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STUDY TITLE

Report

DHDPS

Prenatal Developmental Toxicity Study in Wistar Rats
Oral Administration (Gavage)

TEST GUIDELINES

Commission Regulation (EC) No. 440/2008
OECD 414
U.S. EPA OPPTS 870.3700

AUTHORS

[REDACTED]

STUDY COMPLETION DATE

05 Nov 2014

TEST FACILITY

BASF SE
Experimental Toxicology and Ecology
67056 Ludwigshafen, Germany

TEST FACILITY PROJECT IDENTIFICATION

Project No.: 30R0066/05R018

SPONSORS

BASF SE
67056 Ludwigshafen, Germany

Jiangsu Aolunda High-Tech. Industry Co., Ltd.
Zhoutie Town, Yixing, Jiangsu, China / 214262

Konishi Chemical Ind. Co., Ltd.

LANXESS Deutschland GmbH
D-50569 Köln, Germany

PART I OF III (REPORT SECTION AND SUMMARY TABLES)

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GLP COMPLIANCE STATEMENT

This study was conducted in accordance with the OECD Principles of Good Laboratory Practice and the GLP Principles of the German "Chemikaliengesetz" (Chemicals Act) which meet the United States Environmental Protection Agency Good Laboratory Practice Standards [40 CFR Part 160 (FIFRA) and Part 792 (TSCA)], with the exception that recognized differences exist between the GLP Principles/Standards of OECD and the Principles/Standards of FIFRA and TSCA.

Study Director

Date: 05 Nov, 2018

Typed name of Study Director:

Typed name of Laboratory:

BASF SE
Experimental Toxicology and Ecology
67056 Ludwigshafen
Germany

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I have applied the criteria of 40 CFR 158.34/40 CFR161.34 for flagging studies for potential adverse effects to the results of the attached study. This study neither meets nor exceeds any of the applicable criteria.

Submitter: Date:

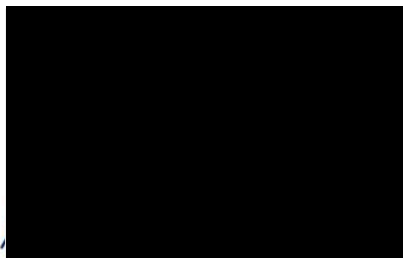
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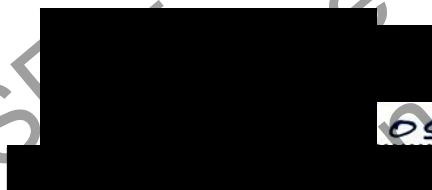
05 Nov. 2014

Analytical Chemistry:



05 Nov. 2014

Test Facility Management:



05 Nov. 2014

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CONTRIBUTORS TO THE STUDY / SUPERVISORY LABORATORY PERSONNEL

Head of Experimental Toxicology and Ecology:

[REDACTED]

Study Director:

[REDACTED]

Analytical Chemistry:

[REDACTED]

Data Processing:

[REDACTED]

Statistics:

[REDACTED]

Quality Assurance Unit (QAU):

[REDACTED]

Coordination QAU:

[REDACTED]

Reproduction Toxicology Laboratory:

[REDACTED]

Analytical Chemistry Laboratory:

[REDACTED]

STATEMENT OF THE QUALITY ASSURANCE UNIT

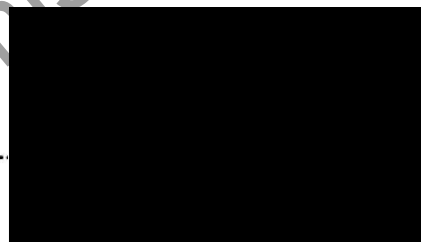
The Quality Assurance Unit (QAU) inspected the study and reported any inspection results to the Study Director and to Management.

The final report reflects the raw data.

