Eye and skin irritant. Not e Specified Substances Chemical Name | Test Results Dermal LD ₅₀ (Rabbit): >2000 mg/kg Oral LD ₅₀ (Rat): >5000 mg/kg Inhalation LC ₅₀ : >5000 mg/m³, 4 H Skin (Rabbit): 500 mg (Severe Irritation) Skin (Rabbit): 100%/24 H (Moderate Irritation) Oral LD ₅₀ (Rat): 3000 mg/kg | | | |
| 1 | 200000000000000000000000000000000000000 | Eye and skin irritant. Not e Specified Substances Chemical Name Kerosene | Test Results Dermal LD ₅₀ (Rabbit): >2000 mg/kg Oral LD ₅₀ (Rat): >5000 mg/kg Inhalation LC ₅₀ : >5000 mg/m³, 4 H Skin (Rabbit): 500 mg (Severe Irritation) Skin (Rabbit): 100%/24 H (Moderate Irritation) | | | |

11.2 Neither ingredient is listed by NTP, IARC or OSHA as a carcinogen. Kerosene (Non-Aerosol), as total hydrocarbon vapor, is listed by ACGIH as A3 (Confirmed Animal Carcinogen).

Eye (Rabbit): Moderate

Skin (Rabbit): Slight

Skin Occluded (Rabbit): Moderate to Severe

12 ECOLOGICAL INFORMATION 12.1 Ecotoxicity: No data available. 12.2 Persistence and degradability: No data available. 12.3 Bioaccumulation potential: No data available. 12.4 Mobility in soil: No data available. 12.5 Other adverse effects: No data available.

13 DISPOSAL CONSIDERATIONS

Generators of waste material are responsible for evaluating materials for compliance with all applicable procedures and regulations. Disposal of unused materials must be in accordance with all local, state and federal regulations. Containers should be cleaned of residual product and rinsed according to all local, state and federal regulations prior to disposal.

REVISION 4: March 28, 2008

MATERIAL SAFETY DATA SHEET

Page 5/6

Naphthenic Acids (Carboxylic Acids, Fatty Acids)

14 TRANSPORT INFORMATION

| | UN Number | Proper Shipping Name | Hazard Class(es) | Packing Group |
|-----------|---------------|--|------------------|---------------|
| ADR/RID: | Not regulated | | , , | |
| IMO/IMDG: | Not regulated | Fatty Acids (Saturated C13+) | | |
| IATA: | Not regulated | | | |
| DOT: | NA3082 | Other Regulated Substances, Liquid, n.o.s., (Naphthenic Acid) | 9 | III |

Note: Material is regulated by DOT only if shipped in a container containing an amount equal to or greater than the Reportable Quantity (RQ) of 100-pounds.

15 REGULATORY INFORMATION

Warning symbol:

×

Warning words:

Harmful. Irritant.

Risk phrases:

R36/38: Irritating to eyes and skin

R65: May cause lung damage if swallowed

Safety phrases:

S23: Do not breathe vapor

S24/25: Avoid contact with eyes and skin

S62: If swallowed, do not induce vomiting. Seek medical advice immediately and show this container or label

HMIS ratings (estimated):



NFPA ratings (estimated):



SARA:

Section 302:

None

Section 311/312: Section 313: Immediate Health Hazard

None

WHMIS:

D2B

Inventories:

CAS Number 1338-24-5 8008-20-6 TSCA Yes DSL Yes

EINECS Yes Yes REVISION 4: March 28, 2008

MATERIAL SAFETY DATA SHEET

Page 6/6

Naphthenic Acids (Carboxylic Acids, Fatty Acids)

16 OTHER INFORMATION

Revision Date:

March 28, 2008

Supercedes Revision Date:

November, 2004

Revisions:

The latest informational changes are indicated by 20% shading.

The information on this form is furnished solely for the purpose of enabling those who transport, handle or use our products to ensure the safety and health of their employees and to comply and local). This information is offered in good faith and is believed to be accurate.

o, nowever, makes no guarantee or warranty, expressed or implied, regarding the accuracy or these data or the results to be obtained from the use hereof.

171

GUIDE SUBSTANCES (LOW TO MODERATE HAZARD)

ERG2004

POTENTIAL HAZARDS

FIRE OR EXPLOSION

- · Some may burn but none ignite readily.
- · Containers may explode when heated.
- · Some may be transported hot.

HEALTH

- · Inhalation of material may be harmful.
- · Contact may cause burns to skin and eyes.
- · Inhalation of Asbestos dust may have a damaging effect on the lungs.
- · Fire may produce irritating, corrosive and/or toxic gases.
- Some liquids produce vapors that may cause dizziness or suffocation.
- · Runoff from fire control may cause pollution.

PUBLIC SAFETY

- · CALL Emergency Response Telephone Number on Shipping Paper first. If Shipping Paper not available or no answer, refer to appropriate telephone number listed on the inside back cover.
- · As an immediate precautionary measure, isolate spill or leak area in all directions for at least 50 meters (150 feet) for liquids and at least 25 meters (75 feet) for solids.
- · Keep unauthorized personnel away.
- · Stay upwind.

PROTECTIVE CLOTHING

- · Wear positive pressure self-contained breathing apparatus (SCBA).
- · Structural firefighters' protective clothing will only provide limited protection.

EVACUATION

Spill

 See the Table of Initial Isolation and Protective Action Distances for highlighted substances. For non-highlighted substances, increase, in the downwind direction, as necessary, the isolation distance shown under "PUBLIC SAFETY"

. If tank, rail car or tank truck is involved in a fire, ISOLATE for 800 meters (1/2 mile) in all directions; also, consider initial evacuation for 800 meters (1/2 mile) in all directions.

ERG2004

SUBSTANCES (LOW TO MODERATE HAZARD) GUIDE

EMERGENCY RESPONSE

FIRE

Small Fires

· Dry chemical, CO, water spray or regular foam.

Large Fires

- · Water spray, fog or regular foam.
- · Move containers from fire area if you can do it without risk
- · Do not scatter spilled material with high pressure water streams.
- · Dike fire-control water for later disposal.

Fire involving Tanks

- · Cool containers with flooding quantities of water until well after fire is out.
- · Withdraw immediately in case of rising sound from venting safety devices or discoloration of tank.
- · ALWAYS stay away from tanks engulfed in fire.

SPILL OR LEAK

- · Do not touch or walk through spilled material.
- · Stop leak if you can do it without risk.
- · Prevent dust cloud.
- · Avoid inhalation of asbestos dust.

Small Dry Spills

· With clean shovel place material into clean, dry container and cover loosely; move containers from spill area.

Small Spills

· Take up with sand or other non-combustible absorbent material and place into containers for later disposal.

Large Spills

- · Dike far ahead of liquid spill for later disposal.
- · Cover powder spill with plastic sheet or tarp to minimize spreading.
- · Prevent entry into waterways, sewers, basements or confined areas.

FIRSTAID

- Move victim to fresh air.
 Call 911 or emergency medical service.
- · Give artificial respiration if victim is not breathing.
- · Administer oxygen if breathing is difficult.
- · Remove and isolate contaminated clothing and shoes.
- In case of contact with substance, immediately flush skin or eyes with running water for at least 20 minutes.
- · Ensure that medical personnel are aware of the material(s) involved and take precautions to protect themselves.

Page 290

Page 291





APPENDIX D

Analyses of Dosing Formulations (WIL Research Laboratories, LLC)

An Oral (Gavage) 14-Day and Pilot Prenatal Developmental Toxicity Study of Naphthenic Acid in Rats

Analyses of Dosing Formulations

Analytical Chemistry Department

WIL Research Laboratories, LLC



TABLE OF CONTENTS

| | | <u>Page</u> |
|-------|--|-------------|
| | Table of Contents | 2 |
| | Index of Figures | 3 |
| | Index of Tables | 3 |
| | Index of Attachments | 4 |
| 1. | Summary | 5 |
| 2. | Introduction | 6 |
| 3. | Key Study Personnel | 6 |
| 4. | Experimental Procedures - Materials and Methods | 7 |
| 4.1. | Internal Standard Identification | 7 |
| 4.2. | Gas Chromatography | 7 |
| 4.3. | Mass Spectrometry | 8 |
| 4.4. | Preparation of Internal Standard Stock Solution | 8 |
| 4.5. | Preparation of Diluent | 8 |
| 4.6. | Preparation of Calibration Stock Solution and Samples | 8 |
| 4.7. | Preparation of Quality Control Stock Solution and Samples | 9 |
| 4.8. | Processing of Formulation Samples | 10 |
| 4.9. | Sample Processing | 11 |
| 4.10. | Calibration and Quantitation | 11 |
| 5. | Results and Discussion | 12 |
| 5.1. | Specificity/Selectivity | 17 |
| 5.2. | Assay Validation: Calibration Reproducibility | 17 |
| 5.3. | Assay Validation: Precision and Accuracy | 18 |
| 5.4. | Assay Ruggedness | 18 |
| 5.5. | Assay Acceptability | 18 |
| 5.6. | Test Article Homogeneity and Resuspension Homogeneity Assessment of Formulations | 19 |
| 5.7. | Test Article Stability in Formulations | 20 |

| | <u>P</u> | age |
|-----------|---|-----|
| 5.8. Te | est Article Concentration in Formulations | .20 |
| 6. Co | onclusion | .21 |
| 7. Re | eport Review and Approval | .22 |
| 8. Al | obreviations | .23 |
| | INDEX OF FIGURES | |
| Figure 1: | Representative Chromatogram of a 150 µg Naphthenic Acid/mL Calibration Sample | .13 |
| Figure 2: | Representative Chromatogram of a Processed 125 mg Naphthenic Acid/mL Quality Control Sample | .14 |
| Figure 3: | Representative Chromatogram of a Processed 100 mg Naphthenic Acid/mL Formulation Sample | .15 |
| Figure 4: | Chromatogram of a Processed Vehicle Blank Sample | .16 |
| | INDEX OF TABLES | |
| Table 1: | Back-Calculated Concentrations of the Validation Calibration Standards | .26 |
| Table 2: | Calculated Concentrations of the Validation Quality Control Samples | .27 |
| Table 3: | Homogeneity Assessment of the 27 May 2010 Formulations | .28 |
| Table 4: | 11-Day Room Temperature Resuspension and Stability Analysis of the 27 May 2010 Formulations | .29 |
| Table 5: | Concentration Assessment of the 1 June 2010 Formulations | .30 |
| Table 6: | Concentration Assessment of the 16 June 2010 Formulations (Back up Samples) | .31 |

INDEX OF ATTACHMENTS

| | | <u>Page</u> |
|----|-------------------------|-------------|
| I: | Certificate of Analysis | 32 |

1. SUMMARY

A gas chromatography method using mass spectrometric detection for the determination of naphthenic acid concentration in corn oil formulations containing test substance ranging in concentration from 2.50 to 125 mg/mL was validated in this study. In addition, test article homogeneity and, following 11 days of room temperature storage, resuspension homogeneity and stability were assessed in formulations prepared at target concentrations of 3 and 100 mg naphthenic acid/mL. Finally, formulations used for dose administration were analyzed to verify test article concentration acceptability.

The naphthenic acid assay procedure was validated in this study with 3 validation sessions. Quantitation was performed using matrix-based calibration samples (*i.e.*, containing corn oil) ranging in test article concentration from 150 to 350 µg/mL. The mean back-calculated calibration sample concentrations had inter-session variability ranging from 2.0% to 6.8% relative standard deviation (RSD) and percent relative error (%RE) ranging from -1.3% to 2.3%, which met the WIL standard operating procedure (SOP) acceptance criteria for calibration samples, *i.e.*, RSD \leq 10% and %RE within \pm 10% (except at the lowest level where RSD \leq 15% and %RE within \pm 15% were acceptable). Assay precision and accuracy were verified by the analysis of QC samples prepared at 2.50, 25.0, and 125 mg naphthenic acid/mL. The mean calculated QC concentrations had inter-session variability (precision) ranging from 4.5% to 6.9% RSD and %RE (accuracy) ranging from -6.3% to -3.4%. The results met the WIL SOP acceptance criteria for precision and accuracy, *i.e.*, RSD \leq 15% and %RE within \pm 15%.

The results of the test article homogeneity assessment in formulations prepared at target concentrations of 3 and 100 mg naphthenic acid/mL met the WIL SOP acceptance criteria, *i.e.*, the RSD for the mean concentration was ≤10% at a concentration within the acceptable limits (85% to 115% of target). Assessment of test article resuspension homogeneity and stability in formulations prepared at target concentrations of 3 and 100 mg naphthenic acid/mL and following 11 days of room temperature storage met the

WIL SOP acceptance criteria for resuspension homogeneity, *i.e.*, the RSD for the mean concentration was 10% or less, and stability, *i.e.*, the post-storage concentration was not <90% of the pre-storage value.

The analyzed formulations used for dose administration met the WIL SOP requirement for concentration acceptability for suspension formulations, *i.e.*, the analyzed concentration was 85% to 115% of the target concentration. No quantifiable amount of test article was detected in the analyzed vehicle administered to the control group.

2. Introduction

This report provides a detailed description and validation of a gas chromatography (GC) method using mass spectrometric detection for the determination of naphthenic acid concentration in corn oil formulations containing test article ranging in concentration from 2.50 to 125 mg/mL. Assay specificity/selectivity, calibration reproducibility, precision, accuracy, and ruggedness were assessed. In addition, formulations prepared at target concentrations of 3 and 100 mg naphthenic acid/mL were analyzed to assess test article homogeneity and, following 11 days of room temperature storage, resuspension homogeneity and stability. Finally, formulations used for dose administration were analyzed to verify test article concentration acceptability.

A list of abbreviations that may have been used in this report is located in Section 8. (Abbreviations).

3. KEY STUDY PERSONNEL



Assistant Director, Analytical Chemistry Chemist III, Analytical Chemistry Chemist II, Analytical Chemistry Data Coordinator, Reporting & Technical Support Services Group Manager, Reporting & Technical Support Services

4. EXPERIMENTAL PROCEDURES - MATERIALS AND METHODS

4.1. Internal Standard Identification

The internal standard (IS) used for calibration standards, QC samples, and formulation samples was received from Sigma Aldrich, St. Louis, MO, on 15 April 2010 as follows:

| Identification | Quantity Received | Physical Description |
|---|------------------------------|----------------------|
| Fluorine-9-carboxylic acid, 97% (9-FCA) Lot No. 03706KH [WIL log no. ACC 13364] | 1 Bottle Net weight: 10 g | Light Yellow Powder |

A Certificate of Analysis for the IS was provided by the supplier and is presented in Attachment I. The IS was stored at room temperature.

4.2. GAS CHROMATOGRAPHY

Instrument: Varian CP-3800 gas chromatograph (GC) equipped with a

Varian 4000 ion-trap mass spectrometer (MS), an

autosampler, and Varian MS Workstation data system, or

equivalent.

Column: Varian VF-5MS column, $30 \text{ m} \times 0.25 \text{ mm ID}$,

0.25-µm film-thickness.

Temperature Program: Initial temperature 100°C, hold for 3.0 minutes.

Ramp at 20°C/min to 300°C, hold for 2.0 minutes

Carrier Gas: Helium
Carrier Gas Pressure: 10.0 psi
Injector Temperature 250°C

Injection Volume: 1.0 μL splitless
Purge Flow: 50 psi at 1.0 min

Retention Time: Approximately 10.3 minutes for naphthenic acid

Approximately 11.9 minutes for 9-FCA (IS)

Run Time: 15 minutes

4.3. MASS SPECTROMETRY

Acquisition Parameters

Solvent Delay: 5.0 minutes

Ion Source: Electron Impact

Scan Range: 250-350 m/z

Transfer Line Temperature: 280°C
Trap Temperature: 210°C
Manifold Temperature: 60°C

| Analyte | Mass (m/z) |
|-----------------|------------|
| Naphthenic Acid | 257.3 |
| 9-FCA | 267.4 |

4.4. PREPARATION OF INTERNAL STANDARD STOCK SOLUTION

The internal standard stock solution was prepared at a concentration of 0.100 mg 9-fluorenecarboxylic acid (9-FCA)/mL as follows. Approximately 10 mg of 9-FCA (Sigma Aldrich, St Louis, MO) was accurately weighed in a tared glass weigh funnel and transferred to a 100-mL volumetric flask with rinses of methylene chloride. Approximately 75 mL of methylene chloride was added to the flask, and the preparation was mixed and sonicated as needed to achieve complete dissolution of the internal standard. Additional methylene chloride was added to yield the desired concentration and the solution was thoroughly mixed.

4.5. PREPARATION OF DILUENT

The diluent was prepared by thoroughly mixing methylene chloride and corn oil in a 90:10 (v/v) ratio.

4.6. Preparation of Calibration Stock Solution and Samples

The calibration stock solution was prepared at a concentration of 1.00 mg naphthenic acid/mL as follows. Approximately 50 mg of naphthenic acid

(WIL ID. 10006B, no correction for purity) was accurately weighed in a tared glass weigh funnel and transferred to a 50-mL volumetric flask with rinses of methylene chloride. Approximately 40 mL of methylene chloride was added to the flask, and the preparation was mixed and sonicated as needed to achieve complete dissolution of the test substance. Additional methylene chloride was added to yield the desired concentration, and the solution was thoroughly mixed.

As detailed in the following table, matrix-based calibration samples were prepared by combining aliquots of the calibration stock solution, corn oil (vehicle), and methylene chloride in polypropylene tubes. The tubes were capped and mixed with vortex action. The samples were further processed as described in Section 4.9. (Sample Processing). Triplicate calibration samples at each concentration were prepared for the validation sessions; at least single calibration samples at each concentration were prepared for routine analyses.

| Test Substance Concentration (µg/mL) | Calibration Stock Volume (mL) | Methylene Chloride Volume (mL) | Corn Oil Volume (mL) |
|--|-------------------------------------|--------------------------------------|----------------------------|
| 150 | 0.150 | 0.750 | 0.100 |
| 200 | 0.200 | 0.700 | 0.100 |
| 250 | 0.250 | 0.650 | 0.100 |
| 300 | 0.300 | 0.600 | 0.100 |
| 350 | 0.350 | 0.550 | 0.100 |

4.7. PREPARATION OF QUALITY CONTROL STOCK SOLUTION AND SAMPLES

The quality control stock solution was prepared at a concentration of 125 mg naphthenic acid/mL as follows. Approximately 0.625 g of naphthenic acid (WIL ID. 10006B, no correction for purity) was accurately weighed in a tared glass weigh funnel and transferred to a 5-mL volumetric flask with rinses of methylene chloride. Approximately 2 mL of methylene chloride was added, and the preparation was

mixed and sonicated as needed to achieve complete dissolution of the test substance. Additional methylene chloride was added to yield the desired concentration and the solution was thoroughly mixed.

As detailed in the following table, QC samples were prepared to simulate the processing of formulations at concentrations of 2.50, 25.0, and 125 mg/mL (nominal QC concentration) by combining aliquots of the QC stock solution, corn oil (vehicle), and methylene chloride in polypropylene tubes. The samples were mixed with vortex action and/or inversion. The samples were further diluted with diluent as necessary. The samples were processed as described in Section 4.9. (Sample Processing). Triplicate QC samples at each concentration were prepared; a single blank sample was prepared.

| QC Level | Nominal QC Concentration (mg/mL) | Vehicle Volume (mL) | QC Stock Volume (mL) | Methylene Chloride Volume (mL) | Secondary Dilution | Theoretical Final Concentration (µg/mL) |
|-------------|--|---------------------------|-------------------------------|---|-----------------------|--|
| Blank | 0 | 1.0 | 0 | 9.00 | NA | 0 |
| 1 | 2.50 | 1.0 | 0.020 | 8.98 | NA | 250 |
| 2 | 25.0 | 1.0 | 0.200 | 8.80 | 10-fold | 250 |
| 3 | 125 | 1.0 | 1.00 | 8.00 | 50-fold | 250 |

NA = Not applicable

4.8. Processing of Formulation Samples

Quadruplicate formulations samples were collected using a syringe and cannula and placed in polypropylene tubes. Two samples from each quadruplicate set were processed for analysis and the remaining 2 samples (back-up samples) were stored frozen (approximately -20°C) and discarded, if not processed for analysis, after the Study Director's approval of the results. As detailed in the following table, methylene chloride was added to the formulations samples and the samples were mixed with vortex action and/or inversion. The samples were further diluted with diluent as necessary. The samples were further processed as described in Section 4.9. (Sample Processing).

| Sample Concentration (mg/mL) | Sample Volume (mL) | Methylene Chloride Volume (mL) | Secondary Dilution | Theoretical Final Concentration (µg/mL) |
|------------------------------------|--------------------------|--------------------------------------|-----------------------|---|
| 0 | 1.00 | 11.0 | NA | 0 |
| 3 | 1.00 | 11.0 | NA | 250 |
| 10 | 1.00 | 39.0 | NA | 250 |
| 30 | 1.00 | 39.0 | 3-fold | 250 |
| 100 | 1.00 | 19.0 | 20-fold | 250 |

NA = Not applicable

4.9. <u>Sample Processing</u>

Calibration, QC, and formulation samples were processed by combining $50.0~\mu L$ of each sample, $10.0~\mu L$ of the IS stock solution, and $50.0~\mu L$ of N-methyl-N-(t-butyldimethylsiyl)trifluoroacetamide (MTBSTFA) containing 1% t-butyldimethylsilylchloride in a glass conical autosampler vial. The samples were capped and mixed with vortex action for approximately 20~minutes at approximately $60^{\circ}C$. The caps were removed and the samples were evaporated to dryness under a stream of nitrogen. The samples were reconstituted with 0.500~mL of methylene chloride and mixed with vortex action.

4.10. CALIBRATION AND QUANTITATION

Single injections were made of each calibration, QC, and formulation sample. A calibration curve was constructed for each set of analyses. The naphthenic acid/IS peak area ratios (y) and the theoretical concentrations (x) of the calibration samples were fit with least-squares regression to the quadratic function:

$$y = ax^2 + bx + c$$

Concentrations were back-calculated from the results of the regression analysis using Microsoft[®] Excel. The concentration data were transferred to another Excel spreadsheet, where appropriate summary statistics, *i.e.*, mean, standard deviation (SD), relative standard deviation (RSD), percent relative error (%RE), and concentration as a percent of

WIL-402010 Naphthenic Acid American Petroleum Institute

target, were calculated and presented in tabular form. The concentrations of the formulation and QC samples were calculated by applying any necessary multiplication factors to correct for dilution and/or unit conversions.

5. RESULTS AND DISCUSSION

Under the described chromatographic conditions, the retention time of the test substance and IS were approximately 10.3 and 11.9 minutes, respectively. Figure 1, Figure 2, Figure 3, and Figure 4 are typical chromatograms of a calibration sample, a processed QC sample, a processed formulation sample, and a processed vehicle blank sample, respectively. The total analysis time required for each run was 15 minutes.

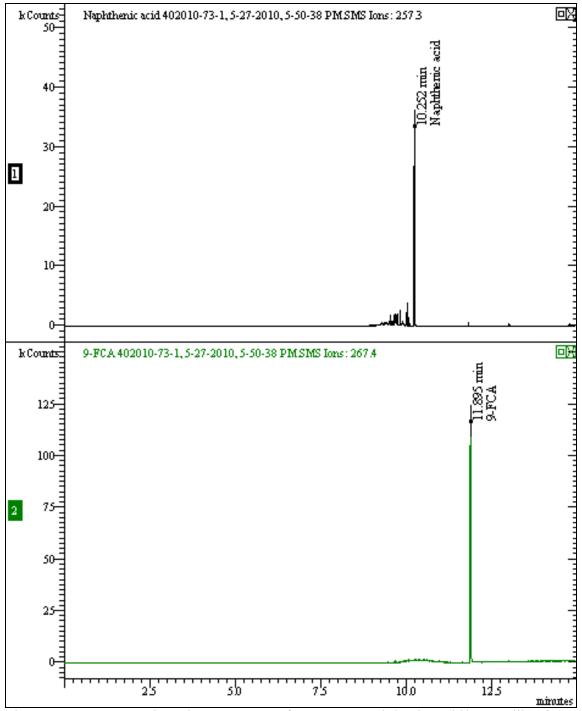


Figure 1: Representative Chromatogram of a 150 μg Naphthenic Acid/mL Calibration Sample

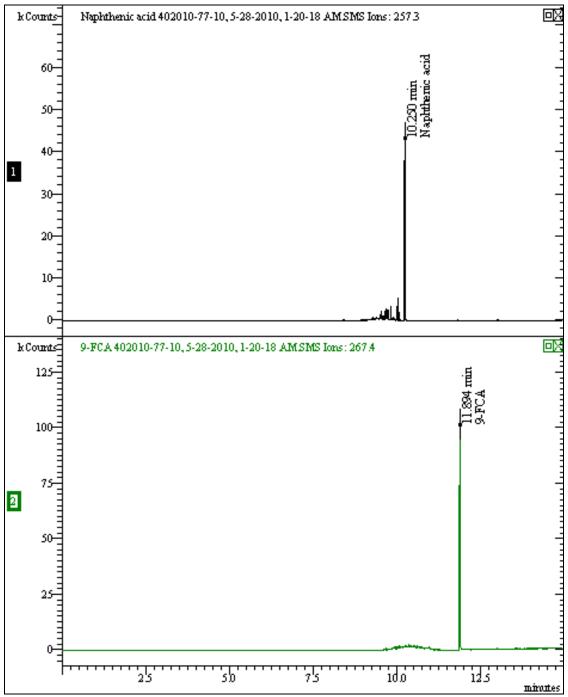


Figure 2: Representative Chromatogram of a Processed 125 mg Naphthenic Acid/mL Quality Control Sample

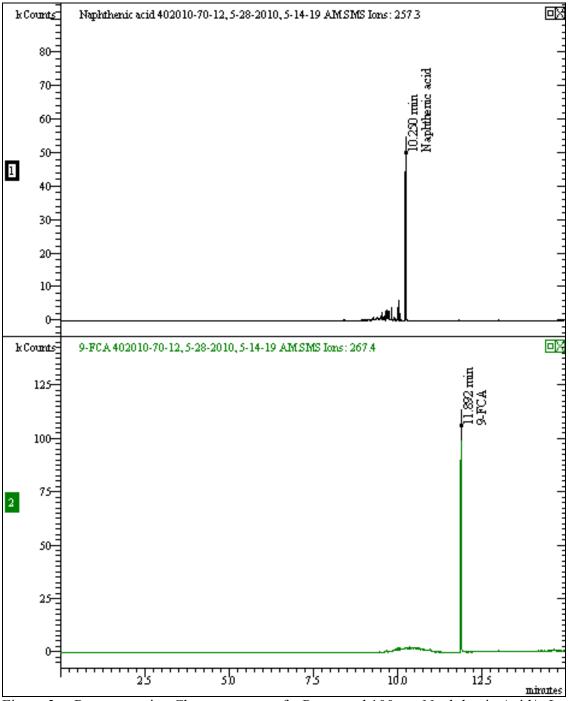


Figure 3: Representative Chromatogram of a Processed 100 mg Naphthenic Acid/mL Formulation Sample

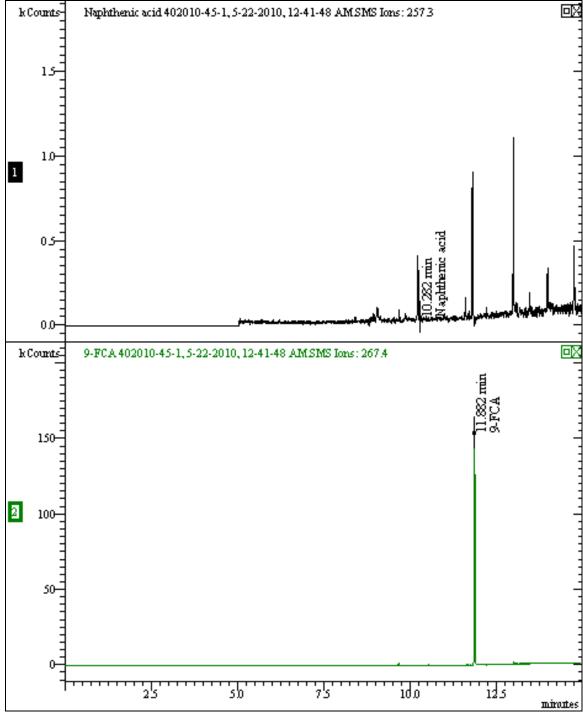


Figure 4: Chromatogram of a Processed Vehicle Blank Sample

5.1. SPECIFICITY/SELECTIVITY

As shown in Figure 4 (and in contrast to the chromatograms shown in Figure 1, Figure 2, and Figure 3), assay specificity/selectivity was confirmed when GC/MS analysis of processed vehicle blank samples revealed that there were no quantifiable peaks at or near the retention time for the test substance and IS (approximately 10.3 and 11.9 minutes, respectively).

5.2. Assay Validation: Calibration Reproducibility

During each of 3 validation sessions, triplicate matrix-based calibration samples at 5 concentrations were prepared and analyzed as described previously. Single injections were made of each calibration sample. The resulting naphthenic acid/IS peak area ratio versus theoretical naphthenic acid concentration data were fit to the quadratic function using least-squares regression analysis. The results of the regression analyses were used to back-calculate the corresponding concentrations from the peak area ratio data. As per WIL standard operating procedure (SOP), the reproducibility of the calibration curve data was considered valid when 1) the inter-session variability, expressed as RSD, of the back-calculated concentrations at each calibration level was \leq 10% RSD, except at the lowest calibration level where \leq 15% was acceptable and; 2) the mean back-calculated concentrations at each calibration level were within \pm 10% of the theoretical values (%RE within \pm 10%), except at the lowest calibration level where %RE within \pm 15% was acceptable.

The back-calculated concentrations and the associated intra- and inter-session statistics for the naphthenic acid assay calibration samples are summarized in Table 1. The inter-session variability (RSD) of the back-calculated concentrations ranged from 2.0% to 6.8% RSD. The inter-session mean concentrations had %RE values ranging from -1.3% to 2.3%. Based on the stated criteria, the reproducibility of the calibration data was acceptable.

5.3. ASSAY VALIDATION: PRECISION AND ACCURACY

During each of 3 validation sessions, triplicate QC samples at 3 concentrations were prepared and analyzed as described previously. Single injections were made of each processed QC sample. The results of the regression analyses were used to calculate the corresponding concentrations from the QC peak area ratio data. The variability (RSD) of the calculated QC concentration data was used as a measure of assay precision, and the difference between the theoretical and calculated mean QC concentrations (%RE) was used as a measure of assay accuracy. According to WIL SOP, the precision of the method was considered acceptable when the inter-session RSD of the calculated concentrations at each QC level was $\leq 15\%$, and the accuracy of the method was considered acceptable when the inter session calculated mean concentration at each QC level had a %RE value within \pm 15%.

The calculated concentrations and the associated intra- and inter-session statistics for the naphthenic acid assay QC samples are summarized in Table 2. The inter-session variability (RSD) of the calculated concentrations of each QC sample (precision) ranged from 4.5% to 6.9% RSD. The inter-session mean concentrations of the QC samples had %RE values (accuracy) ranging from -6.3% to -3.4%. Based on the stated criteria, the precision and accuracy of the naphthenic acid assay were acceptable.

5.4. ASSAY RUGGEDNESS

Assay ruggedness, as required by WIL SOP, was successfully demonstrated for this method because at least 2 of the 3 validation sessions were performed by different analysts.

5.5. ASSAY ACCEPTABILITY

In addition to the experimental samples, each analytical session consisted of (but was not limited to) calibration samples at 5 concentrations and triplicate QC samples prepared at each of 3 concentrations. In this study, the formulations were prepared at target concentrations of 0, 3, 10, 30, and 100 mg naphthenic acid/mL, and the QC samples were

prepared at nominal concentrations of 2.50, 25.0, and 125 mg naphthenic acid/mL. For an analytical session to be considered valid, at least two-thirds of the calculated QC concentrations with at least 1 sample at each concentration had to be 85% to 115% of the nominal QC concentration. All reported results were from analytical sessions that met the acceptance criteria.

5.6. TEST ARTICLE HOMOGENEITY AND RESUSPENSION HOMOGENEITY ASSESSMENT OF FORMULATIONS

Duplicate samples from the top, middle, and bottom strata of the formulations prepared on 27 May 2010 at target test article concentrations of 3 and 100 mg/mL were analyzed to assess test article homogeneity. The formulations that remained after sampling were divided into aliquots as would be used for daily dispensation. Representative aliquots were stored at room temperature for 11 days, at which time the test article was resuspended by stirring. Duplicate samples were collected from the top and bottom strata of the aliquots and analyzed to assess 11-day resuspension homogeneity. The results of the homogeneity and resuspension homogeneity analyses are presented in Table 3 and Table 4, respectively, with the overall statistics summarized as follows:

| Homogeneity Assessment of the 27 May 2010 Formulations | | | |
|--|-------------------|--------------------|--|
| | Group 2 (3 mg/mL) | Group 5 (100mg/mL) | |
| Mean Concentration (mg/mL) | 3.04 | 101 | |
| SD | 0.060 | 5.0 | |
| RSD (%) | 2.0 | 5.0 | |
| Mean Concentration % of Target | 101 | 101 | |

| 11-Day Room Temperature Resuspension Homogeneity | |
|--|--|
| Assessment of the 27 May 2010 Formulations | |

| | Group 2 (3 mg/mL) | Group 5 (100 mg/mL) |
|--------------------------------|----------------------|------------------------|
| Mean Concentration (mg/mL) | 2.94 | 101 |
| SD | 0.14 | 6.1 |
| RSD (%) | 4.7 | 6.0 |
| Mean Concentration % of Target | 97.9 | 101 |

The homogeneity assessment of the 27 May 2010 formulations met the WIL SOP requirement, *i.e.*, the RSD for the mean concentration was \leq 10% at a concentration within the acceptable limits (within 85% to 115% of target concentration). The resuspension homogeneity assessments of the 27 May 2010 formulations met the WIL SOP requirement, *i.e.*, the RSD for the mean concentration was \leq 10%.

5.7. TEST ARTICLE STABILITY IN FORMULATIONS

Formulations prepared on 27 May 2010 at target concentrations of 3 and 100 mg naphthenic acid/mL were analyzed on the day of preparation. Aliquots of the formulations were stored at room temperature for 11 days and analyzed to assess test article stability. The 11-day room temperature stability results are presented in Table 4. The mean post-storage concentrations ranged from 96.6% to 100% of the pre-storage values which met the WIL SOP requirement for stability, *i.e.*, the mean post-storage concentration was not <90% of the pre-storage value.

5.8. TEST ARTICLE CONCENTRATION IN FORMULATIONS

Formulations used for dose administration were analyzed to assess test article concentration acceptability. The results of the concentration acceptability assessments are presented in Table 5 and Table 6 with the mean concentration and percent of target values summarized in the following table.

| | Mean Concentration, mg/mL (Percent of Target) | | | | | | | | | | | |
|---------------------|---|-------------------|--------------------|--------------------|---------------------|--|--|--|--|--|--|--|
| Date of Preparation | Group 1 (0 mg/mL) | Group 2 (3 mg/mL) | Group 3 (10 mg/mL) | Group 4 (30 mg/mL) | Group 5 (100 mg/mL) | | | | | | | |
| 1 June 2010 | NQ | 3.34 (111) | 9.22 (92.2) | 28.7 (95.8) | 97.4 (97.4) | | | | | | | |
| 16 June 2010 | NQ | 3.04 (101) | 9.87 (98.7) | 27.9 (92.9) | 89.7 (89.7) | | | | | | | |

NQ = Not quantifiable

The analyzed formulations used for dose administration met the WIL SOP requirement for concentration acceptability for suspension formulations, *i.e.*, the analyzed concentration was 85% to 115% of the target concentration. No quantifiable amount of test article was detected in the analyzed vehicle administered to the control group (Group 1).

6. CONCLUSION

A GC/MS method for the determination of naphthenic acid concentration in corn oil formulations containing test article ranging in concentration from 2.50 to 125 mg/mL was validated in this study. Method specificity/selectivity, ruggedness, calibration reproducibility, precision, and accuracy were assessed and validated, satisfying WIL SOP criteria.

Formulations prepared at target test article concentrations of 3 and 100 mg naphthenic acid/mL met the WIL SOP requirement for homogeneity and, after 11 days of room temperature storage, resuspension homogeneity and stability. Analyzed formulations used for dose administration met the applicable WIL SOP acceptance criteria for test article concentration acceptability. No quantifiable amount of test article was detected in the analyzed vehicle administered to the control group.

8 Dec 2010

Date

7. REPORT REVIEW AND APPROVAL

Report Approved by:

Staff Toxicologist, Developmental and Reproductive Toxicology Study Director

Report Prepared by:

Assistant Director, Analytical Chemistry

8. ABBREVIATIONS

The following abbreviations may apply to this report:

μ - micro

uL - microliter

ACN - acetonitrile

amu - atomic mass unit

btm - bottom

cm - centimeter

conc. - concentration

DI - deionized

DMSO - dimethylsulfoxide

EPA - Environmental Protection Agency

ESI+ - positive electrospray ionization

FA - formic acid

FDA - Food and Drug Administration

FIFRA - Federal Insecticide, Fungicide and Rodenticide Act

g - gram

GLP - Good Laboratory Practices

GMP - Good Manufacturing Practices

HPLC - high performance liquid chromatography

hr - hour(s)

IS - internal standard

kg - kilogram

L - liter

LLOQ - lower limit of quantitation

M - molar

MC - methylcellulose

MeOH - methanol

mg - milligram

mL - milliliter

mm - millimeter

msec - milliseconds

MS - mass spectrometry

mM - millimolar

NA - not applicable

ND - not detected

ng - nanogram

nm - nanometer

ppm - parts per million

QC - quality control

Naphthenic Acid WIL-402010

American Petroleum Institute

%RE - percent relative error

RSD - relative standard deviation

SD - standard deviation

SOP - standard operating procedure

SPE - solid phase extraction

UV - ultraviolet

v - volume

w - weight

WIL - WIL Research Laboratories, LLC
WTDMSTM - WIL Toxicology Data Management System

TABLES 1 - 6

AN ORAL (GAVAGE) 14-DAY AND PILOT PRENATAL DEVELOPMENTAL TOXICITY STUDY OF NAPHTHENIC ACID IN RATS

Table 1: Back-Calculated Concentrations of the Validation Calibration Standards

| Theoretical Concentration | 150 | 200 | 250 | 300 | 350 |
|------------------------------|------|-----|-------|-------|-------|
| (μg/mL) | | | | | |
| Session 1 | 146 | 206 | 250 | 300 | 365 |
| (21 May 2010) | 152 | 202 | 237 | 304 | 335 |
| Ruggedness | 147 | 208 | 243 | 305 | 349 |
| Mean | 148 | 205 | 244 | 303 | 349 |
| SD | 3.5 | 3.2 | 6.8 | 2.5 | 15 |
| %RSD | 2.4 | 1.6 | 2.8 | 0.81 | 4.3 |
| %RE | -1.1 | 2.7 | -2.6 | 1.1 | -0.14 |
| | | | | | |
| Session 2 | 143 | 206 | 212 | 289 | 350 |
| (25-26 May 2010) | 156 | 199 | 264 | 301 | 355 |
| | 144 | 214 | 265 | 302 | 353 |
| Mean | 148 | 206 | 247 | 297 | 353 |
| SD | 7.1 | 7.8 | 30 | 7.5 | 2.6 |
| %RSD | 4.8 | 3.8 | 12 | 2.5 | 0.73 |
| %RE | -1.5 | 3.2 | -1.1 | -0.91 | 0.77 |
| | | | | | |
| Session 3 | 158 | 200 | 243 | 294 | 335 |
| (27 May 2010) | 138 | 204 | 258 | 291 | 361 |
| | 147 | 202 | 259 | 296 | 360 |
| Mean | 148 | 202 | 253 | 294 | 352 |
| SD | 10 | 2.2 | 9.2 | 2.3 | 14 |
| %RSD | 7.0 | 1.1 | 3.6 | 0.79 | 4.1 |
| %RE | -1.4 | 1.1 | 1.4 | -2.1 | 0.63 |
| Inter-Session Statistic | s | | | | |
| n | 9 | 9 | 9 | 9 | 9 |
| Mean | 148 | 205 | 248 | 298 | 351 |
| SD | 6.5 | 4.7 | 17 | 5.8 | 11 |
| %RSD | 4.4 | 2.3 | 6.8 | 2.0 | 3.0 |
| %RE | -1.3 | 2.3 | -0.76 | -0.63 | 0.42 |

402010 results.xls I Printed: 20Aug2010 9:00 AM

> I 402010 results.xls Printed: 8/20/2010 9:02 AM

AN ORAL (GAVAGE) 14-DAY AND PILOT PRENATAL DEVELOPMENTAL TOXICITY STUDY OF NAPHTHENIC ACID IN RATS

Table 2: Calculated Concentrations of the Validation Quality Control Samples

| Theoretical Concentration (mg/mL) | 2.50 | 25.0 | 125 |
|---|-------|-------|------|
| Session 1 | 2.49 | 24.8 | 119 |
| (21 May 2010) | 2.42 | 23.1 | 119 |
| Ruggedness | 2.42 | 23.8 | 116 |
| Mean | 2.44 | 23.9 | 118 |
| SD | 0.037 | 0.88 | 1.7 |
| %RSD | 1.5 | 3.7 | 1.4 |
| %RE | -2.2 | -4.4 | -5.5 |
| | | | |
| Session 2 | 2.14 | 22.6 | 111 |
| (25-26 May 2010) | 2.27 | 24.1 | 107 |
| | 2.19 | 24.1 | 109 |
| Mean | 2.20 | 23.6 | 109 |
| SD | 0.063 | 0.83 | 1.8 |
| %RSD | 2.8 | 3.5 | 1.7 |
| %RE | -12 | -5.6 | -13 |
| | | | |
| Session 3 | 2.50 | 23.4 | 131 |
| (27 May 2010) | 2.56 | 25.7 | * |
| | 2.56 | 25.7 | 125 |
| Mean | 2.54 | 24.9 | 128 |
| SD | 0.035 | 1.3 | 4.0 |
| %RSD | 1.4 | 5.3 | 3.1 |
| %RE | 1.7 | -0.33 | 2.3 |
| Inter-Session Statistics | | | |
| n | 9 | 9 | 8 |
| Mean | 2.40 | 24.1 | 117 |
| SD | 0.16 | 1.1 | 8.1 |
| %RSD | 6.6 | 4.5 | 6.9 |
| %RE | -4.1 | -3.4 | -6.3 |

^{*} Sample not included in summary statistics, suspected sample preparation error

402010 results.xls II Printed: 20Aug2010 9:00 AM

II 402010 results.xls Printed: 8/20/2010 9:02 AM

28

AN ORAL (GAVAGE) 14-DAY AND PILOT PRENATAL DEVELOPMENTAL TOXICITY STUDY OF NAPHTHENIC ACID IN RATS

Table 3: Homogeneity Assessment of the 27 May 2010 Formulations (Analyzed 27 May 2010)

| Group/ <u>Strata</u> | Dose Conc. (mg/mL) | <u>Ref#</u> (402010 -) | <u>Line #</u> (402010E) | Analyzed Conc. (mg/mL) | Percent of <u>Target</u> (%) | Mean Conc. (mg/mL) | <u>SD</u> | <u>RSD</u> (%) | Mean Conc % of Target (%) |
|-------------------------|----------------------|-------------------------|-------------------------|--------------------------|------------------------------------|----------------------|-----------|-------------------|---------------------------------|
| 2/Top | 3 | 70 - 1 | 61 | 3.01 | 100 | 3.04 | 0.060 | 2.0 | 101 |
| | | 70 - 2 | 62 | 2.96 | 98.7 | | | | |
| 2/Mid | | 70 - 3 | 63 | 3.04 | 101 | | | | |
| | | 70 - 4 | 64 | 3.02 | 101 | | | | |
| 2/Btm | | 70 - 5 | 65 | 3.05 | 102 | | | | |
| | | 70 - 6 | 66 | 3.14 | 105 | | | | |
| 5/Top | 100 | 70 - 7 | 67 | 93.4 | 93.4 | 101 | 5.0 | 5.0 | 101 |
| - | | 70 - 8 | 68 | 104 | 104 | | | | |
| 5/Mid | | 70 - 9 | 69 | 96.2 | 96.2 | | | | |
| | | 70 - 10 | 70 | 104 | 104 | | | | |
| 5/Btm | | 70 - 11 | 71 | 105 | 105 | | | | |
| | | 70 - 12 | 72 | 105 | 105 | | | | |

402010 results.xls 1H Printed: 09Aug2010 2:38 PM

AN ORAL (GAVAGE) 14-DAY AND PILOT PRENATAL DEVELOPMENTAL TOXICITY STUDY OF NAPHTHENIC ACID IN RATS

Table 4: 11-Day Room Temperature Resuspension and Stability Analysis of the 27 May 2010 Formulations (Analyzed 7 June 2010)

| Group/ <u>Strata</u> | Dose Conc. (mg/mL) | <u>Ref#</u> (402010 -) | <u>Line #</u> (402010G -) | Analyzed Conc. (mg/mL) | Percent of Target (%) | Mean <u>Conc.</u> (mg/mL) | <u>SD</u> | <u>RSD</u> (%) | Mean Conc % of Target (%) | Mean Conc % of Time Zero |
|-------------------------|----------------------|-------------------------|--------------------------------|--------------------------|-----------------------------|-----------------------------------|-----------|-------------------|---------------------------------|--------------------------|
| 2 Top | 3 | 119 - 1 | 33 | 2.78 | 92.6 | 2.94 | 0.14 | 4.7 | 97.9 | 96.6 |
| - | | 119 - 2 | 34 | 2.97 | 99.2 | | | | | |
| 2 Btm | | 119 - 3 | 35 | 2.89 | 96.4 | | | | | |
| | | 119 - 4 | 36 | 3.10 | 103 | | | | | |
| 5 Top | 100 | 119 - 5 | 37 | 92.1 | 92.1 | 101 | 6.1 | 6.0 | 101 | 100 |
| | | 119 - 6 | 38 | 106 | 106 | | | | | |
| 5 Btm | | 119 - 7 | 39 | 103 | 103 | | | | | |
| | | 119 - 8 | 40 | 104 | 104 | | | | | |

| Time Zero Concentrations | | | | | | |
|--------------------------|---------------|--|--|--|--|--|
| Group | Conc. (mg/mL) | | | | | |
| 2 | 3.04 | | | | | |
| 5 | 101 | | | | | |

402010 results.xls 2RH Printed: 09Aug2010 2:38 PM

30

Naphthenic Acid

AN ORAL (GAVAGE) 14-DAY AND PILOT PRENATAL DEVELOPMENTAL TOXICITY STUDY OF NAPHTHENIC ACID IN RATS

Table 5: Concentration Assessment of the 1 June 2010 Formulations(Analyzed 1 June 2010)

| Group/ <u>Strata</u> | Dose Conc. (mg/mL) | <u>Ref#</u> (402010 -) | <u>Line #</u> (402010F) | Analyzed Conc. (mg/mL) | Percent of Target (%) | Mean <u>Conc.</u> (mg/mL) | <u>SD</u> | <u>RSD</u> (%) | Mean Conc % of Target (%) |
|-------------------------|----------------------|-------------------------|-------------------------|--------------------------|------------------------|---|-----------|-------------------|---------------------------------|
| 1/Mid | 0 | 90 - 1 | 34 | | | <not quanti<="" td=""><td></td><td></td><td></td></not> | | | |
| | | 90 - 2 | 35 | | • | <not quanti<="" td=""><td>fiable></td><td></td><td></td></not> | fiable> | | |
| 2/Mid | 3 | 90 - 3 | 36 | 3.33 | 111 | 3.34 | 0.013 | 0.38 | 111 |
| | | 90 - 4 | 37 | 3.35 | 112 | | | | |
| 3/Mid | 10 | 90 - 5 | 38 | 8.65 | 86.5 | 9.22 | 0.80 | 8.7 | 92.2 |
| | | 90 - 6 | 39 | 9.79 | 97.9 | | | | |
| 4/Mid | 30 | 90 - 7 | 40 | 29.2 | 97.5 | 28.7 | 0.70 | 2.5 | 95.8 |
| 1/ 1 /11 C | 30 | 90 - 8 | 41 | 28.2 | 94.1 | 20.7 | 0.70 | 2.3 | 73.0 |
| 5/Mid | 100 | 90 - 9 | 42 | 101 | 101 | 97.4 | 5.6 | 5.7 | 97.4 |
| 3/1VIIU | 100 | 90 - 9 90 - 10 | 42 | 93.5 | 93.5 | 71. 4 | 5.0 | 5.7 | 97.4 |

402010 results.xls 1C Printed: 09Aug2010 2:38 PM

Naphthenic Acid

AN ORAL (GAVAGE) 14-DAY AND PILOT PRENATAL DEVELOPMENTAL TOXICITY STUDY OF NAPHTHENIC ACID IN RATS

Table 6: Concentration Assessment of the 16 June 2010 Formulations (Back up Samples) (Analyzed 18 June 2010)

| Group/ <u>Strata</u> | Dose <u>Conc.</u> (mg/mL) | <u>Ref #</u> (402010 -) | <u>Line #</u> (402010H2) | Analyzed Conc. (mg/mL) | Percent of Target (%) | Mean Conc. (mg/mL) | <u>SD</u> | <u>RSD</u> (%) | Mean Conc % of Target (%) |
|-------------------------|-----------------------------|---------------------------|-----------------------------|--------------------------|-------------------------|---|-----------|-------------------|---------------------------|
| 1/Mid | 0 | 151 - 1 | 35 | | | <not quanti<="" td=""><td>fiable></td><td></td><td></td></not> | fiable> | | |
| | | 151 - 2 | 36 | | | <not quanti<="" td=""><td>fiable></td><td></td><td></td></not> | fiable> | | |
| 2/Mid | 3 | 151 - 3 | 37 | 2.91 | 97.1 | 3.04 | 0.18 | 5.9 | 101 |
| | | 151 - 4 | 38 | 3.17 | 106 | | | | |
| 3/Mid | 10 | 151 - 5 | 39 | 10.1 | 101 | 9.87 | 0.38 | 3.9 | 98.7 |
| | | 151 - 6 | 40 | 9.60 | 96.0 | | | | |
| 4/Mid | 30 | 151 - 7 | 41 | 27.8 | 92.7 | 27.9 | 0.087 | 0.31 | 92.9 |
| 1/11114 | 30 | 151 - 8 | 42 | 27.9 | 93.1 | 27.9 | 0.007 | 0.51 | 72.7 |
| 5/Mid | 100 | 151 - 9 | 43 | 94.9 | 94.9 | 89.7 | 7.3 | 8.1 | 89.7 |
| J/ WIIU | 100 | 151 - 9 | 44 | 84.6 | 84.6 | 67.7 | 1.5 | 0.1 | 69.7 |

402010 results.xls 3C Printed: 09Aug2010 2:38 PM

ATTACHMENT I

Certificate of Analysis

Certificate of Analysis

Fluorene-9-carboxylic acid, **Product Name** 97%

F1409 **Product Number Product Brand** ALDRICH **CAS Number** 1989-33-9 Molecular Formula C₁₄H₁₀O₂ **Molecular Weight** 210.23

TEST

SPECIFICATION

WHITE TO TAN POWDER, **APPEARANCE** CRYSTALS, CRYSTALLINE

TITRATION 96.5%-103.5% (WITH NAOH)

96.5% (MINIMUM)

HIGH PRESSURE LIQUID CHROMATOGRAPHY

LOT 03706KH RESULTS

LIGHT YELLOW POWDER

CONFORMS TO STRUCTURE.

98.7% (WITH NAOH)

99.9%

AUGUST 2007

Barbara Rajzer, Supervisor

Quality Control

INFRARED SPECTRUM

QUALITY CONTROL

ACCEPTANCE DATE

Milwaukee, Wisconsin USA

APPENDIX E

Animal Room Environmental Conditions

PROJECT NO.:WIL- 402010 TEMPERATURE/HUMIDITY - STUDY SUMMARY REPORT

SPONSOR: 402 - AMERICAN PETROLEUM Page 1 of 5

STUDY SPECIFICATIONS: 402010 DATE IN 05/18/10 TIME IN 11:00

DATE OUT 06/24/10 TIME OUT 08:00

Page 159 of 360

ROOM SPECIFICATIONS: B ROOM 61 LOW TEMPERATURE °F: 66.0 HIGH TEMPERATURE °F: 76.0 LOW HUMIDITY %RH: 30.0

TEST SYSTEM: Rat LOW TEMPERATURE °C: 18.9 HIGH TEMPERATURE °C: 24.4 HIGH HUMIDITY %RH: 70.0

| | PRIMARY TEMP | | SECONDARY TEM | ſP | PRIMARY HUM | SECONDARY HUM |
|----------|--------------|-----------|---------------|-----------|-------------|---------------|
| DATE | MEAN (°F) | MEAN (°C) | MEAN (°F) | MEAN (°C) | MEAN (%RH) | MEAN (%RH) |
| 05/18/10 | 70.9 | 21.6 | 70.8 | 21.6 | 46.3 | 46.3 |
| 05/19/10 | 70.8 | 21.6 | 70.5 | 21.4 | 46.2 | 46.4 |
| 05/20/10 | 72.0 | 22.2 | 71.1 | 21.7 | 44.2 | 44.6 |
| 05/21/10 | 70.8 | 21.6 | 70.4 | 21.3 | 46.2 | 46.5 |
| 05/22/10 | 71.1 | 21.7 | 70.6 | 21.4 | 46.5 | 47.0 |
| 05/23/10 | 72.1 | 22.3 | 71.1 | 21.7 | 45.1 | 45.8 |
| 05/24/10 | 72.1 | 22.3 | 71.1 | 21.7 | 45.8 | 46.6 |
| 05/25/10 | 72.4 | 22.4 | 71.2 | 21.8 | 46.2 | 47.1 |
| 05/26/10 | 72.7 | 22.6 | 72.1 | 22.3 | 45.8 | 46.9 |
| 05/27/10 | 71.8 | 22.1 | 72.6 | 22.6 | 42.4 | 43.3 |
| 05/28/10 | 72.5 | 22.5 | 73.1 | 22.8 | 42.3 | 43.2 |
| 05/29/10 | 72.4 | 22.4 | 73.0 | 22.8 | 42.4 | 43.3 |
| 05/30/10 | 72.6 | 22.6 | 73.5 | 23.1 | 44.2 | 45.1 |
| 05/31/10 | 71.8 | 22.1 | 73.8 | 23.2 | 45.4 | 46.2 |
| 06/01/10 | 71.0 | 21.7 | 73.2 | 22.9 | 43.7 | 44.5 |
| 06/02/10 | 71.4 | 21.9 | 73.6 | 23.1 | 44.8 | 45.4 |
| 06/03/10 | 71.5 | 21.9 | 73.4 | 23.0 | 45.4 | 44.8 |
| 06/04/10 | 71.1 | 21.7 | 73.4 | 23.0 | 47.7 | 44.0 |
| 06/05/10 | 71.4 | 21.9 | 73.6 | 23.1 | 51.3 | 44.8 |

ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS TEMPERATURE/HUMIDITY - STUDY SUMMARY REPORT

SPONSOR: 402 - AMERICAN PETROLEUM Page 2 of 5

| | | PRIMARY TEMP | | SECONDARY TEM | IP | PRIMARY HUM | SECONDARY HUM |
|----|----------|--------------|-----------|---------------|-----------|-------------|---------------|
| | DATE | MEAN (°F) | MEAN (°C) | MEAN (°F) | MEAN (°C) | MEAN (%RH) | MEAN (%RH) |
| | 06/06/10 | 70.6 | 21.4 | 73.0 | 22.8 | 47.4 | 44.3 |
| | 06/07/10 | 70.8 | 21.6 | 73.3 | 22.9 | 43.4 | 43.4 |
| | 06/08/10 | 71.0 | 21.7 | 73.6 | 23.1 | 43.0 | 42.7 |
| | 06/09/10 | 70.9 | 21.6 | 73.6 | 23.1 | 46.4 | 42.9 |
| | 06/10/10 | 72.2 | 22.3 | 73.9 | 23.3 | 44.3 | 42.6 |
| | 06/11/10 | 70.9 | 21.6 | 73.2 | 22.9 | 48.4 | 42.5 |
| | 06/12/10 | 72.5 | 22.5 | 73.9 | 23.3 | 53.0 | 44.7 |
| d | 06/13/10 | 71.7 | 22.1 | 73.5 | 23.1 | 52.8 | 43.3 |
| 3 | 06/14/10 | 71.6 | 22.0 | 73.5 | 23.1 | 52.3 | 43.2 |
| | 06/15/10 | 71.1 | 21.7 | 73.3 | 22.9 | 57.7 | 44.3 |
| 5 | 06/16/10 | 71.5 | 21.9 | 73.1 | 22.8 | 47.1 | 42.0 |
| + | 06/17/10 | 71.1 | 21.7 | 73.2 | 22.9 | 42.9 | 39.3 |
| 76 | 06/18/10 | 71.9 | 22.2 | 73.6 | 23.1 | 44.0 | 40.2 |
|) | 06/19/10 | 72.0 | 22.2 | 73.4 | 23.0 | 45.6 | 40.6 |
| | 06/20/10 | 71.7 | 22.1 | 73.3 | 22.9 | 43.8 | 38.7 |
| | 06/21/10 | 71.9 | 22.2 | 73.5 | 23.1 | 47.4 | 40.4 |
| | 06/22/10 | 73.5 | 23.1 | 74.6 | 23.7 | 51.6 | 43.8 |
| | 06/23/10 | 73.6 | 23.1 | 74.7 | 23.7 | 57.2 | 43.0 |
| | 06/24/10 | 72.3 | 22.4 | 74.3 | 23.5 | 55.1 | 45.9 |

PROJECT NO.:WIL- 402010

PROJECT NO.:WIL- 402010 TEMPERATURE/HUMIDITY - STUDY SUMMARY REPORT

SPONSOR: 402 - AMERICAN PETROLEUM Page 3 of 5

| | PRIMARY TEMP | | SECONDARY TEM | P | PRIMARY HUM | SECONDARY HUM | |
|------------------------|--------------|---------------|---------------|-----------|-------------|---------------|------------|
| DATE | MEAN (°F |) ME <i>A</i> | AN (°C) | MEAN (°F) | MEAN (°C) | MEAN (%RH) | MEAN (%RH) |
| SUMMARY OF DAILY MEANS | MEAN | MIN | MAX | | | | |
| PRIMARY TEMP °F: | 71.7 | 70.6 | 73.6 | | | | |
| PRIMARY TEMP °C: | 22.1 | 21.4 | 23.1 | | | | |
| SECONDARY TEMP °F: | 72.9 | 70.4 | 74.7 | | | | |
| SECONDARY TEMP °C: | 22.7 | 21.3 | 23.7 | | | | |
| PRIMARY HUM %RH: | 46.8 | 42.3 | 57.7 | | | | |
| SECONDARY HUM %RH: | 43.9 | 38.7 | 47.1 | | | | |
| N DAYS | 38 | | | | | | |

PROJECT NO.:WIL- 402010

TEMPERATURE/HUMIDITY - STUDY SUMMARY REPORT

SPONSOR: 402 - AMERICAN PETROLEUM Page 4 of 5

B ROOM 61 SUMMARY OF HOURLY VALUES

| | PRIMARY TEMP | | | | SECONDARY TEMP | | | | PRIMARY HUM | | SECONDARY HUM | |
|-----------|--------------|----|------|----|----------------|----|------|----|-------------|-----|---------------|-----|
| MEAN | 71.7 | °F | 22.1 | °C | 72.9 | °F | 22.7 | °C | 46.8 | %RH | 43.9 | %RH |
| MIN | 68.8 | ۰F | 20.4 | °C | 69.1 | ۰F | 20.6 | °C | 37.8 | %RH | 33.1 | %RH |
| MAX | 82.2 | ۰F | 27.9 | °C | 79.9 | ۰F | 26.6 | °C | 70.3 | %RH | 64.5 | %RH |
| SD | 1.79 | | 0.99 | | 1.54 | | 0.86 | | 5.15 | | 3.70 | |
| SE | 0.06 | | 0.03 | | 0.05 | | 0.03 | | 0.17 | | 0.12 | |
| N SAMPLES | 883 | | | | 883 | | | | 883 | | 883 | |
| FIRST DAY | 05/18/ | 10 | | | | | | | | | | |

N DAYS

LAST DAY 06/24/10 38

PROJECT NO.:WIL- 402010

TEMPERATURE/HUMIDITY - STUDY SUMMARY REPORT

SPONSOR: 402 - AMERICAN PETROLEUM Page 5 of 5

STUDY 402010 SUMMARY OF HOURLY VALUES

| | PRIMARY TEME | | SECONDARY TE | MP | PRIMARY HUM | SECONDARY HUM |
|-----------|--------------|---------|--------------|---------|-------------|---------------|
| MEAN | 71.7 °F | 22.1 °C | 72.9 °F | 22.7 °C | 46.8 %RH | 43.9 %RH |
| MIN | 68.8 °F | 20.4 °C | 69.1 °F | 20.6 °C | 37.8 %RH | 33.1 %RH |
| MAX | 82.2 °F | 27.9 °C | 79.9 °F | 26.6 °C | 70.3 %RH | 64.5 %RH |
| SD | 1.79 | 0.99 | 1.54 | 0.86 | 5.15 | 3.70 |
| SE | 0.06 | 0.03 | 0.05 | 0.03 | 0.17 | 0.12 |
| N SAMPLES | 883 | | 883 | | 883 | 883 |
| DIDOM DAI | 05/10/10 | | | | | |

FIRST DAY 05/18/10
LAST DAY 06/24/10
N DAYS 38

APPENDIX F

WIL Developmental Historical Control Data Version 3.9 [Crl:CD(SD) Rats]

Study Date Range: 21 Apr 1998 - 27 Feb 2009 Mean of Study Means

| Endpoint | Total | Mean | S.D. | SEM | Median | Min | Max | 25th Quartile | 75th Quartile |
|---|-------|------|------|------|--------|------|-------|------------------|------------------|
| NO. OF DATASETS | 152 | | | | | | | | |
| Total No. of Animals in the Control Group | 3767 | | | | | | | | |
| No. of Animals That Died | 6 | | | | | | | | |
| No. of Animals That Aborted | 0 | | | | | | | | |
| No. of Animals That Delivered | 0 | | | | | | | | |
| Percent Pregnant | | 96.3 | 5.16 | 0.42 | 98.0 | 69.2 | 100.0 | 95.9 | 100.0 |
| No. Gravid | 3631 | | | | | | | | |
| No. With Only Resorptions | 5 | | | | | | | | |
| No. of Dams With Live Fetuses | 3621 | | | | | | | | |
| No. Nongravid | 136 | | | | | | | | |
| No. of Animals Examined at Laparohysterectomy | 3761 | | | | | | | | |
| Mean Gravid Uterine Weight (g) | | 85.4 | 3.70 | 0.30 | 85.7 | 72.2 | 95.4 | 82.8 | 88.1 |
| Mean No. Viable Fetuses/Dam | | 15.2 | 0.67 | 0.05 | 15.2 | 12.2 | 17.1 | 14.8 | 15.6 |
| Total No. Viable Fetuses | 54997 | | | | | | | | |
| Viable Fetuses (%/Litter) | | 95.2 | 1.54 | 0.13 | 95.5 | 90.1 | 98.0 | 94.1 | 96.2 |
| Mean No. Postimplantation Loss/Dam | | 0.7 | 0.22 | 0.02 | 0.7 | 0.3 | 1.4 | 0.6 | 0.9 |
| Total No. of Postimplantation Losses | 2703 | | | | | | | | |
| Postimplantation Loss (%/Litter) | | 4.8 | 1.54 | 0.13 | 4.5 | 2.0 | 9.9 | 3.8 | 5.9 |
| Early Resorptions (%/Litter) | | 4.7 | 1.56 | 0.13 | 4.5 | 1.5 | 9.9 | 3.6 | 5.8 |
| Late Resorptions (%/Litter) | | 0.1 | 0.19 | 0.02 | 0.0 | 0.0 | 0.8 | 0.0 | 0.2 |
| Dead Fetuses (%/Litter) | | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.5 | 0.0 | 0.0 |
| Mean No. Implantations/Dam | | 15.9 | 0.67 | 0.05 | 15.9 | 13.0 | 17.6 | 15.5 | 16.3 |
| Mean No. Corpora Lutea/Dam | | 17.3 | 0.82 | 0.07 | 17.3 | 14.5 | 19.4 | 16.7 | 17.8 |
| Mean No. Preimplantation Loss/Dam | | 1.4 | 0.60 | 0.05 | 1.3 | 0.3 | 3.1 | 0.9 | 1.7 |
| Total No. Preimplantation Losses | 4967 | | | | | | | | |
| Preimplantation Loss (%/Litter) | | 7.4 | 3.04 | 0.25 | 7.3 | 1.5 | 15.7 | 5.3 | 9.2 |
| Total No. Male Fetuses | 27427 | | | | | | | | |
| Total No. Female Fetuses | 27570 | | | | | | | | |
| % Males/Litter | | 49.8 | 2.72 | 0.22 | 49.7 | 43.1 | 56.7 | 47.9 | 51.7 |
| % Females/Litter | | 50.2 | 2.72 | 0.22 | 50.3 | 43.3 | 56.9 | 48.3 | 52.1 |
| Mean Fetal Body Weight (g) | | 3.7 | 0.11 | 0.01 | 3.6 | 3.4 | 3.9 | 3.6 | 3.7 |
| Mean Male Body Weight (g) | | 3.7 | 0.12 | 0.01 | 3.7 | 3.5 | 4.0 | 3.7 | 3.8 |
| Mean Female Body Weight (g) | | 3.6 | 0.11 | 0.01 | 3.5 | 3.4 | 3.8 | 3.5 | 3.6 |

| NO. OF DATASETS | 149 | |
|--|-------|------|
| Total No. of Fetuses/Litters Examined Externally | 53902 | 3548 |
| Total No. of Fetuses/Litters Examined Viscerally | 53708 | 3548 |
| Total No. of Fetuses/Litters Examined Skeletally | 53695 | 3547 |

| MALFORMATIONS (% Per Litter) | Mean | S.D. | SEM | Median | Min | Max | 25th Quartile | 75th Quartile |
|--|------|------|------|--------|-----|-----|------------------|------------------|
| | | | | | | | | |
| TOTAL EXTERNAL MALFORMATIONS | 0.1 | 0.21 | 0.02 | 0.0 | 0.0 | 1.3 | 0.0 | 0.2 |
| TOTAL VISCERAL MALFORMATIONS | 0.1 | 0.17 | 0.01 | 0.0 | 0.0 | 0.9 | 0.0 | 0.0 |
| TOTAL SKELETAL MALFORMATIONS | 0.1 | 0.24 | 0.02 | 0.0 | 0.0 | 1.1 | 0.0 | 0.2 |
| TOTAL MALFORMATIONS | 0.3 | 0.38 | 0.04 | 0.0 | 0.0 | 1.6 | 0.0 | 0.3 |
| EXTERNAL | | | | | | | | |
| Aglossia | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Anal Atresia | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Apodia | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Astomia | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Carpal and/or Tarsal Flexure | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Cleft Face | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Cleft Lip | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Cleft Palate | 0.0 | 0.04 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Craniorachischisis | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Cyclopia | 0.0 | 0.04 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Ectromelia | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Exencephaly with or without Open Eyelid(s) | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Fetal Anasarca | 0.0 | 0.12 | 0.01 | 0.0 | 0.0 | 1.3 | 0.0 | 0.0 |
| Filamentous Tail | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Gastroschisis | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Hydrocephaly with or without Dome Head | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| Mandibular Agnathia | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Mandibular Micrognathia | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Maxillary Agnathia | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| Meningoencephalocele | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |

NO. OF DATASETS149Total No. of Fetuses/Litters Examined Externally539023548Total No. of Fetuses/Litters Examined Viscerally537083548Total No. of Fetuses/Litters Examined Skeletally536953547

| MALFORMATIONS (% Per Litter) | Mean | S.D. | SEM | Median | Min | Max | 25th Quartile | 75th Quartile |
|---------------------------------------|------|------|------|--------|-----|-----|------------------|------------------|
| | | | | | | | Q | Ç 1 |
| EXTERNAL | | | | | | | | |
| Micromelia | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| Microphthalmia and/or Anophthalmia | 0.0 | 0.09 | 0.01 | 0.0 | 0.0 | 0.5 | 0.0 | 0.0 |
| Omphalocele | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Open Eyelid(s) | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| Proboscis-Like Nose | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| Tail- Short | 0.0 | 0.04 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Umbilical Herniation of the Intestine | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| VISCERAL | | | | | | | | |
| Aorta- Narrowed | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Epididymis- Absent | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Heart and/or Great Vessel Anomaly | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| Hydrocephaly | 0.0 | 0.08 | 0.01 | 0.0 | 0.0 | 0.8 | 0.0 | 0.0 |
| Interrupted Aortic Arch | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Kidney(s)- Absent | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.4 | 0.0 | 0.0 |
| Kidney(s) and/or Ureter(s) Absent | 0.0 | 0.04 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Lung(s)- Lobular Agenesis | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.6 | 0.0 | 0.0 |
| Lung(s)- Lobular Dysgenesis | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Retroesophageal Aortic Arch | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Situs Inversus | 0.0 | 0.09 | 0.01 | 0.0 | 0.0 | 0.4 | 0.0 | 0.0 |
| Testis- Absent | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Thyroid Gland(s)- Absent | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Transposition of the Great Vessels | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Vessel(s)- Malpositioned | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| | | | | | | | | |

| NO. OF DATASETS | 149 | |
|--|-------|------|
| Total No. of Fetuses/Litters Examined Externally | 53902 | 3548 |
| Total No. of Fetuses/Litters Examined Viscerally | 53708 | 3548 |
| Total No. of Fetuses/Litters Examined Skeletally | 53695 | 3547 |

| MALFORMATIONS (% Per Litter) | Mean | S.D. | SEM | Median | Min | Max | 25th Quartile | 75th Quartile |
|--|------|------|------|--------|-----|-----|------------------|------------------|
| SKELETAL | | | | | | | | |
| Bent Limb Bone(s) | 0.0 | 0.09 | 0.01 | 0.0 | 0.0 | 1.0 | 0.0 | 0.0 |
| Costal Cartilage Anomaly | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Rib Anomaly | 0.0 | 0.08 | 0.01 | 0.0 | 0.0 | 0.4 | 0.0 | 0.0 |
| Rib(s)- Only 11 Pairs Present | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| Rib(s)- Only 12 Pairs Present | 0.0 | 0.06 | 0.01 | 0.0 | 0.0 | 0.5 | 0.0 | 0.0 |
| Skull Anomaly | 0.0 | 0.04 | 0.00 | 0.0 | 0.0 | 0.4 | 0.0 | 0.0 |
| Sternebra(e)- Fused | 0.0 | 0.08 | 0.01 | 0.0 | 0.0 | 1.0 | 0.0 | 0.0 |
| Sternebra(e)- Malaligned (Severe) | 0.0 | 0.06 | 0.00 | 0.0 | 0.0 | 0.4 | 0.0 | 0.0 |
| Sternoschisis | 0.0 | 0.07 | 0.01 | 0.0 | 0.0 | 0.6 | 0.0 | 0.0 |
| Vertebral Agenesis | 0.0 | 0.07 | 0.01 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Vertebral Anomaly with or without Associated Rib Anomaly | 0.0 | 0.11 | 0.01 | 0.0 | 0.0 | 0.7 | 0.0 | 0.0 |
| Vertebral Centra Anomaly | 0.0 | 0.06 | 0.00 | 0.0 | 0.0 | 0.5 | 0.0 | 0.0 |

| NO. OF DATASETS | 149 | |
|--|-------|------|
| Total No. of Fetuses/Litters Examined Externally | 53902 | 3548 |
| Total No. of Fetuses/Litters Examined Viscerally | 53708 | 3548 |
| Total No. of Fetuses/Litters Examined Skeletally | 53695 | 3547 |

| VARIATIONS (% Per Litter) | Mean | S.D. | SEM | Median | Min | Max | 25th Quartile | 75th Quartile |
|---|------|------|------|--------|------|------|------------------|------------------|
| | | | | | | | | |
| TOTAL EXTERNAL VARIATIONS | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| TOTAL VISCERAL VARIATIONS | 0.4 | 0.70 | 0.06 | 0.2 | 0.0 | 4.0 | 0.0 | 0.6 |
| TOTAL SKELETAL VARIATIONS | 33.6 | 6.79 | 0.56 | 33.4 | 18.0 | 50.4 | 29.6 | 37.2 |
| TOTAL VARIATIONS | 33.4 | 6.91 | 0.71 | 31.6 | 18.0 | 51.1 | 19.7 | 36.2 |
| EXTERNAL | | | | | | | | |
| Twinning | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| VISCERAL | | | | | | | | |
| Adrenal Gland(s)- Enlarged | 0.0 | 0.12 | 0.01 | 0.0 | 0.0 | 1.4 | 0.0 | 0.0 |
| Adrenal Gland(s)- Pale | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| Diaphragm- Thin | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Hemorrhagic Iris | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Hemorrhagic Ring Around the Iris | 0.0 | 0.07 | 0.01 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Kidney(s)- Small | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Liver- Accessory Lobule(s) | 0.0 | 0.09 | 0.01 | 0.0 | 0.0 | 0.8 | 0.0 | 0.0 |
| Liver- Pale | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Liver- Swollen | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Major Blood Vessel Variation | 0.1 | 0.15 | 0.01 | 0.0 | 0.0 | 0.8 | 0.0 | 0.0 |
| Renal Papilla(e) not Developed and/or Distended Ureter(s) | 0.2 | 0.62 | 0.05 | 0.0 | 0.0 | 4.0 | 0.0 | 0.3 |
| Spleen- Accessory | 0.0 | 0.09 | 0.01 | 0.0 | 0.0 | 1.0 | 0.0 | 0.0 |
| Spleen- Pale | 0.0 | 0.07 | 0.01 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Spleen- Small | 0.0 | 0.13 | 0.01 | 0.0 | 0.0 | 1.4 | 0.0 | 0.0 |
| Testis- Small | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| SKELETAL | | | | | | | | |
| 14th Full Rib(s) | 0.1 | 0.17 | 0.01 | 0.0 | 0.0 | 0.9 | 0.0 | 0.0 |

| NO. OF DATASETS | 149 | |
|--|-------|------|
| Total No. of Fetuses/Litters Examined Externally | 53902 | 3548 |
| Total No. of Fetuses/Litters Examined Viscerally | 53708 | 3548 |
| Total No. of Fetuses/Litters Examined Skeletally | 53695 | 3547 |

| ARIATIONS (% Per Litter) | Mean | S.D. | SEM | Median | Min | Max | 25th Quartile | 75th Quartile |
|--|------|------|------|--------|-----|------|------------------|------------------|
| SKELETAL | | | | | | | | |
| 14th Rudimentary Rib(s) | 6.8 | 2.84 | 0.23 | 6.5 | 0.0 | 15.1 | 4.8 | 8.6 |
| 25 Presacral Vertebrae | 0.1 | 0.31 | 0.03 | 0.0 | 0.0 | 2.0 | 0.0 | 0.0 |
| 27 Presacral Vertebrae | 0.2 | 0.28 | 0.02 | 0.0 | 0.0 | 1.8 | 0.0 | 0.3 |
| 7th Cervical Rib(s) | 0.8 | 0.78 | 0.06 | 0.7 | 0.0 | 3.5 | 0.3 | 1.1 |
| 7th Sternebra | 0.0 | 0.25 | 0.02 | 0.0 | 0.0 | 2.5 | 0.0 | 0.0 |
| Bent Rib(s) | 0.2 | 0.48 | 0.04 | 0.0 | 0.0 | 4.0 | 0.0 | 0.3 |
| Cervical Centrum #1 Ossified | 20.2 | 5.87 | 0.48 | 19.7 | 6.6 | 35.8 | 16.3 | 24.2 |
| Entire Sternum Unossified | 0.0 | 0.06 | 0.01 | 0.0 | 0.0 | 0.4 | 0.0 | 0.0 |
| Extra Site of Ossification Anterior to Cervical Centrum #2 | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Extra Site of Ossification Anterior to Rib #5 | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| Extra Site of Ossification Anterior to Sternebra #1 | 0.0 | 0.04 | 0.00 | 0.0 | 0.0 | 0.4 | 0.0 | 0.0 |
| Extra Site of Ossification Ventral to Cervical Centrum #2 | 0.0 | 0.02 | 0.00 | 0.0 | 0.0 | 0.3 | 0.0 | 0.0 |
| Hyoid Unossified | 1.3 | 0.98 | 0.08 | 1.2 | 0.0 | 4.4 | 0.5 | 2.0 |
| Ischium Unossified | 0.0 | 0.06 | 0.01 | 0.0 | 0.0 | 0.6 | 0.0 | 0.0 |
| Pubis Unossified | 0.1 | 0.22 | 0.02 | 0.0 | 0.0 | 2.3 | 0.0 | 0.0 |
| Reduced Ossification of the 13th Rib(s) | 0.6 | 0.69 | 0.06 | 0.5 | 0.0 | 3.6 | 0.2 | 0.8 |
| Reduced Ossification of the Limb Bone(s) | 0.0 | 0.03 | 0.00 | 0.0 | 0.0 | 0.2 | 0.0 | 0.0 |
| Reduced Ossification of the Rib(s) | 0.0 | 0.13 | 0.01 | 0.0 | 0.0 | 1.2 | 0.0 | 0.0 |
| Reduced Ossification of the Skull | 0.1 | 0.20 | 0.02 | 0.0 | 0.0 | 1.0 | 0.0 | 0.0 |
| Reduced Ossification of the Vertebral Arches | 0.1 | 0.20 | 0.02 | 0.0 | 0.0 | 1.1 | 0.0 | 0.2 |
| Skull Bone(s)- Accessory | 0.0 | 0.08 | 0.01 | 0.0 | 0.0 | 0.7 | 0.0 | 0.0 |
| Sternebra(e) #1, #2, #3 and/or #4 Unossified | 0.2 | 0.27 | 0.02 | 0.0 | 0.0 | 1.3 | 0.0 | 0.3 |
| Sternebra(e) #5 and/or #6 Unossified | 6.6 | 6.03 | 0.49 | 4.0 | 0.0 | 26.1 | 2.5 | 9.3 |
| Sternebra(e)- Malaligned (Slight or Moderate) | 0.4 | 0.49 | 0.04 | 0.3 | 0.0 | 3.0 | 0.0 | 0.6 |
| Sternebrae with Thread-Like Attachment | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.6 | 0.0 | 0.0 |

| NO. OF DATASETS | 149 | |
|--|-------|------|
| Total No. of Fetuses/Litters Examined Externally | 53902 | 3548 |
| Total No. of Fetuses/Litters Examined Viscerally | 53708 | 3548 |
| Total No. of Fetuses/Litters Examined Skeletally | 53695 | 3547 |

| VARIATIONS (% Per Litter) | Mean | S.D. | SEM | Median | Min | Max | 25th Quartile | 75th Quartile |
|--------------------------------|------|------|------|--------|-----|-----|------------------|------------------|
| SKELETAL | | | | | | | | |
| Unco-Ossified Vertebral Centra | 0.0 | 0.06 | 0.00 | 0.0 | 0.0 | 0.5 | 0.0 | 0.0 |
| Vertebral Centra Unossified | 0.0 | 0.05 | 0.00 | 0.0 | 0.0 | 0.4 | 0.0 | 0.0 |

Modal Distribution of Fetal Body Weights

No. of Datasets in Historical Control152Range of Study Dates4/21/1998 - 2/27/2009No. of Dams in Historical Control3767No. of Dams with Live Fetuses3621Mean Fetal Body Weight Range (g)3.4 - 3.9

| Mean Fetal Body Weight (g) | 3.4 | 3.5 | 3.6 | 3.7 | 3.8 | 3.9 |
|----------------------------|-------------|-------------|-------------|-------------|-------------|-------------|
| Total No. Datasets | 2 | 27 | 52 | 43 | 20 | 8 |
| Mean Litter Size | 15.1 | 15.4 | 15.2 | 15.2 | 14.7 | 14.6 |
| Litter Size Range | 15.0 - 15.1 | 14.4 - 17.1 | 13.7 - 16.7 | 13.7 - 16.6 | 12.2 - 15.8 | 13.8 - 15.3 |

| NO. OF DATASETS | 149 | |
|--|-------|------|
| Total No. of Fetuses/Litters Examined Externally | 53902 | 3548 |
| Total No. of Fetuses/Litters Examined Viscerally | 53708 | 3548 |
| Total No. of Fetuses/Litters Examined Skeletally | 53695 | 3547 |

| | Nun | nber |
|--|---------|---------|
| MALFORMATIONS | Fetuses | Litters |
| EXTERNAL | | |
| Microphthalmia and/or Anophthalmia | 18 | 18 |
| Fetal Anasarca | 7 | 5 |
| Mandibular Micrognathia | 6 | 6 |
| Omphalocele | 6 | 6 |
| Anal Atresia | 5 | 5 |
| Filamentous Tail | 5 | 5 |
| Cleft Palate | 4 | 4 |
| Umbilical Herniation of the Intestine | 4 | 4 |
| Tail- Short | 3 | 3 |
| Aglossia | 2 | 2 |
| Carpal and/or Tarsal Flexure | 2 | 2 |
| Cyclopia | 2 | 2 |
| Exencephaly with or without Open Eyelid(s) | 2 | 2 |
| Mandibular Agnathia | 2 | 2 |
| Apodia | 1 | 1 |
| Astomia | 1 | 1 |
| Cleft Face | 1 | 1 |
| Cleft Lip | 1 | 1 |
| Craniorachischisis | 1 | 1 |
| Ectromelia | 1 | 1 |
| Gastroschisis | 1 | 1 |
| Hydrocephaly with or without Dome Head | 1 | 1 |
| Maxillary Agnathia | 1 | 1 |
| | | |

| NO. OF DATASETS | 149 | |
|--|-------|------|
| Total No. of Fetuses/Litters Examined Externally | 53902 | 3548 |
| Total No. of Fetuses/Litters Examined Viscerally | 53708 | 3548 |
| Total No. of Fetuses/Litters Examined Skeletally | 53695 | 3547 |

| | Nun | ıber |
|--|---------|---------|
| MALFORMATIONS | Fetuses | Litters |
| EXTERNAL | | |
| Meningoencephalocele | 1 | 1 |
| Micromelia | 1 | 1 |
| Open Eyelid(s) | 1 | 1 |
| Proboscis-Like Nose | 1 | 1 |
| VISCERAL | | |
| Situs Inversus | 13 | 13 |
| Hydrocephaly | 8 | 8 |
| Lung(s)- Lobular Dysgenesis | 5 | 5 |
| Retroesophageal Aortic Arch | 5 | 5 |
| Kidney(s) and/or Ureter(s) Absent | 4 | 4 |
| Interrupted Aortic Arch | 2 | 2 |
| Thyroid Gland(s)- Absent | 2 | 2 |
| Transposition of the Great Vessels | 2 | 2 |
| Aorta- Narrowed | 1 | 1 |
| Epididymis- Absent | 1 | 1 |
| Heart and/or Great Vessel Anomaly | 1 | 1 |
| Kidney(s)- Absent | 1 | 1 |
| Lung(s)- Lobular Agenesis | 1 | 1 |
| Testis- Absent | 1 | 1 |
| Vessel(s)- Malpositioned | 1 | 1 |
| SKELETAL | | |
| Vertebral Anomaly with or without Associated Rib Anomaly | 20 | 17 |
| Rib Anomaly | 11 | 11 |

| NO. OF DATASETS | 149 | |
|--|-------|------|
| Total No. of Fetuses/Litters Examined Externally | 53902 | 3548 |
| Total No. of Fetuses/Litters Examined Viscerally | 53708 | 3548 |
| Total No. of Fetuses/Litters Examined Skeletally | 53695 | 3547 |

| | Nun | Number | | |
|-----------------------------------|---------|---------|--|--|
| IALFORMATIONS | Fetuses | Litters | | |
| SKELETAL | | | | |
| Vertebral Agenesis | 10 | 10 | | |
| Sternoschisis | 10 | 9 | | |
| Vertebral Centra Anomaly | 6 | 6 | | |
| Bent Limb Bone(s) | 6 | 4 | | |
| Costal Cartilage Anomaly | 5 | 5 | | |
| Rib(s)- Only 12 Pairs Present | 5 | 5 | | |
| Sternebra(e)- Malaligned (Severe) | 5 | 5 | | |
| Sternebra(e)- Fused | 2 | 2 | | |
| Rib(s)- Only 11 Pairs Present | 1 | 1 | | |
| Skull Anomaly | 1 | 1 | | |

| NO. OF DATASETS | 149 | |
|--|-------|------|
| Total No. of Fetuses/Litters Examined Externally | 53902 | 3548 |
| Total No. of Fetuses/Litters Examined Viscerally | 53708 | 3548 |
| Total No. of Fetuses/Litters Examined Skeletally | 53695 | 3547 |

| | Nun | ıber |
|---|---------|---------|
| ARIATIONS | Fetuses | Litters |
| EXTERNAL | | |
| Twinning | 1 | 1 |
| VISCERAL | | |
| Renal Papilla(e) not Developed and/or Distended Ureter(s) | 121 | 75 |
| Major Blood Vessel Variation | 42 | 41 |
| Hemorrhagic Ring Around the Iris | 11 | 11 |
| Spleen- Pale | 8 | 8 |
| Spleen- Accessory | 8 | 5 |
| Liver- Accessory Lobule(s) | 7 | 5 |
| Spleen- Small | 6 | 6 |
| Adrenal Gland(s)- Enlarged | 5 | 1 |
| Hemorrhagic Iris | 2 | 2 |
| Kidney(s)- Small | 2 | 2 |
| Liver- Pale | 2 | 2 |
| Testis- Small | 2 | 2 |
| Adrenal Gland(s)- Pale | 1 | 1 |
| Diaphragm- Thin | 1 | 1 |
| Liver- Swollen | 1 | 1 |
| SKELETAL | | |
| Cervical Centrum #1 Ossified | 10707 | 2698 |
| 14th Rudimentary Rib(s) | 3655 | 1519 |
| Sternebra(e) #5 and/or #6 Unossified | 3593 | 1287 |
| Hyoid Unossified | 705 | 448 |
| 7th Cervical Rib(s) | 433 | 335 |

| NO. OF DATASETS | 149 | |
|--|-------|------|
| Total No. of Fetuses/Litters Examined Externally | 53902 | 3548 |
| Total No. of Fetuses/Litters Examined Viscerally | 53708 | 3548 |
| Total No. of Fetuses/Litters Examined Skeletally | 53695 | 3547 |

| | Nun | nber |
|--|---------|---------|
| VARIATIONS | Fetuses | Litters |
| SKELETAL | | |
| Reduced Ossification of the 13th Rib(s) | 328 | 220 |
| Sternebra(e)- Malaligned (Slight or Moderate) | 201 | 180 |
| Sternebra(e) #1, #2, #3 and/or #4 Unossified | 103 | 95 |
| Bent Rib(s) | 88 | 67 |
| 27 Presacral Vertebrae | 82 | 64 |
| Reduced Ossification of the Vertebral Arches | 59 | 54 |
| 25 Presacral Vertebrae | 57 | 34 |
| Reduced Ossification of the Skull | 47 | 40 |
| 14th Full Rib(s) | 40 | 34 |
| Pubis Unossified | 37 | 31 |
| 7th Sternebra | 24 | 10 |
| Reduced Ossification of the Rib(s) | 19 | 16 |
| Entire Sternum Unossified | 8 | 8 |
| Skull Bone(s)- Accessory | 8 | 8 |
| Vertebral Centra Unossified | 5 | 5 |
| Unco-Ossified Vertebral Centra | 4 | 4 |
| Ischium Unossified | 4 | 3 |
| Extra Site of Ossification Anterior to Sternebra #1 | 3 | 3 |
| Reduced Ossification of the Limb Bone(s) | 2 | 2 |
| Sternebrae with Thread-Like Attachment | 2 | 1 |
| Extra Site of Ossification Anterior to Cervical Centrum #2 | 1 | 1 |
| Extra Site of Ossification Anterior to Rib #5 | 1 | 1 |
| Extra Site of Ossification Ventral to Cervical Centrum #2 | 1 | 1 |
| | | |

FINAL REPORT



<u>Contents</u>: Volume 2 of 2

Appendix G

Study Title: An Oral (Gavage) 14-Day and Pilot Prenatal

Developmental Toxicity Study of Naphthenic Acid in Rats

Study Number: WIL-402010

Study Director:

<u>Data Requirements</u>: Not Applicable

Study Initiation Date: 11 May 2010

Study Completion Date: 8 December 2010

Performing Laboratory: WIL Research Laboratories, LLC

1407 George Road

Ashland, OH 44805-8946

Sponsor: American Petroleum Institute

1220 L Street, NW

Washington, DC 20005

APPENDIX G

Individual Animal Data

TABLE I1 (MALES - DAILY EXAMINATIONS) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 1
SPONSOR:AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE PANCE: 06-02-10 TO 06-16-10

| | | | TABLE RANGE: 06-02-10 TO 06-16-10 |
|--------------------|----------------------------|-----------------------|--|
| ANIMAL SEX | GROUP | CATEGORY | DATE TIME GRADE OBSERVATIONS |
| 73665 M | 0 MG/KG/DAY | NORMAL | 06-02-10 8:44 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 8:19 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 8:07 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 8:32 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 8:54 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:43 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 7:41 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 8:28 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 7:42 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 8:12 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 7:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 7:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 7:41 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-16-10 7:43 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73665 M 73666 M | 0 MG/KG/DAY 0 MG/KG/DAY | DISPOSITION NORMAL | 06-16-10 8:44 P SCHEDULED EUTHANASIA 06-02-10 8:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 8:08 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 8:32 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 8:32 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 8:54 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:43 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 7:42 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 8:28 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 7:42 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 8:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 7:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 7:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 7:42 P NO SIGNIFICANT CLINICAL OBSERVATIONS |