Phase of study	Date of inspection (mm-dd-yyyy)	Reported to Study Director and to Test Facility Management (mm-dd-yyyy)
Study Plan:	09-10-2013	09-10-2013
Conduct of study:	09-26-2013 10-01-2013	09-26-2013 10-01-2013
Report:	09-22-2014	09-22-2014

Ludwigshafen,

05 November 2014



GLP CERTIFICATE (FROM THE COMPETENT AUTHORITY)



Rheinland-Pfalz

LANDESAMT FÜR UMWELT,
WASSERWIRTSCHAFT UND
GEWERBEAUFICHT

GUTE LABORPRAXIS – GOOD LABORATORY PRACTICE
GLP-BESCHEINIGUNG
STATEMENT OF GLP COMPLIANCE
gemäß/according to § 19b Abs. 1 Chemikaliengesetz

Eine GLP-Inspektion zur Überwachung der Einhaltung der GLP-Grundsätze gemäß Chemikaliengesetz bzw. Richtlinie 2004/9/EG wurde durchgeführt in:

Assessment of conformity with GLP according to Chemikaliengesetz and Directive 2004/9/EC at:

Prüfeinrichtung / Test facility

BASF SE
Experimentelle Toxikologie und Ökologie
67056 Ludwigshafen

BASF SE
Experimental Toxicology and Ecology
67056 Ludwigshafen, Germany

Prüfung nach Kategorien / Areas of Expertise

(gemäß / according ChemVwV-GLP Nr. 5.3/OECD guidance)

1,2,3,4,5,8,9

Kat. 9 – Biochemische und pathologische Untersuchungen zu Wirkmechanismen /
Biochemical and pathological examinations concerning mode of action

Datum der Inspektion / Date of Inspection

(Tag.Monat.Jahr / day.month.year)

03. bis 05.09.2013

Die genannte Prüfeinrichtung befindet sich im nationalen GLP-Überwachungsverfahren und wird regelmäßig auf Einhaltung der GLP-Grundsätze überwacht.

Auf der Grundlage des Inspektionsberichtes wird hiermit bestätigt, dass in dieser Prüfeinrichtung die oben genannten Prüfungen unter Einhaltung der GLP-Grundsätze durchgeführt werden können.

Eine erneute behördliche Überprüfung der Einhaltung der GLP-Grundsätze durch die Prüfeinrichtung ist spätestens drei Jahre nach der letzten Inspektion zu beantragen. Ohne diesen Antrag wird die Prüfeinrichtung nach Ablauf der Frist aus dem deutschen GLP-Überwachungsprogramm genommen und diese GLP-Bescheinigung verliert ihre Gültigkeit.

The above mentioned test facility is included in the national GLP Compliance Programme and is inspected on a regular basis.

Based on the inspection report it can be confirmed, that the test facility is able to conduct the aforementioned studies in compliance with the Principles of GLP.

Verification of the compliance of the test facility with the Principles of the GLP has to be applied for not later than three years after the last inspection. Elapsing this term, the test facility will be taken out of the German GLP-Monitoring Programme and this GLP Certificate becomes invalid.

Unterschrift, Datum / Signature, Date

Pia Hirsch 18.12.2013

Dr.-Ing. Pia Hirsch - Stellvertretung Präsident -
(Name und Funktion der verantwortlichen Person /
name and function of responsible person)



MESSEN
BEWERTEN
BERATEN 

Landesamt für Umwelt, Wasserwirtschaft und Gewerbeaufsicht
Kaiser-Friedrich-Straße 7, 55116 Mainz

(Name und Adresse der GLP-Überwachungsbehörde /
name and address of the GLP Monitoring Authority)

CONTENTS OF PART I (REPORT SECTION AND SUMMARY TABLES)

TITLE PAGE

STATEMENT OF NO DATA CONFIDENTIALITY CLAIMS

GLP COMPLIANCE STATEMENT

FLAGGING CRITERIA

SIGNATURE PAGE

CONTRIBUTORS TO THE STUDY / SUPERVISORY LABORATORY PERSONNEL

STATEMENT OF THE QUALITY ASSURANCE UNIT

GLP CERTIFICATE (FROM THE COMPETENT AUTHORITY)

CONTENTS OF PART I (REPORT SECTION AND SUMMARY TABLES)**1. SUMMARY****1.1. METHODS****1.2. OBSERVATIONS****1.3. RESULTS****1.3.1. Analytics****1.3.2. Effects****1.4. CONCLUSION****2. INTRODUCTION****2.1. OBJECTIVES****2.2. SELECTION OF DOSES****2.3. TEST GUIDELINES****2.4. STUDY DATES****2.5. RETENTION OF RECORDS****2.6. ANIMAL WELFARE****3. MATERIAL AND METHODS**