GRADE CODE: 1 - SLIGHT 2 - MODERATE 3 - SEVERE P - PRESENT

PAGE 2

PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10

| | | | | | IABLE KA | ANGE: U | 5-02- | 10 10 06-16-10 |
|---|-------------------------|-------------|-------------|--|--|---|---|--|
| _ | ANIMAL | SEX | GROUP | CATEGORY | DATE | TIME | GRAD | E OBSERVATIONS |
| | 73666 73666 73666 | M | 0 MG/KG/DAY | NORMAL DISPOSITION BODY/INTEGUMENT | 06-16-3 06-03-3 06-03-3 | 10 8:44 10 8:21 10 8:21 | 4 P 1 1 1 1 | NO SIGNIFICANT CLINICAL OBSERVATIONS SCHEDULED EUTHANASIA WET YELLOW MATERIAL RIGHT INGUINAL AREA WET YELLOW MATERIAL LEFT INGUINAL AREA WET YELLOW MATERIAL INGOENITAL AREA |
| | 73676 73676 73691 | M M M | 0 MG/KG/DAY | DISPOSITION | 06-03- 06-02- 06-04- 06-05- 06-06- 06-07- 06-10- 06-11- 06-13- 06-14- 06-15- 06-16- 06-02- 06-03- 06-04- 06-05- 06-06- | 10 8:41 10 8:21 10 8:31 10 8:31 10 7:41 10 7:41 10 7:41 10 7:41 10 7:41 10 7:41 10 7:41 10 7:41 10 7:41 10 8:41 10 8:41 10 8:41 10 8:41 10 8:41 10 8:41 10 8:41 10 8:41 10 8:41 10 8:41 | 1 P P P P P P P P P P P P P P P P P P P | WET YELLOW MATERIAL UROGENITAL AREA NO SIGNIFICANT CLINICAL OBSERVATIONS |
| _ | | | | | 06-07-1 06-08-1 06-09-1 | 10 7:3 | | NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS |

PAGE 3 PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10

| ANIMAL | SEX | GROUP | CATEGORY | DATE 7 | TIME | GRAD | E OBSERVATIONS | | | |
|--------|-----|--------------|-------------|----------|------|------|----------------|----------|--------------|------|
| 73691 | M | 0 MG/KG/DAY | NORMAL | 06-10-10 | 8:29 |) P | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | , -, | | 06-11-10 | | | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-12-10 | 8:13 | 3 P | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-13-10 | 7:50 |) P | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-14-10 | 7:47 | 7 P | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-15-10 | 7:42 | 2 P | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-16-10 | 7:44 | ł P | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| 73691 | M | 0 MG/KG/DAY | DISPOSITION | 06-16-10 | 8:44 | ł P | SCHEDULED EUTH | ANASIA | | |
| 73692 | M | 0 MG/KG/DAY | NORMAL | 06-02-10 | 8:46 | 5 P | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-03-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-04-10 | 8:08 | 3 P | NO SIGNIFICANT | | | |
| | | | | 06-05-10 | 8:33 | 3 P | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-06-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-07-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-08-10 | 7:38 | 3 P | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-09-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-10-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-11-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-12-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-13-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-14-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-15-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-16-10 | | | NO SIGNIFICANT | | OBSERVATIONS | |
| 73692 | M | 0 MG/KG/DAY | DISPOSITION | 06-16-10 | | | SCHEDULED EUTH | | | |
| 73668 | M | 30 MG/KG/DAY | NORMAL | 06-02-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-04-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-05-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-06-10 | | | NO SIGNIFICANT | | | |
| | | | | 06-07-10 | 7:44 | ŀ P | NO SIGNIFICANT | CLINICAL | OBSERVATIONS | |

TABLE I1 (MALES - DAILY EXAMINATIONS) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RAT

PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 4
SPONSOR:AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10

| ANIMAI | SEX | GROUP | CATEGORY | DATE | TIME | | 2-10 10 06-16-10 ADE OBSERVATIONS |
|--------|-----|----------------|-----------------|--------------------|------|-----|---|
| 73668 | M | 30 MG/KG/DAY | NORMAL | 06-08- | | | |
| | | | | 06-09- | | 5 P | |
| | | | | 06-10- | | 9 P | |
| | | | | 06-11- | | | |
| | | | | 06-12- | | | |
| | | | | 06-13- | | | |
| | | | | 06-14- | | | |
| | | | | 06-15- | | | |
| | | | | 06-16- | | | |
| 73668 | M | 30 MG/KG/DAY | | 06-16- | | | |
| 73668 | M | 30 MG/KG/DAY | BODY/INTEGUMENT | | | 2 1 | |
| 73669 | M | 30 MG/KG/DAY | NORMAL | 06-02- | | | |
| | | | | 06-03- | | | |
| | | | | 06-04- | | | |
| | | | | 06-05- | | | |
| | | | | 06-06- | | | |
| | | | | 06-07- | | | |
| | | | | 06-08- | | | |
| | | | | 06-09- | | | |
| | | | | 06-10-1 06-11-1 | | | |
| | | | | 06-11- | | | |
| | | | | 06-12- | | | |
| | | | | 06-13- | | | |
| | | | | 06-14- | | | |
| | | | | 06-15- | | | |
| 73669 | М | 30 MG/KG/DAY | DISPOSITION | 06-16- | | | |
| 73682 | M | , , | NORMAL | 06-10- | | | |
| 13002 | 141 | 30 MG/ NG/ DAI | NORMALI | 06-02- | | | P NO SIGNIFICANT CLINICAL OBSERVATIONS P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | |

PAGE 5

PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10

| | | | | | | | 02 - | | . • | | | |
|----------------|-----|------------------------------|-------------|--|--|--|-----------------------|--|---|--|------|--|
| ANIMAL | SEX | GROUP | CATEGORY | DATE | TIM | IE (| GRADE | E OBSERVATION | is | | | |
| 73682 | M | 30 MG/KG/DAY | | 06-04- 06-05- 06-06- 06-07- 06-08- 06-09- 06-10- 06-11- 06-12- | 10 10 10 10 10 10 | 8:33 8:56 7:45 7:38 7:47 8:30 7:44 8:14 | P P P P P | NO SIGNIFICA NO SIGNIFICA NO SIGNIFICA NO SIGNIFICA NO SIGNIFICA NO SIGNIFICA NO SIGNIFICA | NT CLINICAL | OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS | | |
| 73682 73687 | | 30 MG/KG/DAY 30 MG/KG/DAY | DISPOSITION | 06-13- 06-14- 06-15- 06-16- 06-16- 06-02- 06-03- 06-04- | 10 10 10 10 10 | 7:51 7:48 7:43 7:47 8:45 8:50 8:23 8:09 | P P P P | NO SIGNIFICA NO SIGNIFICA SCHEDULED EU NO SIGNIFICA NO SIGNIFICA | NT CLINICAL NT CLINICAL NT CLINICAL THANASIA NT CLINICAL NT CLINICAL | OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS | | |
| | | | | 06-05- 06-06- 06-07- 06-08- 06-09- 06-10- 06-11- 06-12- 06-13- | 10 10 10 10 10 10 10 | 8:34 8:56 7:45 7:39 7:47 8:30 7:44 8:15 7:51 | P P P P P | NO SIGNIFICA NO SIGNIFICA NO SIGNIFICA NO SIGNIFICA NO SIGNIFICA NO SIGNIFICA NO SIGNIFICA | NT CLINICAL | OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS | | |
| | | | | 06-14- 06-15- 06-16- | 10 10 10 | 7:49 7:44 7:47 | P P P | NO SIGNIFICA | NT CLINICAL NT CLINICAL NT CLINICAL | OBSERVATIONS OBSERVATIONS OBSERVATIONS | | |

PAGE 6 PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10

| ANIMAL SEX GROUP CATEGORY DATE TIME GRADE OBSERVATIONS 73687 M 30 MG/KG/DAY DISPOSITION 06-16-10 8:45 P SCHEDULED EUTHANASIA 73688 M 30 MG/KG/DAY NORMAL 06-02-10 8:51 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 8:23 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 8:10 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 8:34 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 8:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 7:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | TABLE RAN | GE: 06 | -02- | ·10 TO 06-16-10 |
|---|--------|-----|---------------|----------|-----------|--------|------|--------------------------------------|
| 73688 M 30 MG/KG/DAY NORMAL 06-02-10 8:51 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 8:23 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 8:10 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 8:34 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-01 8:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | ANIMAL | SEX | GROUP | CATEGORY | DATE I | IME | GRAD | DE OBSERVATIONS |
| 06-03-10 8:23 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 8:10 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 8:34 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 8:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-04-10 8:10 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 8:34 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 8:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | 73688 | M | 30 MG/KG/DAY | NORMAL | | | | |
| 06-05-10 8:34 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 8:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-06-10 8:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-07-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-08-10 7:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| | | | | | | | | |
| 16-19-10 7.48 D NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| | | | | | | | | |
| 06-10-10 8:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-11-10 7:44 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-12-10 8:15 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-13-10 7:52 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-14-10 7:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-15-10 7:44 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-16-10 7:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | // | | | | | |
| 73688 M 30 MG/KG/DAY DISPOSITION 06-16-10 8:45 P SCHEDULED EUTHANASIA | | | | | | | | |
| 73663 M 100 MG/KG/DAY NORMAL 06-02-10 8:52 P NO SIGNIFICANT CLINICAL OBSERVATIONS | 73663 | M | 100 MG/KG/DAY | NORMAL | | | | |
| 06-03-10 8:23 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-04-10 8:10 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-05-10 8:34 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-06-10 8:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-07-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-08-10 7:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-09-10 7:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-10-10 8:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-11-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-12-10 8:15 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | |
| 06-13-10 7:52 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | 06-13-10 | 7:52 | Р | NO SIGNIFICANT CLINICAL OBSERVATIONS |

PAGE 7

PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10

| | | | | | | ICAIVO | | - 02 - 1 | |
|----------------|-----|-----|------------------------|-----------------------|---|--|--|-----------------------|--|
| ANIMAL | SEX | | GROUP | CATEGORY | DATE | TI | IME (| GRADE | E OBSERVATIONS |
| 73663 | М | 100 | MG/KG/DAY | NORMAL | 06-14 06-15 06-16 | 5-10 | 7:49 7:44 7:48 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73663 73670 | | | MG/KG/DAY MG/KG/DAY | DISPOSITION NORMAL | 06-16 06-02 06-04 06-05 06-06 06-07 06-10 06-12 06-13 06-14 06-15 | 5-10 2-10 4-10 5-10 5-10 7-10 3-10 9-10 0-10 1-10 2-10 3-10 | 8:45 8:54 8:10 8:35 8:57 7:46 7:50 8:31 7:45 7:52 7:50 7:45 | P P P P P P P P P P P | SCHEDULED EUTHANASIA NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73670 | M | 100 | MG/KG/DAY | DISPOSITION | 06-16 06-16 | | 7:49 8:45 | | NO SIGNIFICANT CLINICAL OBSERVATIONS SCHEDULED EUTHANASIA |
| 73670 | | | MG/KG/DAY | | 06-03 | 3-10 | 8:24 | 1 | WET YELLOW MATERIAL RIGHT INGUINAL AREA WET YELLOW MATERIAL LEFT INGUINAL AREA |
| 73680 | М | 100 | MG/KG/DAY | NORMAL | 06 - 02 06 - 03 06 - 04 06 - 05 06 - 05 06 - 08 | 2-10 3-10 4-10 5-10 5-10 7-10 | 8:55 8:24 8:10 8:35 8:57 7:46 7:40 | P P P P P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | | |

PAGE

8

PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10 ANIMAL SEX GROUP CATEGORY DATE TIME GRADE OBSERVATIONS 73680 M 100 MG/KG/DAY NORMAL 06-10-10 8:32 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 8:16 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 7:52 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 7:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-16-10 7:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-16-10 8:45 P SCHEDULED EUTHANASIA 73680 M 100 MG/KG/DAY DISPOSITION 73683 M 100 MG/KG/DAY NORMAL 06-02-10 8:59 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 8:24 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 8:11 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 8:35 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 8:57 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:46 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 7:52 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 8:32 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 8:16 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 7:53 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 7:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 7:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-16-10 7:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS M 100 MG/KG/DAY DISPOSITION 06-16-10 8:45 P SCHEDULED EUTHANASIA 73683 73684 M 100 MG/KG/DAY NORMAL 06-02-10 8:59 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 8:25 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 8:11 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 8:35 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 8:57 P NO SIGNIFICANT CLINICAL OBSERVATIONS

PAGE 9 PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10

| ANIMAL | SEX | GROUP | CATEGORY | DATE | TIME | G | RADE | OBSERVATIONS |
|--------|-----|---------------|----------|--------|-----------|--------|------|---|
| 73684 | | 100 MG/KG/DAY | NORMAL | 06-07- | 10 7: | 47 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-08- | | 40 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-09- | | 52 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-10- | 10 8: | 32 | Р | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-11- | 10 7: | 46 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-12- | 10 8: | 16 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-13- | 10 7: | 53 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-14- | 10 7: | 50 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-15- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-16- | | 51 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73684 | | 100 MG/KG/DAY | | 06-16- | | | | SCHEDULED EUTHANASIA |
| 73667 | M | 300 MG/KG/DAY | NORMAL | 06-02- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-03- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-04- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-05- | | 35 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-06- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-07- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-08- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-09- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-10- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-11- | | 46 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-12- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-13- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-14- | | 51 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-15- | | 46 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 72667 | 3.4 | 200 MG/KG/DAY | | 06-16- | | 51 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73667 | | 300 MG/KG/DAY | | 06-16- | | | | SCHEDULED EUTHANASIA |
| 73674 | IvI | 300 MG/KG/DAY | NORMAL | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-03- | TO 8: | ∠⊃ | | NO SIGNIFICANI CLINICAL OBSERVALIONS |
| | | | | | | | | |

TABLE I1 (MALES - DAILY EXAMINATIONS)

ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 10 PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10

| ANIMAL | SEX | | GROUP | CATEGORY | DATE | TIN | ΊE | GRADE | OB | SERVATIONS | | | |
|--------|-----|-----|-----------|----------|--------|-----|------|-------|----|--------------|----------|--------------|--|
| 73674 | M | 300 | MG/KG/DAY | NORMAL | 06-04- | -10 | 8:11 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | | 06-05- | -10 | 8:36 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | | 06-06- | -10 | 8:58 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | | 06-07- | -10 | 7:47 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | | 06-08- | -10 | 7:41 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | | 06-09- | -10 | 7:54 | | | | | OBSERVATIONS | |
| | | | | | 06-10- | -10 | 8:33 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | | 06-11- | | 7:46 | | | | | OBSERVATIONS | |
| | | | | | 06-12- | -10 | 8:17 | | | | | OBSERVATIONS | |
| | | | | | 06-13- | | 7:53 | | | | | OBSERVATIONS | |
| | | | | | 06-14- | | 7:51 | | | | | OBSERVATIONS | |
| | | | | | 06-15- | | 7:46 | | | | | OBSERVATIONS | |
| | | | | | 06-16- | | 7:52 | | | | | OBSERVATIONS | |
| 73674 | | | MG/KG/DAY | | 06-16- | | 8:45 | | | EDULED EUTHA | | | |
| 73675 | M | 300 | MG/KG/DAY | NORMAL | 06-02- | | 9:01 | | | | | OBSERVATIONS | |
| | | | | | 06-03- | | 8:25 | | | | | OBSERVATIONS | |
| | | | | | 06-04- | | 8:12 | | | | | OBSERVATIONS | |
| | | | | | 06-05- | | 8:36 | | | | | OBSERVATIONS | |
| | | | | | 06-06- | | 8:58 | | | | | OBSERVATIONS | |
| | | | | | 06-07- | | 7:47 | | | | | OBSERVATIONS | |
| | | | | | 06-08- | | 7:41 | | | | | OBSERVATIONS | |
| | | | | | 06-09- | | 7:55 | | | | | OBSERVATIONS | |
| | | | | | 06-10- | | 8:34 | | | | | OBSERVATIONS | |
| | | | | | 06-11- | | 7:46 | | | | | OBSERVATIONS | |
| | | | | | 06-12- | | 8:17 | | | | | OBSERVATIONS | |
| | | | | | 06-13- | | 7:54 | | | | | OBSERVATIONS | |
| | | | | | 06-14- | | 7:51 | | | | | OBSERVATIONS | |
| | | | | | 06-15- | | 7:46 | | | | | OBSERVATIONS | |
| | | | | | 06-16- | -10 | 7:52 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |

PAGE 11 PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10

| | | | | | TABLE RA | ANGE | : 06- | 02-1 | 10 'I'O 06-16-10 |
|--------|-----|-----|-----------|-------------|----------|------|-------|------|--------------------------------------|
| ANIMAL | SEX | | GROUP | CATEGORY | DATE | TIM | E G | RADI | E OBSERVATIONS |
| 73675 | | | | | | | | | SCHEDULED EUTHANASIA |
| 73681 | М | 300 | MG/KG/DAY | NORMAL | 06-02-1 | | 9:03 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-03-1 | | 8:26 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-04-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-05-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-06-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-07-1 | | 7:47 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-08-1 | | 7:41 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-09-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-10-1 | | 8:34 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-11-1 | | 7:47 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-12-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-13-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-14-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-15-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-16-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73681 | | | | DISPOSITION | 06-16-1 | | | | SCHEDULED EUTHANASIA |
| 73685 | M | 300 | MG/KG/DAY | NORMAL | 06-02-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-03-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-04-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-05-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-06-1 | | 8:58 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-07-1 | | 7:48 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-08-1 | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-09-1 | 10 | 7:56 | Ρ | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-10-1 | 10 | 8:34 | Ρ | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-11-1 | | 7:47 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-12-1 | | 8:18 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-13-3 | 10 | 7:54 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | | |

PAGE 12 PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| | | | | | TABLE R | ANGE: | 06- | 02- | 10 TO 06-16-10 |
|---|--------------|-----|----------------------------------|----------|-------------------------------|----------------|----------------------|-------------|--|
| | ANIMAL | SEX | GROUP | CATEGORY | DATE | TIME | G1 | RAD: | E OBSERVATIONS |
| 7 | 3685 | М | 300 MG/KG/DAY | NORMAL | 06-14-1 06-15-1 06-16-1 | 10 7: | | P | NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | 3685 3671 | | 300 MG/KG/DAY 1000 MG/KG/DAY | NORMAL | 06-16-3 06-02-3 06-03-3 | 10 9: | :04 | | SCHEDULED EUTHANASIA NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | 3671 3671 | | 1000 MG/KG/DAY 1000 MG/KG/DAY | | 06-16- 06-04- | 10 8: 10 8: | :46 :13 | P 1 | SCHEDULED EUTHANASIA HAIR LOSS VENTRAL ABDOMINAL AREA |
| | | | | | 06-05-1 06-06-1 06-06-1 | 10 8: 10 8: | :59 | 2 1 | HAIR LOSS VENTRAL ABDOMINAL AREA HAIR LOSS VENTRAL ABDOMINAL AREA HAIR LOSS RIGHT HINDLIMB |
| | | | | | 06-06-1 06-07-1 | 10 7: | : 59 : 48 : 49 | 2 | HAIR LOSS LEFT HINDLIMB HAIR LOSS VENTRAL ABDOMINAL AREA HAIR LOSS RIGHT HINDLIMB |
| | | | | | 06-07- 06-08- | 10 7: 10 7: | 49:43:43 | 1 2 | HAIR LOSS LEFT HINDLIMB HAIR LOSS VENTRAL ABDOMINAL AREA HAIR LOSS RIGHT HINDLIMB |
| | | | | | 06-08- 06-09- | 10 7: 10 7: | :43 | 1 2 | HAIR LOSS LEFT HINDLIMB HAIR LOSS VENTRAL ABDOMINAL AREA |
| | | | | | 06-09-1 06-09-1 06-10-1 | 10 7: | | 1 1 1 | HAIR LOSS RIGHT HINDLIMB HAIR LOSS LEFT HINDLIMB HAIR LOSS RIGHT HINDLIMB |
| | | | | | 06-10- | 10 8: | :35 | 1 | HAIR LOSS LEFT HINDLIMB |

06-10-10 8:35 2 HAIR LOSS VENTRAL ABDOMINAL AREA 06-11-10 7:47 2 HAIR LOSS VENTRAL ABDOMINAL AREA

06-12-10 8:18 2 HAIR LOSS VENTRAL ABDOMINAL AREA

06-11-10 7:48 1 HAIR LOSS RIGHT HINDLIMB 06-11-10 7:48 1 HAIR LOSS LEFT HINDLIMB

06-12-10 8:18 1 HAIR LOSS RIGHT HINDLIMB

PAGE 13 PROJECT NO.:WIL-402010M INDIVIDUAL CLINICAL OBSERVATIONS SPONSOR: AMERICAN PETROLEUM

TABLE RANGE: 06-02-10 TO 06-16-10

| | | | | | TADLE K | | | - 0 2 - 1 | -10 10 00-10-10 |
|--------|-----|------|-----------|-----------------|--|--|--|--|--|
| ANIMAL | SEX | | GROUP | CATEGORY | DATE | TIME | | GRADE | DE OBSERVATIONS |
| 73671 | М | 1000 | MG/KG/DAY | BODY/INTEGUMENT | 06-12- 06-13- 06-13- 06-14- 06-14- | 10 7 10 7 10 7 10 7 | | 2 1 1 2 | HAIR LOSS LEFT HINDLIMB HAIR LOSS VENTRAL ABDOMINAL AREA |
| | | | | | 06-14- 06-15- 06-15- 06-15- 06-16- | 10 7 10 7 10 7 10 7 10 7 | 7:52 7:48 7:48 7:48 7:48 7:54 | 1 2 1 1 2 | HAIR LOSS LEFT HINDLIMB HAIR LOSS VENTRAL ABDOMINAL AREA HAIR LOSS RIGHT HINDLIMB HAIR LOSS LEFT HINDLIMB HAIR LOSS LEFT LATERAL ABDOMINAL AREA HAIR LOSS VENTRAL ABDOMINAL AREA |
| 73677 | М | 1000 | MG/KG/DAY | | 06-16- 06-16- 06-02- 06-03- 06-04- 06-05- 06-06- 06-07- 06-08- 06-10- 06-11- 06-12- 06-13- 06-14- | 10 7 10 9 10 8 10 8 10 8 10 7 10 7 10 7 10 8 10 7 10 8 10 7 | | 1 P P P P P P P P P P P P P P P P P P P | HAIR LOSS LEFT HINDLIMB NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-15- 06-16- | 10 7 | :49 :55 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS |

TABLE I1 (MALES - DAILY EXAMINATIONS)

PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 14 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-16-10 ______ ANIMAL SEX GROUP CATEGORY DATE TIME GRADE OBSERVATIONS 73677 M 1000 MG/KG/DAY DISPOSITION 06-16-10 8:46 P SCHEDULED EUTHANASIA 73678 M 1000 MG/KG/DAY NORMAL 06-02-10 9:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 8:27 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 7:44 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 7:59 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 8:36 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 7:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 8:19 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 7:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 7:53 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 7:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 73678 M 1000 MG/KG/DAY DISPOSITION 06-16-10 8:46 P SCHEDULED EUTHANASIA M 1000 MG/KG/DAY BODY/INTEGUMENT 06-04-10 8:14 2 WET YELLOW MATERIAL UROGENITAL AREA 73678 06-05-10 8:38 1 WET YELLOW MATERIAL UROGENITAL AREA 06-06-10 9:00 1 WET YELLOW MATERIAL UROGENITAL AREA 06-16-10 7:56 1 WET YELLOW MATERIAL UROGENITAL AREA M 1000 MG/KG/DAY NORMAL 73686 06-02-10 9:06 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 8:27 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 8:14 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 7:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 8:36 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 8:19 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 7:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 7:53 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 7:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 73686 M 1000 MG/KG/DAY DISPOSITION 06-16-10 8:46 P SCHEDULED EUTHANASIA 73686 M 1000 MG/KG/DAY BODY/INTEGUMENT 06-05-10 8:39 1 WET YELLOW MATERIAL UROGENITAL AREA

06-06-10 9:01 1 WET YELLOW MATERIAL UROGENITAL AREA

TABLE I1 (MALES - DAILY EXAMINATIONS) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

TABLE DANCE. 06 02 10 TO 06 16 10

| ANIMAL | SEX | | GROUP | CATEGORY | DATE T | IME | GRADI | E OBSERVATIONS |
|--------|-----|------|-----------|-----------------|----------|------|-------|--------------------------------------|
| 73686 | M | 1000 | MG/KG/DAY | BODY/INTEGUMENT | | | | WET YELLOW MATERIAL UROGENITAL AREA |
| | | | | | 06-09-10 | | | WET YELLOW MATERIAL UROGENITAL AREA |
| | | | | | 06-11-10 | | | |
| | | | | | 06-16-10 | | | WET YELLOW MATERIAL UROGENITAL AREA |
| 73690 | M | 1000 | MG/KG/DAY | NORMAL | 06-02-10 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-03-10 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-04-10 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-05-10 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-06-10 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-08-10 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-09-10 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-10-10 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-11-10 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | / / | | 06-12-10 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73690 | | | | | 06-16-10 | | | SCHEDULED EUTHANASIA |
| 73690 | M | 1000 | MG/KG/DAY | BODY/INTEGUMENT | | | | WET YELLOW MATERIAL UROGENITAL AREA |
| | | | | | 06-13-10 | | | WET YELLOW MATERIAL UROGENITAL AREA |
| | | | | | 06-14-10 | | | WET YELLOW MATERIAL UROGENITAL AREA |
| | | | | | 06-15-10 | | | WET YELLOW MATERIAL UROGENITAL AREA |
| | | | | | 06-16-10 | 7:58 | 3 1 | WET YELLOW MATERIAL UROGENITAL AREA |

GRADE CODE: 1 - SLIGHT 2 - MODERATE 3 - SEVERE P - PRESENT

PCRDv4.17 07/15/2010

PAGE 15

Page 194 of 360

PROJECT NO.:WIL-402010M

SPONSOR: AMERICAN PETROLEUM

TABLE I2 (MALES - AT TIME OF DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-15-10

| ANIMAL | SEX | GROUP | CATEGORY | DATE TIME GRADE OBSERVATIONS |
|--------|-----|----------------|------------------|---|
| | | | | AT TIME OF DOSING |
| 73670 | M | 100 MG/KG/DAY | ORAL/DENTAL | 06-09-10 9:59 1 SALIVATION PRIOR TO DOSING |
| | | | | 06-10-10 9:51 1 SALIVATION PRIOR TO DOSING |
| 73671 | M | 1000 MG/KG/DAY | ORAL/DENTAL | 06-08-10 10:10 2 SALIVATION PRIOR TO DOSING |
| 73677 | M | 1000 MG/KG/DAY | CARDIO-PULMONARY | 06-12-10 9:59 P GASPING |
| 73677 | M | 1000 MG/KG/DAY | ORAL/DENTAL | 06-10-10 10:04 2 SALIVATION PRIOR TO DOSING |
| | | | | 06-11-10 10:24 1 SALIVATION PRIOR TO DOSING |
| | | | | 06-12-10 9:56 2 SALIVATION PRIOR TO DOSING |
| 73678 | M | 1000 MG/KG/DAY | ORAL/DENTAL | 06-12-10 9:57 2 SALIVATION PRIOR TO DOSING |
| | | | • | 06-15-10 10:17 1 SALIVATION PRIOR TO DOSING |
| 73686 | M | 1000 MG/KG/DAY | ORAL/DENTAL | 06-08-10 10:12 2 SALIVATION PRIOR TO DOSING |
| | | | • | 06-11-10 10:25 2 SALIVATION PRIOR TO DOSING |
| | | | | 06-12-10 9:57 1 SALIVATION PRIOR TO DOSING |
| | | | | 06-13-10 9:42 1 SALIVATION PRIOR TO DOSING |
| | | | | 06-14-10 10:16 1 SALIVATION PRIOR TO DOSING |
| | | | | 06-15-10 10:18 2 SALIVATION PRIOR TO DOSING |
| 73690 | M | 1000 MG/KG/DAY | ORAL/DENTAL | 06-10-10 10:06 2 SALIVATION PRIOR TO DOSING |
| | | | , | 06-15-10 10:19 1 SALIVATION PRIOR TO DOSING |

GRADE CODE: 1 - SLIGHT 2 - MODERATE 3 - SEVERE P - PRESENT

PCRDv4.17 07/15/2010

PAGE 1

Page 195 of 360

TABLE I3 (MALES - 1 HOUR POST-DOSING)

ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| ANIMAI | SEX | GROUP | CATEGORY | DATE TIME GRADE OBSERVATIONS |
|--------|-----|-------------|----------|---|
| | | | | POST-DOSING OBSERVATIONS |
| 73665 | M | 0 MG/KG/DAY | NORMAL | 06-02-10 10:36 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | ,, | | 06-03-10 10:26 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-04-10 10:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-05-10 10:34 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-06-10 10:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-07-10 10:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-08-10 10:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-09-10 10:42 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-10-10 10:28 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-11-10 10:57 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-12-10 10:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-13-10 10:25 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-14-10 10:46 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-15-10 10:43 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73666 | M | 0 MG/KG/DAY | NORMAL | 06-02-10 10:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-03-10 10:27 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-04-10 10:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-05-10 10:35 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-06-10 10:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-07-10 10:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-08-10 10:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-09-10 10:42 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-10-10 10:28 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-11-10 10:57 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-12-10 10:38 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-13-10 10:25 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-14-10 10:46 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-15-10 10:44 P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73676 | M | 0 MG/KG/DAY | NORMAL | 06-02-10 10:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS |

PAGE 1

TABLE I3 (MALES - 1 HOUR POST-DOSING)

PAGE

2

ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-15-10 ANIMAL SEX GROUP CATEGORY DATE TIME GRADE OBSERVATIONS POST-DOSING OBSERVATIONS 73676 M 0 MG/KG/DAY NORMAL 06-03-10 10:27 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 10:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS

06-05-10 10:35 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 10:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 10:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 10:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 10:43 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:28 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 10:57 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 10:38 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:25 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 10:46 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 10:44 P NO SIGNIFICANT CLINICAL OBSERVATIONS 73691 M 0 MG/KG/DAY NORMAL 06-02-10 10:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 10:27 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 10:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 10:35 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 10:57 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 10:38 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 10:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 10:43 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:28 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 10:57 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 10:38 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:25 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 10:46 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 10:44 P NO SIGNIFICANT CLINICAL OBSERVATIONS 73692 M 0 MG/KG/DAY NORMAL 06-02-10 10:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 10:27 P NO SIGNIFICANT CLINICAL OBSERVATIONS

PAGE 3 PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-15-10

| ANIMAL | SEX | GROUP | CATEGORY | | TIME | | E OF | BSERVATIONS | | | |
|--------|-----|--------------|----------|--------|---------|-----|--------------|-------------|------------------|--------------|------|
| | | | | POS | | | ERV <i>I</i> | ATIONS | | | |
| 73692 | M | 0 MG/KG/DAY | NORMAL | 06-04- | 10 10:4 | 8 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-05- | 10 10:3 | 5 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-06- | 10 10:5 | 7 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-07- | 10 10:3 | 8 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-08- | 10 10:4 | 0 P | NO | SIGNIFICANT | ${\tt CLINICAL}$ | OBSERVATIONS | |
| | | | | 06-09- | 10 10:4 | 3 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-10- | 10 10:2 | 8 P | | | | OBSERVATIONS | |
| | | | | 06-11- | 10 10:5 | 7 P | NO | SIGNIFICANT | ${\tt CLINICAL}$ | OBSERVATIONS | |
| | | | | 06-12- | 10 10:3 | 9 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-13- | 10 10:2 | 6 P | | | | OBSERVATIONS | |
| | | | | 06-14- | | | | | | OBSERVATIONS | |
| | | | | 06-15- | | | | | | OBSERVATIONS | |
| 73668 | M | 30 MG/KG/DAY | NORMAL | 06-02- | | | | | | OBSERVATIONS | |
| | | | | 06-03- | | | | | | OBSERVATIONS | |
| | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | OBSERVATIONS | |
| | | | | 06-07- | | | | | | OBSERVATIONS | |
| | | | | 06-08- | | | | | | OBSERVATIONS | |
| | | | | 06-09- | | | | | | OBSERVATIONS | |
| | | | | 06-10- | | | | | | OBSERVATIONS | |
| | | | | 06-11- | | | | | | OBSERVATIONS | |
| | | | | 06-12- | | | | | | OBSERVATIONS | |
| | | | | 06-13- | | | | | | OBSERVATIONS | |
| | | | | 06-14- | | | | | | OBSERVATIONS | |
| | | | | | | | | | | OBSERVATIONS | |
| 73669 | M | 30 MG/KG/DAY | NORMAL | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | OBSERVATIONS | |
| | | | | 06-04- | 10 10:4 | 8 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |

PAGE 4

PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| | | | TABLE RANGE: 06 | 5-02- | 2-10 TO 06-15-10 |
|------------|--------------|----------|--|---|---|
| ANIMAL SEX | GROUP | CATEGORY | DATE TIME | GRAD: | ADE OBSERVATIONS |
| 73669 M | 30 MG/KG/DAY | NORMAL | 06-06-10 10:57 06-07-10 10:38 06-08-10 10:41 06-09-10 10:42 06-10-10 10:34 06-11-10 10:58 06-12-10 10:39 06-13-10 10:26 | F P P P P P P P P P P P P P P P P P P P | BSERVATIONS P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73682 M | 30 MG/KG/DAY | NORMAL | 06-02-10 10:38 06-03-10 10:28 06-04-10 10:48 06-05-10 10:38 06-07-10 10:39 06-08-10 10:44 06-09-10 10:44 06-10-10 10:38 06-11-10 10:58 06-12-10 10:40 06-13-10 10:40 | 8 P P P P P P P P P P P P P P P P P P P | P NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73687 M | 30 MG/KG/DAY | | 06-02-10 10:39 06-03-10 10:28 06-04-10 10:49 | 5 P 9 P 8 P | P NO SIGNIFICANT CLINICAL OBSERVATIONS |

PAGE 5

PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-15-10 ANIMAL SEX GROUP CATEGORY DATE TIME GRADE OBSERVATIONS POST-DOSING OBSERVATIONS 73687 M 30 MG/KG/DAY NORMAL 06-06-10 10:58 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 10:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 10:41 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 10:44 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:35 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 10:58 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 10:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:27 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 10:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 10:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 73688 M 30 MG/KG/DAY NORMAL 06-02-10 10:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 10:28 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 10:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 10:58 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 10:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 10:42 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 10:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:35 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 10:58 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 10:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:27 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 10:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 10:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 73663 M 100 MG/KG/DAY NORMAL 06-02-10 10:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-03-10 10:37 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-04-10 10:53 P NO SIGNIFICANT CLINICAL OBSERVATIONS

06-05-10 10:41 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:03 P NO SIGNIFICANT CLINICAL OBSERVATIONS

Page 201 of 360

PROJECT NO.:WIL-402010M SPONSOR:AMERICAN PETROLEUM

TABLE I3 (MALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

PAGE 6

TABLE RANGE: 06-02-10 TO 06-15-10

| | | | | | | | | | | | | |
|--------|-----|------------------|---------------|--------|--------|------|-------|------|-------------|----------|------------------------------|---|
| ANIMAL | SEX | GROUP | CATEGORY | DATE | J. T M | E (| JRADE | S OF | BSERVATIONS | | | |
| | | | | POS' | T-DO: | STNG | OBSE | :PVZ | TTONS | | | _ |
| 73663 | M | 100 MG/KG/DAY | | | | | | | | CLINICAL | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | 06-10- | 10 1 | 0:37 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-11- | 10 1 | 0:59 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-12- | 10 1 | 0:41 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-13- | 10 1 | 0:37 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-14- | 10 1 | 0:48 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-15- | | | | | | | OBSERVATIONS | |
| 73670 | M | 100 MG/KG/DAY | NORMAL | 06-02- | | | | | | | OBSERVATIONS | |
| | | | | 06-03- | | | | | | | OBSERVATIONS | |
| | | | | 06-04- | | | | | | | OBSERVATIONS | |
| | | | | 06-05- | | | | | | | OBSERVATIONS | |
| | | | | 06-06- | | | | | | | OBSERVATIONS | |
| | | | | 06-07- | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | 06-09- | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | 06-11- | | | | | | | OBSERVATIONS | |
| | | | | 06-12- | | | | | | | OBSERVATIONS | |
| | | | | 06-13- | | | | | | | OBSERVATIONS | |
| 50650 | | 100 100/110/1211 | 0037 /0037037 | 06-15- | | | | | | | OBSERVATIONS | |
| | | 100 MG/KG/DAY | | 06-14- | | | | | CLEAR MATER | | | |
| 73680 | M | 100 MG/KG/DAY | NORMAL | 06-02- | | | | | | | OBSERVATIONS | |
| | | | | 06-03- | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS OBSERVATIONS | |
| | | | | | | | | | SIGNIFICANI | | OBSERVATIONS | |
| | | | | | | | | | | | | |

Page 202 of 360

TABLE I3 (MALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 7 PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| | | | | | TABLE RANG | GE: 06 | -02-1 | 10 1 | 0 06-15-10 | | | | |
|---|--------|-----|---------------|----------|--|---|------------------|--|---|---|--|------|--|
| | ANIMAL | SEX | GROUP | CATEGORY | DATE T | IME | GRADI | E OE | SERVATIONS | | | | |
| | 73680 | М | 100 MG/KG/DAY | NORMAL | 06-09-10 06-10-10 06-11-10 06-12-10 06-13-10 | 10:48 10:46 10:37 10:59 10:41 10:38 | P P P P | NO NO NO NO NO | SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT | CLINICAL CLINICAL CLINICAL CLINICAL | OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS | | |
| ı | 73683 | М | 100 MG/KG/DAY | | 06-02-10 06-03-10 06-04-10 06-05-10 06-06-10 | 10:40 10:38 10:54 10:41 11:04 10:44 | P P P P | NO NO NO NO NO | SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT | CLINICAL CLINICAL CLINICAL CLINICAL CLINICAL CLINICAL | OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS | | |
| | 73684 | М | 100 MG/KG/DAY | | 06-13-10 06-14-10 06-15-10 06-02-10 06-03-10 06-04-10 06-05-10 06-06-10 06-07-10 | 10:38 10:59 10:42 10:38 10:49 10:46 10:40 10:38 10:54 10:42 11:04 | P P P P P P P P | NO N | SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT | CLINICAL | OBSERVATIONS | | |

PAGE 8 PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| TARTE | RANGE: | 06-02- | .1∩ ⊤ | 0 00 | 5 _ 1 5 | -10 | |
|-------|--------|--------|-------|------|---------|-----|--|
| | | | | | | | |

| ANIMAL | SEX | | GROUP | CATEGORY | DATE | 11T | ИЕ: | GRADE | | SERVATIONS | | |
|--------|-----|-----|-----------|----------|---------|---------|---------|-------|-----|-------------|------------------|--------------|
| | | | | CATEGORY | | | | | | | | |
| | | | | | POST | - DC | OSING | OBSE | RVA | ATIONS | | |
| 73684 | M | 100 | MG/KG/DAY | NORMAL | | | | | | | CLINICAL | OBSERVATIONS |
| | | | | | 06-10-3 | LO 1 | 10:38 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-11-3 | LO 1 | 11:00 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-12-1 | LO 1 | 10:42 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-13-1 | LO 1 | 10:38 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-14-1 | LO 1 | 10:49 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-15-1 | LO 1 | 10:47 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| 73667 | M | 300 | MG/KG/DAY | NORMAL | 06-02-1 | LO 1 | 10:47 | P | NO | SIGNIFICANT | ${\tt CLINICAL}$ | OBSERVATIONS |
| | | | | | 06-03-1 | LO 1 | 10:39 | | | | | OBSERVATIONS |
| | | | | | 06-04-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-05-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-06-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-07-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-08-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-09-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-10-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-11-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-12-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-13-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-14-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-15-1 | | | | | | | OBSERVATIONS |
| 73674 | M | 300 | MG/KG/DAY | | 06-02-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-03-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-04-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-05-1 | | | | | | | OBSERVATIONS |
| | | | | | | | | | | | | OBSERVATIONS |
| | | | | | | | | | | | | OBSERVATIONS |
| | | | | | 06-08-1 | | | | | | | OBSERVATIONS |
| | | | | | 06-09-1 | 10 1 | 10:58 | Ь | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |

PROJECT NO.:WIL-402010M

TABLE I3 (MALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 9

SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| | ANIMAL | SEX | | GROUP | CATEGORY | DATE TIME GRADE OBSERVATIONS | |
|---|--------|-----|-----|-------------|-------------|---|--|
| | | | | | | POST-DOSING OBSERVATIONS | |
| 7 | 3674 | M | 300 | MG/KG/DAY | ΝΟΡΜΔΤ. | 06-10-10 10:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| , | 3074 | 1-1 | 500 | MO/ NO/ DAI | NOIGHAL | 06-11-10 11:09 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-12-10 10:43 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-13-10 10:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-14-10 10:59 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-15-10 11:01 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| 7 | 3675 | M | 300 | MG/KG/DAY | NORMAL | 06-02-10 10:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-03-10 10:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-04-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-05-10 10:46 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-06-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-07-10 10:54 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-08-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-09-10 10:59 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-10-10 10:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-11-10 11:09 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-12-10 10:43 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-13-10 10:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-14-10 10:59 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-15-10 11:01 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| 7 | 3681 | M | 300 | MG/KG/DAY | NORMAL | 06-02-10 10:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-03-10 10:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-04-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-05-10 10:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-06-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-08-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| | | | | | | 06-11-10 11:09 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |
| 7 | 3681 | M | 300 | MG/KG/DAY | ORAL/DENTAL | 06-07-10 10:54 1 DRIED CLEAR MATERIAL AROUND MOUTH | |
| | | | | | | 06-09-10 10:59 1 DRIED CLEAR MATERIAL AROUND MOUTH | |

PROJECT NO.:WIL-402010M

Page 205 of 360

TABLE I3 (MALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

PAGE 10

TABLE RANGE: 06-02-10 TO 06-15-10

| | | | | TABLE F | CANGE: U | 5-02 | 10 10 06-15-10 |
|--------|-----|----------------|---------------------|---------|-----------|------|---|
| ANIMAL | SEX | GROUP | CATEGORY | DATE | TIME | GRAD | E OBSERVATIONS |
| | | | | POS | T-DOSTNO | OBS | ERVATIONS |
| 73681 | М | 300 MG/KG/DAY | ORAL/DENTAL | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | -, -, | , | | | | DRIED CLEAR MATERIAL AROUND MOUTH |
| | | | | 06-13- | 10 10:40 |) 1 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | 06-14- | 10 11:00 |) 1 | DRIED CLEAR MATERIAL AROUND MOUTH |
| | | | | 06-15- | 10 11:01 | L 1 | WET CLEAR MATERIAL AROUND MOUTH |
| 73685 | M | 300 MG/KG/DAY | NORMAL | 06-02- | 10 10:49 |) P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-03- | 10 10:40 |) P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 10 10:56 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 10 11:06 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 10 10:59 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 10 10:40 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73685 | M | 300 MG/KG/DAY | ORAL/DENTAL | | 10 10:47 | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 10 10:55 | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 10 10:56 | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 10 10:49 | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 10 11:09 | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 10 10:44 | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 10 11:00 | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 10 11:02 | | WET CLEAR MATERIAL AROUND MOUTH |
| 73671 | М | 1000 MG/KG/DAY | NORMAL | | 10 10:49 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 10 10:51 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 10 11:20 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 10 11:03 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| F2.6F1 | 3.6 | 1000 MG/RG/DAR | DODY / TAMBECI MENT | | 10 10:59 | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73671 | M | 1000 MG/KG/DAY | BODY/INTEGUMENT | | | | WET CLEAR MATERIAL RIGHT FORELIMB |
| F2.6F1 | 3.6 | 1000 MG/RG/DAR | EVEC /EXPC /NOCE | | 10 11:03 | | WET CLEAR MATERIAL LEFT FORELIMB |
| 73671 | ΙVI | TUUU MG/KG/DAY | EYES/EARS/NOSE | | | | DRIED RED MATERIAL AROUND NOSE WET CLEAR MATERIAL AROUND NOSE |
| 73671 | М | 1000 MC/KC/DAY | ODAT /DENTEAT | | 10 11:03 | | WET CLEAR MATERIAL AROUND MOUTH |
| 13011 | 1*1 | 1000 MG/KG/DAI | ORALI/ DENTAL | 00-04- | .10 10:26 |) I | WEI CHEAR PHIERIAL AROUND PROUIR |

PAGE 11

PROJECT NO.:WIL-402010M SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| ANIMAL SEX GROUP CATEGORY DATE TIME GRADE OBSERVATIONS POST-DOSING OBSERVATIONS 73671 M 1000 MG/KG/DAY ORAL/DENTAL 06-05-10 10:49 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-09-10 11:00 1 DRIED RED MATERIAL AROUND MOUTH 06-10-10 10:49 1 WET CLEAR MATERIAL AROUND MOUTH 06-11-10 11:11 2 WET CLEAR MATERIAL AROUND MOUTH 06-12-10 10:45 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-14-10 11:02 2 WET CLEAR MATERIAL AROUND MOUTH 06-14-10 11:02 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:41 2 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:42 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH | |
|---|--|
| 73671 M 1000 MG/KG/DAY ORAL/DENTAL 06-05-10 10:49 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-09-10 11:00 1 DRIED RED MATERIAL AROUND MOUTH 06-10-10 10:49 1 WET CLEAR MATERIAL AROUND MOUTH 06-11-10 11:11 2 WET CLEAR MATERIAL AROUND MOUTH 06-12-10 10:45 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-12-10 10:45 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:41 2 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 1 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL LEFT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE | |
| 06-09-10 11:00 1 DRIED RED MATERIAL AROUND MOUTH 06-10-10 10:49 1 WET CLEAR MATERIAL AROUND MOUTH 06-11-10 11:11 2 WET CLEAR MATERIAL AROUND MOUTH 06-12-10 10:45 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-13-10 10:45 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-14-10 11:02 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-13-10 10:42 1 WET CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 2 WET CLEAR MATERIAL VENTRAL NECK 06-13-10 10:42 2 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL LEFT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL LEFT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND MOUTH 06-03-10 10:41 2 WET CLEAR MATERIAL AROUND MOUTH 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH 06-03-10 | |
| 06-10-10 10:49 1 WET CLEAR MATERIAL AROUND MOUTH 06-11-10 11:11 2 WET CLEAR MATERIAL AROUND MOUTH 06-11-10 10:45 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-12-10 10:45 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-14-10 11:02 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:41 2 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 1 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL LEFT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND MOUTH 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND MOUTH 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH 06-03-03-03-03-03-03-03-03-03-03-03-03-03- | |
| 06-11-10 11:11 2 WET CLEAR MATERIAL AROUND MOUTH 06-12-10 10:45 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 10:42 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS NORMAL 06-02-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:41 WET CLEAR MATERIAL VENTRAL NECK 06-13-10 10:42 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 WET CLEAR MATERIAL LEFT FORELIMB 06-13-10 10:42 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 WET CLEAR MATERIAL AROUND NOSE 06-03-10 10:51 WET CLEAR MATERIAL AROUND MOUTH WET CLEAR MATERIAL WET | |
| 06-12-10 10:45 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-13-10 10:41 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-14-10 11:02 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 WET CLEAR MATERIAL AROUND MOUTH 06-02-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS O6-13-10 10:41 WET CLEAR MATERIAL VENTRAL NECK 06-13-10 10:42 WET CLEAR MATERIAL RIGHT FORELIMB O6-13-10 10:42 WET CLEAR MATERIAL LEFT FORELIMB O6-13-10 10:42 WET CLEAR MATERIAL AROUND NOSE O6-13-10 10:41 WET CLEAR MATERIAL AROUND NOSE O6-13-10 10:41 WET CLEAR MATERIAL AROUND NOSE O6-03-10 10:51 WET CLEAR MATERIAL AROUND MOUTH O6-03-10 10:51 WET CLEAR MATERIAL AROUND MOUTH OF CLEAR MATERIAL AROUND | |
| 06-13-10 10:41 1 DRIED CLEAR MATERIAL AROUND MOUTH 06-14-10 11:02 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-02-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:41 2 WET CLEAR MATERIAL VENTRAL NECK 06-13-10 10:42 1 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL LEFT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH 06-03-03-03-03-03-03-03-03-03-03-03-03-03- | |
| 06-14-10 11:02 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:02 1 WET CLEAR MATERIAL AROUND MOUTH 06-02-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:41 2 WET CLEAR MATERIAL VENTRAL NECK 06-13-10 10:42 1 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:44 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH 06-03-03-03-03-03-03-03-03-03-03-03-03-03- | |
| 73677 M 1000 MG/KG/DAY NORMAL 06-02-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 73677 M 1000 MG/KG/DAY BODY/INTEGUMENT 06-13-10 10:41 2 WET CLEAR MATERIAL VENTRAL NECK 06-13-10 10:42 1 WET CLEAR MATERIAL VENTRAL NECK 06-13-10 10:42 2 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL LEFT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH | |
| 73677 M 1000 MG/KG/DAY NORMAL 06-02-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 73677 M 1000 MG/KG/DAY BODY/INTEGUMENT 06-13-10 10:41 2 WET CLEAR MATERIAL VENTRAL NECK 06-13-10 10:42 1 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL LEFT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:46 3 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:45 1 WET CLEAR MATERIAL AROUND NOSE | |
| 73677 M 1000 MG/KG/DAY BODY/INTEGUMENT 06-13-10 10:41 2 WET CLEAR MATERIAL VENTRAL NECK 06-13-10 10:42 1 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL LEFT FORELIMB 73677 M 1000 MG/KG/DAY EYES/EARS/NOSE 06-12-10 10:46 3 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 73677 M 1000 MG/KG/DAY ORAL/DENTAL 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-13-10 10:42 1 WET CLEAR MATERIAL RIGHT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL LEFT FORELIMB 06-13-10 10:42 2 WET CLEAR MATERIAL AROUND NOSE 06-12-10 10:46 3 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH | |
| 73677 M 1000 MG/KG/DAY EYES/EARS/NOSE 06-13-10 10:42 2 WET CLEAR MATERIAL LEFT FORELIMB 06-12-10 10:46 3 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 73677 M 1000 MG/KG/DAY ORAL/DENTAL 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH | |
| 73677 M 1000 MG/KG/DAY EYES/EARS/NOSE 06-12-10 10:46 3 WET CLEAR MATERIAL AROUND NOSE 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 73677 M 1000 MG/KG/DAY ORAL/DENTAL 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-13-10 10:41 2 WET CLEAR MATERIAL AROUND NOSE 73677 M 1000 MG/KG/DAY ORAL/DENTAL 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH | |
| 73677 M 1000 MG/KG/DAY ORAL/DENTAL 06-03-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH | |
| | |
| 00-04-10 10.30 1 WEI CHEAR MAIERIAH AROUND MOOIH | |
| 06-05-10 10:49 2 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-06-10 11:21 1 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-07-10 11:04 1 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-08-10 10:59 2 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-09-10 11:01 1 DRIED CLEAR MATERIAL AROUND MOUTH | |
| 06-10-10 10:49 2 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-11-10 11:12 1 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-12-10 10:45 3 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-12-10 10:46 1 SALIVATION | |
| 06-13-10 10:41 3 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-14-10 11:03 2 WET CLEAR MATERIAL AROUND MOUTH | |
| 06-15-10 11:03 1 WET CLEAR MATERIAL AROUND MOUTH | |
| 73678 M 1000 MG/KG/DAY NORMAL 06-02-10 10:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS | |

Page 207 of 360

TABLE I3 (MALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

PAGE 12

TABLE RANGE: 06-02-10 TO 06-15-10

| ANIMAL | SEX | | GROUP | CATEGORY | DATE | TIME | GRAD | E OBSERVATIONS |
|--------|-----|------|-----------|-----------------|--------|----------|------|--------------------------------------|
| | | | | | POS | T-DOSING | OBS | ERVATIONS |
| 73678 | M | 1000 | MG/KG/DAY | NORMAL | 06-04- | 10 10:57 | 7 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-05- | 10 10:49 |) P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-06- | 10 11:21 | . Р | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-07- | 10 11:04 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-08- | 10 10:59 |) P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-10- | 10 10:50 |) P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-13- | 10 10:42 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 3678 | M | 1000 | MG/KG/DAY | BODY/INTEGUMENT | 06-12- | 10 10:47 | 7 3 | WET CLEAR MATERIAL VENTRAL NECK |
| | | | | | 06-12- | 10 10:47 | 7 1 | WET CLEAR MATERIAL RIGHT FORELIMB |
| | | | | | 06-12- | 10 10:47 | 7 2 | WET CLEAR MATERIAL LEFT FORELIMB |
| | | | | | 06-14- | 10 11:04 | 1 | WET CLEAR MATERIAL VENTRAL NECK |
| | | | | | 06-14- | 10 11:04 | 1 | WET CLEAR MATERIAL RIGHT FORELIMB |
| | | | | | 06-14- | 10 11:04 | | WET CLEAR MATERIAL LEFT FORELIMB |
| | | | | | | 10 11:03 | | |
| 3678 | M | 1000 | MG/KG/DAY | EYES/EARS/NOSE | 06-11- | 10 11:12 | 1 | WET CLEAR MATERIAL AROUND NOSE |
| | | | | | | 10 11:04 | | |
| 73678 | M | 1000 | MG/KG/DAY | ORAL/DENTAL | 06-03- | 10 10:51 | . 1 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-09- | 10 11:01 | . 1 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-11- | 10 11:12 | 2 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-12- | 10 10:47 | 7 2 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-14- | 10 11:04 | 2 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-15- | 10 11:03 | 2 | WET CLEAR MATERIAL AROUND MOUTH |
| 73686 | M | 1000 | MG/KG/DAY | NORMAL | 06-02- | 10 10:50 |) P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-04- | 10 10:57 | 7 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-05- | 10 10:50 |) P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-06- | 10 11:21 | . P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-07- | 10 11:05 | 5 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-08- | 10 11:00 |) P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | _ | NO SIGNIFICANT CLINICAL OBSERVATIONS |

| | | | | - | rable R | ANGE: 06 | -02- | 10 TO 06-15-10 |
|----------------|-----|------|-----------|----------------|---|--|---|--|
| ANIMAL | SEX | | GROUP | CATEGORY | DATE | TIME | GRAD: | E OBSERVATIONS |
| 73686 73686 | | | | NORMAL | 06-11-3 06-14-3 | 10 11:13 10 11:05 | P P | REVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS WET CLEAR MATERIAL AROUND NOSE |
| 73686 | M | 1000 | MG/KG/DAY | ORAL/DENTAL | 06-03-1 06-09-1 06-12-1 06-13-1 | 10 10:52 10 11:02 10 10:48 10 10:42 | 1 1 1 | WET CLEAR MATERIAL AROUND MOUTH WET CLEAR MATERIAL AROUND MOUTH DRIED CLEAR MATERIAL AROUND MOUTH DRIED CLEAR MATERIAL AROUND MOUTH WET CLEAR MATERIAL AROUND MOUTH |
| 73690 | M | 1000 | MG/KG/DAY | | 06-04-3 06-07-3 | 10 10:57 10 11:05 | P P | NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73690 | М | 1000 | MG/KG/DAY | | 06-05-1 06-05-1 06-12-1 06-12-1 06-13-1 06-13-1 06-15-1 | 10 10:51 10 10:52 10 10:49 10 10:49 10 10:49 10 10:43 10 10:43 10 11:04 10 11:05 | 1 1 3 3 2 1 1 2 2 | WET CLEAR MATERIAL VENTRAL NECK WET CLEAR MATERIAL RIGHT FORELIMB WET CLEAR MATERIAL LEFT FORELIMB WET CLEAR MATERIAL RIGHT FORELIMB WET CLEAR MATERIAL RIGHT FORELIMB WET CLEAR MATERIAL VENTRAL NECK WET CLEAR MATERIAL RIGHT FORELIMB WET CLEAR MATERIAL LEFT FORELIMB WET CLEAR MATERIAL LEFT FORELIMB WET CLEAR MATERIAL VENTRAL NECK WET CLEAR MATERIAL VENTRAL NECK WET CLEAR MATERIAL LEFT FORELIMB WET CLEAR MATERIAL LEFT FORELIMB |
| 73690 | M | 1000 | MG/KG/DAY | EYES/EARS/NOSE | 06-05-1 06-06-1 06-12-1 06-13-1 | 10 10:51 10 11:23 10 10:49 10 10:43 10 11:04 | 1 1 3 2 2 | WET CLEAR MATERIAL AROUND NOSE |
| 73690 | M | 1000 | MG/KG/DAY | ORAL/DENTAL | 06-03-3 | 10 10:52 | 2 | WET CLEAR MATERIAL AROUND MOUTH |

GRADE CODE: 1 - SLIGHT 2 - MODERATE 3 - SEVERE P - PRESENT

PROJECT NO.:WIL-402010M SPONSOR:AMERICAN PETROLEUM

Page 208 of 360

PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN SPONSOR:AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| | | | | | E RANGE: 06-02-10 TO 06-15-10 | |
|--------|-----|------|-----------|-------------|--|---|
| ANIMAL | SEX | | GROUP | CATEGORY | E TIME GRADE OBSERVATIONS | |
| 73690 | М | 1000 | MG/KG/DAY | ORAL/DENTAL | POST-DOSING OBSERVATIONS 05-10 10:50 3 WET CLEAR MATERIAL ARG 06-10 11:23 3 WET CLEAR MATERIAL ARG 08-10 11:01 2 WET CLEAR MATERIAL ARG 10-10 10:51 2 WET CLEAR MATERIAL ARG 11-10 11:13 3 WET CLEAR MATERIAL ARG 12-10 10:49 3 SALIVATION 12-10 10:49 3 WET CLEAR MATERIAL ARG 13-10 10:43 2 WET CLEAR MATERIAL ARG 13-10 10:43 2 WET CLEAR MATERIAL ARG 14-10 11:05 2 WET CLEAR MATERIAL ARG | OUND MOUTH OUND MOUTH OUND MOUTH OUND MOUTH OUND MOUTH OUND MOUTH |
| | | | | | L5-10 11:04 3 WET CLEAR MATERIAL ARG | OUND MOUTH |

GRADE CODE: 1 - SLIGHT 2 - MODERATE 3 - SEVERE P - PRESENT

PCRDv4.17 07/15/2010

TABLE 14 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL BODY WEIGHTS [G]

| | | | | MALES |
|-----------|---------------|----------|------|-------|
| | DAY 0 | 7 | 14 | |
| ANIMALS 1 | FROM GROUP 1: | 0 MG/KG/ | DAY | |
| 73665 | 327. | 371. | 421. | |
| 73666 | 318. | 354. | 398. | |
| 73676 | 326. | 379. | 433. | |
| 73691 | 304. | 343. | 383. | |
| 73692 | 302. | 347. | 393. | |
| MEAN | 315. | 359. | 406. | |
| S.D. | 11.9 | 15.3 | 20.6 | |
| S.E. | 5.3 | 6.9 | 9.2 | |
| N | 5 | 5 | 5 | |

TABLE I4 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL BODY WEIGHTS [G]

PAGE 2

------MALES DAY 0 7 14

| ANIMALS | FROM GROUP 2: | 30 MG/KG | /DAY |
|---------|---------------|----------|------|
| 73668 | 304. | 341. | 371. |
| 73669 | 324. | 367. | 410. |
| 73682 | 331. | 377. | 416. |
| 73687 | 307. | 349. | 394. |
| 73688 | 322. | 358. | 400. |
| MEAN | 317. | 359. | 398. |
| S.D. | 11.8 | 14.2 | 17.5 |
| S.E. | 5.3 | 6.3 | 7.8 |
| N | 5 | 5 | 5 |

TABLE 14 (MALES) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL BODY WEIGHTS [G]

| | DAY 0 | 7 | 14 | MALES |
|-----------|---------------|-----------|------|-------|
| ANIMALS E | FROM GROUP 3: | 100 MG/KG | /DAY | |
| 73663 | 335. | 373. | 414. | |
| 73670 | 327. | 373. | 423. | |
| 73680 | 308. | 341. | 375. | |
| 73683 | 300. | 337. | 372. | |
| 73684 | 315. | 349. | 394. | |
| MEAN | 317. | 354. | 396. | |
| S.D. | 13.9 | 17.6 | 22.8 | |
| S.E. | 6.2 | 7.9 | 10.2 | |
| N | 5 | 5 | 5 | |

TABLE 14 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL BODY WEIGHTS [G]

PAGE 4

| | DAY 0 | 7 | 14 | MALES |
|-----------|--------------|-----------|------|-------|
| ANIMALS F | ROM GROUP 4: | 300 MG/KG | /DAY | |
| 73667 | 291. | 325. | 350. | |
| 73674 | 311. | 346. | 385. | |
| 73675 | 310. | 355. | 393. | |
| 73681 | 331. | 393. | 447. | |
| 73685 | 339. | 380. | 423. | |
| MEAN | 316. | 360. | 399. | |
| S.D. | 19.0 | 27.1 | 37.3 | |
| S.E. | 8.5 | 12.1 | 16.7 | |
| N | 5 | 5 | 5 | |

TABLE 14 (MALES)
ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010M INDIVIDUAL BODY WEIGHTS [G] SPONSOR: AMERICAN PETROLEUM

| | DAY 0 | 7 | 14 | MALES |
|-----------|--------------|-----------|-------|------------|
| ANIMALS F | ROM GROUP 5: | 1000 MG/K | G/DAY | |
| 73671 | 330. | 340. | 365. | |
| 73677 | 308. | 350. | 374. | |
| 73678 | 352. | 371. | 427. | |
| 73686 | 318. | 337. | 343. | |
| 73690 | 281. | 308. | 337. | |
| MEAN | 318. | 341. | 369. | |
| S.D. | 26.2 | 23.1 | 35.8 | |
| S.E. | 11.7 | 10.3 | 16.0 | |
| N | 5 | 5 | 5 | |
| | | | | PJTBWv4.23 |
| | | | | 07/15/2010 |

07/15/2010

SPONSOR: AMERICAN PETROLEUM

TABLE I5 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL BODY WEIGHT CHANGES [G]

| DA | AY 0- 7 | 7- 14 | 0- 14 | MALES |
|------------------------------------|--------------------------|--------------------------|---------------------------|-------|
| ANIMALS FROM | GROUP 1: | 0 MG/KG/I | DAY | |
| 73665 73666 73676 73691 | 44. 36. 53. 40. | 50. 44. 55. 40. | 93. 79. 108. 80. | |
| 73692 MEAN S.D. S.E. N | 45. 43. 6.5 2.9 | 46. 47. 5.6 2.5 | 91. 90. 11.6 5.2 | |

SPONSOR: AMERICAN PETROLEUM

TABLE 15 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL BODY WEIGHT CHANGES [G]

| | | | | MALES |
|--------------|----------|-----------|-------|-------|
| D. | AY 0- 7 | 7- 14 | 0- 14 | |
| ANIMALS FROM | GROUP 2: | 30 MG/KG/ | DAY | |
| 73668 | 38. | 30. | 67. | |
| 73669 | 43. | 43. | 86. | |
| 73682 | 46. | 39. | 85. | |
| 73687 | 42. | 45. | 87. | |
| 73688 | 37. | 42. | 79. | |
| MEAN | 41. | 40. | 81. | |
| S.D. | 3.9 | 6.1 | 8.2 | |
| S.E. | 1.7 | 2.7 | 3.7 | |
| N | 5 | 5 | 5 | |

TABLE I5 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL BODY WEIGHT CHANGES [G]

| | | | | MALES |
|-------------|------------|------------|-------|-------|
| | DAY 0- 7 | 7- 14 | 0- 14 | |
| ANIMALS FRO | M GROUP 3: | 100 MG/KG/ | DAY | |
| 73663 | 38. | 41. | 79. | |
| 73670 | 46. | 50. | 96. | |
| 73680 | 33. | 35. | 67. | |
| 73683 | 36. | 36. | 72. | |
| 73684 | 34. | 45. | 79. | |
| MEAN | 37. | 41. | 79. | |
| S.D. | 5.3 | 6.5 | 11.1 | |
| S.E. | 2.4 | 2.9 | 4.9 | |
| N | 5 | 5 | 5 | |

SPONSOR: AMERICAN PETROLEUM

TABLE I5 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL BODY WEIGHT CHANGES [G]

| | DAY 0- 7 | 7- 14 | 0- 14 | MALES |
|-----------|---------------|-----------|-------|-------|
| ANIMALS I | FROM GROUP 4: | 300 MG/KG | J/DAY | |
| 73667 | 34. | 24. | 58. | |
| 73674 | 35. | 39. | 74. | |
| 73675 | 45. | 38. | 83. | |
| 73681 | 62. | 54. | 116. | |
| 73685 | 41. | 43. | 83. | |
| MEAN | 43. | 40. | 83. | |
| S.D. | 11.6 | 10.6 | 21.3 | |
| S.E. | 5.2 | 4.7 | 9.5 | |
| N | 5 | 5 | 5 | |

TABLE I5 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RAT SPONSOR:AMERICAN PETROLEUM INDIVIDUAL BODY WEIGHT CHANGES [G]

| • | | | | | |
|------------|-------|--------|----------|---------------|---------|
| | MALES | 0- 14 | 7- 14 | DAY 0- 7 | |
| | | KG/DAY | 1000 MG/ | FROM GROUP 5: | ANIMALS |
| | | 35. | 25. | 11. | 73671 |
| | | 65. | 24. | 41. | 73677 |
| | | 76. | 56. | 20. | 73678 |
| | | 25. | 6. | 19. | 73686 |
| | | 56. | 29. | 27. | 73690 |
| | | 52. | 28. | 24. | MEAN |
| | | 20.9 | 17.9 | 11.5 | S.D. |
| | | 9.3 | 8.0 | 5.1 | S.E. |
| | | 5 | 5 | 5 | N |
| PJTBWv4.23 | | | | | |
| 07/15/2010 | | | | | |

07/15/2010

SPONSOR: AMERICAN PETROLEUM

TABLE I6 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FOOD CONSUMPTION [G/ANIMAL/DAY]

PAGE 1

MALES DAY 0- 7 7- 14 0- 14 ______ ANIMALS FROM GROUP 1: 0 MG/KG/DAY 73665 22. 21. 21. 18. 19. 22. 22. 17. 17. 19. 20. 19. 22. 17. 73666 73676 73691 20. 73692 20. MEAN 19. 20. 2.2 2.1 2.1 S.D. 1.0 1.0 S.E. 0.9 N 5 5 5

S.E.

N

0.8

5

0.7

5

0.8

5

TABLE I6 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FOOD CONSUMPTION [G/ANIMAL/DAY]

PAGE 2

______ MALES DAY 0- 7 7-14 0-14 ______ ANIMALS FROM GROUP 2: 30 MG/KG/DAY 73668 18. 16. 17. 22. 20. 21. 21. 20. 21. 21. 20. 20. 73669 22. 20. 73682 73687 73688 20. MEAN 20. 1.7 20. 20. 1.9 1.7 S.D.

SPONSOR: AMERICAN PETROLEUM

5

5

5

N

TABLE I6 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FOOD CONSUMPTION [G/ANIMAL/DAY]

PAGE 3

______ MALES DAY 0- 7 7-14 0-14 ______ ANIMALS FROM GROUP 3: 100 MG/KG/DAY 73663 20. 19. 19. 22. 22. 16. 18. 17. 18. 19. 19. 22. 19. 19. 73670 73680 73683 20. 73684 20. MEAN 18. 19. 2.0 1.7 1.4 S.D. 0.9 0.8 0.6 S.E.

Page 222 of 360

SPONSOR: AMERICAN PETROLEUM

N

5

5

5

TABLE I6 (MALES) PROJECT NO.: WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FOOD CONSUMPTION [G/ANIMAL/DAY]

PAGE 4

______ MALES DAY 0- 7 7-14 0-14 ______ ANIMALS FROM GROUP 4: 300 MG/KG/DAY 73667 17. 14. 16. 18. 19. 23. 19. 19. 23. 20. 20. 23. 73674 73675 73681 73685 19. MEAN 20. 18. 19. 2.4 3.1 2.7 2.4 S.D. 1.4 1.1 1.2 S.E.

Page 223 of 360

TABLE I6 (MALES)
ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010M INDIVIDUAL FOOD CONSUMPTION [G/ANIMAL/DAY] SPONSOR: AMERICAN PETROLEUM

| D | DAY 0- 7 | 7- 14 | 0- 14 | MALES |
|--------------|------------|-----------|-------|-------------|
| ANIMALS FROM | GROUP 5: 1 | 000 MG/KG | /DAY | |
| 73671 | 15. | 14. | 14. | |
| 73677 | 19. | 16. | 18. | |
| 73678 | 18. | 20. | 19. | |
| 73686 | 15. | 15. | 15. | |
| 73690 | 17. | 16. | 16. | |
| MEAN | 17. | 16. | 16. | |
| S.D. | 2.0 | 2.4 | 2.0 | |
| S.E. | 0.9 | 1.1 | 0.9 | |
| N | 5 | 5 | 5 | |
| | | | | DTTEWs:4 21 |

PJTFWv4.21 07/15/2010

SPONSOR:AMERICAN PETROLEUM

TABLE 17 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FOOD CONSUMPTION [G/KG/DAY]

| I | DAY 0- 7 | 7- 14 | 0- 14 | MALES |
|--------------|------------|-----------|-------|-------|
| ANIMALS FROM | M GROUP 1: | 0 MG/KG/D |)AY | |
| 73665 | 62. | 52. | 56. | |
| 73666 | 57. | 48. | 52. | |
| 73676 | 63. | 55. | 59. | |
| 73691 | 53. | 46. | 49. | |
| 73692 | 63. | 51. | 57. | |
| MEAN | 59. | 50. | 55. | |
| S.D. | 4.6 | 3.5 | 4.0 | |
| S.E. | 2.1 | 1.6 | 1.8 | |
| N | 5 | 5 | 5 | |

SPONSOR:AMERICAN PETROLEUM

TABLE 17 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FOOD CONSUMPTION [G/KG/DAY]

| I | DAY 0- 7 | 7- 14 | 0- 14 | MALES |
|--------------|------------|-----------|-------|-------|
| ANIMALS FROM | M GROUP 2: | 30 MG/KG/ | /DAY | |
| 73668 | 55. | 46. | 50. | |
| 73669 | 63. | 52. | 57. | |
| 73682 | 62. | 52. | 57. | |
| 73687 | 61. | 55. | 58. | |
| 73688 | 60. | 53. | 56. | |
| MEAN | 60. | 52. | 56. | |
| S.D. | 3.0 | 3.7 | 3.1 | |
| S.E. | 1.3 | 1.6 | 1.4 | |
| N | 5 | 5 | 5 | |

SPONSOR: AMERICAN PETROLEUM

TABLE I7 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FOOD CONSUMPTION [G/KG/DAY]

PAGE 3

------MALES DAY 0- 7 7-14 0-14

| ANIMALS | FROM | GROUP | 3: | 100 | MG/KG/DAY | Z |
|---|------|-------|--------------------------|-----|--------------------------|--------------------------|
| 73663 73670 73680 73683 73684 | | | 56. 64. 59. 58. | | 48. 54. 46. 48. | 52. 59. 52. 53. |
| MEAN S.D. S.E. N | | | 60. 2.8 1.3 5 | | 49. 3.1 1.4 5 | 54. 2.8 1.3 5 |

Page 227 of 360

PROJECT NO.:WIL-402010M SPONSOR:AMERICAN PETROLEUM

TABLE I7 (MALES) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FOOD CONSUMPTION [G/KG/DAY]

PAGE 4

.....

| | DAY 0- 7 | 7- 14 | 0- 14 | MALES |
|------------|-------------|------------|-------|-------|
| ANIMALS FR | OM GROUP 4: | 300 MG/KG/ | DAY | |
| 73667 | 55. | 43. | 48. | |
| 73674 | 60. | 49. | 54. | |
| 73675 | 61. | 50. | 55. | |
| 73681 | 65. | 55. | 59. | |
| 73685 | 53. | 43. | 48. | |
| MEAN | 59. | 48. | 53. | |
| S.D. | 4.8 | 5.0 | 4.9 | |
| S.E. | 2.1 | 2.2 | 2.2 | |
| N | 5 | 5 | 5 | |

PROJECT NO.:WIL-402010M

TABLE I7 (MALES) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FOOD CONSUMPTION [G/KG/DAY]

| PONSOR: AMERICA | | Olds | | INDIVIDUAL FOOD CONSUMPTION [G/KG/DAY] | CD | J |
|-----------------|--------------|------------|-------|--|--------|---|
| | | 7 14 | 0- 14 | MALES | | |
| | DAY 0- 7 | | | | | |
| ANIMALS FROM | M GROUP 5: 1 | 1000 MG/KG | /DAY | | | |
| 73671 | 43. | 40. | 41. | | | |
| 73677 | 58. | 45. | 51. | | | |
| 73678 | 50. | 51. | 50. | | | |
| 73686 | 44. | 44. | 45. | | | |
| 73690 | 57. | 50. | 53. | | | |
| MEAN | 50. | 46. | 48. | | | |
| S.D. | 6.8 | 4.4 | 4.9 | | | |
| S.E. | 3.1 | 2.0 | 2.2 | | | |
| N | 5 | 5 | 5 | | | |
| | | | | | JTFWv4 | |

07/15/2010

Page 230 of 360

TABLE I8 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

ANIMAL NO. 73665 GROUP 1: 0 MG/KG/DAY MALE SCHEDULED EUTH 06/16/10 DATE OF DEATH: 06/16/10 GRADE ORGAN WEIGHT ABS.(G) REL. NO SIGNIFICANT EPIDIDYMIDES INTESTINE LIVER 19.20 4.561 CHANGES OBSERVED GROSS:ADRENAL GLANDS BRAIN ESOPHAGUS 3.57 0.848 EYES INTESTINE KIDNEYS HEART KIDNEYS 1.56 0.371 1.55 0.368 0.4311 0.102 TESTIS, RIGHT LIVER LN, MESENTERIC LUNGS MAMMARY GLAND PANCREAS TESTIS, LEFT PITUITARY PROSTATE SPINAL CORD SAL. GLAND MAND SKIN SPLEEN STOMACH
SEMINAL VESICLES THYMUS THYROID GLANDS TRACHEA
URINARY BLADDER DIAPHRAGM TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) 421.

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

Page 231 of 360

TABLE I8 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

| ANIMAL NO. 736 | 666 GROUP | 1: 0 M | G/KG/DAY | MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: (| 06/16/10 | GRADE |
|------------------|-----------|--------|-----------|-----------|----------------------|----------------|------------------|---------------|-------|
| ORGAN WEIGHT | ABS.(G) | REL. | NO SIGNIF | ICANT | | | | | |
| LIVER | 18.27 | 4.590 | CHANGES O | BSERVED | GROSS:ADRENAL GLANDS | BRAIN | EPIDIDYMIDES | ESOPHAGUS | |
| KIDNEYS | 2.98 | 0.749 | | | EYES | HEART | INTESTINE | KIDNEYS | |
| TESTIS, RIGHT | 1.68 | 0.422 | | | LIVER | LN, MESENTERIC | LUNGS | MAMMARY GLAND | |
| TESTIS, LEFT | 1.72 | 0.432 | | | PANCREAS | PITUITARY | PROSTATE | SPINAL CORD | |
| THYMUS | 0.4517 | 0.113 | | | SAL. GLAND MAND | SKIN | SPLEEN | STOMACH | |
| FINAL BODY WT(G) |) 398. | | | | SEMINAL VESICLE | S THYMUS | THYROID GLANDS | S TRACHEA | |
| | , | | | | URINARY BLADDER | DIAPHRAGM | TESTIS, RIGHT | TESTIS, LEFT | |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

TABLE 18 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

| ANIMAL NO. 7367 | 6 GROUP | 1: 0 M | G/KG/DAY | MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: | 06/16/10 | GRADE |
|------------------|---------|--------|------------|--------|----------------------|----------------|----------------|---------------|-------|
| ORGAN WEIGHT | ABS.(G) | REL. | NO SIGNIFI | CANT | | | | | |
| LIVER | 21.50 | 4.965 | CHANGES OB | SERVED | GROSS:ADRENAL GLANDS | BRAIN | EPIDIDYMIDES | ESOPHAGUS | |
| KIDNEYS | 3.41 | 0.788 | | | EYES | HEART | INTESTINE | KIDNEYS | |
| TESTIS, RIGHT | 1.93 | 0.446 | | | LIVER | LN, MESENTERIC | LUNGS | MAMMARY GLAND | |
| TESTIS, LEFT | 1.94 | 0.448 | | | PANCREAS | PITUITARY | PROSTATE | SPINAL CORD | |
| THYMUS | 0.7997 | 0.185 | | | SAL. GLAND MAND | SKIN | SPLEEN | STOMACH | |
| FINAL BODY WT(G) | 433. | | | | SEMINAL VESICLE | S THYMUS | THYROID GLAND | S TRACHEA | |
| , , , | | | | | URINARY BLADDER | DIAPHRAGM | TESTIS, RIGHT | TESTIS, LEFT | |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

| ANIMAL NO. 7369 | 1 GROUP | 1: 0 M | G/KG/DAY MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: 0 | 6/16/10 GRADE |
|------------------|---------|--------|------------------|----------------------|----------------|------------------|------------------|
| ORGAN WEIGHT | ABS.(G) | REL. | NO SIGNIFICANT | | | | |
| LIVER | 15.25 | 3.982 | CHANGES OBSERVED | GROSS:ADRENAL GLANDS | BRAIN | EPIDIDYMIDES | ESOPHAGUS |
| KIDNEYS | 2.78 | 0.726 | | EYES | HEART | INTESTINE | KIDNEYS |
| TESTIS, RIGHT | 1.64 | 0.428 | | LIVER | LN, MESENTERIC | LUNGS | MAMMARY GLAND |
| TESTIS, LEFT | 1.62 | 0.423 | | PANCREAS | PITUITARY | PROSTATE | SPINAL CORD |
| THYMUS | 0.4888 | 0.128 | | SAL. GLAND MAND | SKIN | SPLEEN | STOMACH |
| FINAL BODY WT(G) | 383. | | | SEMINAL VESICLE | S THYMUS | THYROID GLANDS | TRACHEA |
| | | | | URINARY BLADDER | DIAPHRAGM | TESTIS, RIGHT | TESTIS, LEFT |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

| ANIMAL NO. 73692 | GROUP | 1: 0 MG | /KG/DAY | MALE S | CHEDULED EUTH (| 06/16/10 | DATE OF DEATH: (| 06/16/10 | GRADE |
|--|--|---|----------------------------|--------|---|--|---|---|-------|
| ORGAN WEIGHT ILIVER KIDNEYS TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) | ABS.(G) 17.80 3.22 1.89 1.72 0.5043 393. | REL. 4.529 0.819 0.481 0.438 0.128 | NO SIGNIFIC CHANGES OBS | | S:ADRENAL GLANDS EYES LIVER PANCREAS SAL. GLAND MAND SEMINAL VESICLES | BRAIN HEART LN, MESENTERIC PITUITARY SKIN THYMUS | EPIDIDYMIDES INTESTINE LUNGS PROSTATE SPLEEN THYROID GLANDS | ESOPHAGUS KIDNEYS MAMMARY GLAND SPINAL CORD STOMACH S TRACHEA | |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

TABLE I8 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

ANIMAL NO. 73668 GROUP 2: 30 MG/KG/DAY MALE SCHEDULED EUTH 06/16/10 DATE OF DEATH: 06/16/10 GRADE ORGAN WEIGHT ABS.(G) REL. NO SIGNIFICANT EPIDIDYMIDES INTESTINE LIVER 16.87 4.547 CHANGES OBSERVED GROSS:ADRENAL GLANDS BRAIN ESOPHAGUS 3.07 0.827 EYES INTESTINE KIDNEYS HEART KIDNEYS 1.76 0.474 1.68 0.453 TESTIS, RIGHT LIVER LN, MESENTERIC LUNGS MAMMARY GLAND TESTIS, LEFT 0.7645 0.206 PANCREAS PITUITARY PROSTATE SPINAL CORD SAL. GLAND MAND SKIN SPLEEN STOMACH
SEMINAL VESICLES THYMUS THYROID GLANDS TRACHEA
URINARY BLADDER DIAPHRAGM TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) 371.

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

TABLE 18 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

| ANIMAL NO. | 73669 | GROUP | 2: | 30 N | MG/KG/DAY | MALE | SC | HEDULED EU | JTH 0 | 6/16/10 | DATE (| OF DEATH: | 06/16/ | 10 | GRADE |
|---------------|-------|--------|-----|------|-----------|----------|-------|------------|----------|----------------|--------|------------|--------|-------------|-------|
| ORGAN WEIGHT | A | BS.(G) | RE | | NO SIGNI | | | | | | | | | | |
| LIVER | | 18.77 | 4.5 | 78 | CHANGES | OBSERVED | GROSS | :ADRENAL (| GLANDS | BRAIN | EPII | DIDYMIDES | ES | OPHAGUS | |
| KIDNEYS | | 3.08 | 0.7 | 51 | | | | EYES | | HEART | INT | ESTINE | KI | DNEYS | |
| TESTIS, RIGHT | 1 | 1.64 | 0.4 | 00 | | | | LIVER | | LN, MESENTERIC | LUNG | 3S | MZ | MMARY GLAND | |
| TESTIS, LEFT | | 1.62 | 0.3 | 95 | | | | PANCREAS | | PITUITARY | PROS | STATE | SE | INAL CORD | |
| THYMUS | | 0.5729 | 0.1 | 40 | | | | SAL. GLAN | ND MAND | SKIN | SPL | EEN | SI | OMACH | |
| FINAL BODY WT | '(G) | 410. | | | | | | SEMINAL V | JESICLES | THYMUS | THY | ROID GLANI | DS TF | ACHEA | |
| | | | | | | | | URINARY E | BLADDER | DIAPHRAGM | TES: | ris, Right | T TE | STIS, LEFT | |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

PAGE 7

Page 236 of 360

TABLE 18 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

| ANIMAL NO. 7368 | 32 GROUP | 2: 30 | MG/KG/DAY MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: 0 | 6/16/10 GRADE |
|---|--------------------------------|-------------------------|-------------------------|--|--|---|---|
| ORGAN WEIGHT LIVER KIDNEYS | ABS.(G) 19.05 3.02 | REL. 4.579 0.726 | KIDNEYS NO SIGNIFICANT | GROSS: PALE BILATERAL | | | P |
| TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) | 1.95 1.90 0.5944 416. | 0.469 0.457 0.143 | CHANGES OBSERVED | GROSS:ADRENAL GLANDS EYES LN, MESENTERIC PITUITARY SKIN THYMUS DIAPHRAGM | BRAIN HEART LUNGS PROSTATE SPLEEN THYROID GLANDS TESTIS, RIGHT | EPIDIDYMIDES INTESTINE MAMMARY GLAND SPINAL CORD STOMACH TRACHEA TESTIS, LEFT | ESOPHAGUS LIVER PANCREAS SAL. GLAND MAND SEMINAL VESICLES URINARY BLADDER |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

| ANIMAL NO. 7368 | 37 GROUP | 2: 30 | MG/KG/DAY MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: 0 | 6/16/10 GRADE |
|---|--|---|------------------------------------|---|----------|---|--|
| ORGAN WEIGHT LIVER KIDNEYS TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) | ABS.(G) 19.02 3.48 1.86 1.87 0.6234 394. | REL. 4.827 0.883 0.472 0.475 0.158 | NO SIGNIFICANT CHANGES OBSERVED | GROSS:ADRENAL GLANDS EYES LIVER PANCREAS SAL. GLAND MAND SEMINAL VESICLE: URINARY BLADDER | S THYMUS | EPIDIDYMIDES INTESTINE LUNGS PROSTATE SPLEEN THYROID GLANDS TESTIS, RIGHT | ESOPHAGUS KIDNEYS MAMMARY GLAND SPINAL CORD STOMACH TRACHEA TESTIS, LEFT |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

TABLE I8 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

ANIMAL NO. 73688 GROUP 2: 30 MG/KG/DAY MALE SCHEDULED EUTH 06/16/10 DATE OF DEATH: 06/16/10 GRADE ORGAN WEIGHT ABS.(G) REL. NO SIGNIFICANT EPIDIDYMIDES INTESTINE LIVER 16.77 4.193 CHANGES OBSERVED GROSS:ADRENAL GLANDS BRAIN ESOPHAGUS 3.59 0.897 EYES INTESTINE KIDNEYS HEART KIDNEYS 1.94 0.485 1.91 0.477 0.6785 0.170 TESTIS, RIGHT LIVER LN, MESENTERIC LUNGS MAMMARY GLAND TESTIS, LEFT PANCREAS PITUITARY PROSTATE SPINAL CORD SAL. GLAND MAND SKIN SPLEEN STOMACH
SEMINAL VESICLES THYMUS THYROID GLANDS TRACHEA
URINARY BLADDER DIAPHRAGM TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) 400.

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

PAGE 10

Page 239 of 360

| ANIMAL NO. 736 | 663 GROUP | 3: 100 I | MG/KG/DAY MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: 06 | 6/16/10 GRADE |
|--|---|---|------------------------------------|--|----------|---|--|
| ORGAN WEIGHT LIVER KIDNEYS TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G | ABS. (G) 17.99 3.81 2.07 2.12 0.7315 | REL. 4.345 0.920 0.500 0.512 0.177 | NO SIGNIFICANT CHANGES OBSERVEI | GROSS:ADRENAL GLANDS EYES LIVER PANCREAS SAL. GLAND MAND SEMINAL VESICLE URINARY BLADDER | S THYMUS | EPIDIDYMIDES INTESTINE LUNGS PROSTATE SPLEEN THYROID GLANDS TESTIS, RIGHT | ESOPHAGUS KIDNEYS MAMMARY GLAND SPINAL CORD STOMACH TRACHEA TESTIS, LEFT |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

| ANIMAL NO. 736 | 70 GROUP | 3: 100 1 | MG/KG/DAY | MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: | 06/16/10 | GRADE |
|------------------|----------|----------|-------------|--------|----------------------|----------------|----------------|---------------|-------|
| ORGAN WEIGHT | ABS.(G) | REL. | NO SIGNIFIC | CANT | | | | | |
| LIVER | 18.68 | 4.416 | CHANGES OB | SERVED | GROSS:ADRENAL GLANDS | BRAIN | EPIDIDYMIDES | ESOPHAGUS | |
| KIDNEYS | 3.27 | 0.773 | | | EYES | HEART | INTESTINE | KIDNEYS | |
| TESTIS, RIGHT | 1.70 | 0.402 | | | LIVER | LN, MESENTERIC | LUNGS | MAMMARY GLAND | |
| TESTIS, LEFT | 1.68 | 0.397 | | | PANCREAS | PITUITARY | PROSTATE | SPINAL CORD | |
| THYMUS | 0.5558 | 0.131 | | | SAL. GLAND MAND | SKIN | SPLEEN | STOMACH | |
| FINAL BODY WT(G) | 423. | | | | SEMINAL VESICLE | S THYMUS | THYROID GLAND | S TRACHEA | |
| | | | | | URINARY BLADDER | DIAPHRAGM | TESTIS, RIGHT | TESTIS, LEFT | |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

| ANIMAL NO. | 73680 GROUP | 3: 100 | MG/KG/DAY MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: 06 | 7/16/10 GRADE |
|--|---|---|-----------------------------------|---|--|---|--|
| ORGAN WEIGHT LIVER KIDNEYS TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT | ABS. (G) 17.72 3.06 1.56 1.60 0.8100 (G) 375. | REL. 4.725 0.816 0.416 0.427 0.216 | NO SIGNIFICANT CHANGES OBSERVE | GROSS:ADRENAL GLANDS EYES LIVER PANCREAS SAL. GLAND MAND SEMINAL VESICLE: URINARY BLADDER | BRAIN HEART LN, MESENTERIC PITUITARY SKIN S THYMUS DIAPHRAGM | EPIDIDYMIDES INTESTINE LUNGS PROSTATE SPLEEN THYROID GLANDS TESTIS, RIGHT | ESOPHAGUS KIDNEYS MAMMARY GLAND SPINAL CORD STOMACH TRACHEA TESTIS, LEFT |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

ANIMAL NO. 73683 GROUP 3: 100 MG/KG/DAY MALE SCHEDULED EUTH 06/16/10 DATE OF DEATH: 06/16/10 GRADE ORGAN WEIGHT ABS.(G) REL. NO SIGNIFICANT EPIDIDYMIDES INTESTINE LIVER 16.91 4.546 CHANGES OBSERVED GROSS:ADRENAL GLANDS BRAIN ESOPHAGUS 3.19 0.858 INTESTINE EYES KIDNEYS HEART KIDNEYS 1.68 0.452 1.67 0.449 TESTIS, RIGHT LIVER LN, MESENTERIC LUNGS MAMMARY GLAND PANCREAS TESTIS, LEFT 0.5300 0.142 PITUITARY PROSTATE SPINAL CORD SAL. GLAND MAND SKIN SPLEEN STOMACH
SEMINAL VESICLES THYMUS THYROID GLANDS TRACHEA
URINARY BLADDER DIAPHRAGM TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) 372.

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

TABLE I8 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

ANIMAL NO. 73684 GROUP 3: 100 MG/KG/DAY MALE SCHEDULED EUTH 06/16/10 DATE OF DEATH: 06/16/10 GRADE ORGAN WEIGHT ABS.(G) REL. NO SIGNIFICANT EPIDIDYMIDES INTESTINE LIVER 18.54 4.706 CHANGES OBSERVED GROSS:ADRENAL GLANDS BRAIN ESOPHAGUS 2.89 0.734 1.52 0.386 1.68 0.426 EYES INTESTINE KIDNEYS HEART KIDNEYS TESTIS, RIGHT LIVER LN, MESENTERIC LUNGS MAMMARY GLAND PANCREAS TESTIS, LEFT 0.5175 0.131 PITUITARY PROSTATE SPINAL CORD SAL. GLAND MAND SKIN SPLEEN STOMACH
SEMINAL VESICLES THYMUS THYROID GLANDS TRACHEA
URINARY BLADDER DIAPHRAGM TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) 394.