- 3.1. TEST ITEM
- 3.2. TEST SYSTEM
 - 3.2.1. Species and strain
 - 3.2.2. Animal identification
 - 3.2.3. Reason for species selection
- 3.3. HOUSING AND DIET
- 3.4. TEST GROUPS AND DOSES
- 3.5. TEST SUBSTANCE PREPARATIONS
 - 3.5.1. Test substance preparations and preparation frequency
- 3.6. ANALYSES
 - 3.6.1. Analyses of the test substance preparations
 - 3.6.2. Analytical methods
 - 3.6.3. Food analyses
 - 3.6.4. Drinking water analyses
 - 3.6.5. Bedding and enrichment analyses
- 3.7. EXPERIMENTAL PROCEDURE
- 3.8. CLINICAL EXAMINATIONS OF THE DAMS
 - 3.8.1. Mortality
 - 3.8.2. Clinical symptoms
 - 3.8.3. Food consumption
 - 3.8.4. Body weight data
 - 3.8.5. Corrected (net) body weight gain
 - 3.8.6. Terminal examinations of the dams
 - 3.8.6.1. Necropsy
- 3.9. EXAMINATIONS OF THE FETUSES
 - 3.9.1. Examinations of the fetuses after dissection from the uterus
 - 3.9.2. Soft tissue examination of the fetuses
 - 3.9.3. Skeletal examination of the fetuses
 - 3.9.4. Evaluation criteria for assessing the fetuses
- 3.10. STATISTICS
 - 3.10.1. Statistics of clinical and fetal examinations
- 4. RESULTS
 - 4.1. ANALYSES
 - 4.1.1. Stability analysis
 - 4.1.2. Homogeneity analysis of the test substance preparations
 - 4.1.3. Concentration control analyses of the test substance preparations
 - 4.1.4. Food analyses
 - 4.1.5. Drinking water analyses

- 4.1.6. Bedding and enrichment analyses
- 4.2. EXAMINATION OF THE DAMS
 - 4.2.1. Clinical examinations of the dams
 - 4.2.1.1. Mortality
 - 4.2.1.2. Clinical symptoms
 - 4.2.1.3. Food consumption
 - 4.2.1.4. Body weight data
 - 4.2.1.5. Corrected (net) body weight gain
 - 4.2.2. Terminal examinations of the dams
 - 4.2.2.1. Uterus weight
 - 4.2.2.2. Necropsy findings
 - 4.2.2.3. Reproduction data
- 4.3. EXAMINATION OF THE FETUSES
 - 4.3.1. Data recorded after cesarean section
 - 4.3.1.1. Sex distribution of the fetuses
 - 4.3.1.2. Weight of the placentae
 - 4.3.1.3. Weight of the fetuses
 - 4.3.2. External examination of the fetuses
 - 4.3.2.1. Fetal external malformations
 - 4.3.2.2. Fetal external variations
 - 4.3.2.3. Fetal external unclassified observations
 - 4.3.3. Soft tissue examination of the fetuses
 - 4.3.3.1. Fetal soft tissue malformations
 - 4.3.3.2. Fetal soft tissue variations
 - 4.3.3.3. Fetal soft tissue unclassified observations
 - 4.3.4. Skeletal examination of the fetuses
 - 4.3.4.1. Fetal skeletal malformations
 - 4.3.4.2. Fetal skeletal variations
 - 4.3.4.3. Fetal skeletal unclassified cartilage observations
 - 4.3.5. Assessment of all fetal external, soft tissue and skeletal observations
- 5. DISCUSSION
- 6. CONCLUSION
- 7. REFERENCES
- 8. APPENDIX
 - 8.1. LIST OF ABBREVIATIONS USED IN PART IA, IB, IIA AND IIB

PART A:**EXAMINATIONS OF THE DAMS (MEAN VALUES AND SUMMARY TABLES)**

	Tables
Summary of maternal clinical observations	IA-001
Mean maternal food consumption	IA-002 - IA-003
Mean maternal body weights	IA-004
Mean maternal body weight change	IA-005 - IA-006
Mean gravid uterine weights and net maternal body weight change	IA-007
Summary of maternal necropsy observations	IA-008
Summary of reproduction data	IA-009 - IA-011

Part B:**EXAMINATIONS OF THE FETUSES (MEAN VALUES AND SUMMARY TABLES)**

	Tables
Mean placental and fetal body weights (on a litter basis)	IB-001
Summary of all classified fetal external observations	IB-002 - IB-004
Summary of fetal external unclassified observations	IB-005
Summary of all classified fetal soft tissue observations	IB-006 - IB-009
Summary of fetal soft tissue unclassified observations	IB-010
Summary of all classified fetal skeletal observations	IB-011 - IB-027
Summary of fetal skeletal unclassified cartilage observations	IB-028 - IB-029
Summary of all classified external, soft tissue and skeletal observations	IB-030

The tables with the individual values/observations are to be found in PART II.