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

TABLE 18 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

| ANIMAL NO. | 73667 | GROUP | 4: 300 | MG/KG/DAY | MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: | 06/16/10 | GRADE |
|---------------|-------|--------|--------|-----------|----------|----------------------|----------------|----------------|---------------|-------|
| ORGAN WEIGHT | A | BS.(G) | REL. | NO SIGNI | FICANT | | | | | |
| LIVER | | 17.00 | 4.857 | CHANGES | OBSERVED | GROSS:ADRENAL GLANDS | BRAIN | EPIDIDYMIDES | ESOPHAGUS | |
| KIDNEYS | | 2.91 | 0.831 | | | EYES | HEART | INTESTINE | KIDNEYS | |
| TESTIS, RIGHT | 1 | 1.54 | 0.440 | | | LIVER | LN, MESENTERIC | LUNGS | MAMMARY GLAND | |
| TESTIS, LEFT | | 1.52 | 0.434 | | | PANCREAS | PITUITARY | PROSTATE | SPINAL CORD | |
| THYMUS | | 0.4702 | 0.134 | | | SAL. GLAND MAND | SKIN | SPLEEN | STOMACH | |
| FINAL BODY WT | '(G) | 350. | | | | SEMINAL VESICLE | S THYMUS | THYROID GLAND | S TRACHEA | |
| | , | | | | | URINARY BLADDER | | TESTIS. RIGHT | | |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

TABLE I8 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

ANIMAL NO. 73674 GROUP 4: 300 MG/KG/DAY MALE SCHEDULED EUTH 06/16/10 DATE OF DEATH: 06/16/10 GRADE ORGAN WEIGHT ABS.(G) REL. NO SIGNIFICANT EPIDIDYMIDES INTESTINE LIVER 18.98 4.930 CHANGES OBSERVED GROSS:ADRENAL GLANDS BRAIN ESOPHAGUS 3.34 0.868 EYES INTESTINE KIDNEYS HEART KIDNEYS 1.73 0.449 1.60 0.416 0.4546 0.118 TESTIS, RIGHT LIVER LN, MESENTERIC LUNGS MAMMARY GLAND PANCREAS TESTIS, LEFT PITUITARY PROSTATE SPINAL CORD SAL. GLAND MAND SKIN SPLEEN STOMACH
SEMINAL VESICLES THYMUS THYROID GLANDS TRACHEA
URINARY BLADDER DIAPHRAGM TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) 385.

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

PAGE 17

Page 246 of 360

| ANIMAL NO. 7367 | 5 GROUP | 4: 300 | MG/KG/DAY MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: 0 | | RADE |
|------------------|---------|--------|------------------|----------------------|----------------|------------------|---------------|------|
| | | | | | | | | |
| ORGAN WEIGHT | ABS.(G) | REL. | NO SIGNIFICANT | | | | | |
| LIVER | 20.97 | 5.336 | CHANGES OBSERVED | GROSS:ADRENAL GLANDS | BRAIN | EPIDIDYMIDES | ESOPHAGUS | |
| KIDNEYS | 3.22 | 0.819 | | EYES | HEART | INTESTINE | KIDNEYS | |
| TESTIS, RIGHT | 1.62 | 0.412 | | LIVER | LN, MESENTERIC | LUNGS | MAMMARY GLAND | |
| TESTIS, LEFT | 1.59 | 0.405 | | PANCREAS | PITUITARY | PROSTATE | SPINAL CORD | |
| THYMUS | 0.8064 | 0.205 | | SAL. GLAND MAND | SKIN | SPLEEN | STOMACH | |
| FINAL BODY WT(G) | 393. | | | SEMINAL VESICLE | S THYMUS | THYROID GLANDS | TRACHEA | |
| | | | | URINARY BLADDER | DIAPHRAGM | TESTIS, RIGHT | TESTIS, LEFT | |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

| ANIMAL NO. 7368 | 31 GROUP | 4: 300 | MG/KG/DAY MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: 0 | 6/16/10 GRADE |
|---|--------------------------------|-------------------------|-------------------------|--|--|---|---|
| ORGAN WEIGHT LIVER KIDNEYS | ABS.(G) 24.27 3.89 | REL. 5.430 0.870 | KIDNEYS NO SIGNIFICANT | GROSS: PALE BILATERAL | | | P |
| TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) | 1.75 1.75 0.7538 447. | 0.383 0.391 0.169 | CHANGES OBSERVED | GROSS:ADRENAL GLANDS EYES LN, MESENTERIC PITUITARY SKIN THYMUS DIAPHRAGM | BRAIN HEART LUNGS PROSTATE SPLEEN THYROID GLANDS TESTIS, RIGHT | EPIDIDYMIDES INTESTINE MAMMARY GLAND SPINAL CORD STOMACH TRACHEA TESTIS, LEFT | ESOPHAGUS LIVER PANCREAS SAL. GLAND MAND SEMINAL VESICLES URINARY BLADDER |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

TABLE I8 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

ANIMAL NO. 73685 GROUP 4: 300 MG/KG/DAY MALE SCHEDULED EUTH 06/16/10 DATE OF DEATH: 06/16/10 GRADE ORGAN WEIGHT ABS.(G) REL. NO SIGNIFICANT EPIDIDYMIDES INTESTINE LIVER 22.67 5.359 CHANGES OBSERVED GROSS:ADRENAL GLANDS BRAIN ESOPHAGUS 3.54 0.837 EYES INTESTINE KIDNEYS HEART KIDNEYS 1.95 0.461 1.98 0.468 TESTIS, RIGHT LIVER LN, MESENTERIC LUNGS MAMMARY GLAND PANCREAS TESTIS, LEFT 0.8258 0.195 PITUITARY PROSTATE SPINAL CORD SAL. GLAND MAND SKIN SPLEEN STOMACH
SEMINAL VESICLES THYMUS THYROID GLANDS TRACHEA
URINARY BLADDER DIAPHRAGM TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) 423.

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

PAGE 20

Page 249 of 360

TABLE I8 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM INDIVIDUAL MACROSCOPIC FINDINGS

| i | ANIMAL NO. 73 | 671 GROUP | 5: 1000 | MG/KG/DAY MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: 06/ | , | ADE |
|---|-----------------|-----------|---------|------------------|----------------------|-------------|--------------------|-----------------|-----|
| | ORGAN WEIGHT | ABS.(G) | REL. | KIDNEYS | GROSS: PALE | | | | P |
| | LIVER | 22.79 | 6.244 | | BILATERAL | | | | |
| | KIDNEYS | 4.26 | 1.167 | SKIN | GROSS: HAIR LOSS | | | | P |
| | TESTIS, RIGHT | 1.68 | 0.460 | | VENTRAL ABDO | MINAL | | | |
| | TESTIS, LEFT | 1.65 | 0.452 | NO SIGNIFICANT | | | | | |
| | THYMUS | 0.4357 | 0.119 | CHANGES OBSERVED | GROSS:ADRENAL GLANDS | BRAIN | EPIDIDYMIDES | ESOPHAGUS | |
| | FINAL BODY WT(G | 365. | | | EYES | HEART | INTESTINE | LIVER | |
| | | | | | LN, MESENTERIC | LUNGS | MAMMARY GLAND | PANCREAS | |
| | | | | | PITUITARY | PROSTATE | SPINAL CORD | SAL. GLAND MAND | |
| | | | | | SPLEEN | STOMACH | SEMINAL VESICLES | THYMUS | |
| | | | | | THYROID GLANDS | TRACHEA | URINARY BLADDER | DIAPHRAGM | |
| | | | | | TESTIS RIGHT | TESTIS LEFT | | | |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

| ANIMAL NO. 7367 | 77 GROUP | 5: 1000 | MG/KG/DAY MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: 06 | /16/10 GRADE |
|---|--|---|------------------------------------|--|----------|---|--|
| ORGAN WEIGHT LIVER KIDNEYS TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) | ABS.(G) 25.81 3.77 1.90 1.75 0.3647 374. | REL. 6.901 1.008 0.508 0.468 0.098 | NO SIGNIFICANT CHANGES OBSERVEI | GROSS:ADRENAL GLANDS EYES LIVER PANCREAS SAL. GLAND MAND SEMINAL VESICLE URINARY BLADDER | S THYMUS | EPIDIDYMIDES INTESTINE LUNGS PROSTATE SPLEEN THYROID GLANDS TESTIS, RIGHT | ESOPHAGUS KIDNEYS MAMMARY GLAND SPINAL CORD STOMACH TRACHEA TESTIS, LEFT |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

TABLE 18 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

| ANIMAL NO. 7367 | 8 GROUP | 5: 1000 | MG/KG/DAY | MALE | SCHEDULED EUTH | 06/16/10 | DATE OF DEATH: | 06/16/10 | GRADE |
|------------------|---------|---------|-----------|-----------|----------------------|----------------|----------------|---------------|-------|
| ORGAN WEIGHT | ABS.(G) | REL. | NO SIGNIF | ICANT | | | | | |
| LIVER | 27.28 | 6.389 | CHANGES O | BSERVED | GROSS:ADRENAL GLANDS | BRAIN | EPIDIDYMIDES | ESOPHAGUS | |
| KIDNEYS | 3.84 | 0.899 | | | EYES | HEART | INTESTINE | KIDNEYS | |
| TESTIS, RIGHT | 1.49 | 0.349 | | | LIVER | LN, MESENTERIC | LUNGS | MAMMARY GLANI |) |
| TESTIS, LEFT | 1.46 | 0.342 | | | PANCREAS | PITUITARY | PROSTATE | SPINAL CORD | |
| THYMUS | 0.4200 | 0.098 | | | SAL. GLAND MAND | SKIN | SPLEEN | STOMACH | |
| FINAL BODY WT(G) | 427. | | | | SEMINAL VESICLE | S THYMUS | THYROID GLANI | OS TRACHEA | |
| | | | | | URINARY BLADDER | DIAPHRAGM | TESTIS, RIGHT | TESTIS, LEFT | |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

TABLE I8 (MALES) PROJECT NO.:WIL-402010M ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

ANIMAL NO. 73686 GROUP 5: 1000 MG/KG/DAY MALE SCHEDULED EUTH 06/16/10 DATE OF DEATH: 06/16/10 GRADE ORGAN WEIGHT ABS.(G) REL. NO SIGNIFICANT EPIDIDYMIDES INTESTINE LIVER 21.08 6.146 CHANGES OBSERVED GROSS:ADRENAL GLANDS BRAIN ESOPHAGUS 2.93 0.854 EYES INTESTINE KIDNEYS HEART KIDNEYS 1.69 0.493 1.69 0.493 TESTIS, RIGHT LIVER LN, MESENTERIC LUNGS MAMMARY GLAND PANCREAS TESTIS, LEFT 0.5671 0.165 PITUITARY PROSTATE SPINAL CORD SAL. GLAND MAND SKIN SPLEEN STOMACH
SEMINAL VESICLES THYMUS THYROID GLANDS TRACHEA
URINARY BLADDER DIAPHRAGM TESTIS, RIGHT TESTIS, LEFT THYMUS FINAL BODY WT(G) 343.

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

PAGE 24

TABLE 18 (MALES) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MACROSCOPIC FINDINGS

| ANIMAL NO. 7 | 3690 GROUP | 5: 1000 | MG/KG/DAY MA | LE SCHEI | ULED EUTH | 06/16/10 | DATE OF DEATH: | 06/16/10 | GRADE |
|---|----------------------------------|---------------------------------|--------------------------------|--------------------|---|----------------------------------|--|---------------------------------------|-------|
| ORGAN WEIGHT LIVER KIDNEYS TESTIS, RIGHT | ABS.(G) 22.39 3.60 2.00 | REL. 6.644 1.068 0.593 | NO SIGNIFICAN CHANGES OBSER | VED GROSS:AI EY | RENAL GLANDS ES VER | BRAIN HEART LN, MESENTERIC | EPIDIDYMIDES INTESTINE LUNGS | ESOPHAGUS KIDNEYS MAMMARY GLAND |) |
| TESTIS, LEFT THYMUS FINAL BODY WT(| 2.07 0.1993 G) 337. | 0.614 0.059 | | S <i>I</i> SE | NCREAS L. GLAND MAND MINAL VESICLE INARY BLADDER | PITUITARY SKIN S THYMUS | PROSTATE SPLEEN THYROID GLAND TESTIS, RIGHT | | |

GROSS GRADE CODE: 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT

PGRHv4.64 07/16/2010

PAGE 25

Page 254 of 360

PAGE 1

MALE GROUP: 0 MG/KG/DAY

| ANIMAL | FBW(G) | LIVER | KIDNEYS | TESTIS , RIGHT | TESTIS , LEFT | THYMUS |
|---|--------------------------------------|---|--------------------------------------|--------------------------------------|--------------------------------------|--|
| 73665 73666 73676 73691 73692 | 421. 398. 433. 383. 393. | 19.20 18.27 21.50 15.25 17.80 | 3.57 2.98 3.41 2.78 3.22 | 1.56 1.68 1.93 1.64 1.89 | 1.55 1.72 1.94 1.62 1.72 | 0.4311 0.4517 0.7997 0.4888 0.5043 |
| MEAN S.D. S.E. N | 406. 20.7 9.3 | 18.40 2.267 1.014 | 3.19 0.319 0.142 5 | 1.74 0.162 0.072 5 | 1.71 0.147 0.066 5 | 0.5351 0.15073 0.06741 5 |

FBW = FINAL BODY WEIGHT

Page 255 of 360

PAGE 2

MALE GROUP: 30 MG/KG/DAY

| ANIMAL | FBW(G) | LIVER | KIDNEYS | TESTIS , RIGHT | TESTIS , LEFT | THYMUS |
|--------|--------|-------|---------|-------------------|------------------|---------|
| 73668 | 371. | 16.87 | 3.07 | 1.76 | 1.68 | 0.7645 |
| 73669 | 410. | 18.77 | 3.08 | 1.64 | 1.62 | 0.5729 |
| 73682 | 416. | 19.05 | 3.02 | 1.95 | 1.90 | 0.5944 |
| 73687 | 394. | 19.02 | 3.48 | 1.86 | 1.87 | 0.6234 |
| 73688 | 400. | 16.77 | 3.59 | 1.94 | 1.91 | 0.6785 |
| MEAN | 398. | 18.10 | 3.25 | 1.83 | 1.80 | 0.6467 |
| S.D. | 17.4 | 1.170 | 0.266 | 0.131 | 0.136 | 0.07683 |
| S.E. | 7.8 | 0.523 | 0.119 | 0.058 | 0.061 | 0.03436 |
| N | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

Page 256 of 360

PAGE 3

MALE GROUP: 100 MG/KG/DAY

| ANIMAL | FBW(G) | LIVER | KIDNEYS | TESTIS , RIGHT | TESTIS , LEFT | THYMUS |
|--------|--------|-------|---------|-------------------|------------------|---------|
| 73663 | 414. | 17.99 | 3.81 | 2.07 | 2.12 | 0.7315 |
| 73670 | 423. | 18.68 | 3.27 | 1.70 | 1.68 | 0.5558 |
| 73680 | 375. | 17.72 | 3.06 | 1.56 | 1.60 | 0.8100 |
| 73683 | 372. | 16.91 | 3.19 | 1.68 | 1.67 | 0.5300 |
| 73684 | 394. | 18.54 | 2.89 | 1.52 | 1.68 | 0.5175 |
| MEAN | 396. | 17.97 | 3.24 | 1.71 | 1.75 | 0.6290 |
| S.D. | 22.8 | 0.710 | 0.348 | 0.217 | 0.210 | 0.13310 |
| S.E. | 10.2 | 0.317 | 0.155 | 0.097 | 0.094 | 0.05952 |
| N | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

Page 257 of 360

PAGE 4

MALE GROUP: 300 MG/KG/DAY

| ANIMAL | FBW(G) | LIVER | KIDNEYS | TESTIS , RIGHT | TESTIS , LEFT | THYMUS |
|--------|--------|-------|---------|-------------------|------------------|---------|
| 73667 | 350. | 17.00 | 2.91 | 1.54 | 1.52 | 0.4702 |
| 73674 | 385. | 18.98 | 3.34 | 1.73 | 1.60 | 0.4546 |
| 73675 | 393. | 20.97 | 3.22 | 1.62 | 1.59 | 0.8064 |
| 73681 | 447. | 24.27 | 3.89 | 1.71 | 1.75 | 0.7538 |
| 73685 | 423. | 22.67 | 3.54 | 1.95 | 1.98 | 0.8258 |
| MEAN | 400. | 20.78 | 3.38 | 1.71 | 1.69 | 0.6622 |
| S.D. | 37.1 | | 0.365 | 0.154 | 0.183 | 0.18433 |
| S.E. | 16.6 | 1.291 | 0.163 | 0.069 | 0.082 | 0.08244 |
| N | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

Page 258 of 360

PROJECT NO.:WIL-402010M SPONSOR:AMERICAN PETROLEUM

TABLE I9 (MALES) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL ORGAN WEIGHTS AND FINAL BODY WEIGHTS [G]

MALE GROUP: 1000 MG/KG/DAY

| ANIMAL | FBW(G) | LIVER | KIDNEYS | TESTIS , RIGHT | TESTIS , LEFT | THYMUS |
|--------|--------|-------|---------|-------------------|------------------|---------|
| 73671 | 365. | 22.79 | 4.26 | 1.68 | 1.65 | 0.4357 |
| 73677 | 374. | 25.81 | 3.77 | 1.90 | 1.75 | 0.3647 |
| 73678 | 427. | 27.28 | 3.84 | 1.49 | 1.46 | 0.4200 |
| 73686 | 343. | 21.08 | 2.93 | 1.69 | 1.69 | 0.5671 |
| 73690 | 337. | 22.39 | 3.60 | 2.00 | 2.07 | 0.1993 |
| MEAN | 369. | 23.87 | 3.68 | 1.75 | 1.72 | 0.3974 |
| S.D. | 35.7 | 2.575 | 0.485 | 0.201 | 0.222 | 0.13331 |
| S.E. | 16.0 | 1.152 | 0.217 | 0.090 | 0.099 | 0.05962 |
| N | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

POFBWv4.21 07/16/2010

PAGE 5

Page 259 of 360

PAGE 1

MALE GROUP: 0 MG/KG/DAY

| ANIMAL | FBW(G) | LIVER | KIDNEYS | TESTIS , RIGHT | TESTIS , LEFT | THYMUS |
|--------|--------|--------|---------|-------------------|------------------|--------|
| 73665 | 421. | 4.561 | 0.848 | 0.371 | 0.368 | 0.102 |
| 73666 | 398. | 4.590 | 0.749 | 0.422 | 0.432 | 0.113 |
| 73676 | 433. | 4.965 | 0.788 | 0.446 | 0.448 | 0.185 |
| 73691 | 383. | 3.982 | 0.726 | 0.428 | 0.423 | 0.128 |
| 73692 | 393. | 4.529 | 0.819 | 0.481 | 0.438 | 0.128 |
| MEAN | 406. | 4.530 | 0.790 | 0.430 | 0.420 | 0.131 |
| S.D. | 20.7 | 0.3517 | 0.0499 | 0.0401 | 0.0313 | 0.0320 |
| S.E. | 9.3 | 0.1573 | 0.0223 | 0.0179 | 0.0140 | 0.0143 |
| N | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

Page 260 of 360

PAGE 2

MALE GROUP: 30 MG/KG/DAY

| ANIMAL | FBW(G) | LIVER | KIDNEYS | TESTIS , RIGHT | TESTIS , LEFT | THYMUS |
|--------|--------|--------|---------|-------------------|------------------|--------|
| 73668 | 371. | 4.547 | 0.827 | 0.474 | 0.453 | 0.206 |
| 73669 | 410. | 4.578 | 0.751 | 0.400 | 0.395 | 0.140 |
| 73682 | 416. | 4.579 | 0.726 | 0.469 | 0.457 | 0.143 |
| 73687 | 394. | 4.827 | 0.883 | 0.472 | 0.475 | 0.158 |
| 73688 | 400. | 4.193 | 0.897 | 0.485 | 0.477 | 0.170 |
| MEAN | 398. | 4.540 | 0.820 | 0.460 | 0.450 | 0.163 |
| S.D. | 17.4 | 0.2271 | 0.0768 | 0.0341 | 0.0332 | 0.0267 |
| S.E. | 7.8 | 0.1016 | 0.0343 | 0.0153 | 0.0149 | 0.0119 |
| N | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

Page 261 of 360

PAGE 3

MALE GROUP: 100 MG/KG/DAY

| ANIMAL | FBW(G) | LIVER | KIDNEYS | TESTIS , RIGHT | TESTIS , LEFT | THYMUS |
|--------|--------|--------|---------|-------------------|------------------|--------|
| 73663 | 414. | 4.345 | 0.920 | 0.500 | 0.512 | 0.177 |
| 73670 | 423. | 4.416 | 0.773 | 0.402 | 0.397 | 0.131 |
| 73680 | 375. | 4.725 | 0.816 | 0.416 | 0.427 | 0.216 |
| 73683 | 372. | 4.546 | 0.858 | 0.452 | 0.449 | 0.142 |
| 73684 | 394. | 4.706 | 0.734 | 0.386 | 0.426 | 0.131 |
| MEAN | 396. | 4.550 | 0.820 | 0.430 | 0.440 | 0.159 |
| S.D. | 22.8 | 0.1694 | 0.0727 | 0.0456 | 0.0431 | 0.0368 |
| S.E. | 10.2 | 0.0757 | 0.0325 | 0.0204 | 0.0193 | 0.0165 |
| N | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

Page 262 of 360

PAGE 4

MALE GROUP: 300 MG/KG/DAY

| ANIMAL | FBW(G) | LIVER | KIDNEYS | TESTIS , RIGHT | TESTIS , LEFT | THYMUS |
|--------|--------|--------|---------|-------------------|------------------|--------|
| 73667 | 350. | 4.857 | 0.831 | 0.440 | 0.434 | 0.134 |
| 73674 | 385. | 4.930 | 0.868 | 0.449 | 0.416 | 0.118 |
| 73675 | 393. | 5.336 | 0.819 | 0.412 | 0.405 | 0.205 |
| 73681 | 447. | 5.430 | 0.870 | 0.383 | 0.391 | 0.169 |
| 73685 | 423. | 5.359 | 0.837 | 0.461 | 0.468 | 0.195 |
| MEAN | 400. | 5.180 | 0.850 | 0.430 | 0.420 | 0.164 |
| S.D. | 37.1 | 0.2672 | 0.0227 | 0.0316 | 0.0298 | 0.0377 |
| S.E. | 16.6 | 0.1195 | 0.0101 | 0.0141 | 0.0133 | 0.0169 |
| N | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

Page 263 of 360

MALE GROUP: 1000 MG/KG/DAY

| ANIMAL | FBW(G) | LIVER | KIDNEYS | TESTIS , RIGHT | TESTIS , LEFT | THYMUS |
|--------|--------|--------|---------|-------------------|------------------|--------|
| 73671 | 365. | 6.244 | 1.167 | 0.460 | 0.452 | 0.119 |
| 73677 | 374. | 6.901 | 1.008 | 0.508 | 0.468 | 0.098 |
| 73678 | 427. | 6.389 | 0.899 | 0.349 | 0.342 | 0.098 |
| 73686 | 343. | 6.146 | 0.854 | 0.493 | 0.493 | 0.165 |
| 73690 | 337. | 6.644 | 1.068 | 0.593 | 0.614 | 0.059 |
| MEAN | 369. | 6.460 | 1.000 | 0.480 | 0.470 | 0.108 |
| S.D. | 35.7 | 0.3077 | 0.1265 | 0.0886 | 0.0974 | 0.0386 |
| S.E. | 16.0 | 0.1376 | 0.0566 | 0.0396 | 0.0436 | 0.0173 |
| N | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

POFBWv4.21 07/16/2010

PAGE 5

Page 264 of 360

INDIVIDUAL CLINICAL OBSERVATIONS

PAGE 1

| OBSERVATION | ANIMAL GROUP | GESTATIONAL DAY |
|---|--------------|-------------------------|
| NO GEOGRAPHICAL AND | 50510 | |
| NO SIGNIFICANT CLINICAL OBSERVATIONS | 73710 1 | P |
| | 73704 1 | |
| | 73694 1 | |
| | 73713 1 | P |
| | 73720 1 | P |
| | 73702 2 | PPP |
| | 73697 2 | PPPPPP |
| | 73706 2 | P |
| | 73709 2 | P |
| | 73719 2 | PPPPPPPP |
| | 73696 3 | P |
| | 73714 3 | |
| | 73705 3 | P P P P P P P P P P P P |
| | 73708 4 | PPPP P PPPPPPP |
| | 73712 4 | P |
| | 73698 4 | P |
| | 73717 4 | P |
| | 73693 4 | PPP |
| | 73718 5 | PPPPPP |
| | 73701 5 | PP |
| | 73721 5 | PPPP |
| | 73721 5 | |
| | | PPPPPPPP P PPPPPPP |
| | 13122 3 | |
| SCHEDULED EUTHANASIA; GESTATION DAY 20 | 73710 1 | Р |
| COMPOSED POLITICATION DAT 20 | 73710 1 | P |
| | 73694 1 | p D |
| | /3034 I | r |

GRADE CODE: P = PRESENT 1 = SLIGHT 2 = MODERATE 3 = SEVERE

_______ 1- 0 MG/KG/DAY 2- 30 MG/KG/DAY 3- 100 MG/KG/DAY 4- 300 MG/KG/DAY 5- 1000 MG/KG/DAY

Page 265 of 360

Page 266 of 360

TABLE I11 (FEMALES - DAILY EXAMINATIONS) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

PAGE 2

GESTATIONAL DAY 1 1 1 1 1 1 1 1 1 2 OBSERVATION ANIMAL GROUP 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 SCHEDULED EUTHANASIA; GESTATION DAY 20 73713 1 Ρ 73720 1 73702 2 73697 73706 73709 73719 73696 3 73699 3 73714 3 73705 3 73700 3 73708 73712 4 73698 4 73717 73693 73718 73701 73721 5 73707 5 Ρ Ρ 73722 5 PΡ HUNCHED POSTURE 73701 5 HAIR LOSS RIGHT FORELIMB 73694 1 11 1111 73702 2 11 1111111111111

GRADE CODE: P = PRESENT 1 = SLIGHT 2 = MODERATE 3 = SEVERE _______

Page 267 of 360

TABLE I11 (FEMALES - DAILY EXAMINATIONS) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

GESTATIONAL DAY 1 1 1 1 1 1 1 1 2 OBSERVATION ANIMAL GROUP 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 HAIR LOSS RIGHT FORELIMB 73697 1 1 1 1 1 1 1 1 1 1 1 1 1 2 1 1 1 1 1 1 73709 2 73696 3 73699 73705 1 1 1 1 1 1 3 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 73700 73708 1 1 1 1 1 HAIR LOSS LEFT FORELIMB 1 1 1 1 1 1 73694 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 73702 73697 1 1 1 1 1 1 1 1 1 1 1 1 1 73709 1 1 1 1 1 1 73696 1 1 73705 3 73708 HAIR LOSS RIGHT HINDLIMB 73697 73700 3 1 1 2 1 1 1 1 1 73693 1 1 1 1 1 1 1 1 2 2 2 2 2 2 2 1 1 1 1 1 1 1 2 1 1 1 73718 5 73701 5 1 1 1 1 2 2 2 2 2 1 1 1 1 1 1 1 1 1 HAIR LOSS VENTRAL ABDOMINAL AREA 73697 2 1 73714 3 1 1 3 73700 1 1 1 1 1 2 2 2 2 2 2 3 3 2 2 2 2 GRADE CODE: P = PRESENT 1 = SLIGHT 2 = MODERATE 3 = SEVERE

Page 268 of 360

TABLE I11 (FEMALES - DAILY EXAMINATIONS) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

| OBSERVATION | ANIMAL GR | OUP | GES' 0 1 | | | | | | | | | | | | | | | | | |
|--|-----------|-----|-------------|---|-----|---|---|---|---|-----|-----|-----|---|---|---|---|---|---|---|--|
| HAIR LOSS VENTRAL ABDOMINAL AREA | 73693 | 4 | | | | | | | | | | | | 1 | 1 | 1 | 1 | 1 | 1 | |
| MAIN DOSS VENTRAL ADDONINAL AREA | 73718 | 5 | | | | | | 1 | 1 | 1 1 | 1 2 | 2 | 1 | | | | | | | |
| | 73701 | 5 | | 1 | | | | | 1 | | | | | | | | | | | |
| | 73707 | 5 | | _ | | | | | - | | | _ | _ | - | 1 | | | | | |
| WET YELLOW MATERIAL ANOGENITAL AREA | 73709 | 2 | | | | | | | | | | | | | | | | | 2 | |
| | 73700 | 3 | | 1 | | | | | | | | | | | | | | | | |
| | 73701 | 5 | | | 1 | 2 | 2 | 2 | | | | | | | | | | | | |
| | 73721 | 5 | | | | | | 3 | 1 | | | | | | | | | | | |
| HAIR LOSS VENTRAL NECK | 73708 | 4 | | | | 1 | 1 | 1 | | | | | | | | | | | | |
| | 73707 | 5 | | | | | | | | 1 | L | | | | | | | | | |
| WET YELLOW MATERIAL VENTRAL ABDOMINAL AREA | 73693 | 4 | | 1 | . 1 | | | | | | | | | | | | | | | |
| | 73701 | 5 | | | 1 | | | 1 | | | | | | | | | | | | |
| | 73707 | 5 | | 1 | . 1 | 1 | 1 | | | | | | | | | | | | | |
| NET YELLOW MATERIAL UROGENITAL AREA | 73709 | 2 | | | | | | | | | | | | | | | | | 1 | |
| | 73705 | 3 | | | | | | | | | | | | | | | | 1 | | |
| | 73693 | 4 | | | . 1 | | | | | | | | | | | | | | | |
| | 73701 | 5 | | | 2 | | | 2 | | | | | | | | | | | | |
| | 73721 | 5 | | | | | | | | | | | | | | | | | | |
| | 73707 | 5 | | 1 | . 1 | 1 | 1 | | | | | | | | | | | | | |
| | 73722 | 5 | | | | | | | | | 1 | . 1 | | | | | | | | |
| WET YELLOW MATERIAL RIGHT INGUINAL AREA | 73700 | 3 | | | | | | | | | | | | | 1 | | | | | |
| | 73701 | 5 | | 2 | 1 | 1 | 1 | 2 | | | | | | | | | | | | |

GRADE CODE: P = PRESENT 1 = SLIGHT 2 = MODERATE 3 = SEVERE

Page 269 of 360

TABLE I11 (FEMALES - DAILY EXAMINATIONS) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

GESTATIONAL DAY 1 1 1 1 1 1 1 1 1 2 OBSERVATION ANIMAL GROUP 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 3 73700 WET YELLOW MATERIAL LEFT INGUINAL AREA 1 5 73718 73701 5 2 1 1 1 HAIR LOSS LEFT HINDLIMB 73719 1 1 1 1 1 1 73700 73708 4 1 73693 4 1 1 2 2 1 1 1 1 1 2 1 1 1 73718 5 73701 1 1112111 5 WET YELLOW MATERIAL RIGHT HINDLIMB 73701 5 1 1 2 1 1 WET YELLOW MATERIAL LEFT HINDLIMB 73701 5 1 1 1 HAIR LOSS RUMP 73719 1 11111 1111111 1111 73700 3 73693 111 11111 73718 1 11111 73701 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 73700 2 1 2 2 1 1 1 1 1 1 1 1 1 HAIR LOSS LEFT LATERAL ABDOMINAL AREA 3 73718 5 1 1 1 DRIED RED MATERIAL VENTRAL ABDOMINAL AREA 73707 5 2 1 1 1 1 1 1 1 DRIED RED MATERIAL UROGENITAL AREA 73721 5 GRADE CODE: P = PRESENT 1 = SLIGHT 2 = MODERATE 3 = SEVERE

^{1- 0} MG/KG/DAY 2- 30 MG/KG/DAY 3- 100 MG/KG/DAY 4- 300 MG/KG/DAY 5- 1000 MG/KG/DAY

TABLE I11 (FEMALES - DAILY EXAMINATIONS) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 6 INDIVIDUAL CLINICAL OBSERVATIONS SPONSOR: AMERICAN PETROLEUM GESTATIONAL DAY 1 1 1 1 1 1 1 1 2 OBSERVATION ANIMAL GROUP 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 DRIED RED MATERIAL UROGENITAL AREA 73707 5 2 1 1 1 1 1 DRIED RED MATERIAL RIGHT INGUINAL AREA 73707 5 1 DRIED RED MATERIAL ANOGENITAL AREA 73721 5 12111 11 HAIR LOSS VENTRAL THORACIC AREA 73700 3 1 1 1 73693 4 73718 1 5 1 73707 5 SCABBING VENTRAL ABDOMINAL AREA 73700 3 PPPP SCABBING LEFT HINDLIMB 73700 3 PΡ 1 HAIR LOSS UROGENITAL AREA 73700 3 1 2 2 2 73701 1 73721 5 1 1 1 1 HAIR LOSS BASE OF TAIL 73700 3 2 1 1 1 1 73693 1 1 1 1 1 73701 5 1 1 1 1 73700 HAIR LOSS ANOGENITAL AREA 3 1 1 2 2 2 73701 5 1 1 DRIED RED MATERIAL AROUND RIGHT EYE 73701 5 1 1

GRADE CODE: P = PRESENT 1 = SLIGHT 2 = MODERATE 3 = SEVERE ______

PROJECT NO.:WIL-402010F

SPONSOR: AMERICAN PETROLEUM

TABLE I11 (FEMALES - DAILY EXAMINATIONS) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

GESTATIONAL DAY 1 1 1 1 1 1 1 1 2 OBSERVATION ANIMAL GROUP 0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 DRIED RED MATERIAL AROUND LEFT EYE 73701 5 1 1 1 DRIED RED MATERIAL AROUND NOSE 73693 4 1 1 1 73701 5 73707 5 73722 1 1 1 5 DECREASED DEFECATION PPPP 73701 PPP 73707 5 HAIR LOSS LEFT INGUINAL AREA 73700 3 2 2 1 2 2 2 2 73701 1 73721 5 1 1 HAIR LOSS RIGHT INGUINAL AREA 73700 3 2 2 2 73693 4 1 1 1 1 1 73701 73721 5 1 1 DRIED RED MATERIAL AROUND MOUTH 73693 4 1 73701 5 1 2 1 1 73707 5 SALIVATION 73707 5 1 GRADE CODE: P = PRESENT 1 = SLIGHT 2 = MODERATE 3 = SEVERE

1- 0 MG/KG/DAY 2- 30 MG/KG/DAY 3- 100 MG/KG/DAY 4- 300 MG/KG/DAY 5- 1000 MG/KG/DAY

PCOv3.13 07/28/2010

PAGE 7

| PROJECT NO :: WIL- | 402010F |
|--------------------|-----------|
| SPONSOR: AMERICAN | PETROLEUM |
| | |

TABLE I12 (FEMALES - AT TIME OF DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

| | | | | TABLE RANGE: 06-02-10 TO 06-24-10 |
|--------|-----|----------------|-------------|---|
| ANIMAL | SEX | GROUP | CATEGORY | DATE TIME GRADE OBSERVATIONS |
| | | | | AT TIME OF DOSING |
| 73721 | F | 1000 MG/KG/DAY | ORAL/DENTAL | 06-13-10 9:46 1 SALIVATION PRIOR TO DOSING |
| 73707 | F | 1000 MG/KG/DAY | ORAL/DENTAL | 06-13-10 9:46 2 SALIVATION PRIOR TO DOSING |
| | | | | 06-14-10 10:19 1 SALIVATION PRIOR TO DOSING |
| | | | | 06-17-10 10:02 1 SALIVATION PRIOR TO DOSING |
| | | | | 06-18-10 9:57 2 SALIVATION PRIOR TO DOSING |
| | | | | 06-21-10 9.54 2 SALTVATION PRIOR TO DOSING |

GRADE CODE: 1 - SLIGHT 2 - MODERATE 3 - SEVERE P - PRESENT

PCRDv4.17 07/15/2010

PAGE 1

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010F

PAGE 1 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| _ | | | | | TABLE I | RANGE: (| 06-02-10 TO 06-24-10 | | |
|---|--------|-----|-------|----------|---------|----------|----------------------|------|--|
| _ | ANIMAL | SEX | GROUP | CATEGORY | DATE | TIME | GRADE OBSERVATIONS | | |
| _ | | _ | | | | | NG OBSERVATIONS | | |

| | ANIMAL | SEX | GROUP | CATEGORY | DATE | TIME | GRADI | E OB | SERVATIONS | | | | |
|---|--------|-----|-------------|----------|--------|----------|-------|-----------|-------------|----------|--------------|------|--|
| - | | | | | POS' | Γ-DOSINO | OBSE | ERVA' | TIONS | | | | |
| | 73710 | F | 0 MG/KG/DAY | NORMAL | | | | | | CLINICAL | OBSERVATIONS | | |
| | | | , -, | | 06-03- | 10 10:29 |) P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-04- | 10 10:49 |) P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-05- | 10 10:37 | 7 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-06- | 10 10:59 |) P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-07- | 10 10:40 |) P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-08- | 10 10:43 | 3 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-09- | 10 10:47 | 7 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-10- | 10 10:38 | 3 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-11- | 10 11:00 |) P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-12- | 10 10:21 | L P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | | 10 10:28 | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:50 | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:47 | | | | | OBSERVATIONS | | |
| | | | | | 06-16- | 10 10:45 | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:48 | | | | | OBSERVATIONS | | |
| | | | | | 06-18- | 10 10:38 | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:19 | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:36 | | | | | OBSERVATIONS | | |
| | | | | | 06-21- | | | | | | OBSERVATIONS | | |
| | 73704 | F | 0 MG/KG/DAY | NORMAL | 06-03- | | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:50 | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:37 | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:59 | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:40 | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:43 | | | | | OBSERVATIONS | | |
| | | | | | | 10 10:47 | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | 06-11- | 10 11:00 |) P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| _ | | | | | | | | | | | | | |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING)

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 2 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10 ANIMAL SEX GROUP CATEGORY DATE TIME GRADE OBSERVATIONS POST-DOSING OBSERVATIONS 73704 F 0 MG/KG/DAY NORMAL 06-12-10 10:21 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:28 P NO SIGNIFICANT CLINICAL OBSERVATIONS

06-14-10 10:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 10:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-16-10 10:45 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-18-10 10:38 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-19-10 10:19 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-20-10 10:36 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-21-10 10:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-22-10 9:46 P NO SIGNIFICANT CLINICAL OBSERVATIONS 73694 F 0 MG/KG/DAY NORMAL 06-04-10 10:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 10:38 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 10:59 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 10:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 10:43 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 10:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:39 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 11:00 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 10:22 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:28 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 10:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 10:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-16-10 10:46 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:48 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-18-10 10:38 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-19-10 10:20 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-20-10 10:36 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-21-10 10:32 P NO SIGNIFICANT CLINICAL OBSERVATIONS

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 3 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10

| ANIMAL | SEX | | GROUP | CATEGORY | DATE | TI | ME (| GRADI | E OI | SSERVATIONS | | |
|------------|-----|---|--------------|----------|--------|---------|-------|-------|------|-------------|----------|--------------|
| | | | | | POS | T-D | OSTNG | OBSE | : | TIONS | | |
| 73694 | F | 0 | MG/KG/DAY | NORMAL | | | | | | | CLINICAL | OBSERVATIONS |
| , 5 6 5 1 | - | ŭ | 110/110/2111 | 11014111 | 06-23- | 10 | 9:44 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| 73713 | F | 0 | MG/KG/DAY | NORMAL | 06-04- | 10 | 10:50 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | -, -, | | | | | | | | | OBSERVATIONS |
| | | | | | 06-06- | 10 | 10:59 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-07- | 10 | 10:40 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-08- | 10 | 10:43 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-09- | 10 | 10:48 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-10- | 10 | 10:39 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-11- | 10 | 11:01 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-12- | 10 | 10:22 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-13- | 10 | 10:29 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | 06-14- | | | | | | | OBSERVATIONS |
| | | | | | 06-15- | 10 | 10:48 | | | | | OBSERVATIONS |
| | | | | | 06-16- | | | | | | | OBSERVATIONS |
| | | | | | 06-17- | | | | | | | OBSERVATIONS |
| | | | | | | | | | | | | OBSERVATIONS |
| | | | | | | | | | | | | OBSERVATIONS |
| | | | | | | | | | | | | OBSERVATIONS |
| | | | | | 06-21- | | | | | | | OBSERVATIONS |
| | | | | | | | | | | | | OBSERVATIONS |
| | | | | | | | | | | | | OBSERVATIONS |
| 73720 | F | 0 | MG/KG/DAY | NORMAL | | | | | | | | OBSERVATIONS |
| | | | | | 06-06- | | | | | | | OBSERVATIONS |
| | | | | | 06-07- | | | | | | | OBSERVATIONS |
| | | | | | 06-08- | | | | | | | OBSERVATIONS |
| | | | | | | | | | | | | OBSERVATIONS |
| | | | | | | | | | | | | OBSERVATIONS |
| | | | | | 06-11- | Τ0 | 11:01 | Р | ИО | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | | | | | | | | |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010F

PAGE 4 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| TABLE RANGE: 06-0 | 2 - 10 TC |) 06-24-10 |
|-------------------|-----------|------------|

| ANIMAL | SEX | GROUP | CATEGORY | DATE | TIME | GRAD | DE C | BSERVATIONS | | | | |
|--------|-----|-----------------|----------|--------|----------|------|------|-------------|------------|---------------------------|------|--|
| | | | | | | | | | | | | |
| F2F20 | - | 0 MG /17G /D337 | MODMAT | | T-DOSING | | | | CT TNITCHT | OD GEDITA EL ONG | | |
| 73720 | r | 0 MG/KG/DAY | NORMAL | | | | | | | OBSERVATIONS OBSERVATIONS | | |
| | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | OBSERVATIONS | | |
| | | | | 06-10- | | | | | | OBSERVATIONS | | |
| | | | | 06-17- | | | | | | OBSERVATIONS | | |
| | | | | 06-19- | | | | | | OBSERVATIONS | | |
| | | | | 06-20- | | | | | | OBSERVATIONS | | |
| | | | | 06-21- | | | | | | OBSERVATIONS | | |
| | | | | 06-22- | | | | | | OBSERVATIONS | | |
| | | | | 06-23- | | | | | | OBSERVATIONS | | |
| | | | | 06-24- | | | | | | OBSERVATIONS | | |
| 3702 | F | 30 MG/KG/DAY | | 06-03- | | | | | | OBSERVATIONS | | |
| | | | | 06-04- | | | | | | OBSERVATIONS | | |
| | | | | 06-05- | | | | | | OBSERVATIONS | | |
| | | | | 06-06- | | | | | | OBSERVATIONS | | |
| | | | | 06-07- | 10 10:4 | 1 P | NC | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | 06-08- | 10 10:4 | 4 P | NC | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | 06-09- | 10 10:4 | 8 P | NC | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | 06-10- | 10 10:4 |) P | NC | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | 06-11- | 10 11:0 | 1 P | NC | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | 06-12- | 10 10:3 | 2 P | NC | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | 06-13- | 10 10:2 | 9 P | NC | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | 06-14- | 10 10:5 | | | | | OBSERVATIONS | | |
| | | | | 06-15- | 10 10:4 | 9 P | | | | OBSERVATIONS | | |
| | | | | 06-16- | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | OBSERVATIONS | | |
| | | | | 06-18- | 10 10:4 |) P | NC | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 5 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10

| | | | IADLE I | | | - 0 2 - 1 | | | | |
|------------|-----------------|-------------|---------|-----|-------|-----------|--------|-------------|------------|--------------|
| ANIMAL SEX | GROUP | CATEGORY | DATE | | | | E OF | BSERVATIONS | | |
| | | | POS | | | | יוזסק. | \TTONG | | |
| 73702 F | 30 MG/KG/DAY | NORMAT. | 06-19- | -10 | 10.21 | P | MO | SIGNIFICANT | CT.TNTCAT. | OBSERVATIONS |
| 73702 1 | 30 110/110/2111 | 14014 11111 | | | | | | | | OBSERVATIONS |
| | | | | | | | | | | OBSERVATIONS |
| | | | | | | | | | | OBSERVATIONS |
| 73697 F | 30 MG/KG/DAY | | | | | | | | | OBSERVATIONS |
| | ,, | | | | | | | | | OBSERVATIONS |
| | | | | | | | | | | OBSERVATIONS |
| | | | 06-06- | -10 | 11:00 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-07- | -10 | 10:41 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-08- | -10 | 10:44 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-09- | -10 | 10:48 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-10- | -10 | 10:40 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-11- | -10 | 11:01 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-12- | -10 | 10:32 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-13- | -10 | 10:29 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-14- | -10 | 10:51 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-15- | -10 | 10:49 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-16- | -10 | 10:46 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-17- | -10 | 10:49 | | | | | OBSERVATIONS |
| | | | 06-18- | -10 | 10:40 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | 06-19- | | | | | | | OBSERVATIONS |
| | | | 06-20- | | | | | | | OBSERVATIONS |
| | | | 06-21- | | | | | | | OBSERVATIONS |
| | | | 06-22- | | | | | | | OBSERVATIONS |
| 73706 F | 30 MG/KG/DAY | NORMAL | 06-03- | | | | | | | OBSERVATIONS |
| | | | 06-04- | | | | | | | OBSERVATIONS |
| | | | | | | | | | | OBSERVATIONS |
| | | | | | | | | | | OBSERVATIONS |
| | | | 06-07- | -10 | 10:41 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS |
| | | | | | | | | | | |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 6
SPONSOR:AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10 ANIMAL SEX GROUP CATEGORY DATE TIME GRADE OBSERVATIONS POST-DOSING OBSERVATIONS 73706 F 30 MG/KG/DAY NORMAL 06-08-10 10:44 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 11:02 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 10:33 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:30 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-14-10 10:51 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-16-10 10:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS

06-18-10 10:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-19-10 10:21 P NO SIGNIFICANT CLINICAL OBSERVATIONS

06-15-10 10:49 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-16-10 10:47 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-18-10 10:41 P NO SIGNIFICANT CLINICAL OBSERVATIONS

| 06-20-10 | 10:37 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-21-10 | 10:33 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-22-10 | 9:48 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-04-10 | 10:51 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-05-10 | 10:39 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-05-10 | 10:39 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-05-10 | 10:40 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-07-10 | 10:42 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-09-10 | 10:49 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-10-10 | 10:40 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-11-10 | 11:02 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-11-10 | 11:02 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-12-10 | 10:33 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-13-10 | 10:30 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS | 06-14-10 | 10:52 | P | NO SIGNIFICANT CLINICAL OBSERVATIONS |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PROJECT NO.:WIL-402010F PAGE 7 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10

| | | | | | | | | | | | | |
|-----------|-----|------------------------|-----------------------|---|--|-------------|---|--|--|---|------|--|
| ANIMAL SE | X | GROUP | CATEGORY | DATE ' | TIME | GRADE | E OBS | SERVATIONS | | | | |
| 73709 F | 30 | MG/KG/DAY | NORMAL | 06-20-1 06-21-1 06-22-1 | 0 10:22 0 10:37 0 10:33 0 9:48 | P P P | NO S NO S NO S | SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT | CLINICAL CLINICAL CLINICAL | OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS | | |
| | | MG/KG/DAY MG/KG/DAY | ORAL/DENTAL NORMAL | $\begin{array}{c} 06-23-1 \\ 06-08-1 \\ 06-05-1 \\ 06-06-1 \\ 06-07-1 \\ 06-08-1 \\ 06-09-1 \\ 06-10-1 \\ 06-12-1 \\ 06-12-1 \\ 06-13-1 \\ 06-14-1 \\ 06-15-1 \\ 06-15-1 \\ 06-15-1 \\ 06-19-1 \\ 06-22-1 \\ 06-22-1 \end{array}$ | 9:45 0 9:45 10:45 10:39 10:42 10:42 10:42 10:42 10:42 10:42 10:42 10:42 10:43 10 | | NO S | SIGNIFICANT ED RED MATER SIGNIFICANT | CLINICAL RIAL AROU CLINICAL | OBSERVATIONS ND MOUTH OBSERVATIONS | | |
| 73696 F | 100 | MG/KG/DAY | NORMAL | 06-03-1 | 0 9:54 0 10:41 0 10:30 | P P P | NO S NO S | SIGNIFICANT SIGNIFICANT SIGNIFICANT | CLINICAL CLINICAL CLINICAL | OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS OBSERVATIONS | | |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010F

PAGE 8 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| TARLE | PANCE. | 06-02-10 | TΟ | 06-24-10 |
|-------|--------|----------|----|----------|
| IADLE | KANGE: | 00-02-10 | 10 | 00-24-10 |

| ANIMAL | SEX | GROUP | CATEGORY | DATE | TIME | GRAD | E OB | SERVATIONS | | | |
|--------|-----|---------------|-------------|--------|---------|-------|------|-------------|-----------|--------------|------|
| | | | | POS | T-DOSIN | G OBS | ERVA | TIONS | | | |
| 73696 | F | 100 MG/KG/DAY | NORMAL | 06-05- | 10 10:3 | 9 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | , , | | | | | | | | OBSERVATIONS | |
| | | | | 06-07- | 10 10:5 | 5 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-08- | 10 10:4 | 5 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-09- | 10 10:5 | 0 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-10- | 10 10:4 | 1 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-11- | 10 11:0 | 2 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-12- | 10 10:3 | 4 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-13- | 10 10:3 | 1 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-14- | 10 10:5 | 2 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-15- | 10 10:5 | 0 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-16- | 10 10:4 | 7 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | | 10 10:5 | | | | | OBSERVATIONS | |
| | | | | 06-18- | 10 10:4 | 1 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-19- | 10 10:2 | 2 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-20- | 10 10:3 | 7 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| 73696 | F | 100 MG/KG/DAY | ORAL/DENTAL | 06-21- | 10 10:3 | 4 1 | WET | CLEAR MATER | RIAL AROU | ND MOUTH | |
| 73699 | F | 100 MG/KG/DAY | NORMAL | 06-03- | 10 10:3 | 0 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-04- | 10 10:5 | 1 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-05- | 10 10:4 | 0 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-06- | 10 11:0 | 2 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-07- | 10 10:5 | 6 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-08- | 10 10:4 | 6 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-09- | 10 10:5 | 0 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-10- | 10 10:4 | 1 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-11- | 10 11:0 | 2 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-12- | 10 10:3 | 4 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-13- | 10 10:3 | 1 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-14- | 10 10:5 | 2 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 9 PROJECT NO.:WIL-402010F SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| | | | | TABLE R | ANG | E: 06 | -02- | 10 7 | TO 06-24-10 | | | |
|--------|-----|---------------|-----------------|---------|-----|-------|------|------|---------------|-----------|--------------|--|
| ANIMAL | SEX | GROUP | CATEGORY | DATE | TI | ME (| GRAD | E OI | BSERVATIONS | | | |
| | | | | POS | | | | | | | | |
| 73699 | F | 100 MG/KG/DAY | NORMAL | 06-15- | 10 | 10:50 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-16- | 10 | 10:47 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | 06-18- | 10 | 10:42 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-19- | | | | | | | OBSERVATIONS | |
| | | | | 06-20- | 10 | 10:38 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | 06-21- | 10 | 10:34 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| 73714 | F | 100 MG/KG/DAY | NORMAL | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | 06-06- | | | | | | | OBSERVATIONS | |
| | | | | 06-07- | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | 06-12- | 10 | 10:34 | P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| | | | | | | | | | | | OBSERVATIONS | |
| 73714 | F | 100 MG/KG/DAY | BODY/INTEGUMENT | | | | | | | | | |
| | | | | | | | | | Г CLEAR MATEF | | | |
| 73714 | F | 100 MG/KG/DAY | EYES/EARS/NOSE | 06-22- | 10 | 9:49 | 1 | WE: | r clear mater | RIAL AROU | ND NOSE | |

PAGE 10

TABLE RANGE: 06-02-10 TO 06-24-10 ANIMAL SEX GROUP CATEGORY DATE TIME GRADE OBSERVATIONS POST-DOSING OBSERVATIONS 73714 F 100 MG/KG/DAY ORAL/DENTAL 06-14-10 10:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:49 2 WET CLEAR MATERIAL AROUND MOUTH 73705 F 100 MG/KG/DAY NORMAL 06-04-10 10:52 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-05-10 10:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:02 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 10:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 10:46 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 10:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:42 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 11:03 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 10:34 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-13-10 10:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-15-10 10:51 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:51 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-18-10 10:43 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-19-10 10:23 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-20-10 10:38 P NO SIGNIFICANT CLINICAL OBSERVATIONS 73705 F 100 MG/KG/DAY ORAL/DENTAL 06-14-10 10:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-16-10 10:48 1 WET CLEAR MATERIAL AROUND MOUTH 06-21-10 10:35 1 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:45 1 WET CLEAR MATERIAL AROUND MOUTH 73700 F 100 MG/KG/DAY NORMAL 06-05-10 10:40 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:02 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-10 10:56 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 10:51 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:42 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-11-10 11:03 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-12-10 10:35 P NO SIGNIFICANT CLINICAL OBSERVATIONS

GRADE CODE: 1 - SLIGHT 2 - MODERATE 3 - SEVERE P - PRESENT

PROJECT NO.:WIL-402010F

SPONSOR: AMERICAN PETROLEUM

Page 282 of 360

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 11 PROJECT NO.:WIL-402010F SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

| та | AT.F | RANGE: | 06- | 02-10 | TΟ | 06-2 | 4-10 | 1 |
|-------|------|--------|-----|-------|----|------|-------|---|
| IA IA | лошь | KANGE: | 00- | 02-10 | 10 | 00-2 | ・モーエし | J |

| | | | | IADLE I | KANGE: U | 5-02- | 10 10 06-24-10 |
|----------|----|---------------|------------------------------|--|---|---|---|
| ANIMAL S | EX | GROUP | CATEGORY | DATE | TIME | GRAD | E OBSERVATIONS |
| 73700 | F | 100 MG/KG/DAY | NORMAL | POS 06-13- 06-14- 06-16- 06-18- 06-19- 06-20- 06-21- 06-22- | ST-DOSING -10 10:32 -10 10:54 -10 10:48 -10 10:24 -10 10:38 -10 10:38 | G OBS P P P P P P P P P P P P P P P P P P P | ERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73700 | F | | BODY/INTEG II ORAL/DENTAL | 06-23- 06-15- 06-08- 06-15- 06-17- | -10 9:46 -10 10:51 -10 10:47 -10 10:51 | 5 1 L 1 7 1 L 1 L 2 | WET CLEAR MATERIAL AROUND NOSE WET CLEAR MATERIAL VENTRAL NECK WET CLEAR MATERIAL AROUND MOUTH WET CLEAR MATERIAL AROUND MOUTH WET CLEAR MATERIAL AROUND MOUTH |
| 73708 | F | 300 MG/KG/DAY | NORMAL | 06-03- 06-04- 06-05- 06-06- 06-07- 06-10- 06-11- 06-13- 06-14- 06-17- 06-18- | -10 10:43 | 5 P 2 P 2 P 3 P 7 P 6 P 6 P 6 P 6 P 7 P 6 P 7 P 6 P 7 P 7 P 8 P 8 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 12 SPONSOR:AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10

| | | | | | | TADDD 10 | mod. oc | 02. | 10 10 00 24 10 |
|---|--------|-----|-----|--------------|-----------------|----------|----------|--------|--------------------------------------|
| - | ANIMAL | SEX | | GROUP | CATEGORY | DATE | TIME | GRAD | E OBSERVATIONS |
| - | | | | | | DOGT | י-חספדאנ | I ORGI | ERVATIONS |
| | 73708 | F | 300 | MG/KG/DAV | ΝΟΡΜΔΙ. | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | 73708 | | | | | | | | WET CLEAR MATERIAL AROUND NOSE |
| | 73700 | - | 500 | 110/110/2111 | ETES, EMO, NOOE | | | | WET CLEAR MATERIAL AROUND NOSE |
| | 73708 | F | 300 | MG/KG/DAY | BODY/INTEG II | | | | WET CLEAR MATERIAL VENTRAL NECK |
| | 73708 | | | | | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | ,5,00 | - | 500 | 110/110/2111 | 01012/ 2211112 | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | | 06-21-1 | 0 10:36 | 5 2 | WET CLEAR MATERIAL AROUND MOUTH |
| | 73712 | F | 300 | MG/KG/DAY | NORMAL | 06-03-1 | 0 10:47 | 7 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | 06-04-1 | .0 10:52 | 2 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | 06-05-1 | 0 10:53 | 3 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | 06-06-1 | .0 11:18 | 3 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | 06-07-1 | 0 10:57 | 7 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | 06-08-1 | 0 10:53 | 3 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | 06-13-1 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | 06-14-1 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | 06-15-1 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | 06-16-1 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | _ | | / / | | 06-20-1 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | | WET CLEAR MATERIAL AROUND NOSE |
| | 73712 | F. | 300 | MG/KG/DAY | ORAL/DENTAL | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | | 06-71-1 | .u 10:3 | ′ ⊥ | DRIED CLEAR MATERIAL AROUND MOUTH |
| _ | | | | | | | | | |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 13 PROJECT NO.:WIL-402010F INDIVIDUAL CLINICAL OBSERVATIONS SPONSOR: AMERICAN PETROLEUM

TABLE RANGE: 06-02-10 TO 06-24-10

| | | | | IADLE K | ANGE: | J6-U2- | | 10 06-24-10 | | | |
|----------------|-----|--------------------------------|----------|--|---|---|--|---|--|--|------|
| ANIMAL | SEX | GROUP | CATEGORY | DATE | TIME | GRAI | DE O | BSERVATIONS | | | |
| 73712 73698 | | 300 MG/KG/DAY 300 MG/KG/DAY | | 06-22- | | 50 1 | DR | IED RED MATER | | ND MOUTH OBSERVATIONS | |
| 73030 | r | JUU NG/NG/DAT | NOME | 06-04- 06-05- 06-06- | 10 10:9 10 10:9 10 10:9 10 10:9 10 10:9 10 10:9 10 10:9 10 10:9 10 10:9 10 10:9 | 52 P 53 P L9 P 58 P 552 P 16 P 04 P 36 P | NO NO NO NO NO NO NO NO | SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT SIGNIFICANT | CLINICAL | OBSERVATIONS | |
| 73717 | F | 300 MG/KG/DAY | | 06-15- 06-16- 06-17- 06-18- 06-19- 06-20- 06-21- 06-04- 06-05- 06-06- 06-07- 06-08- 06-09- 06-10- | 10 10:: | 52 P 49 P 52 P 44 P 25 P 89 P 53 P 553 P 553 P 553 P 553 P 553 P 553 P 553 P | NO NO NO NO NO NO NO NO NO NO NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 14 INDIVIDUAL CLINICAL OBSERVATIONS SPONSOR: AMERICAN PETROLEUM

| TARLE | PANCE. | 06-02-10 | TΟ | 06-24-10 |
|---------|-----------|----------|----|----------|
| ייועטאי | IVAINGE . | 00-02-10 | 10 | 00-24-10 |

| | | | | GAEDGODY | | | | | | | | | |
|--------|-----|-----|-----------|-----------------|---------|---------|------|-----|---------------|-----------|--------------|------|--|
| ANIMAL | SEX | | GROUP | CATEGORY | DA'I'E | TIME | GRAD | E O | BSERVATIONS | | | | |
| | | | | | POST | | | | | | | | |
| 73717 | F | 300 | MG/KG/DAY | NORMAL | 06-12-1 | 0 10:3 | 6 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | 06-14-1 | 0 10:5 | 5 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-15-1 | 0 10:5 | 3 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-16-1 | 0 10:4 | 9 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-17-1 | 0 10:5 | 3 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-18-1 | 0 10:4 | 4 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-19-1 | .0 10:2 | 5 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-20-1 | .0 10:3 | 9 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | 06-21-1 | .0 10:3 | 7 P | NO | SIGNIFICANT | CLINICAL | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| 73693 | F | 300 | MG/KG/DAY | | | | | | | | OBSERVATIONS | | |
| | | | | | 06-06-1 | .0 11:1 | | | | | OBSERVATIONS | | |
| | | | | | 06-07-1 | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | 06-18-1 | | | | | | OBSERVATIONS | | |
| | | | | | 06-19-1 | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| | | | | | | | | | | | OBSERVATIONS | | |
| 73693 | F | 300 | MG/KG/DAY | BODY/INTEGUMENT | 06-24-1 | .0 9:5 | 55 2 | WE: | r yellow mate | ERIAL ANO | GENITAL AREA | | |
| | | | | | | | | | | | | | |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PROJECT NO.:WIL-402010F PAGE 15 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10

| | | | | | | | 10 10 00-24-10 |
|--------|-----|----------------|----------------|---------|---------|------|---|
| ANIMAL | SEX | GROUP | CATEGORY | DATE | TIME | GRAD | E OBSERVATIONS |
| | | | | POST | -DOSING | OBS | ERVATIONS |
| 73693 | F | 300 MG/KG/DAY | EYES/EARS/NOSE | | | | WET CLEAR MATERIAL AROUND NOSE |
| 73693 | F | 300 MG/KG/DAY | | | | | WET CLEAR MATERIAL VENTRAL NECK |
| 73693 | F | 300 MG/KG/DAY | ORAL/DENTAL | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | • | 06-22-1 | 0 9:52 | 2 2 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | 06-23-1 | 0 9:48 | 3 1 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | 06-24-1 | 0 9:5 | 5 1 | WET CLEAR MATERIAL AROUND MOUTH |
| 73718 | F | 1000 MG/KG/DAY | NORMAL | 06-03-1 | 0 10:4 | 7 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-06-1 | 0 11:24 | l P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-07-1 | 0 10:59 |) P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-08-1 | 0 11:02 | 2 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-09-1 | 0 11:03 | 3 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-10-1 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-14-1 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-16-1 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-17-1 | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73718 | F | 1000 MG/KG/DAY | ORAL/DENTAL | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | 06-11-1 | | | |
| | | | | 06-12-1 | | | |
| | | | | | | | DRIED CLEAR MATERIAL AROUND MOUTH |
| | | | | | | | WET RED MATERIAL AROUND MOUTH |
| | | | | | | | DRIED CLEAR MATERIAL AROUND MOUTH |
| 72701 | - | 1000 Mg/Kg/DAX | | | | | WET CLEAR MATERIAL AROUND MOUTH |
| 73701 | F. | 1000 MG/KG/DAY | NORMAL | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 00-10-1 | O IU:54 | : P | NO SIGNIFICANI CHINICAL ODSERVATIONS |
| | | | | | | | |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 16 SPONSOR:AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10

| ANIMAL | SEX | | GROUP | CATEGORY | DATE | TIME | GRAD | DE OBSERVATIONS |
|--------|-----|------|-----------|-----------------|--------|---------|-------|--------------------------------------|
| | | | | | POS | T-DOSIN | G OBS | ERVATIONS |
| 73701 | F | 1000 | MG/KG/DAY | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-17- | 10 10:5 | 4 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-18- | 10 10:4 | 6 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-19- | 10 10:2 | 6 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-20- | 10 10:3 | 9 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-21- | 10 10:4 | 0 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 3701 | F | 1000 | MG/KG/DAY | BEHAVIOR/CNS | 06-05- | 10 11:0 | 4 P | HUNCHED POSTURE |
| | | | | | 06-05- | 10 11:0 | 5 1 | ROCKS, LURCHES, OR SWAYS AS IT WALKS |
| | | | | | 06-06- | 10 11:2 | 6 P | HUNCHED POSTURE |
| | | | | | 06-07- | 10 11:0 | 0 P | HUNCHED POSTURE |
| 3701 | F | 1000 | MG/KG/DAY | BODY/INTEGUMENT | 06-13- | 10 10:3 | 5 1 | WET CLEAR MATERIAL RIGHT FORELIMB |
| | | | | | 06-13- | 10 10:3 | 6 1 | WET CLEAR MATERIAL LEFT FORELIMB |
| | | | | | 06-14- | 10 11:0 | 8 1 | WET CLEAR MATERIAL RIGHT FORELIMB |
| | | | | | | 10 11:0 | | WET CLEAR MATERIAL LEFT FORELIMB |
| | | | | | | 10 11:0 | | |
| | | | | | | 10 11:0 | | WET CLEAR MATERIAL LEFT FORELIMB |
| | | | | | | 10 9:4 | | WET CLEAR MATERIAL RIGHT FORELIMB |
| | | | | | | 10 9:4 | | WET CLEAR MATERIAL LEFT FORELIMB |
| 3701 | F | 1000 | MG/KG/DAY | EYES/EARS/NOSE | | 10 11:0 | | WET CLEAR MATERIAL AROUND NOSE |
| | | | | | | 10 10:5 | | DRIED RED MATERIAL AROUND NOSE |
| | | | | | | 10 9:4 | | WET CLEAR MATERIAL AROUND NOSE |
| 3701 | F | 1000 | MG/KG/DAY | BODY/INTEG II | | 10 11:0 | | WET CLEAR MATERIAL VENTRAL NECK |
| | | | | | | 10 9:4 | | WET CLEAR MATERIAL VENTRAL NECK |
| 3701 | F | 1000 | MG/KG/DAY | ORAL/DENTAL | | 10 11:0 | | DRIED RED MATERIAL AROUND MOUTH |
| | | | | | | 10 10:3 | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | | 10 11:0 | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-16- | 10 10:5 | 1 1 | WET CLEAR MATERIAL AROUND MOUTH |

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PAGE 17 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10

| ANIMAL | SEX | GROUP | CATEGORY | | | | | E OBSERVATIONS |
|--------|-----|----------------|-----------------|--------|-----|-------|------|--------------------------------------|
| | | | | | | | | |
| | | | | POS | T-D | OSING | OBSI | ERVATIONS |
| 73701 | F | 1000 MG/KG/DAY | ORAL/DENTAL | 06-22- | 10 | 9:53 | 1 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | 06-23- | 10 | 9:48 | 2 | WET CLEAR MATERIAL AROUND MOUTH |
| 73721 | F | 1000 MG/KG/DAY | NORMAL | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-09- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-10- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-11- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-16- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-19- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-20- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-21- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-22- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-23- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73721 | F | 1000 MG/KG/DAY | BODY/INTEGUMENT | | | | | WET CLEAR MATERIAL RIGHT FORELIMB |
| | | | | | | | | WET CLEAR MATERIAL LEFT FORELIMB |
| 73721 | F | 1000 MG/KG/DAY | ORAL/DENTAL | | | | | DRIED RED MATERIAL AROUND MOUTH |
| | | | | | | | | DRIED CLEAR MATERIAL AROUND MOUTH |
| | | | | 06-14- | | | | DRIED RED MATERIAL AROUND MOUTH |
| | | | | 06-15- | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | 06-17- | | | | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | 06-18- | | | | DRIED CLEAR MATERIAL AROUND MOUTH |
| 73707 | F | 1000 MG/KG/DAY | NORMAL | 06-04- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-09- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-10- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-11- | | | | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | 06-12- | 10 | 10:53 | Р | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | | | | |

GRADE CODE: 1 - SLIGHT 2 - MODERATE 3 - SEVERE P - PRESENT

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

PROJECT NO.:WIL-402010F PAGE 18 SPONSOR: AMERICAN PETROLEUM INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10

| POST-DOSING POST-DOSING POST-DOSING POST-DOSING POSTATE | ANIMAL | SEX | | GROUP | CATEGORY | | | | OBSERVATIONS |
|--|--------|-----|------|--------------|-----------------|---------|---------|-----|--------------------------------------|
| 73707 F 1000 MG/KG/DAY | | | | | | | | | PDVNTTONG |
| 73707 F 1000 MG/KG/DAY | 73707 | F | 1000 | MG/KG/DAV | ΝΟΡΜΔΤ. | | | | |
| 06-06-10 11:28 P | | | | | | | | | |
| 06-06-10 11:28 1 ROCKS, LURCHES, OR SWAYS AS IT WALKS | 73707 | - | 1000 | 110/110/2111 | BBIBIVION, CIVE | | | | |
| 73707 F 1000 MG/KG/DAY BODY/INTEGUMENT 06-15-10 11:10 1 WET CLEAR MATERIAL LEFT FORELIMB 73707 F 1000 MG/KG/DAY EYES/EARS/NOSE 06-05-10 11:13 2 LACRIMATION RIGHT EYE 06-05-10 11:13 2 LACRIMATION LEFT EYE 06-15-10 11:10 1 WET CLEAR MATERIAL AROUND NOSE 73707 F 1000 MG/KG/DAY ORAL/DENTAL 06-06-10 11:30 1 DRIED RED MATERIAL AROUND MOUTH 06-07-10 11:02 2 DRIED RED MATERIAL AROUND MOUTH 06-08-10 11:09 1 WET CLEAR MATERIAL AROUND MOUTH 06-14-10 11:09 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:10 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:10 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 10:51 2 SALIVATION 06-17-10 10:54 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 2 SALIVATION 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-08-10 11:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:15 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-07-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| 73707 F 1000 MG/KG/DAY EYES/EARS/NOSE 06-05-10 11:13 2 LACRIMATION RIGHT EYE 06-05-10 11:13 2 LACRIMATION LEFT EYE 06-05-10 11:13 2 LACRIMATION LEFT EYE 06-05-10 11:13 2 LACRIMATION LEFT EYE WET CLEAR MATERIAL AROUND NOSE MET CLEAR MATERIAL AROUND MOUTH O6-05-01 11:05 DRIED RED MATERIAL AROUND MOUTH O6-07-01 11:05 DRIED RED MATERIAL AROUND MOUTH O6-14-10 11:05 DRIED RED MATERIAL AROUND MOUTH O6-14-10 11:10 WET CLEAR MATERIAL AROUND MOUTH O6-15-10 10:11 WET CLEAR MATERIAL AROUND MOUTH O6-16-10 10:15 SALIVATION WET CLEAR MATERIAL AROUND MOUTH O6-18-10 10:47 WET CLEAR MATERIAL AROUND MOUTH O6-19-10 10:47 WET CLEAR MATERIAL AROUND MOUTH O6-20-10 10:40 WET CLEAR MATERIAL AROUND MOUTH O6-20-10 10:40 WET CLEAR MATERIAL AROUND MOUTH O6-21-10 00:40 WET CLEAR MATERIAL AROUND MOUTH WET CLEAR MATERIAL AROUND MOUTH O6-21-10 00:40 WET CLEAR MATERIAL AROUND MOUTH O6-22-10 9:53 WET CLEAR MATERIAL AROUND MOUTH O6-22-10 9:50 WET CLEAR MATERIAL AROUND MOUTH O6-22-10 00:40 WET CLEAR MATERIAL AROUND MOUTH O6-22 | 73707 | F | 1000 | MG/KG/DAY | BODY/INTEGUMENT | | | | |
| 73707 F 1000 MG/KG/DAY ORAL/DENTAL 06-05-10 11:13 2 LACRIMATION LEFT EVE 06-15-10 11:10 1 WET CLEAR MATERIAL AROUND NOSE 06-06-10 11:30 1 DRIED RED MATERIAL AROUND MOUTH 06-07-10 11:02 2 DRIED RED MATERIAL AROUND MOUTH 06-07-10 11:05 1 DRIED RED MATERIAL AROUND MOUTH 06-15-10 11:10 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:10 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 10:51 2 SALIVATION 06-17-10 10:54 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 2 SALIVATION 06-18-10 10:47 2 SALIVATION WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:27 1 WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET CLEAR MATERIAL AROUND MO | | | | | | | | | |
| 73707 F 1000 MG/KG/DAY ORAL/DENTAL | | | | -, -, | ., ., | | | | |
| 06-07-10 11:02 2 DRIED RED MATERIAL AROUND MOUTH 06-08-10 11:05 1 DRIED RED MATERIAL AROUND MOUTH 06-14-10 11:09 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:10 2 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 10:51 2 SALVATION 06-16-10 10:51 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 2 SALVATION 06-18-10 10:47 2 SALVATION 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:27 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 1 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-30-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | 06-15-1 | 0 11:10 | 1 | WET CLEAR MATERIAL AROUND NOSE |
| 06-08-10 11:05 1 DRIED RED MATERIAL AROUND MOUTH 06-14-10 11:09 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:10 2 WET CLEAR MATERIAL AROUND MOUTH 06-16-10 10:51 2 SALIVATION 06-16-10 10:51 2 SALIVATION 06-17-10 10:54 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 2 SALIVATION 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 06-17 | 73707 | F | 1000 | MG/KG/DAY | ORAL/DENTAL | 06-06-1 | 0 11:30 | 1 | DRIED RED MATERIAL AROUND MOUTH |
| 06-14-10 11:09 1 WET CLEAR MATERIAL AROUND MOUTH 06-15-10 11:10 2 WET CLEAR MATERIAL AROUND MOUTH 06-16-10 10:51 2 SALIVATION 06-17-10 10:54 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 2 SALIVATION 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:27 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-05-10 11:131 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | 06-07-1 | 0 11:02 | 2 | DRIED RED MATERIAL AROUND MOUTH |
| 06-15-10 11:10 2 WET CLEAR MATERIAL AROUND MOUTH 06-16-10 10:51 2 SALIVATION 06-17-10 10:54 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 2 SALIVATION 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:27 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-21-10 10:50 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | 06-08-1 | 0 11:05 | 1 | DRIED RED MATERIAL AROUND MOUTH |
| 06-16-10 10:51 2 SALIVATION 06-17-10 10:54 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 2 SALIVATION 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:27 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-33-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | 06-14-1 | 0 11:09 | 1 | WET CLEAR MATERIAL AROUND MOUTH |
| 06-17-10 10:54 1 WET CLEAR MATERIAL AROUND MOUTH 06-18-10 10:47 2 SALIVATION 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:27 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-21-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-05-10 11:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | 06-15-1 | 0 11:10 | | |
| 06-18-10 10:47 2 SALIVATION 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:27 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 11:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | 06-16-1 | 0 10:51 | . 2 | SALIVATION |
| 06-18-10 10:47 1 WET CLEAR MATERIAL AROUND MOUTH 06-19-10 10:27 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-03-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-03-10 11:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| 06-19-10 10:27 1 WET CLEAR MATERIAL AROUND MOUTH 06-20-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-05-10 11:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| 06-20-10 10:40 2 WET RED MATERIAL AROUND MOUTH 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 06-05-10 11:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| 06-21-10 10:40 2 WET CLEAR MATERIAL AROUND MOUTH 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 73722 F 1000 MG/KG/DAY NORMAL 06-05-10 11:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| 06-22-10 9:53 1 WET CLEAR MATERIAL AROUND MOUTH 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 73722 F 1000 MG/KG/DAY NORMAL 06-05-10 11:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| 06-23-10 9:50 2 WET CLEAR MATERIAL AROUND MOUTH 73722 F 1000 MG/KG/DAY NORMAL 06-05-10 11:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| 73722 F 1000 MG/KG/DAY NORMAL 06-05-10 11:13 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| 06-06-10 11:31 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | _ | | / / | | | | | |
| 06-08-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | 73722 | F. | 1000 | MG/KG/DAY | NORMAL | | | | |
| 06-09-10 11:05 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-10-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| 06-10-10 $10:55$ P NO SIGNIFICANT CLINICAL OBSERVATIONS $06-17-10$ $10:55$ P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| 06-17-10 10:55 P NO SIGNIFICANT CLINICAL OBSERVATIONS | | | | | | | | | |
| | | | | | | | | | |
| 00-19-10 10:27 P NO SIGNIFICANT CHINICAL OBSERVATIONS | | | | | | | | | |
| | | | | | | 06-19-1 | 0 10:27 | Р | NO SIGNIFICANI CHINICAL OBSERVATIONS |

GRADE CODE: 1 - SLIGHT 2 - MODERATE 3 - SEVERE P - PRESENT

TABLE I13 (FEMALES - 1 HOUR POST-DOSING) ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL CLINICAL OBSERVATIONS

TABLE RANGE: 06-02-10 TO 06-24-10

| | | | | | יו מחמעו | ANGE. 0 | | 10 10 06-24-10 |
|--------|-----|------|-----------|-----------------|----------|---------|-------|--------------------------------------|
| ANIMAL | SEX | | GROUP | CATEGORY | DATE | TIME | GRAD | E OBSERVATIONS |
| | | | | | POS | T-DOSIN | G OBS | ERVATIONS |
| 73722 | F | 1000 | MG/KG/DAY | NORMAL | 06-20- | 10 10:4 | 0 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-21- | 10 10:4 | 1 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-23- | 10 9:5 | 0 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| | | | | | 06-24- | 10 9:5 | 5 P | NO SIGNIFICANT CLINICAL OBSERVATIONS |
| 73722 | F | 1000 | MG/KG/DAY | BODY/INTEGUMENT | 06-14- | 10 11:1 | 0 1 | WET CLEAR MATERIAL RIGHT FORELIMB |
| | | | | | 06-14- | 10 11:1 | 0 1 | WET CLEAR MATERIAL LEFT FORELIMB |
| 73722 | F | 1000 | MG/KG/DAY | EYES/EARS/NOSE | 06-22- | 10 9:5 | 4 1 | WET CLEAR MATERIAL AROUND NOSE |
| 73722 | F | 1000 | MG/KG/DAY | ORAL/DENTAL | 06-07- | 10 11:0 | 2 1 | DRIED RED MATERIAL AROUND MOUTH |
| | | | | | 06-11- | 10 11:1 | 5 1 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-12- | 10 10:5 | 3 1 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-13- | 10 10:3 | 7 1 | DRIED CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-14- | 10 11:0 | 9 1 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-15- | 10 11:1 | 0 1 | DRIED RED MATERIAL AROUND MOUTH |
| | | | | | 06-16- | 10 10:5 | 2 1 | DRIED RED MATERIAL AROUND MOUTH |
| | | | | | 06-18- | 10 10:4 | 8 2 | WET CLEAR MATERIAL AROUND MOUTH |
| | | | | | 06-22- | 10 9:5 | 4 2 | WET CLEAR MATERIAL AROUND MOUTH |

GRADE CODE: 1 - SLIGHT 2 - MODERATE 3 - SEVERE P - PRESENT

PCRDv4.17 07/15/2010

PAGE 19

Page 291 of 360

| | GNANCY TATUS | DAY 0 | 6 | 7 | 10 | 14 | 17 | 20 |
|--------|-----------------|-------------|----------|------|------|------|------|--------------------------------|
| DAMS 1 | FROM GR | OUP 1: 0 MC | G/KG/DAY | | | | | |
| 73710 | G | 251. | 266. | 271. | 280. | 305. | 334. | 375. SCHEDULED NECROPSY DAY 20 |
| 73704 | G | 257. | 280. | 276. | 297. | 317. | 351. | 397. SCHEDULED NECROPSY DAY 20 |
| 73694 | G | 243. | 255. | 263. | 274. | 282. | 310. | 345. SCHEDULED NECROPSY DAY 20 |
| 73713 | G | 247. | 268. | 271. | 291. | 311. | 350. | 385. SCHEDULED NECROPSY DAY 20 |
| 73720 | G | 268. | 289. | 292. | 293. | 302. | 344. | 392. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 253. | 272. | 275. | 287. | 303. | 338. | 379. |
| S.D. | | 9.8 | 13.2 | 10.8 | 9.6 | 13.3 | 16.9 | 20.6 |
| S.E. | | 4.4 | 5.9 | 4.8 | 4.3 | 5.9 | 7.6 | 9.2 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| | | | | | | | | |

TABLE 114

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS
SDONSOR-AMERICAN DETROLEUM INDIVIDUAL BODY WEIGHTS DURING GESTATION [G]

| | GNANCY TATUS | DAY 0 | 6 | 7 | 10 | 14 | 17 | 20 |
|-------|-----------------|-------------|-----------|------|------|------|------|--------------------------------|
| DAMS | FROM GR | OUP 2: 30 M | IG/KG/DAY | | | | | |
| 73702 | G | 257. | 280. | 286. | 302. | 322. | 356. | 414. SCHEDULED NECROPSY DAY 20 |
| 73697 | G | 251. | 254. | 263. | 276. | 300. | 327. | 373. SCHEDULED NECROPSY DAY 20 |
| 73706 | G | 225. | 249. | 250. | 263. | 279. | 308. | 353. SCHEDULED NECROPSY DAY 20 |
| 73709 | G | 257. | 280. | 276. | 291. | 307. | 347. | 380. SCHEDULED NECROPSY DAY 20 |
| 73719 | G | 252. | 272. | 269. | 295. | 306. | 359. | 406. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 248. | 267. | 269. | 285. | 303. | 339. | 385. |
| S.D. | | 13.4 | 14.6 | 13.6 | 15.7 | 15.6 | 21.5 | 24.9 |
| S.E. | | 6.0 | 6.5 | 6.1 | 7.0 | 7.0 | 9.6 | 11.1 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |

PREGNANCY

PAGE 3

| | GNANCY FATUS | DAY 0 | 6 | 7 | 10 | 14 | 17 | 20 |
|--------|-----------------|---------------|-----------|------|------|------|------|--------------------------------|
| DAMS I | FROM GE | ROUP 3: 100 N | MG/KG/DAY | | | | | |
| 73696 | G | 251. | 281. | 280. | 301. | 323. | 357. | 393. SCHEDULED NECROPSY DAY 20 |
| 73699 | G | 251. | 289. | 295. | 299. | 316. | 343. | 400. SCHEDULED NECROPSY DAY 20 |
| 73714 | G | 244. | 242. | 259. | 267. | 285. | 306. | 344. SCHEDULED NECROPSY DAY 20 |
| 73705 | G | 272. | 297. | 298. | 314. | 330. | 371. | 424. SCHEDULED NECROPSY DAY 20 |
| 73700 | G | 290. | 294. | 290. | 308. | 314. | 357. | 401. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 262. | 281. | 284. | 298. | 314. | 347. | 392. |
| S.D. | | 19.0 | 22.4 | 15.8 | 18.2 | 17.2 | 24.9 | 29.5 |
| S.E. | | 8.5 | 10.0 | 7.0 | 8.1 | 7.7 | 11.1 | 13.2 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |

PREGNANCY STATUS DAY 0 6 7 10 14 17 20

PAGE 4

| 5 | TATUS | DAY 0 | 6 | / | 10 | 14 | 1/ | 20 | | | |
|----------|---------|--------------|----------|------|------|------|------|------------------|---------|-----|----|
| DAMS | FROM GR | OUP 4: 300 M | G/KG/DAY | | | | | | | | |
| 73708 | G | 290. | 305. | 305. | 316. | 341. | 372. | 418. SCHEDULED N | ECROPSY | DAY | 20 |
| 73712 | G | 246. | 259. | 264. | 280. | 293. | 305. | 349. SCHEDULED N | ECROPSY | DAY | 20 |
| 73698 | G | 235. | 244. | 261. | 277. | 298. | 331. | 363. SCHEDULED N | ECROPSY | DAY | 20 |
| 73717 | G | 262. | 275. | 274. | 297. | 301. | 332. | 370. SCHEDULED N | ECROPSY | DAY | 20 |
| 73693 | G | 249. | 257. | 256. | 279. | 289. | 335. | 366. SCHEDULED N | ECROPSY | DAY | 20 |
| MEAN | | 256. | 268. | 272. | 290. | 304. | 335. | 373. | | | |
| S.D. | | 21.1 | 23.4 | 19.6 | 16.7 | 21.0 | 23.9 | 26.3 | | | |
| S.E. | | 9.4 | 10.5 | 8.8 | 7.5 | 9.4 | 10.7 | 11.7 | | | |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 | | | |
| | | | | | | | | | | | |

SPONSOR: AMERICAN PETROLEUM

TABLE 114

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 5 INDIVIDUAL BODY WEIGHTS DURING GESTATION [G]

| | EGNANCY STATUS | DAY 0 | 6 | 7 | 10 | 14 | 17 | 20 |
|-------|-------------------|-------------|----------|------|------|------|------|--------------------------------|
| DAMS | FROM GR | OUP 5: 1000 | MG/KG/DA | Y | | | | |
| 73718 | G | 273. | 291. | 293. | 300. | 319. | 302. | 300. SCHEDULED NECROPSY DAY 20 |
| 73701 | G | 245. | 225. | 231. | 255. | 246. | 273. | 280. SCHEDULED NECROPSY DAY 20 |
| 73721 | G | 254. | 259. | 263. | 281. | 298. | 324. | 360. SCHEDULED NECROPSY DAY 20 |
| 73707 | G | 279. | 252. | 276. | 290. | 310. | 316. | 319. SCHEDULED NECROPSY DAY 20 |
| 73722 | G | 247. | 259. | 257. | 265. | 278. | 305. | 339. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 260. | 257. | 264. | 278. | 290. | 304. | 320. |
| S.D. | | 15.5 | 23.5 | 23.0 | 18.3 | 29.1 | 19.4 | 31.5 |
| S.E. | | 6.9 | 10.5 | 10.3 | 8.2 | 13.0 | 8.7 | 14.1 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |

G = GRAVID NG = NONGRAVID - WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

PGBWv4.09 07/15/2010 N

PAGE 1

PREGNANCY STATUS DAY 0- 6 6- 7 7- 10 10- 14 14- 17 17- 20 0- 20 ______ DAMS FROM GROUP 1: 0 MG/KG/DAY 25. 29. 41. 124. SCHEDULED NECROPSY DAY 20
20. 34. 46. 140. SCHEDULED NECROPSY DAY 20
8. 28. 35. 102. SCHEDULED NECROPSY DAY 20
20. 39. 35. 138. SCHEDULED NECROPSY DAY 20
9. 42. 48. 124. SCHEDULED NECROPSY DAY 20 73710 G 15. 5. 9. 21. 11. 20. -4. 8. 3. 73704 G 23. 73694 G 21. 12. 73713 G 21. 73720 G MEAN 16. 34. 41. 126. 18. 3. 12. 4.4 8.3 7.5 15.2 S.D. 4.7 6.1 6.0 3.4 S.E. 2.1 2.0 3.7 2.7 2.7 6.8

5

5

5

G = GRAVID NG = NONGRAVID - WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

5

5

5

5

PREGNANCY

| STA | TUS | DAY 0- | 6 6- 7 | 7- 10 | 10- 14 | 14- 17 | 17- 20 | 0- 20 | | | | |
|-------|------|----------|-------------|--------|--------|--------|--------|-------|-----------|----------|-------|---|
| DAMS | FROM | GROUP 2: | 30 MG/KG/DA | .Y | | | | | | | | |
| 73702 | G | 23 | . 6. | 16. | 20. | 34. | 58. | 157. | SCHEDULED | NECROPSY | DAY 2 | 0 |
| 73697 | G | 3 | . 9. | 13. | 24. | 27. | 46. | 122. | SCHEDULED | NECROPSY | DAY 2 | 0 |
| 73706 | G | 24 | . 1. | 13. | 16. | 29. | 45. | 128. | SCHEDULED | NECROPSY | DAY 2 | 0 |
| 73709 | G | 23 | 4. | 15. | 16. | 40. | 33. | 123. | SCHEDULED | NECROPSY | DAY 2 | 0 |
| 73719 | G | 20 | 3. | 26. | 11. | 53. | 47. | 154. | SCHEDULED | NECROPSY | DAY 2 | 0 |
| MEAN | | 19 | . 2. | 17. | 17. | 37. | 46. | 137. | | | | |
| S.D. | | 8. | 8 5.6 | 5.4 | 4.9 | 10.5 | 8.9 | 17.3 | | | | |
| S.E. | | 4. | 0 2.5 | 2.4 | 2.2 | 4.7 | 4.0 | 7.7 | | | | |
| N | | | 5 5 | 5 | 5 | 5 | 5 | 5 | | | | |

5

5

PAGE 3

PREGNANCY STATUS DAY 0- 6 6- 7 7- 10 10- 14 14- 17 17- 20 0- 20 ______ DAMS FROM GROUP 3: 100 MG/KG/DAY 22. 34. 36. 142. SCHEDULED NECROPSY DAY 20
17. 27. 57. 149. SCHEDULED NECROPSY DAY 20
18. 21. 38. 100. SCHEDULED NECROPSY DAY 20
16. 41. 53. 152. SCHEDULED NECROPSY DAY 20
6. 43. 44. 111. SCHEDULED NECROPSY DAY 20 -1. 73696 G 30. 21. 38. 4. 8. 6. 73699 G 17. 1. -4. 73714 G -2. 8. 16. 25. 73705 G 73700 G 4. 18. MEAN 19. 46. 131. 4. 13. 16. 33. 5.9 8.2 9.2 23.7 S.D. 17.2 7.1 9.3 3.2 S.E. 7.7 3.7 2.7 4.2 4.1 10.6

5

G = GRAVID NG = NONGRAVID - WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

5

5

5

5

N

PREGNANCY STATUS DAY 0- 6 6- 7 7- 10 10- 14 14- 17 17- 20 0- 20 ______ DAMS FROM GROUP 4: 300 MG/KG/DAY 73708 G 15. 0. 11. 25. 31. 46. 128. SCHEDULED NECROPSY DAY 20 73712 G 13. 5. 16. 13. 12. 44. 103. SCHEDULED NECROPSY DAY 20 SCHEDULED NECROPSY DAY 20

> SCHEDULED NECROPSY DAY 20 SCHEDULED NECROPSY DAY 20

| 73698 | G | 9. | 17. | 16. | 21. | 33. | 32. | 128. |
|-------|---|-----|-----|-----|-----|------|-----|------|
| 73717 | G | 13. | -1. | 23. | 4. | 31. | 38. | 108. |
| 73693 | G | 8. | -1. | 23. | 10. | 46. | 31. | 117. |
| MEAN | | 12. | 4. | 18. | 15. | 31. | 38. | 117. |
| S.D. | | 3.0 | 7.7 | 5.2 | 8.4 | 12.1 | 6.8 | 11.4 |
| S.E. | | 1.3 | 3.4 | 2.3 | 3.8 | 5.4 | 3.0 | 5.1 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| | | | | | | | | |

SPONSOR: AMERICAN PETROLEUM

TABLE 115
PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 5

INDIVIDUAL BODY WEIGHT CHANGES DURING GESTATION [G]

| | | | | | | | | | | | | | | |
|---------------|------|------------|-----------|---------|--------|--------|--------|-------|-----------|----------|-----|----|------|--|
| PREGN. STA | | DAY 0- | 6 6- | 7 7- 10 | 10- 14 | 14- 17 | 17- 20 | 0- 20 | | | | | | |
| DAMS | FROM | GROUP 5: 1 | 000 MG/KG | /DAY | | | | | | | | | | |
| 73718 | G | 18 | . 2 | . 7. | 19. | -17. | -2. | 27. | SCHEDULED | NECROPSY | DAY | 20 | | |
| 73701 | G | -20 | . 6 | . 24. | -9. | 27. | 7. | 35. | SCHEDULED | NECROPSY | DAY | 20 | | |
| 73721 | G | 5 | . 4 | . 18. | 17. | 26. | 36. | 106. | SCHEDULED | NECROPSY | DAY | 20 | | |
| 73707 | G | -27 | . 24 | . 14. | 20. | 6. | 3. | 40. | SCHEDULED | NECROPSY | DAY | 20 | | |
| 73722 | G | 12 | 2 | . 8. | 13. | 27. | 34. | 92. | SCHEDULED | NECROPSY | DAY | 20 | | |
| MEAN | | -2 | . 7 | . 14. | 12. | 14. | 16. | 60. | | | | | | |
| S.D. | | 20. | 0 10. | 1 7.1 | 12.0 | 19.4 | 18.0 | 36.2 | | | | | | |
| S.E. | | 8. | 9 4. | 5 3.2 | 5.4 | 8.7 | 8.1 | 16.2 | | | | | | |
| N | | | 5 | 5 5 | 5 | 5 | 5 | 5 | | | | | | |