Further information (detailed analytical results and historical control data) is included in PART III (SUPPLEMENT).

**CONTENTS OF PART II
(TABLES SECTION, INDIVIDUAL VALUES AND OBSERVATIONS)**

Title Page
Table of contents

**Part A:
EXAMINATIONS OF THE DAMS (INDIVIDUAL VALUES)**

Tables

Individual maternal clinical observations	IIA-001 - IIA-008
Individual maternal food consumption	IIA-009 - IIA-012
Individual maternal body weights	IIA-013 - IIA-016
Individual maternal body weight change	IIA-017 - IIA-020
Individual uterine weights and net maternal body weight change	IIA-021 - IIA-024
Individual maternal necropsy observations	IIA-025 - IIA-032
Individual reproduction data	IIA-033 - IIA-036

**Part B:
EXAMINATIONS OF THE FETUSES (INDIVIDUAL VALUES)**

Tables

Individual fetal status and uterine location	IIB-001 - IIB-004
Individual placental weights	IIB-005 - IIB-008
Individual fetal body weights	IIB-009 - IIB-012
Individual fetal external, visceral, skeletal and/or cartilage observations	IIB-013 - IIB-130

CONTENTS OF PART III (SUPPLEMENT)

Title Page
Table of contents

1. Analysis of the test substance

- Final Report: Characterization of "4,4'-Dihydroxydiphenylsulfon"

2. Analyses of the test substance preparations

- DHDPS: Stability Analysis in 1% Carboxymethylcellulose in Drinking Water
- Homogeneity and Concentration Control Analysis of DHDPS in 1% Carboxymethylcellulose in Drinking water

3. Historical control data

- Tables

This report consists of PARTS I, II and III.

1. SUMMARY

1.1. METHODS

DHDPS was tested for its prenatal developmental toxicity in Wistar rats. The test substance was administered as an aqueous suspension to groups of 25 time-mated female Wistar rats by gavage at doses of 30, 100, and 300 mg/kg body weight/day (mg/kg bw/d) on gestation days (GD) 6 through 19. The control group, consisting of 25 females, was dosed with the vehicle (1% Carboxymethylcellulose in drinking water (1% CMC)) in parallel. A standard dose volume of 10 mL/kg body weight was used for each test group.

At terminal sacrifice on GD 20, 24-25 females per group had implantation sites.

1.2. OBSERVATIONS

Food consumption and body weights of the animals were recorded regularly throughout the study period. The state of health of the animals was checked each day.

On GD 20, all females were sacrificed by cervical dislocation (under isoflurane anesthesia) and assessed by gross pathology (including weight determinations of the unopened uterus and the placenta). For each dam, corpora lutea were counted and number and distribution of implantation sites (differentiated between resorptions, live and dead fetuses) were determined. The fetuses were removed from the uterus, sexed, weighed and further investigated for external findings. Thereafter, one half of the fetuses of each litter were examined for soft tissue findings and the remaining fetuses for skeletal (inclusive cartilage) findings.

1.3. RESULTS

1.3.1. Analytics

- The stability of the test substance preparations was demonstrated over a period of 4 days at room temperature and over 7 days in the refrigerator.
- The homogeneous distribution of the test substance in the vehicle (1% CMC) was confirmed.
- The correctness of the prepared concentrations was shown.