G = GRAVID NG = NONGRAVID - WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

PGBWv4.09 07/15/2010

TABLE 116
ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010F PAGE 1 INDIVIDUAL GRAVID UTERINE WTS. AND NET BODY WT. CHANGES [G]

| | PREGNAN STATUS | CY INITIAL BODY WT. | TERMINAL BODY WT. | GRAVID UTERINE WT. | NET BODY WT. | NET BODY WT. CHANGE | |
|-------|-------------------|------------------------|----------------------|-----------------------|-----------------|------------------------|--|
| DAM # | | GROUP 1: 0 | MG/KG/DAY | | | | |
| 73710 | G | 251. | 375. | 66.1 | 308.9 | 57.9 | |
| 73704 | G | 257. | 397. | 81.6 | 315.4 | 58.4 | |
| 73694 | G | 243. | 345. | 74.0 | 271.0 | 28.0 | |
| 73713 | G | 247. | 385. | 83.2 | 301.8 | 54.8 | |
| 73720 | G | 268. | 392. | 68.6 | 323.4 | 55.4 | |
| MEAN | | 253. | 379. | 74.7 | 304.1 | 50.9 | |
| S.D. | | 9.8 | 20.6 | 7.61 | 20.15 | 12.89 | |
| S.E. | | 4.4 | 9.2 | 3.40 | 9.01 | 5.77 | |
| N | | 5 | 5 | 5 | 5 | 5 | |

G = GRAVID

TABLE I16
PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 2

INDIVIDUAL GRAVID UTERINE WTS. AND NET BODY WT. CHANGES [G]

| | PREGNAN STATUS | ICY INITIAL BODY WI | | GRAVID UTERINE WT | NET BODY . WT. | NET BODY WT. CHANGE | |
|-------|-------------------|------------------------|--------------|----------------------|----------------|------------------------|--|
| DAM # | | GROUP 2: | 30 MG/KG/DAY | | | | |
| 73702 | G | 257. | 414. | 76.6 | 337.4 | 80.4 | |
| 73697 | G | 251. | 373. | 79.8 | 293.2 | 42.2 | |
| 73706 | G | 225. | 353. | 66.4 | 286.6 | 61.6 | |
| 73709 | G | 257. | 380. | 79.8 | 300.2 | 43.2 | |
| 73719 | G | 252. | 406. | 90.3 | 315.7 | 63.7 | |
| MEAN | | 248. | 385. | 78.6 | 306.6 | 58.2 | |
| S.D. | | 13.4 | 24.9 | 8.55 | 20.32 | 15.93 | |
| S.E. | | 6.0 | 11.1 | 3.82 | 9.09 | 7.13 | |
| N | | 5 | 5 5 | 5 | 5 | 5 | |

G = GRAVID

TABLE I16
PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

INDIVIDUAL GRAVID UTERINE WTS. AND NET BODY WT. CHANGES [G]

PAGE 3

| | PREGNANC STATUS | Y INITIAL BODY WT. | TERMINAL BODY WT. | GRAVID UTERINE WT. | NET BODY . WT. | NET BODY WT. CHANGE | |
|-------|--------------------|-----------------------|----------------------|-----------------------|-------------------|------------------------|--|
| DAM # | | GROUP 3: 100 | MG/KG/DAY | | | | |
| 73696 | G | 251. | 393. | 85.1 | 307.9 | 56.9 | |
| 73699 | G | 251. | 400. | 84.2 | 315.8 | 64.8 | |
| 73714 | G | 244. | 344. | 63.7 | 280.3 | 36.3 | |
| 73705 | G | 272. | 424. | 93.0 | 331.0 | 59.0 | |
| 73700 | G | 290. | 401. | 85.8 | 315.2 | 25.2 | |
| MEAN | | 262. | 392. | 82.4 | 310.0 | 48.4 | |
| S.D. | | 19.0 | 29.5 | 11.00 | 18.63 | 16.87 | |
| S.E. | | 8.5 | 13.2 | 4.92 | 8.33 | 7.54 | |
| N | | 5 | 5 | 5 | 5 | 5 | |

G = GRAVID

TABLE 116
ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PROJECT NO.:WIL-402010F PAGE 4 INDIVIDUAL GRAVID UTERINE WTS. AND NET BODY WT. CHANGES [G]

| | PREGNANCY STATUS | INITIAL BODY WT. | TERMINAL BODY WT. | GRAVID UTERINE WT. | NET BODY | NET BODY WT. CHANGE | |
|-------|---------------------|---------------------|----------------------|-----------------------|----------|------------------------|--|
| DAM # | G | ROUP 4: 300 | MG/KG/DAY | | | | |
| 73708 | G | 290. | 418. | 83.4 | 334.6 | 44.6 | |
| 73712 | G | 246. | 349. | 59.9 | 289.1 | 43.1 | |
| 73698 | G | 235. | 363. | 73.4 | 289.6 | 54.6 | |
| 73717 | G | 262. | 370. | 75.5 | 294.5 | 32.5 | |
| 73693 | G | 249. | 366. | 81.3 | 284.7 | 35.7 | |
| MEAN | Ī | 256. | 373. | 74.7 | 298.5 | 42.1 | |
| S.D. | | 21.1 | 26.3 | 9.23 | 20.48 | 8.61 | |
| S.E. | | 9.4 | 11.7 | 4.13 | 9.16 | 3.85 | |
| N | | 5 | 5 | 5 | 5 | 5 | |

G = GRAVID

SPONSOR: AMERICAN PETROLEUM

TABLE I16
PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 5

INDIVIDUAL GRAVID UTERINE WTS. AND NET BODY WT. CHANGES [G]

| | PREGNANCY STATUS | INITIAL BODY WT. | TERMINAL BODY WT. | GRAVID UTERINE WT. | NET BODY WT. | NET BODY WT. CHANGE | |
|-------|---------------------|---------------------|----------------------|-----------------------|-----------------|------------------------|--|
| DAM # | GI | ROUP 5: 100 | 0 MG/KG/DAY | | | | |
| 73718 | G | 273. | 300. | 1.0 | 299.0 | 26.0 | |
| 73701 | G | 245. | 280. | 3.9 | 276.1 | 31.1 | |
| 73721 | G | 254. | 360. | 63.2 | 296.8 | 42.8 | |
| 73707 | G | 279. | 319. | 1.3 | 317.7 | 38.7 | |
| 73722 | G | 247. | 339. | 70.9 | 268.1 | 21.1 | |
| MEAN | | 260. | 320. | 28.1 | 291.5 | 31.9 | |
| S.D. | | 15.5 | 31.5 | 35.71 | 19.72 | 8.90 | |
| S.E. | | 6.9 | 14.1 | 15.97 | 8.82 | 3.98 | |
| N | | 5 | 5 | 5 | 5 | 5 | |

G = GRAVID

PUTv4.06
07/15/2010

| | GNANCY TATUS DA | AY 0-6 | 6- 7 | 7-10 | 10-14 | 14-17 | 17-20 | 0-20 |
|-------|--------------------|-----------|--------|------|-------|-------|-------|-------------------------------|
| DAMS | FROM GROUE | P 1: 0 MG | KG/DAY | | | | | |
| 73710 | G | 11. | 13. | 13. | 16. | 17. | 18. | 14. SCHEDULED NECROPSY DAY 20 |
| 73704 | G | 15. | 12. | 13. | 18. | 19. | 20. | 17. SCHEDULED NECROPSY DAY 20 |
| 73694 | G | 12. | 16. | 12. | 12. | 14. | 14. | 13. SCHEDULED NECROPSY DAY 20 |
| 73713 | G | 14. | 12. | 15. | 15. | 19. | 16. | 15. SCHEDULED NECROPSY DAY 20 |
| 73720 | G | 16. | 15. | 11. | 11. | 19. | 21. | 15. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 14. | 14. | 13. | 14. | 18. | 18. | 15. |
| S.D. | | 2.1 | 1.8 | 1.5 | 2.9 | 2.2 | 2.9 | 1.5 |
| S.E. | | 0.9 | 0.8 | 0.7 | 1.3 | 1.0 | 1.3 | 0.7 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |

PROJECT NO.:WIL-402010F

PAGE 2

| | GNANCY TATUS | DAY 0-6 | 6- 7 | 7-10 | 10-14 | 14-17 | 17-20 | 0-20 |
|-------|-----------------|--------------|----------|------|-------|-------|-------|-------------------------------|
| DAMS | FROM GI | ROUP 2: 30 M | G/KG/DAY | | | | | |
| 73702 | G | 14. | 21. | 17. | 17. | 17. | 22. | 17. SCHEDULED NECROPSY DAY 20 |
| 73697 | G | 11. | 15. | 15. | 16. | 16. | 18. | 15. SCHEDULED NECROPSY DAY 20 |
| 73706 | G | 13. | 15. | 14. | 17. | 17. | 20. | 16. SCHEDULED NECROPSY DAY 20 |
| 73709 | G | 17. | 16. | 17. | 17. | 20. | 20. | 18. SCHEDULED NECROPSY DAY 20 |
| 73719 | G | 15. | 4. | 15. | 16. | 22. | 21. | 17. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 14. | 14. | 16. | 17. | 18. | 20. | 17. |
| S.D. | | 2.2 | 6.2 | 1.3 | 0.5 | 2.5 | 1.5 | 1.1 |
| S.E. | | 1.0 | 2.8 | 0.6 | 0.2 | 1.1 | 0.7 | 0.5 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |

SPONSOR: AMERICAN PETROLEUM

TABLE I17
PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 3

INDIVIDUAL FOOD CONSUMPTION DURING GESTATION [G/ANIMAL/DAY]

| | GNANCY TATUS | DAY 0-6 | 6- 7 | 7-10 | 10-14 | 14-17 | 17-20 | 0-20 |
|-------|---------------------|---------------|----------|------|-------|-------|-------|-------------------------------|
| DAMS | FROM G | ROUP 3: 100 M | G/KG/DAY | | | | | |
| 73696 | G | 15. | 13. | 16. | 17. | 19. | 15. | 16. SCHEDULED NECROPSY DAY 20 |
| 73699 | G | 15. | 16. | 12. | 14. | 17. | 19. | 15. SCHEDULED NECROPSY DAY 20 |
| 73714 | G | 10. | 16. | 14. | 13. | 14. | 15. | 13. SCHEDULED NECROPSY DAY 20 |
| 73705 | G | 17. | 16. | 17. | 17. | 20. | 21. | 18. SCHEDULED NECROPSY DAY 20 |
| 73700 | G | 15. | 11. | 15. | 14. | 19. | 18. | 16. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 14. | 14. | 15. | 15. | 18. | 18. | 16. |
| S.D. | | 2.6 | 2.3 | 1.9 | 1.9 | 2.4 | 2.6 | 1.8 |
| S.E. | | 1.2 | 1.0 | 0.9 | 0.8 | 1.1 | 1.2 | 0.8 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |

PROJECT NO.:WIL-402010F

SPONSOR: AMERICAN PETROLEUM

PAGE 4

| | GNANCY TATUS D |)AY 0-6 | 6- 7 | 7-10 | 10-14 | 14-17 | 17-20 | 0-20 |
|-------|-------------------|-------------|-----------|------|-------|-------|-------|-------------------------------|
| DAMS | FROM GROU | JP 4: 300 M | IG/KG/DAY | | | | | |
| 73708 | G | 15. | 15. | 14. | 17. | 19. | 18. | 16. SCHEDULED NECROPSY DAY 20 |
| 73712 | G | 12. | 16. | 16. | 14. | 13. | 16. | 14. SCHEDULED NECROPSY DAY 20 |
| 73698 | G | 11. | 18. | 15. | 16. | 18. | 14. | 15. SCHEDULED NECROPSY DAY 20 |
| 73717 | G | 11. | 12. | 15. | 12. | 15. | 14. | 13. SCHEDULED NECROPSY DAY 20 |
| 73693 | G | 10. | 11. | 14. | 12. | 21. | 14. | 13. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 12. | 14. | 15. | 14. | 17. | 15. | 14. |
| S.D. | | 1.9 | 2.9 | 0.8 | 2.3 | 3.2 | 1.8 | 1.3 |
| S.E. | | 0.9 | 1.3 | 0.4 | 1.0 | 1.4 | 0.8 | 0.6 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |

TABLE 117

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS
SPONSOR:AMERICAN PETROLEUM INDIVIDUAL FOOD CONSUMPTION DURING GESTATION [G/ANIMAL/DAY]

| | GNANCY | DAY 0-6 | 6- 7 | 7-10 | 10-14 | 14-17 | 17-20 | 0-20 |
|-------|--------|--------------|-----------|------|-------|-------|-------|-------------------------------|
| DAMS | FROM G | ROUP 5: 1000 | MG/KG/DAY | Y | | | | |
| 73718 | G | 13. | 17. | 15. | 18. | 15. | 13. | 15. SCHEDULED NECROPSY DAY 20 |
| 73701 | G | 5. | 8. | 18. | 11. | 18. | NA | NA SCHEDULED NECROPSY DAY 20 |
| 73721 | G | 11. | 9. | 15. | 14. | 16. | 19. | 14. SCHEDULED NECROPSY DAY 20 |
| 73707 | G | 7. | 15. | 18. | NA | 14. | 17. | 13. SCHEDULED NECROPSY DAY 20 |
| 73722 | G | 10. | 11. | 11. | 13. | 16. | 15. | 13. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 9. | 12. | 15. | 14. | 16. | 16. | 14. |
| S.D. | | 3.2 | 3.9 | 2.9 | 2.9 | 1.5 | 2.6 | 1.0 |
| S.E. | | 1.4 | 1.7 | 1.3 | 1.5 | 0.7 | 1.3 | 0.5 |
| N | | 5 | 5 | 5 | 4 | 5 | 4 | 4 |

 ${\tt G} = {\tt GRAVID} \quad {\tt NG} = {\tt NONGRAVID} \mbox{ - WEIGHT(S)} \mbox{ NOT INCLUDED IN CALCULATION OF MEAN NA = NOT APPLICABLE}$

PGFWv4.11 07/16/2010

PAGE 5

| | GNANCY TATUS | DAY 0-6 | 6- 7 | 7-10 | 10-14 | 14-17 | 17-20 | 0-20 |
|-------|-----------------|--------------|---------|------|-------|-------|-------|-------------------------------|
| DAMS | FROM GF | ROUP 1: 0 MG | /KG/DAY | | | | | |
| 73710 | G | 42. | 48. | 47. | 55. | 53. | 51. | 47. SCHEDULED NECROPSY DAY 20 |
| 73704 | G | 56. | 43. | 45. | 59. | 57. | 53. | 55. SCHEDULED NECROPSY DAY 20 |
| 73694 | G | 48. | 62. | 45. | 43. | 47. | 43. | 46. SCHEDULED NECROPSY DAY 20 |
| 73713 | G | 54. | 44. | 53. | 50. | 57. | 43. | 50. SCHEDULED NECROPSY DAY 20 |
| 73720 | G | 57. | 52. | 38. | 37. | 59. | 57. | 48. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 51. | 50. | 46. | 49. | 55. | 49. | 49. |
| S.D. | | 6.3 | 7.7 | 5.4 | 8.9 | 4.8 | 6.2 | 3.6 |
| S.E. | | 2.8 | 3.4 | 2.4 | 4.0 | 2.1 | 2.8 | 1.6 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |

| | FATUS | DAY 0-6 | 6- 7 | 7-10 | 10-14 | 14-17 | 17-20 | 0-20 |
|--------|---------|-------------|----------|------|-------|-------|-------|-------------------------------|
| DAMS 1 | FROM GR | OUP 2: 30 M | G/KG/DAY | | | | | |
| 73702 | G | 52. | 74. | 58. | 54. | 50. | 57. | 54. SCHEDULED NECROPSY DAY 20 |
| 73697 | G | 43. | 58. | 56. | 56. | 51. | 51. | 51. SCHEDULED NECROPSY DAY 20 |
| 73706 | G | 55. | 60. | 54. | 63. | 58. | 60. | 58. SCHEDULED NECROPSY DAY 20 |
| 73709 | G | 63. | 58. | 60. | 57. | 61. | 55. | 59. SCHEDULED NECROPSY DAY 20 |
| 73719 | G | 57. | 15. | 53. | 53. | 66. | 55. | 55. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 54. | 53. | 56. | 57. | 57. | 56. | 55. |
| S.D. | | 7.3 | 22.3 | 2.9 | 3.9 | 6.8 | 3.3 | 3.2 |
| S.E. | | 3.3 | 10.0 | 1.3 | 1.7 | 3.0 | 1.5 | 1.4 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |

| | GNANCY TATUS | DAY 0-6 | 6- 7 | 7-10 | 10-14 | 14-17 | 17-20 | 0-20 |
|-------|-----------------|--------------|----------|---------|-------|-------|-------|-------------------------------|
| DAMS | FROM GRO | OUP 3: 100 M | G/KG/DAY | | | | | |
| 73696 | G | 56. | 46. | 55. | 54. | 56. | 40. | 51. SCHEDULED NECROPSY DAY 20 |
| 73699 | G | 56. | 55. | 40. | 45. | 52. | 51. | 48. SCHEDULED NECROPSY DAY 20 |
| 73714 | G | 41. | 64. | 53. | 47. | 47. | 46. | 47. SCHEDULED NECROPSY DAY 20 |
| 73705 | G | 60. | 54. | 56. | 53. | 57. | 53. | 55. SCHEDULED NECROPSY DAY 20 |
| 73700 | G | 51. | 38. | 50. | 45. | 57. | 47. | 50. SCHEDULED NECROPSY DAY 20 |
| MEAN | | 53. | 51. | 51. | 49. | 54. | 47. | 50. |
| S.D. | | 7.3 | 9.8 | 6.5 | 4.4 | 4.3 | 5.0 | 3.1 |
| S.E. | | 3.3 | 4.4 | 2.9 | 2.0 | 1.9 | 2.2 | 1.4 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 | 5 |

MEAN

S.D.

S.E.

N

PAGE 4

PREGNANCY STATUS DAY 0-6 6-7 7-10 10-14 14-17 17-20 0-20 ______ DAMS FROM GROUP 4: 300 MG/KG/DAY 45. 52. 53. 46. 48. SCHEDULED NECROPSY DAY 20 59. 49. 43. 49. 49. SCHEDULED NECROPSY DAY 20 56. 56. 57. 40. 52. SCHEDULED NECROPSY DAY 20 52. 40. 47. 40. 43. SCHEDULED NECROPSY DAY 20 52. 42. 67. 40. 45. SCHEDULED NECROPSY DAY 20 73708 G 50. 49. 61. 71. 44. 73712 G 47. 59. 56. 52. 52. 46. 41. 73698 G 73717 G 40. 43. 73693 G

53.

9.3

4.2

5

43.

4.2

1.9

5

47.

3.5

1.6

5

G = GRAVID NG = NONGRAVID - WEIGHT(S) NOT INCLUDED IN CALCULATION OF MEAN

53.

5.3

2.4

5

48.

6.7

3.0

5

54.

12.1

5.4

5

45.

4.2

1.9

5

SPONSOR: AMERICAN PETROLEUM

TABLE 118

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 5

INDIVIDUAL FOOD CONSUMPTION DURING GESTATION [G/KG/DAY]

| | REGNANCY STATUS | DAY 0-6 | 6- 7 | 7-10 | 10-14 | 14-17 | 17-20 | 0-20 |
|-------|--------------------|-------------|-----------|------|-------|-------|-------|-------------------------------|
| DAMS | FROM GR | OUP 5: 1000 | MG/KG/DAY | Y | | | | |
| 73718 | G G | 46. | 58. | 51. | 58. | 48. | 43. | 51. SCHEDULED NECROPSY DAY 20 |
| 73701 | . G | 21. | 35. | 74. | 44. | 69. | NA | NA SCHEDULED NECROPSY DAY 20 |
| 73721 | . G | 43. | 34. | 55. | 48. | 51. | 56. | 48. SCHEDULED NECROPSY DAY 20 |
| 73707 | 7 G | 26. | 57. | 64. | NA | 45. | 53. | 45. SCHEDULED NECROPSY DAY 20 |
| 73722 | 2 G | 40. | 43. | 42. | 48. | 55. | 47. | 47. SCHEDULED NECROPSY DAY 20 |
| MEAN | 1 | 35. | 45. | 57. | 50. | 54. | 50. | 48. |
| S.D. | | 11.0 | 11.6 | 12.3 | 6.0 | 9.4 | 5.9 | 2.5 |
| S.E. | | 4.9 | 5.2 | 5.5 | 3.0 | 4.2 | 2.9 | 1.3 |
| N | | 5 | 5 | 5 | 4 | 5 | 4 | 4 |
| | | | | | | | | |

 ${\tt G} = {\tt GRAVID} \quad {\tt NG} = {\tt NONGRAVID} \mbox{ - WEIGHT(S)} \mbox{ NOT INCLUDED IN CALCULATION OF MEAN NA = NOT APPLICABLE}$

PGFWv4.11 07/16/2010

TABLE I19 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SDONGOR-AMERICAN DEPTROLEUM INDIVIDUAL MATERNAL MACROSCOPIC FINDINGS

| DAMS FROM GROUP 1: 0 MG/KG/DAY | MATERNAL GROSS OBSERVATION | |
|---|---|--|
| 73710 73704 73694 73713 73720 | NO SIGNIFICANT CHANGES OBSERVED | |

PAGE 1

Page 317 of 360

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MATERNAL MACROSCOPIC FINDINGS

| DAMS FROM GROUP 2: 30 MG/KG/DAY | MATERNAL GROSS OBSERVATION |
|----------------------------------|--|
| 73702 73697 73706 73709 | NO SIGNIFICANT CHANGES OBSERVED NO SIGNIFICANT CHANGES OBSERVED NO SIGNIFICANT CHANGES OBSERVED NO SIGNIFICANT CHANGES OBSERVED |
| 73719 | NO SIGNIFICANT CHANGES OBSERVED |

PAGE 2

Page 318 of 360

73705

73700

TABLE I19 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM INDIVIDUAL MATERNAL MACROSCOPIC FINDINGS TABLE I19

DAMS FROM GROUP 3: 100 MG/KG/DAY MATERNAL GROSS OBSERVATION ______ 73696 NO SIGNIFICANT CHANGES OBSERVED 73699 NO SIGNIFICANT CHANGES OBSERVED 73714 NO SIGNIFICANT CHANGES OBSERVED

NO SIGNIFICANT CHANGES OBSERVED

NO SIGNIFICANT CHANGES OBSERVED

PAGE 3

Page 319 of 360

TABLE I19 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM INDIVIDUAL MATERNAL MACROSCOPIC FINDINGS TABLE I19

DAMS FROM GROUP 4: 300 MG/KG/DAY MATERNAL GROSS OBSERVATION ______ 73708 NO SIGNIFICANT CHANGES OBSERVED 73712 NO SIGNIFICANT CHANGES OBSERVED 73698 NO SIGNIFICANT CHANGES OBSERVED 73717 NO SIGNIFICANT CHANGES OBSERVED 73693 NO SIGNIFICANT CHANGES OBSERVED

PAGE 4

Page 320 of 360

| Page 321 | |
|----------|--|
| of 360 | |

| PROJECT NO.:WIL-402010F SPONSOR:AMERICAN PETROLEUM | TABLE 119 ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL MATERNAL MACROSCOPIC FINDINGS | PAGE | 5 |
|---|---|------|---|
| | | | |

| DAMS FROM GROUP 5: 1000 MG/KG/DAY | MATERNAL GROSS OBSERVATION |
|-----------------------------------|---|
| 73718 73701 | GRAVID AMMONIUM SULFIDE POSITIVE NO SIGNIFICANT CHANGES OBSERVED |
| 73721 | NO SIGNIFICANT CHANGES OBSERVED |
| 73707 | GRAVID AMMONIUM SULFIDE POSITIVE |
| 73722 | NO SIGNIFICANT CHANGES OBSERVED |

PMFGRDv4.16 07/15/2010

FEMALE GROUP: 0 MG/KG/DAY

| ANIMAL | PREGNANCY STATUS | FBW(G): | LIVER | KIDNEYS | THYMUS | OVARY/O D, LEFT | OVARY/OD , RIGHT |
|--------|---------------------|---------|-------|---------|---------|--------------------|---------------------|
| 73710 | G | 375. | 16.02 | 1.83 | 0.3019 | 0.0649 | 0.0843 |
| 73704 | G | 397. | 15.58 | 2.05 | 0.2845 | 0.0801 | 0.0899 |
| 73694 | G | 345. | 12.33 | 1.51 | 0.2581 | 0.0542 | 0.0693 |
| 73713 | G | 385. | 15.15 | 1.90 | 0.2039 | 0.0638 | 0.1032 |
| 73720 | G | 392. | 15.00 | 1.90 | 0.1920 | 0.0808 | 0.0730 |
| MEAN | | 379. | 14.82 | 1.84 | 0.2481 | 0.0688 | 0.0839 |
| S.D. | | 20.6 | 1.446 | 0.200 | 0.04853 | 0.01146 | 0.01361 |
| S.E. | | 9.2 | 0.646 | 0.090 | 0.02170 | 0.00512 | 0.00609 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

TABLE 120 ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL ORGAN WEIGHTS AND FINAL BODY WEIGHTS [G]

PAGE 2

FEMALE GROUP: 30 MG/KG/DAY

| ANIMAL | PREGNANCY STATUS | FBW(G): | LIVER | KIDNEYS | THYMUS | OVARY/O D, LEFT | OVARY/OD , RIGHT |
|---|---------------------|--------------------------------------|---|--------------------------------------|--|--|--|
| 73702 73697 73706 73709 73719 | G G G G | 414. 373. 353. 380. 406. | 16.35 15.83 13.61 15.05 16.60 | 2.07 1.94 1.65 2.11 2.07 | 0.2791 0.2734 0.2062 0.2463 0.3182 | 0.0669 0.0692 0.0550 0.0845 0.0877 | 0.0726 0.0744 0.0831 0.0809 0.0900 |
| MEAN S.D. S.E. N | | 385. 24.9 11.1 5 | 15.49 1.206 0.539 5 | 1.97 0.189 0.085 5 | 0.2646 0.04155 0.01858 5 | 0.0727 0.01345 0.00601 5 | 0.0802 0.00701 0.00313 5 |

FBW = FINAL BODY WEIGHT

TABLE I20 ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL ORGAN WEIGHTS AND FINAL BODY WEIGHTS [G]

PAGE 3

FEMALE GROUP: 100 MG/KG/DAY

| ANIMAL | PREGNANCY STATUS | FBW(G): | LIVER | KIDNEYS | THYMUS | OVARY/O D, LEFT | OVARY/OD , RIGHT |
|----------------|---------------------|--------------|----------------|--------------|------------------|--------------------|---------------------|
| 73696 73699 | G | 393. | 15.90 16.04 | 1.77 | 0.3802 0.2078 | 0.0756 0.0707 | 0.1070 0.0747 |
| 73699 | G G | 400. 344. | 16.04 | 1.81 1.67 | 0.2078 | 0.0707 | 0.0747 |
| 73705 | G | 424. | 17.73 | 2.18 | 0.2702 | 0.0847 | 0.1073 |
| 73700 | G | 401. | 16.88 | 1.90 | 0.3269 | 0.0767 | 0.0877 |
| MEAN | | 392. | 16.18 | 1.87 | 0.2890 | 0.0745 | 0.0891 |
| S.D. | | 29.5 | 1.266 | 0.194 | 0.06621 | 0.00733 | 0.01790 |
| S.E. | | 13.2 | 0.566 | 0.087 | 0.02961 | 0.00328 | 0.00800 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

G = GRAVID, NG = NONGRAVID - WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

Page 324 of 360

TABLE I20 ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL ORGAN WEIGHTS AND FINAL BODY WEIGHTS [G]

PAGE 4

FEMALE GROUP: 300 MG/KG/DAY

| ANIMAL | PREGNANCY STATUS | FBW(G): | LIVER | KIDNEYS | THYMUS | OVARY/O D, LEFT | OVARY/OD , RIGHT |
|--------|---------------------|---------|-------|---------|---------|--------------------|---------------------|
| 73708 | G | 418. | 17.80 | 2.24 | 0.3266 | 0.1068 | 0.0962 |
| 73712 | G | 349. | 15.18 | 1.87 | 0.1831 | 0.0593 | 0.0739 |
| 73698 | G | 363. | 15.97 | 1.93 | 0.2637 | 0.0882 | 0.0817 |
| 73717 | G | 370. | 16.09 | 1.80 | 0.2265 | 0.0920 | 0.0692 |
| 73693 | G | 366. | 14.99 | 1.96 | 0.2750 | 0.0738 | 0.0553 |
| MEAN | | 373. | 16.01 | 1.96 | 0.2550 | 0.0840 | 0.0753 |
| S.D. | | 26.3 | 1.111 | 0.168 | 0.05381 | 0.01814 | 0.01514 |
| S.E. | | 11.7 | 0.497 | 0.075 | 0.02407 | 0.00811 | 0.00677 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

G = GRAVID, NG = NONGRAVID - WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

SPONSOR: AMERICAN PETROLEUM

TABLE I20 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL ORGAN WEIGHTS AND FINAL BODY WEIGHTS [G]

FEMALE GROUP: 1000 MG/KG/DAY

| ANIMAL | PREGNANCY STATUS | FBW(G): | LIVER | KIDNEYS | THYMUS | OVARY/O D, LEFT | OVARY/OD , RIGHT |
|--------|---------------------|---------|-------|---------|---------|--------------------|---------------------|
| 73718 | G | 300. | 20.19 | 2.41 | 0.2600 | 0.0691 | 0.0667 |
| 73701 | G | 280. | 19.10 | 2.02 | 0.2081 | 0.0650 | 0.0432 |
| 73721 | G | 360. | 21.41 | 2.07 | 0.0890 | 0.0636 | 0.0736 |
| 73707 | G | 319. | 20.80 | 2.14 | 0.2789 | 0.0436 | 0.0685 |
| 73722 | G | 339. | 19.05 | 1.82 | 0.1360 | 0.0713 | 0.0574 |
| MEAN | | 320. | 20.11 | 2.09 | 0.1944 | 0.0625 | 0.0619 |
| S.D. | | 31.5 | 1.039 | 0.214 | 0.08086 | 0.01102 | 0.01197 |
| S.E. | | 14.1 | 0.465 | 0.096 | 0.03616 | 0.00493 | 0.00535 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

G = GRAVID, NG = NONGRAVID - WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

POFBWv4.21 07/16/2010

PAGE 5

Page 326 of 360

TABLE I21 ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL ORGAN WTS. RELATIVE TO FINAL BODY WTS. [G/100 G]

PAGE 1

FEMALE GROUP: 0 MG/KG/DAY

| ANIMAL | PREGNANCY STATUS | FBW(G): | LIVER | KIDNEYS | THYMUS | OVARY/O D, LEFT | OVARY/OD , RIGHT |
|----------------|---------------------|--------------|----------------|----------------|----------------|--------------------|---------------------|
| 73710 73704 | G G | 375. 397. | 4.272 3.924 | 0.488 0.516 | 0.081 0.072 | 0.017 0.020 | 0.022 0.023 |
| 73694 | G | 345. | 3.574 | 0.438 | 0.072 | 0.020 | 0.023 |
| 73713 73720 | G G | 385. 392. | 3.935 3.827 | 0.494 0.485 | 0.053 0.049 | 0.017 0.021 | 0.027 0.019 |
| 73720 | G | 392. | 3.027 | 0.465 | 0.049 | 0.021 | 0.019 |
| MEAN | | 379. | 3.910 | 0.480 | 0.066 | 0.018 | 0.022 |
| S.D. | | 20.6 | 0.2509 | 0.0287 | 0.0140 | 0.0022 | 0.0031 |
| S.E. | | 9.2 | 0.1122 | 0.0128 | 0.0062 | 0.0010 | 0.0014 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

G = GRAVID, NG = NONGRAVID - WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

PAGE 2

FEMALE GROUP: 30 MG/KG/DAY

| ANIMAL | PREGNANCY STATUS | FBW(G): | LIVER | KIDNEYS | THYMUS | OVARY/O D, LEFT | OVARY/OD , RIGHT |
|----------------|---------------------|---------|----------------|----------------|----------------|--------------------|---------------------|
| 73702 73697 | G | 414. | 3.949 | 0.500 | 0.067 0.073 | 0.016 | 0.018 0.020 |
| 73706 | G G | 353. | 4.244 3.856 | 0.520 0.467 | 0.073 | 0.019 0.016 | 0.020 |
| 73709 | G | 380. | 3.961 | 0.555 | 0.065 | 0.022 | 0.021 |
| 73719 | G | 406. | 4.089 | 0.510 | 0.078 | 0.022 | 0.022 |
| MEAN | | 385. | 4.020 | 0.510 | 0.068 | 0.019 | 0.021 |
| S.D. | | 24.9 | 0.1504 | 0.0319 | 0.0077 | 0.0030 | 0.0022 |
| S.E. | | 11.1 | 0.0673 | 0.0143 | 0.0034 | 0.0014 | 0.0010 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

G = GRAVID, NG = NONGRAVID - WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

Page 328 of 360

PAGE 3

FEMALE GROUP: 100 MG/KG/DAY

| ANIMAL | PREGNANCY | FBW(G): | LIVER | KIDNEYS | THYMUS | OVARY/O D, LEFT | OVARY/OD , RIGHT |
|--------|-----------|---------|--------|---------|--------|--------------------|---------------------|
| MILIME | DIATOD | | | | | ъ, шыг | , Righi |
| 73696 | G | 393. | 4.046 | 0.450 | 0.097 | 0.019 | 0.027 |
| 73699 | G | 400. | 4.010 | 0.452 | 0.052 | 0.018 | 0.019 |
| 73714 | G | 344. | 4.166 | 0.485 | 0.076 | 0.019 | 0.020 |
| 73705 | G | 424. | 4.182 | 0.514 | 0.064 | 0.020 | 0.025 |
| 73700 | | | 4.209 | 0.474 | 0.082 | 0.019 | 0.022 |
| MEAN | | 392. | 4.120 | 0.480 | 0.074 | 0.019 | 0.023 |
| S.D. | | 29.5 | 0.0887 | 0.0262 | 0.0171 | 0.0008 | 0.0034 |
| S.E. | | 13.2 | 0.0397 | 0.0117 | 0.0076 | 0.0004 | 0.0015 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

G = GRAVID, NG = NONGRAVID - WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

Page 329 of 360

PAGE 4

FEMALE GROUP: 300 MG/KG/DAY

| ANIMAL | PREGNANCY STATUS | FBW(G): | LIVER | KIDNEYS | THYMUS | OVARY/O D, LEFT | OVARY/OD , RIGHT |
|--------|---------------------|---------|--------|---------|--------|--------------------|---------------------|
| | | | | | | -, | |
| 73708 | G | 418. | 4.258 | 0.536 | 0.078 | 0.026 | 0.023 |
| 73712 | G | 349. | 4.350 | 0.536 | 0.052 | 0.017 | 0.021 |
| 73698 | G | 363. | 4.399 | 0.532 | 0.073 | 0.024 | 0.023 |
| 73717 | G | 370. | 4.349 | 0.486 | 0.061 | 0.025 | 0.019 |
| 73693 | | | 4.096 | 0.536 | 0.075 | 0.020 | 0.015 |
| MEAN | | 373. | 4.290 | 0.530 | 0.068 | 0.022 | 0.020 |
| S.D. | | 26.3 | 0.1202 | 0.0216 | 0.0108 | 0.0037 | 0.0033 |
| S.E. | | 11.7 | 0.0537 | 0.0097 | 0.0048 | 0.0016 | 0.0015 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

G = GRAVID, NG = NONGRAVID - WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

Page 330 of 360

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS

SPONSOR: AMERICAN PETROLEUM INDIVIDUAL ORGAN WTS. RELATIVE TO FINAL BODY WTS. [G/100 G]

FEMALE GROUP: 1000 MG/KG/DAY

| ANIMAL | PREGNANCY STATUS | FBW(G): | LIVER | KIDNEYS | THYMUS | OVARY/O D, LEFT | OVARY/OD , RIGHT |
|----------------|---------------------|---------|----------------|----------------|----------------|--------------------|---------------------|
| 73718 73701 | G | 300. | 6.730 6.821 | 0.803 0.721 | 0.087 0.074 | 0.023 | 0.022 0.015 |
| 73701 | G G | 360. | 5.947 | 0.721 | 0.074 | 0.023 0.018 | 0.015 |
| 73707 | G | 319. | 6.520 | 0.671 | 0.087 | 0.014 | 0.021 |
| 73722 | G | 339. | 5.619 | 0.537 | 0.040 | 0.021 | 0.017 |
| MEAN | | 320. | 6.330 | 0.660 | 0.063 | 0.020 | 0.019 |
| S.D. | | 31.5 | 0.5218 | 0.1082 | 0.0286 | 0.0041 | 0.0029 |
| S.E. | | 14.1 | 0.2334 | 0.0484 | 0.0128 | 0.0018 | 0.0013 |
| N | | 5 | 5 | 5 | 5 | 5 | 5 |

FBW = FINAL BODY WEIGHT

G = GRAVID, NG = NONGRAVID - WEIGHTS NOT INCLUDED IN CALCULATION OF THE MEAN

POFBWv4.21 07/16/2010

PAGE 5

TADIVIDONE TERM DATA IN SCHEDOLD MERCIOT

PAGE 1

| | | | VIA | BLE FE | TUSES | DEA | D FETU | ISES | EARLY | RESORE | PTIONS | LATE | RESORP | TIONS | IMPLAN | TATION | SITES | CORP | ORA LU | JTEA |
|-----------|-----|---------|--------------|---------------|-------|--------------|---------------|-------|--------------|---------------|--------|--------------|---------------|-------|--------------|---------------|-------|------|--------------|------------|
| DAM# | M | EX F | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | | LEFT HORN | RIGHT HORN | TOTAL | | RIGHT ARY | r TOTAI |
| 73710 | | 6 | | 7 | 11 | | | | 1 | 1 | 2 | | | | 5 | | 13 | | | 13 |
| 73704 | 9 | 6 | 7 | 8 | 15 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 7 | 8 | 15 | 7 | 8 | 19 |
| 73694 | 9 | 4 | 5 | 8 | 13 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 5 | 8 | 13 | 5 | 9 | 14 |
| 73713 | 8 | 8 | 4 | 12 | 16 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 4 | 12 | 16 | 12 | 15 | 27 |
| 73720 | 8 | 4 | 5 | 7 | 12 | 0 | 0 | 0 | 1 | 0 | 1 | 0 | 0 | 0 | 6 | 7 | 13 | 6 | 7 | 13 |
| TOTAL | 39 | 28 | 25 | 42 | 67 | 0 | 0 | 0 | 2 | 1 | 3 | 0 | 0 | 0 | 27 | 43 | 70 | 35 | 47 | 82 |
| MEAN | 7.8 | 5.6 | 5.0 | 8.4 | 13.4 | 0.0 | 0.0 | 0.0 | 0.4 | 0.2 | 0.6 | 0.0 | 0.0 | 0.0 | 5.4 | 8.6 | 14.0 | 7.0 | 9.4 | 16. |
| S.D. 1 | .64 | 1.67 | 1.22 | 2.07 | 2.07 | 0.00 | 0.00 | 0.00 | 0.55 | 0.45 | 0.89 | 0.00 | 0.00 | 0.00 | 1.14 | 1.95 | 1.41 | 2.92 | 3.21 | 5.98 |
| S.E. 0 | .73 | 0.75 | 0.55 | 0.93 | 0.93 | 0.00 | 0.00 | 0.00 | 0.24 | 0.20 | 0.40 | 0.00 | 0.00 | 0.00 | 0.51 | 0.87 | 0.63 | 1.30 | 1.44 | 2.68 |

PAGE 2

| | | | VIA | BLE FE | ETUSES | DEA | D FETU | JSES | EARLY | RESORP | TIONS | LATE | RESORF | PTIONS | IMPLAN | TATION | N SITES | CORP | ORA L | JTEA |
|--------|--------|---------|--------------|---------------|--------|--------------|---------------|-------|--------------|---------------|-------|--------------|---------------|--------|--------------|---------------|---------|------------|--------------|------------|
| DAM# | S M | EX F | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | TOTAL | LEFT OV | RIGH 'ARY | T TOTAL |
| 73702 | 3 | 12 | 7 | | 15 | | | 0 | 1 | | 1 | | | 0 | | | 16 | 10 | | 18 |
| 73697 | 7 | 7 | 7 | 7 | 14 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 7 | 7 | 14 | 7 | 8 | 15 |
| 3706 | 6 | 6 | 5 | 7 | 12 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 5 | 8 | 13 | 5 | 8 | 13 |
| 73709 | 5 | 9 | 6 | 8 | 14 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 6 | 8 | 14 | 6 | 8 | 14 |
| 73719 | 9 | 9 | 9 | 9 | 18 | 0 | 0 | 0 | 1 | 0 | 1 | 0 | 0 | 0 | 10 | 9 | 19 | 11 | 11 | 22 |
| TOTAL | 30 | 43 | 34 | 39 | 73 | 0 | 0 | 0 | 2 | 1 | 3 | 0 | 0 | 0 | 36 | 40 | 76 | 39 | 43 | 82 |
| MEAN (| 6.0 | 8.6 | 6.8 | 7.8 | 14.6 | 0.0 | 0.0 | 0.0 | 0.4 | 0.2 | 0.6 | 0.0 | 0.0 | 0.0 | 7.2 | 8.0 | 15.2 | 7.8 | 8.6 | 16.4 |
| S.D. 2 | .24 | 2.30 | 1.48 | 0.84 | 2.19 | 0.00 | 0.00 | 0.00 | 0.55 | 0.45 | 0.55 | 0.00 | 0.00 | 0.00 | 1.92 | 0.71 | 2.39 | 2.59 | 1.34 | 3.65 |
| S.E. 1 | .00 | 1.03 | 0.66 | 0.37 | 0.98 | 0.00 | 0.00 | 0.00 | 0.24 | 0.20 | 0.24 | 0.00 | 0.00 | 0.00 | 0.86 | 0.32 | 1.07 | 1.16 | 0.60 | 1.63 |

SPUNSUR: AMERICAN PETROLEUM INDIVIDUAL FETAL DATA AT SCHEDULED NECROPSI

PAGE 3

| DAMS | FROI | M GROU | JP 3: 1 | .00 MG/ | KG/DAY | | | | | | | | | | | | | | | |
|-----------|------|----------|--------------|---------------|--------|--------------|---------------|-------|--------------|---------------|-------|--------------|---------------|-------|--------------|---------------|---------|------------|--------------|-------|
| | | | VIA | BLE FE | TUSES | DEA | D FETU | SES | EARLY | RESORP | TIONS | LATE | RESORP | TIONS | IMPLAN | TATION | I SITES | CORP | ORA L | JTEA |
| DAM# | M | SEX F | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | TOTAL | LEFT OV | RIGH' ARY | TOTAL |
| 73696 | 3 | 12 | 5 | 10 | 15 | 0 | 0 | 0 | 0 | 1 | 1 | 0 | 0 | 0 | 5 | 11 | 16 | 5 | 11 | 16 |
| 73699 | 9 | 5 | 7 | 7 | 14 | 0 | 0 | 0 | 0 | 2 | 2 | 0 | 0 | 0 | 7 | 9 | 16 | 8 | 10 | 18 |
| 3714 | 8 | 4 | 6 | 6 | 12 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 6 | 6 | 12 | 6 | 9 | 15 |
| 3705 | 9 | 8 | 7 | 10 | 17 | 0 | 0 | 0 | 0 | 2 | 2 | 0 | 0 | 0 | 7 | 12 | 19 | 8 | 13 | 21 |
| 73700 | 8 | 8 | 7 | 9 | 16 | 0 | 0 | 0 | 1 | 0 | 1 | 0 | 0 | 0 | 8 | 9 | 17 | 8 | 9 | 17 |
| TOTAL | 37 | 37 | 32 | 42 | 74 | 0 | 0 | 0 | 1 | 5 | 6 | 0 | 0 | 0 | 33 | 47 | 80 | 35 | 52 | 87 |
| MEAN | 7.4 | 7.4 | 6.4 | 8.4 | 14.8 | 0.0 | 0.0 | 0.0 | 0.2 | 1.0 | 1.2 | 0.0 | 0.0 | 0.0 | 6.6 | 9.4 | 16.0 | 7.0 | 10.4 | 17.4 |
| S.D. 2 | 2.51 | 3.13 | 0.89 | 1.82 | 1.92 | 0.00 | 0.00 | 0.00 | 0.45 | 1.00 | 0.84 | 0.00 | 0.00 | 0.00 | 1.14 | 2.30 | 2.55 | 1.41 | 1.67 | 2.30 |
| S.E. 1 | 1.12 | 1.40 | 0.40 | 0.81 | 0.86 | 0.00 | 0.00 | 0.00 | 0.20 | 0.45 | 0.37 | 0.00 | 0.00 | 0.00 | 0.51 | 1.03 | 1.14 | 0.63 | 0.75 | 1.03 |