1.3.2. Effects

The following test substance-related, adverse effects/findings were noted:

Test group 3 (300 mg/kg bw/d):

Dams

- Decreased food consumption during parts of gestation (15% below control on GD 6-13, 8% below control on GD 6-19)
- Decreased body weight gain (29% below control on GD 8-10, 8% below control on GD 6-19)
- Reduced net body weight gain during treatment (10% below control on GD 6-19)

Fetuses

- No test substance-related adverse effects

Test group 2 (100 mg/kg bw/d):

- No test substance-related adverse effects

Test group 1 (30 mg/kg bw/d):

- No test substance-related adverse effects

1.4. CONCLUSION

Under the conditions of this prenatal developmental toxicity study, the oral administration of **DHDPS** to pregnant Wistar rats from implantation to one day prior to the expected day of parturition (GD 6-19) at a dose of **300 mg/kg bw/d** caused evidence of maternal toxicity, such as reduced food consumption and (net) body weight gain.

In conclusion, the **no observed adverse effect level (NOAEL)** for **maternal toxicity** is **100 mg/kg bw/d**.

There were no toxicologically relevant adverse fetal findings evident. Thus, the **no observed adverse effect level (NOAEL)** for **prenatal developmental toxicity** is **300 mg/kg bw/d**.

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2. INTRODUCTION

2.1. OBJECTIVES

The purpose of this study was to assess the effects of **DHDPS** on embryonic and fetal development in Wistar rats. Moreover, information about an influence of the test substance on the maternal organism was expected. The scope of examinations complied with the respective test guideline requirements (see 2.3. TEST GUIDELINES).

DHDPS was administered to pregnant Wistar rats by gavage, daily as a preparation in 1% CMC from implantation to one day prior to the expected day of parturition (GD 6-19).

2.2. SELECTION OF DOSES

At request of the sponsor, the following doses were chosen for the present prenatal developmental toxicity study in Wistar rats:

30 mg/kg body weight/day:	as low-dose level
100 mg/kg body weight/day:	as mid-dose level
300 mg/kg body weight/day:	as high-dose level

The oral route was selected since this has proven to be suitable for the detection of a toxicological hazard.

2.3. TEST GUIDELINES

The study was conducted according to the following test guidelines:

- Commission Regulation (EC) No 440/2008 of 30 May 2008 laying down test methods pursuant to Regulation (EC) No 1907/2006 of the European Parliament and of the Council on the Registration, Evaluation, Authorisation and Restriction of Chemicals (REACH), Part B: Methods for the determination of toxicity and other health effects: Prenatal Developmental Toxicity Study; Official Journal of the European Union, No. L 142
- OECD Guideline for the Testing of Chemicals; Proposal for Updating Guideline 414, 22 Jan 2001 ("Prenatal Developmental Toxicity Study")
- U.S. EPA, Health Effects Test Guidelines; OPPTS 870.3700, Aug 1998 ("Prenatal Developmental Toxicity Study")

2.4. STUDY DATES

Due to technical reasons, the study was carried out in 2 cohorts. Each dose group was represented in every cohort (see Tab. 2.4.1. Time schedule).

Tab. 2.4.1. Time schedule:

Study initiation date:		10 Sep 2013		
Experimental starting date: (arrival of 1 st cohort of test animals)		10 Sep 2013		
	Supply of animals/ Beginning of acclimatization (GD 0)	Beginning of treatment/ End of acclimatization (GD 6)	End of treatment (GD 19)	Sacrifice of the animals (GD 20)
1 st cohort	10 Sep 2013	16 Sep 2013	29 Sep 2013	30 Sep 2013
2 nd cohort	11 Sep 2013	17 Sep 2013	30 Sep 2013	01 Oct 2013
Experimental completion date: (draft to QAU*)		27 Aug 2014		

* QAU = Quality Assurance Unit

2.5. RETENTION OF RECORDS

GLP – relevant records and materials are archived at BASF SE for at least the period of time specified in the GLP principles. Details concerning responsibilities or locations of archiving can be seen from the respective SOPs and from the raw data.

2.6. ANIMAL WELFARE

This study was performed in an AAALAC-approved laboratory in accordance with the German Animal Welfare Act and the effective European Council Directive.

3. MATERIAL AND METHODS

3.1. TEST ITEM

The analyses of the test item (= test substance) were carried out at Competence Center Analytics, BASF SE, 67056 Ludwigshafen, Germany.

Name of test substance:	DHDPS
Test substance No.:	05/0066-4
Batch No.:	69611767J0
CAS No.:	80-09-1
Purity:	99.3 and 99.5% (HPLC) 99.4 g/100 g (1H NMR) (see Report, Project No.: 12L00002)
Homogeneity:	given (visually)
Storage stability:	Expiry date: 28 Nov 2013

The stability of the test substance under storage conditions over the test period was guaranteed by the sponsor, and the sponsor holds this responsibility.