SPONSOR:AMERICAN PETROLEUM INDIVIDUAL FETAL DATA AT SCHEDULED NECROPSY

PAGE 4

| | | | VIA | BLE FE | TUSES | DEA | D FETU | ISES | EARLY | RESORF | TIONS | LATE | RESORF | TIONS | IMPLAN | TATION | N SITES | CORP | ORA L | JTEA |
|-----------|--------|---------|--------------|---------------|-------|--------------|---------------|-------|--------------|---------------|-------|--------------|---------------|-------|--------------|---------------|------------|------|-------|-------|
| DAM# | S M | EX F | LEFT HORN | RIGHT HORN | TOTAL | LEFT HORN | RIGHT HORN | r TOTAL | | RIGH' | TOTAL |
| 73708 | 10 | | 7 | 9 | 16 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 7 | 9 | 16 | 8 | 9 | 17 |
| 73712 | 6 | 4 | 7 | 3 | 10 | 0 | 0 | 0 | 1 | 0 | 1 | 0 | 0 | 0 | 8 | 3 | 11 | 8 | 4 | 12 |
| 73698 | 5 | 8 | 6 | 7 | 13 | 0 | 0 | 0 | 1 | 0 | 1 | 0 | 0 | 0 | 7 | 7 | 14 | 7 | 9 | 16 |
| 73717 | 9 | 6 | 9 | 6 | 15 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 9 | 6 | 15 | 9 | 6 | 15 |
| 73693 | 10 | 5 | 9 | 6 | 15 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 9 | 6 | 15 | 9 | 6 | 15 |
| TOTAL | 40 | 29 | 38 | 31 | 69 | 0 | 0 | 0 | 2 | 0 | 2 | 0 | 0 | 0 | 40 | 31 | 71 | 41 | 34 | 75 |
| MEAN | 8.0 | 5.8 | 7.6 | 6.2 | 13.8 | 0.0 | 0.0 | 0.0 | 0.4 | 0.0 | 0.4 | 0.0 | 0.0 | 0.0 | 8.0 | 6.2 | 14.2 | 8.2 | 6.8 | 15.0 |
| S.D. 2 | .35 | 1.48 | 1.34 | 2.17 | 2.39 | 0.00 | 0.00 | 0.00 | 0.55 | 0.00 | 0.55 | 0.00 | 0.00 | 0.00 | 1.00 | 2.17 | 1.92 | 0.84 | 2.17 | 1.87 |
| S.E. 1 | .05 | 0.66 | 0.60 | 0.97 | 1.07 | 0.00 | 0.00 | 0.00 | 0.24 | 0.00 | 0.24 | 0.00 | 0.00 | 0.00 | 0.45 | 0.97 | 0.86 | 0.37 | 0.97 | 0.84 |

N = 5

TABLE I22 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FETAL DATA AT SCHEDULED NECROPSY

DAMS FROM GROUP 5: 1000 MG/KG/DAY _____ VIABLE FETUSES DEAD FETUSES EARLY RESORPTIONS LATE RESORPTIONS IMPLANTATION SITES CORPORA LUTEA LEFT RIGHT LEFT RIGHT SEX LEFT RIGHT LEFT RIGHT LEFT RIGHT LEFT RIGHT DAM# M F HORN HORN TOTAL HORN TOTAL HORN HORN TOTAL HORN HORN TOTAL HORN TOTAL OVARY TOTAL 1 2 5 0 3 0 0 0 73718 0 0 0 0 2 5 0 0 10 16 4 5 8 12 4 5 0 0 73707 0 73722 6 8 5 14 TOTAL 13 13 13 13 19 39 MEAN 2.6 2.6 S.D. 3.58 3.71 3.97 3.71 7.16 0.89 0.00 0.89 1.22 1.79 2.59 0.00 0.00 3.00 3.03 4.87 2.19 2.51 3.39 S.E. 1.60 1.66 1.78 1.66 3.20 0.40 0.00 0.40 0.55 0.80 1.16 0.00 0.00 0.00 1.34 1.36 2.18 0.98 1.12 1.52

> PLRDv4.08 07/15/2010

PAGE 5

Page 336 of 360

TABLE I23 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM INDIVIDUAL FETAL DATA AT SCHEDULED NECROPSY [% PER LITTER]

PAGE 1

DAMS FROM GROUP 1: 0 MG/KG/DAY

| DAM ‡ | CORPORA # LUTEA | IMPLANTATION SITES | FET VIABLE | USES DEAD | RES EARLY | ORPTIC LATE | NS TOTAL | PRE- IMPLANTATION LOSS | POST- IMPLANTATION LOSS | MALES | FEMALES | |
|-------|--------------------|-----------------------|---------------|--------------|--------------|----------------|-------------|------------------------------|-------------------------------|-------|---------|--|
| | # | # | % | 8 | % | % | % | 용 | 8 | % | % | |
| 73710 | 13.0 | 13.0 | 84.6 | 0.0 | 15.4 | 0.0 | 15.4 | 0.0 | 15.4 | 45.5 | 54.5 | |
| 73704 | 15.0 | 15.0 | 100.0 | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 | 60.0 | 40.0 | |
| 73694 | 14.0 | 13.0 | 100.0 | 0.0 | 0.0 | 0.0 | 0.0 | 7.1 | 0.0 | 69.2 | 30.8 | |
| 73713 | 3 27.0 | 16.0 | 100.0 | 0.0 | 0.0 | 0.0 | 0.0 | 40.7 | 0.0 | 50.0 | 50.0 | |
| 73720 | 0 13.0 | 13.0 | 92.3 | 0.0 | 7.7 | 0.0 | 7.7 | 0.0 | 7.7 | 66.7 | 33.3 | |
| MEAN | 16.4 | 14.0 | 95.4 | 0.0 | 4.6 | 0.0 | 4.6 | 9.6 | 4.6 | 58.3 | 41.7 | |
| S.D. | 5.98 | 1.41 | 6.89 | 0.00 | 6.89 | 0.00 | 6.89 | 17.68 | 6.89 | 10.32 | 10.32 | |
| S.E. | 2.68 | 0.63 | 3.08 | 0.00 | 3.08 | 0.00 | 3.08 | 7.91 | 3.08 | 4.62 | 4.62 | |
| N | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | |

PAGE 2

DAMS FROM GROUP 2: 30 MG/KG/DAY

| DAM | CORPORA # LUTEA | IMPLANTATION SITES | FET VIABLE | USES DEAD | RES EARLY | ORPTIC LATE | NS TOTAL | PRE- IMPLANTATION LOSS | POST- IMPLANTATION LOSS | MALES | FEMALES | |
|------|--------------------|-----------------------|---------------|--------------|--------------|----------------|-------------|------------------------------|-------------------------------|-------|---------|--|
| | # | # | % | % | % | % | % | 8 | % | % | % | |
| 737 | 02 18.0 | 16.0 | 93.8 | 0.0 | 6.3 | 0.0 | 6.3 | 11.1 | 6.3 | 20.0 | 80.0 | |
| 736 | 97 15.0 | 14.0 | 100.0 | 0.0 | 0.0 | 0.0 | 0.0 | 6.7 | 0.0 | 50.0 | 50.0 | |
| 737 | 06 13.0 | 13.0 | 92.3 | 0.0 | 7.7 | 0.0 | 7.7 | 0.0 | 7.7 | 50.0 | 50.0 | |
| 737 | 09 14.0 | 14.0 | 100.0 | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 | 35.7 | 64.3 | |
| 737 | 19 22.0 | 19.0 | 94.7 | 0.0 | 5.3 | 0.0 | 5.3 | 13.6 | 5.3 | 50.0 | 50.0 | |
| MEAN | 16.4 | 15.2 | 96.2 | 0.0 | 3.9 | 0.0 | 3.9 | 6.3 | 3.9 | 41.1 | 58.9 | |
| S.D. | 3.65 | 2.39 | 3.61 | 0.00 | 3.63 | 0.00 | 3.63 | 6.24 | 3.63 | 13.34 | 13.34 | |
| S.E. | 1.63 | 1.07 | 1.61 | 0.00 | 1.62 | 0.00 | 1.62 | 2.79 | 1.62 | 5.97 | 5.97 | |
| N | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | |

TABLE I23 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM INDIVIDUAL FETAL DATA AT SCHEDULED NECROPSY [% PER LITTER]

PAGE 3

DAMS FROM GROUP 3: 100 MG/KG/DAY

| DAM ‡ | CORPORA LUTEA | IMPLANTATION SITES | FET VIABLE | USES DEAD | RES EARLY | ORPTIC LATE | NS TOTAL | PRE- IMPLANTATION LOSS | POST- IMPLANTATION LOSS | MALES | FEMALES | |
|-------|------------------|-----------------------|---------------|--------------|--------------|----------------|-------------|------------------------------|-------------------------------|-------|---------|--|
| | # | # | % | 8 | % | % | % | 8 | 8 | % | % | |
| 73696 | 16.0 | 16.0 | 93.8 | 0.0 | 6.3 | 0.0 | 6.3 | 0.0 | 6.3 | 20.0 | 80.0 | |
| 73699 | 9 18.0 | 16.0 | 87.5 | 0.0 | 12.5 | 0.0 | 12.5 | 11.1 | 12.5 | 64.3 | 35.7 | |
| 73714 | 15.0 | 12.0 | 100.0 | 0.0 | 0.0 | 0.0 | 0.0 | 20.0 | 0.0 | 66.7 | 33.3 | |
| 73705 | 5 21.0 | 19.0 | 89.5 | 0.0 | 10.5 | 0.0 | 10.5 | 9.5 | 10.5 | 52.9 | 47.1 | |
| 73700 | 17.0 | 17.0 | 94.1 | 0.0 | 5.9 | 0.0 | 5.9 | 0.0 | 5.9 | 50.0 | 50.0 | |
| MEAN | 17.4 | 16.0 | 93.0 | 0.0 | 7.0 | 0.0 | 7.0 | 8.1 | 7.0 | 50.8 | 49.2 | |
| S.D. | 2.30 | 2.55 | 4.83 | 0.00 | 4.83 | 0.00 | 4.83 | 8.42 | 4.83 | 18.62 | 18.62 | |
| S.E. | 1.03 | 1.14 | 2.16 | 0.00 | 2.16 | 0.00 | 2.16 | 3.77 | 2.16 | 8.33 | 8.33 | |
| N | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | |

TABLE I23 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM INDIVIDUAL FETAL DATA AT SCHEDULED NECROPSY [% PER LITTER]

PAGE 4

DAMS FROM GROUP 4: 300 MG/KG/DAY

| DAM ; | CORPORA # LUTEA | IMPLANTATION SITES | FET VIABLE | USES DEAD | | ORPTIC LATE | NS TOTAL | PRE- IMPLANTATION LOSS | POST- IMPLANTATION LOSS | MALES | FEMALES | |
|-------|--------------------|-----------------------|---------------|--------------|------|----------------|-------------|------------------------------|-------------------------------|-------|---------|--|
| | # | # | % | % | % | % | % | 8 | 웅 | % | % | |
| 73708 | 8 17.0 | 16.0 | 100.0 | 0.0 | 0.0 | 0.0 | 0.0 | 5.9 | 0.0 | 62.5 | 37.5 | |
| 73712 | 2 12.0 | 11.0 | 90.9 | 0.0 | 9.1 | 0.0 | 9.1 | 8.3 | 9.1 | 60.0 | 40.0 | |
| 73698 | 8 16.0 | 14.0 | 92.9 | 0.0 | 7.1 | 0.0 | 7.1 | 12.5 | 7.1 | 38.5 | 61.5 | |
| 7371 | 7 15.0 | 15.0 | 100.0 | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 | 60.0 | 40.0 | |
| 73693 | 3 15.0 | 15.0 | 100.0 | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 | 0.0 | 66.7 | 33.3 | |
| MEAN | 15.0 | 14.2 | 96.8 | 0.0 | 3.2 | 0.0 | 3.2 | 5.3 | 3.2 | 57.5 | 42.5 | |
| S.D. | 1.87 | 1.92 | 4.49 | 0.00 | 4.49 | 0.00 | 4.49 | 5.42 | 4.49 | 11.00 | 11.00 | |
| S.E. | 0.84 | 0.86 | 2.01 | 0.00 | 2.01 | 0.00 | 2.01 | 2.42 | 2.01 | 4.92 | 4.92 | |
| N | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | |

TABLE 123

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS
SPONSOR:AMERICAN PETROLEUM INDIVIDUAL FETAL DATA AT SCHEDULED NECROPSY [% PER LITTER]

| DAMS FROM GROUP 5: 1000 MG/KG/DAY | | |
|-----------------------------------|------|-------|
| | DDE- | P09T- |

| DAM # | ORPORA LUTEA | IMPLANTATION SITES | FET VIABLE | USES DEAD | RES EARLY | ORPTIC LATE | NS TOTAL | IMPLANTATION LOSS | IMPLANTATION LOSS | MALES | FEMALES | |
|---|------------------------------------|-----------------------------------|-------------------------------------|-----------------------------|--------------------------------------|--------------------------|--------------------------------------|------------------------------------|---------------------------------------|-----------------------------------|-----------------------------------|--|
| | # | # | % | % | % | % | % | * | 8 | % | % | |
| 73718 73701 73721 73707 73722 | 16.0 9.0 13.0 8.0 14.0 | 5.0 3.0 12.0 5.0 14.0 | 0.0 0.0 100.0 0.0 100.0 | 0.0 66.7 0.0 0.0 | 100.0 33.3 0.0 100.0 0.0 | 0.0 0.0 0.0 0.0 | 100.0 33.3 0.0 100.0 0.0 | 68.8 66.7 7.7 37.5 0.0 | 100.0 100.0 0.0 100.0 0.0 | 0.0 0.0 58.3 0.0 42.9 | 0.0 0.0 41.7 0.0 57.1 | |
| MEAN S.D. S.E. N | 12.0 3.39 1.52 5 | 7.8 4.87 2.18 | 40.0 54.77 24.49 | 13.3 29.83 13.34 5 | 46.7 50.55 22.61 | 0.0 0.00 0.00 5 | 46.7 50.55 22.61 | 36.1 32.08 14.35 5 | 60.0 54.77 24.49 5 | 50.6 10.94 7.74 2 | 49.4 10.94 7.74 2 | |

PILPv4.02 07/15/2010

PAGE 5

S.E. 0.08

PAGE 1

| FET | US # | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | 11 | 12 | 13 | 14 | 15 | 16 | 17 | 18 | 19 | 20 | 21 | 22 | 23 |
|-------|------|------|-------|-----|-------|------|------|-----|-----|-----|-----|-----|-----|-----|-----|-----|-----|----|----|----|----|----|----|----|
| DAM # | MEAN | GROU | IP 1: | 0 | MG/KG | /DAY | | | | | | | | | | | | | | | | | | |
| | 3.9 | | | | | | | | | | | | | | | | | | | | | | | |
| 73704 | 3.7 | | | | | | | | | | | | | | 3.6 | 3.6 | | | | | | | | |
| 73694 | | 3.7 | | | | | | | | | | | | | | | | | | | | | | |
| 73713 | 3.5 | 3.3 | 3.6 | 3.7 | 3.8/ | 3.7 | 3.4 | 3.7 | 3.4 | 3.8 | 3.6 | 3.5 | 3.5 | 3.4 | 3.5 | 3.4 | 2.7 | | | | | | | |
| 73720 | 3.9 | 3.6 | 3.9 | 4.1 | E | 3.6 | 4.1/ | 4.3 | 3.9 | 4.0 | 4.1 | 3.8 | 3.5 | 4.0 | | | | | | | | | | |
| MEAN | 3.8 | | | | | | | | | | | | | | | | | | | | | | | |
| S.D. | 0.18 | | | | | | | | | | | | | | | | | | | | | | | |

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FETAL WEIGHTS [G]

| - | FET | US # | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | 11 | 12 | 13 | 14 | 15 | 16 | 17 | 18 | 19 | 20 | 21 | 22 | 23 | |
|---|---|---------------------------------|------|-------------------|-------------------|-------------------|--------------------|--------------------|--------------------|-----------------|-------------------|-------------------|-------------------|-------------------|-------------------|------------|----|----|-----|-----|-----|----|----|----|----|--|
| _ | DAM # | MEAN | GROU | IP 2: | 30 | MG/F | KG/DAY | | | | | | | | | | | | | | | | | | | |
| | 73702 73697 73706 73709 73719 | 3.4 3.8 3.8 3.9 3.3 | 4.0 | 3.9 3.3 3.9 | 3.7 3.5 4.0 | 4.0 4.0 4.0 | 3.8 4.0/ 4.1 | 3.9 4.0 3.7/ | 1.8/ 3.5 3.7 | 3.6 E 3.8 | 4.0 4.2 3.8 | 3.8 3.7 4.0 | 4.2 4.0 4.0 | 4.2 3.9 3.9 | 3.9 3.5 4.0 | 4.2 3.6 | | | 3.2 | 3.7 | 3.2 | | | | | |
| | MEAN S.D. S.E. N | 0.27 | | | | | | | | | | | | | | | | | | | | | | | | |

PAGE 2

______ FETUS # 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23

PAGE 3

DAM # MEAN GROUP 3: 100 MG/KG/DAY _____ 73696 3.9 4.0 4.0 3.4 3.8 4.2/4.3 4.0 3.3 4.1 E 3.7 3.7 3.9 3.6 4.0 3.9 73699 4.1 4.0 4.2 4.1 4.1 4.1 4.1 4.1/4.3 4.1 E E 4.2 4.1 4.2 3.7 4.1 73714 3.5 3.3 3.3 3.3 3.7 3.8 3.6 3.4 3.5 3.8 3.2 3.6 3.4 73700 3.5 3.8 3.4 3.5 3.8 E 3.5/3.5 3.5 3.5 3.6 3.4 3.7 3.2 3.6 3.6 3.6 3.7 3.7 3.8 3.9 4.0/ E 3.4 3.8 3.5 E 3.7 3.7 3.6 3.6 3.6 3.4 73700 3.5 3.8 3.4 3.5 3.8 E 3.5/3.5 3.5 3.5 3.1 3.3 3.4 3.4 3.7 3.2 3.3

MEAN 3.7 S.D. 0.27 S.E. 0.12 N

Page 344 of 360

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM PAGE 4

| _ | | | | | | | | | | | | | | | | | | | | | | | : | | |
|---|----------------|---------------------------------|-------------------|-------------------|-----------------|-----------------|-------------------|-------------------|-----|--------------------|--------------------|-------------------|-------------------|-----|-----|-----|-----|-----|----|----|----|----|----|----|----|
| | FET | US # | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | 11 | 12 | 13 | 14 | 15 | 16 | 17 | 18 | 19 | 20 | 21 | 22 | 23 |
| _ | DAM # | MEAN | GROU | P 4: | 300 | MG/F | (G/DAY | ζ | | | | | | | | | | | | | | | | | |
| _ | 73712 73698 | 3.6 4.1 3.9 3.3 3.9 | 3.2 4.0 3.3 | 4.2 3.8 3.4 | E 3.8 3.2 | 4.2 E 3.3 | 4.2 3.6 3.4 | 3.9 4.1 3.2 | 4.3 | 3.9/ 4.2 3.4 | 4.1 4.2 3.4/ | 4.6 3.2 3.2 | 4.1 3.6 3.4 | 4.2 | 4.0 | 3.7 | 3.2 | 3.3 | | | | | | | |
| | | 3.8 0.31 0.14 | | | | | | | | | | | | | | | | | | | | | | | |

______ FETUS # 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23

DAM # MEAN GROUP 5: 1000 MG/KG/DAY _____ MEAN 3.5 S.D. 0.21 S.E. 0.15 N

E = EARLY RESORPTION L = LATE RESORPTION D = DEAD FETUS '/' DENOTES POSITION OF CERVIX

PFWTv4.15 07/15/2010

PAGE 5

Page 346 of 360

TABLE 125
PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS
SPONSOR:AMERICAN PETROLEUM INDIVIDUAL FETAL EXTERNAL FINDINGS

| BIONDOK, AMERICAN TETROBLOM | - | INDIVIDOAL | I LIAL BALBANAL LINDINGS |
|--------------------------------|-------------|------------|---|
| DAMS FROM GROUP 1: 0 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
| 73710 | 1 | M | NO REMARKABLE OBSERVATIONS |
| | 2 | F | NO REMARKABLE OBSERVATIONS |
| | 2 3 4 | | EARLY RESORPTION |
| | | F | NO REMARKABLE OBSERVATIONS |
| | 5 6 | F | NO REMARKABLE OBSERVATIONS |
| | | M | NO REMARKABLE OBSERVATIONS |
| | 7 | F | NO REMARKABLE OBSERVATIONS |
| | 8 | M | NO REMARKABLE OBSERVATIONS |
| | 9 | F | NO REMARKABLE OBSERVATIONS |
| | 10 | | EARLY RESORPTION |
| | 11 | M | NO REMARKABLE OBSERVATIONS |
| | 12 | F | NO REMARKABLE OBSERVATIONS |
| | 13 | M | NO REMARKABLE OBSERVATIONS |
| 73704 | 1 | F | NO REMARKABLE OBSERVATIONS |
| | 2 | M | NO REMARKABLE OBSERVATIONS |
| | 3 | M | NO REMARKABLE OBSERVATIONS |
| | 4 | F | NO REMARKABLE OBSERVATIONS |
| | 5 | M | NO REMARKABLE OBSERVATIONS |
| | 6 | M | NO REMARKABLE OBSERVATIONS |
| | 7 | M | NO REMARKABLE OBSERVATIONS |
| | 8 | M | NO REMARKABLE OBSERVATIONS |
| | 9 | M | NO REMARKABLE OBSERVATIONS |
| | 10 11 | F F | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 12 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 13 | F | NO REMARKABLE OBSERVATIONS |
| | 14 | F | NO REMARKABLE OBSERVATIONS |
| | 15 | M | NO REMARKABLE OBSERVATIONS |
| 73694 | 1 | F | NO REMARKABLE OBSERVATIONS |
| 73094 | 2 | M | NO REMARKABLE OBSERVATIONS |
| | 3 | M | NO REMARKABLE OBSERVATIONS |
| | 4 | F | NO REMARKABLE OBSERVATIONS |
| | 5 | M | NO REMARKABLE OBSERVATIONS |
| | - | | |

PAGE 1

Page 348 of 360

PAGE 2

| | DAMS FROM GROUP 1: 0 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|----------|--------------------------------|--------|--------|--|
| | 73694 | 6 | М | NO REMARKABLE OBSERVATIONS |
| | | 7 | M | NO REMARKABLE OBSERVATIONS |
| | | 8 | F | NO REMARKABLE OBSERVATIONS |
| | | 8 9 | M | NO REMARKABLE OBSERVATIONS |
| | | 10 | M | NO REMARKABLE OBSERVATIONS |
| | | 11 | M | NO REMARKABLE OBSERVATIONS |
| | | 12 | M | NO REMARKABLE OBSERVATIONS |
| | | 13 | F | NO REMARKABLE OBSERVATIONS |
| | 73713 | 1 2 | F | NO REMARKABLE OBSERVATIONS |
| | | 2 | F | NO REMARKABLE OBSERVATIONS |
| | | 3 | M | NO REMARKABLE OBSERVATIONS |
| | | 4 | F | NO REMARKABLE OBSERVATIONS |
| | | 5 | M | NO REMARKABLE OBSERVATIONS |
| | | 6 | M | NO REMARKABLE OBSERVATIONS |
| | | 7 | M | NO REMARKABLE OBSERVATIONS |
| 1 | | 8 | M | NO REMARKABLE OBSERVATIONS |
| | | 9 | F | NO REMARKABLE OBSERVATIONS |
| | | 10 | F | NO REMARKABLE OBSERVATIONS |
|) | | 11 | F | NO REMARKABLE OBSERVATIONS |
| _ | | 12 | M | NO REMARKABLE OBSERVATIONS |
|) | | 13 | M | NO REMARKABLE OBSERVATIONS |
| | | 14 | F | NO REMARKABLE OBSERVATIONS |
| , | | 15 | M | NO REMARKABLE OBSERVATIONS |
| ` | 73720 | 16 | F | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| <u> </u> | 13120 | 1 | F | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | 3 | M M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | 3 1 | IvI | EARLY RESORPTION |
| | | 5 | F | NO REMARKABLE OBSERVATIONS |
| | | 5 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | 7 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | 8 | M | NO REMARKABLE OBSERVATIONS |
| | | 9 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | , | 1.1 | NO REPRESENTATIONS |

TABLE I25

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 3

SPONSOR:AMERICAN PETROLEUM INDIVIDUAL FETAL EXTERNAL FINDINGS

| 73720 10 M NO REMARKABLE OBSERVATIONS 11 F NO REMARKABLE OBSERVATIONS 12 F NO REMARKABLE OBSERVATIONS 13 M NO REMARKABLE OBSERVATIONS | DAMS FROM GROUP 1: 0 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|---|--------------------------------|----------|--------|----------------------------|
| 15 II III III ODDINIII ODDINIII ONO | 73720 | 11 12 | F F | NO REMARKABLE OBSERVATIONS |

TABLE 125
PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FETAL EXTERNAL FINDINGS SPONSOR: AMERICAN PETROLEUM

| DAMS FROM GROUP 2: 30 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|---------------------------------|----------|--------|---|
| 73702 | 1 | F | NO REMARKABLE OBSERVATIONS |
| | 2 | F | NO REMARKABLE OBSERVATIONS |
| | 3 4 | | EARLY RESORPTION |
| | 4 | F | NO REMARKABLE OBSERVATIONS |
| | 5 | F | NO REMARKABLE OBSERVATIONS |
| | 6 | F | NO REMARKABLE OBSERVATIONS |
| | 7 | F | NO REMARKABLE OBSERVATIONS |
| | 8 | F | NO REMARKABLE OBSERVATIONS |
| | 9 | F | NO REMARKABLE OBSERVATIONS |
| | 10 | M | NO REMARKABLE OBSERVATIONS |
| | 11 | F | NO REMARKABLE OBSERVATIONS |
| | 12 | F | NO REMARKABLE OBSERVATIONS |
| | 13 | M | NO REMARKABLE OBSERVATIONS |
| | 14 | M | NO REMARKABLE OBSERVATIONS |
| | 15 | F | NO REMARKABLE OBSERVATIONS |
| | 16 | F | NO REMARKABLE OBSERVATIONS |
| 73697 | 1 | F | NO REMARKABLE OBSERVATIONS |
| | 2 | M | NO REMARKABLE OBSERVATIONS |
| | 3 | F | NO REMARKABLE OBSERVATIONS |
| | 4 | F | NO REMARKABLE OBSERVATIONS |
| | 5 | F | NO REMARKABLE OBSERVATIONS |
| | 6 | F | NO REMARKABLE OBSERVATIONS |
| | 7 | M | M OMPHALOCELE |
| | | | SEVERAL LOOPS OF INTESTINE PROTRUDE THROUGH AN OPENING IN THE |
| | • | _ | UMBILICUS, REMNANTS OF A MEMBRANOUS SAC |
| | 8 | F | NO REMARKABLE OBSERVATIONS |
| | 9 | M | NO REMARKABLE OBSERVATIONS |
| | 10 | F | NO REMARKABLE OBSERVATIONS |
| | 11 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 12 13 | M | |
| | | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| 73706 | 14 1 | M M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |

OBSERVATION CODE: M = MALFORMATION V = VARIATION

Page 350 of 360

Page 351 of 360

PAGE 5

| DAMS FROM GROUP 2: 30 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|---------------------------------|--------|-----|----------------------------|
| 73706 | 2 | F | NO REMARKABLE OBSERVATIONS |
| | 3 | F | NO REMARKABLE OBSERVATIONS |
| | 4 | M | NO REMARKABLE OBSERVATIONS |
| | 5 | F | NO REMARKABLE OBSERVATIONS |
| | 6 | M | NO REMARKABLE OBSERVATIONS |
| | 7 | M | NO REMARKABLE OBSERVATIONS |
| | 8 | | EARLY RESORPTION |
| | 9 | M | NO REMARKABLE OBSERVATIONS |
| | 10 | F | NO REMARKABLE OBSERVATIONS |
| | 11 | M | NO REMARKABLE OBSERVATIONS |
| | 12 | F | NO REMARKABLE OBSERVATIONS |
| | 13 | F | NO REMARKABLE OBSERVATIONS |
| 73709 | 1 | M | NO REMARKABLE OBSERVATIONS |
| | 2 | M | NO REMARKABLE OBSERVATIONS |
| | 3 | M | NO REMARKABLE OBSERVATIONS |
| | 4 | F | NO REMARKABLE OBSERVATIONS |
| | 5 | M | NO REMARKABLE OBSERVATIONS |
| | 6 | F | NO REMARKABLE OBSERVATIONS |
| | 7 | F | NO REMARKABLE OBSERVATIONS |
| | 8 | F | NO REMARKABLE OBSERVATIONS |
| | 9 | F | NO REMARKABLE OBSERVATIONS |
| | 10 | F | NO REMARKABLE OBSERVATIONS |
| | 11 | F | NO REMARKABLE OBSERVATIONS |
| | 12 | F | NO REMARKABLE OBSERVATIONS |
| | 13 | M | NO REMARKABLE OBSERVATIONS |
| | 14 | F | NO REMARKABLE OBSERVATIONS |
| 73719 | 1 | M | NO REMARKABLE OBSERVATIONS |
| | 2 | | EARLY RESORPTION |
| | 3 | M | NO REMARKABLE OBSERVATIONS |
| | 4 | F | NO REMARKABLE OBSERVATIONS |
| | 5 | F | NO REMARKABLE OBSERVATIONS |
| | 6 | M | NO REMARKABLE OBSERVATIONS |
| | 7 | F | NO REMARKABLE OBSERVATIONS |
| | | | |

TABLE I25 PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FETAL EXTERNAL FINDINGS

PAGE 6

______ DAMS FROM GROUP 2: 30 MG/KG/DAY FETUS# SEX FETAL GROSS OBSERVATION ______ 8 M NO REMARKABLE OBSERVATIONS
9 F NO REMARKABLE OBSERVATIONS
10 F NO REMARKABLE OBSERVATIONS
11 M NO REMARKABLE OBSERVATIONS
12 M NO REMARKABLE OBSERVATIONS
13 F NO REMARKABLE OBSERVATIONS
14 F NO REMARKABLE OBSERVATIONS
15 F NO REMARKABLE OBSERVATIONS
16 M NO REMARKABLE OBSERVATIONS
16 M NO REMARKABLE OBSERVATIONS
17 F NO REMARKABLE OBSERVATIONS
18 M NO REMARKABLE OBSERVATIONS 73719 M NO REMARKABLE OBSERVATIONS 18 M NO REMARKABLE OBSERVATIONS 19

Page 353 of 360

| - | DAMS FROM GROUP 3: 100 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|--------|----------------------------------|-----------------------|--------|---|
| - | 73696 | 1 | F | NO REMARKABLE OBSERVATIONS |
| | 73030 | 2 | F | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | 2 3 4 5 6 | F | NO REMARKABLE OBSERVATIONS |
| | | 4 | F | NO REMARKABLE OBSERVATIONS |
| | | 5 | M | NO REMARKABLE OBSERVATIONS |
| | | 6 | F | NO REMARKABLE OBSERVATIONS |
| | | 7 | M | NO REMARKABLE OBSERVATIONS |
| | | 8 | F | NO REMARKABLE OBSERVATIONS |
| | | 9 | M | NO REMARKABLE OBSERVATIONS |
| | | 10 | | EARLY RESORPTION |
| | | 11 | F | NO REMARKABLE OBSERVATIONS |
| | | 12 | F | NO REMARKABLE OBSERVATIONS |
| | | 13 | F | NO REMARKABLE OBSERVATIONS |
| | | 14 | F | NO REMARKABLE OBSERVATIONS |
| | | 15 | F | NO REMARKABLE OBSERVATIONS |
| - | | 16 | F | NO REMARKABLE OBSERVATIONS |
| | 73699 | 1 2 | F | NO REMARKABLE OBSERVATIONS |
| | | | M | NO REMARKABLE OBSERVATIONS |
| , | | 3 | F | NO REMARKABLE OBSERVATIONS |
| , 1 | | 4 5 6 | F | NO REMARKABLE OBSERVATIONS |
| ذ | | 5 | M | NO REMARKABLE OBSERVATIONS |
| | | | M | NO REMARKABLE OBSERVATIONS |
| Ś | | 7 | F | NO REMARKABLE OBSERVATIONS |
| , | | 8 | M | NO REMARKABLE OBSERVATIONS |
|) | | 9 | M | NO REMARKABLE OBSERVATIONS |
| | | 10 | | EARLY RESORPTION |
| | | 11 | 3.6 | EARLY RESORPTION |
| | | 12 13 | M | NO REMARKABLE OBSERVATIONS |
| | | 14 | F M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | 15 | M M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | 16 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 73714 | 1 | M M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 13114 | 1 | 141 | NO REPARRADLE ODSERVATIONS |

TABLE I25

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS PAGE 8

SPONSOR:AMERICAN PETROLEUM INDIVIDUAL FETAL EXTERNAL FINDINGS

| DAMS FROM GROUP 3: 100 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|----------------------------------|--------|-----|----------------------------|
| | 2 | F | NO REMARKABLE OBSERVATIONS |
| | 3 | F | NO REMARKABLE OBSERVATIONS |
| | 4 | M | NO REMARKABLE OBSERVATIONS |
| | 5 | M | NO REMARKABLE OBSERVATIONS |
| | 6 | M | NO REMARKABLE OBSERVATIONS |
| | 7 | F | NO REMARKABLE OBSERVATIONS |
| | 8 | M | NO REMARKABLE OBSERVATIONS |
| | 9 | M | NO REMARKABLE OBSERVATIONS |
| | 10 | F | NO REMARKABLE OBSERVATIONS |
| | 11 | M | NO REMARKABLE OBSERVATIONS |
| | 12 | M | NO REMARKABLE OBSERVATIONS |
| 73705 | 1 | M | NO REMARKABLE OBSERVATIONS |
| | 2 | M | NO REMARKABLE OBSERVATIONS |
| | 3 | M | NO REMARKABLE OBSERVATIONS |
| | 4 | F | NO REMARKABLE OBSERVATIONS |
| | 5 | M | NO REMARKABLE OBSERVATIONS |
| | 6 | M | NO REMARKABLE OBSERVATIONS |
| | 7 | M | NO REMARKABLE OBSERVATIONS |
| | 8 | | EARLY RESORPTION |
| | 9 | F | NO REMARKABLE OBSERVATIONS |
| | 10 | F | NO REMARKABLE OBSERVATIONS |
| | 11 | F | NO REMARKABLE OBSERVATIONS |
| | 12 | | EARLY RESORPTION |
| | 13 | F | NO REMARKABLE OBSERVATIONS |
| | 14 | M | NO REMARKABLE OBSERVATIONS |
| | 15 | F | NO REMARKABLE OBSERVATIONS |
| | 16 | M | NO REMARKABLE OBSERVATIONS |
| | 17 | F | NO REMARKABLE OBSERVATIONS |
| | 18 | M | NO REMARKABLE OBSERVATIONS |
| | 19 | F | NO REMARKABLE OBSERVATIONS |
| 73700 | 1 | M | NO REMARKABLE OBSERVATIONS |
| | 2 | F | NO REMARKABLE OBSERVATIONS |
| | 3 | F | NO REMARKABLE OBSERVATIONS |

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS INDIVIDUAL FETAL EXTERNAL FINDINGS

PAGE 9

| DAMS FROM GROUP 3: 100 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|----------------------------------|--------|-----|----------------------------|
| 73700 | 4 | М | NO REMARKABLE OBSERVATIONS |
| | 5 | M | NO REMARKABLE OBSERVATIONS |
| | 6 | M | NO REMARKABLE OBSERVATIONS |
| | 7 | | EARLY RESORPTION |
| | 8 | F | NO REMARKABLE OBSERVATIONS |
| | 9 | M | NO REMARKABLE OBSERVATIONS |
| | 10 | F | NO REMARKABLE OBSERVATIONS |
| | 11 | M | NO REMARKABLE OBSERVATIONS |
| | 12 | F | NO REMARKABLE OBSERVATIONS |
| | 13 | F | NO REMARKABLE OBSERVATIONS |
| | 14 | F | NO REMARKABLE OBSERVATIONS |
| | 15 | M | NO REMARKABLE OBSERVATIONS |
| | 16 | F | NO REMARKABLE OBSERVATIONS |
| | 17 | M | NO REMARKABLE OBSERVATIONS |

Page 356 of 360

PAGE 10

| | DAMS FROM GROUP 4: 300 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|----------|----------------------------------|----------------------------|--------|---|
| | 73708 | 1 | М | NO REMARKABLE OBSERVATIONS |
| | | 2 | M | NO REMARKABLE OBSERVATIONS |
| | | 3 | F | NO REMARKABLE OBSERVATIONS |
| | | 4 | F | NO REMARKABLE OBSERVATIONS |
| | | 5 | M | NO REMARKABLE OBSERVATIONS |
| | | 2 3 4 5 6 7 | F | NO REMARKABLE OBSERVATIONS |
| | | | M | NO REMARKABLE OBSERVATIONS |
| | | 8 | M | NO REMARKABLE OBSERVATIONS |
| | | 9 | M | NO REMARKABLE OBSERVATIONS |
| | | 10 | M | NO REMARKABLE OBSERVATIONS |
| | | 11 | F | NO REMARKABLE OBSERVATIONS |
| | | 12 | F | NO REMARKABLE OBSERVATIONS |
| | | 13 | M | NO REMARKABLE OBSERVATIONS |
| | | 14 | M | NO REMARKABLE OBSERVATIONS |
| | | 15 | M | NO REMARKABLE OBSERVATIONS |
| j | | 16 | F | NO REMARKABLE OBSERVATIONS |
| | 73712 | 1 2 | F | NO REMARKABLE OBSERVATIONS |
| | | | F | NO REMARKABLE OBSERVATIONS |
| , | | 3 | ., | EARLY RESORPTION |
| í | | 4 | M | NO REMARKABLE OBSERVATIONS |
| ١ | | 5 6 | M | NO REMARKABLE OBSERVATIONS |
| | | 6 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| , | | 7 8 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| , | | 8 9 | F M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | 10 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | 11 | F | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 73698 | 1 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 73090 | 2 | F | NO REMARKABLE OBSERVATIONS |
| | | 3 | F | NO REMARKABLE OBSERVATIONS |
| | | 4 | - | EARLY RESORPTION |
| | | 5 | F | NO REMARKABLE OBSERVATIONS |
| | | 6 | F | NO REMARKABLE OBSERVATIONS |
| | | · · | - | |

Page 357 of 360

| PROJECT NO.:WIL-402010F | ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID | IN RA |
|-----------------------------|--|-------|
| SPONSOR: AMERICAN PETROLEUM | INDIVIDUAL FETAL EXTERNAL FINDINGS | |

| DAMS FROM GROUP 4: 300 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|----------------------------------|------------------|--------|--|
| 73698 | 7 | M | NO REMARKABLE OBSERVATIONS |
| | 8 | M | NO REMARKABLE OBSERVATIONS |
| | 9 | M | NO REMARKABLE OBSERVATIONS |
| | 10 | F | NO REMARKABLE OBSERVATIONS |
| | 11 | F | NO REMARKABLE OBSERVATIONS |
| | 12 | M | NO REMARKABLE OBSERVATIONS |
| | 13 | F | NO REMARKABLE OBSERVATIONS |
| | 14 | F | NO REMARKABLE OBSERVATIONS |
| 73717 | 1 2 | M | NO REMARKABLE OBSERVATIONS |
| | 2 | M | NO REMARKABLE OBSERVATIONS |
| | 3 | F | NO REMARKABLE OBSERVATIONS |
| | 4 | F | NO REMARKABLE OBSERVATIONS |
| | 4 5 6 7 | M | NO REMARKABLE OBSERVATIONS |
| | 6 | F | NO REMARKABLE OBSERVATIONS |
| | | F | NO REMARKABLE OBSERVATIONS |
| | 8 | M | NO REMARKABLE OBSERVATIONS |
| | 9 | M | NO REMARKABLE OBSERVATIONS |
| | 10 | F | NO REMARKABLE OBSERVATIONS |
| | 11 | M | NO REMARKABLE OBSERVATIONS |
| | 12 | M | NO REMARKABLE OBSERVATIONS |
| | 13 | F | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 14 | M | |
| 73693 | 15 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| 73693 | 1 2 3 | M M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 2 | F | NO REMARKABLE OBSERVATIONS |
| | 3 1 | r M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 5 6 | F | NO REMARKABLE OBSERVATIONS |
| | 7 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 8 | F | NO REMARKABLE OBSERVATIONS |
| | 9 | M | NO REMARKABLE OBSERVATIONS |
| | 10 | M | NO REMARKABLE OBSERVATIONS |

PROJECT NO.:WIL-402010F ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RATS SPONSOR:AMERICAN PETROLEUM INDIVIDUAL FETAL EXTERNAL EXPENSAL EX

PAGE 12

| DAMS FROM GROUP 4: 300 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|----------------------------------|----------|--------|---|
| 73693 | 11 12 | M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 13 | F | NO REMARKABLE OBSERVATIONS |
| | 14 15 | M M | NO REMARKABLE OBSERVATIONS NO REMARKABLE OBSERVATIONS |
| | 13 | 111 | NO REMARKABLE OBSERVATIONS |

Page 359 of 360

| SPONSOR: AMERICAN PETROLEUM | INDIVIDUAL | FETAL | EXTERNAL | FINDINGS | |
|-----------------------------|------------|-------|----------|----------|--|
| | | | | | |

| DAMS FROM GROUP 5: 1000 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION |
|-----------------------------------|--------|-----|--|
| 73718 | 1 | | EARLY RESORPTION |
| | 2 | | EARLY RESORPTION |
| | 3 | | EARLY RESORPTION |
| | 4 | | EARLY RESORPTION |
| | 5 | | EARLY RESORPTION |
| 73701 | 1 | | EARLY RESORPTION |
| | 2 | | DEAD FETUS |
| | | | CROWN-RUMP LENGTH: 1.2 CM, OPEN EYELID, BILATERAL, |
| | | | VENTRAL MIDLINE NOT CLOSED, MALE, WEIGHT: 0.32 G |
| | 3 | | DEAD FETUS |
| | | | CROWN-RUMP LENGTH: 1.1 CM, OPEN EYELID, BILATERAL, |
| | | | VENTRAL MIDLINE NOT CLOSED, FEMALE, WEIGHT: 0.22 G |
| 73721 | 1 | M | NO REMARKABLE OBSERVATIONS |
| | 2 | F | NO REMARKABLE OBSERVATIONS |
| | 3 | M | NO REMARKABLE OBSERVATIONS |
| | 4 | M | NO REMARKABLE OBSERVATIONS |
| | 5 | M | NO REMARKABLE OBSERVATIONS |
| | 6 | F | NO REMARKABLE OBSERVATIONS |
| | 7 | F | NO REMARKABLE OBSERVATIONS |
| | 8 | F | NO REMARKABLE OBSERVATIONS |
| | 9 | M | NO REMARKABLE OBSERVATIONS |
| | 10 | M | NO REMARKABLE OBSERVATIONS |
| | 11 | F | NO REMARKABLE OBSERVATIONS |
| | 12 | M | NO REMARKABLE OBSERVATIONS |
| 73707 | 1 | | EARLY RESORPTION |
| | 2 | | EARLY RESORPTION |
| | 3 | | EARLY RESORPTION |
| | 4 | | EARLY RESORPTION |
| | 5 | | EARLY RESORPTION |
| 73722 | 1 | M | NO REMARKABLE OBSERVATIONS |
| | 2 | F | NO REMARKABLE OBSERVATIONS |
| | 3 | F | NO REMARKABLE OBSERVATIONS |
| | 4 | F | NO REMARKABLE OBSERVATIONS |

TABLE 125

| PROJECT NO.:WIL-402010F | ORAL 14-DAY/PILOT DEV TOX STUDY OF NAPHTHENIC ACID IN RAIS |
|------------------------------|--|
| SPONSOR · AMERICAN PETROLEUM | INDIVIDIAL FETAL FYTERNAL FINDINGS |

| DAMS FROM GROUP 5: 1000 MG/KG/DAY | FETUS# | SEX | FETAL GROSS OBSERVATION | |
|------------------------------------|-------------|-----|----------------------------|-------------|
| 73722 | 5 | F | NO REMARKABLE OBSERVATIONS | |
| | 6 | F | NO REMARKABLE OBSERVATIONS | |
| | 7 | F | NO REMARKABLE OBSERVATIONS | |
| | 8 | M | NO REMARKABLE OBSERVATIONS | |
| | 9 | F | NO REMARKABLE OBSERVATIONS | |
| | 10 | M | NO REMARKABLE OBSERVATIONS | |
| | 11 | M | NO REMARKABLE OBSERVATIONS | |
| | 12 | M | NO REMARKABLE OBSERVATIONS | |
| | 13 | F | NO REMARKABLE OBSERVATIONS | |
| | 14 | M | NO REMARKABLE OBSERVATIONS | |
| OBSERVATION CODE: M = MALFORMATION | V = VARIATI | ON | | |
| | | | | PMFGRDv4.16 |
| | | | | 07/15/2010 |

PAGE 14