ADDITIONAL TEST SUBSTANCE INFORMATION

IUPAC name:	4,4'-Dihydroxydiphenylsulfon; 4,4'-Sulfonyldiphenol; 4,4'-Dihydroxyphenylsulfon
Physical state/Appearance:	Solid / white
Storage conditions:	Room temperature, moisture protection

3.2. TEST SYSTEM

3.2.1. Species and strain

Time-mated Wistar rats (CrI:WI[Han]) were supplied by Charles River Laboratories, Research Models and Services, Germany GmbH at an age of about 10-12 weeks. Only animals free from clinical signs of disease were used for the investigations.

3.2.2. Animal identification

The animals were paired by the breeder and supplied on GD 0 (= detection of vaginal plug/sperm). After arrival of each cohort, the assignment of the animals to the different test groups was carried out by withdrawing them from the transport boxes and placing them into the cages at random. Each cohort was evenly distributed among the dose groups. After randomization the rats were identified consecutively and uniquely by ear tattoo.

3.2.3. Reason for species selection

The CrI:WI(Han) strain was selected since extensive historical control data is available from the test facility for Wistar rats. This specific strain has been proven to be sensitive to substances with a teratogenic potential.

3.3. HOUSING AND DIET

During the study period, the rats were housed individually in Makrolon type M III cages supplied by BECKER & CO., Castrop-Rauxel, Germany (floor area about 800 cm²).

Dust-free wooden bedding was used in this study (the present supplier is documented in the raw data).

For enrichment, wooden gnawing blocks were offered (Typ NGM E-022, supplied by Abedd[®] Lab. and Vet. Service GmbH, Vienna, Austria).

The animals were accommodated in fully air-conditioned rooms in which central air conditioning maintained a range of temperature of 20-24°C and a range of relative humidity of 30-70%. The air exchange rate was 15 times per hour. There were no deviations from these limits during the entire study.

The day/night cycle was generally 12 hours (12 hours light from 6.00 h to 18.00 h and 12 hours darkness from 18.00 h to 6.00 h).

Before the study started, the animal room was completely disinfected using a disinfectant ("AUTEX" fully automatic, formalin-ammonia-based terminal disinfection). The walls and the floor were cleaned once a week with water containing an appropriate disinfectant.

The food used was ground Kliba maintenance diet mouse/rat "GLP", meal, supplied by Provimi Kliba SA, Kaiseraugst, Switzerland. Food and drinking water (potable tap water in water bottles) were available *ad libitum*.

3.4. TEST GROUPS AND DOSES

Test group	Dose [mg/kg bw/d]	Concentration [mg/100 mL]	Volume [mL/kg]	Number of animals	Animal No.
0	0	0	10 ^{a)}	25	1 – 25
1	30	300	10 ^{b)}	25	26 – 50
2	100	1000	10 ^{b)}	25	51 – 75
3	300	3000	10 ^{b)}	25	76 – 100

a) 1% Carboxymethylcellulose suspension in drinking water

b) Test substance preparations in 1% Carboxymethylcellulose suspension in drinking water

3.5. TEST SUBSTANCE PREPARATIONS

3.5.1. Test substance preparations and preparation frequency

The aqueous test substance preparations were prepared at the beginning of the administration period and thereafter at maximum intervals of 7 days, which took into account the period of established stability. The preparations were kept in a refrigerator.

For the test substance preparations, the specific amount of test substance were weighed, topped up with 1% Carboxymethylcellulose suspension in drinking water in a calibrated beaker and intensely mixed with a homogenizer.

During administration the preparations were kept homogeneous with a magnetic stirrer.

3.6. ANALYSES

3.6.1. Analyses of the test substance preparations

The analyses of the test substance preparations were carried out at the Analytical Chemistry Laboratory of Experimental Toxicology and Ecology of BASF SE, Ludwigshafen, Germany.

Analytical verifications of the stability of the test substance in 1% Carboxymethylcellulose suspension in drinking water over a period of 4 days at room temperature or 7 days in a refrigerator were carried out prior to the start of the study (Project No.: 01Y0066/05Y009, see PART III).

Samples of the test substance preparations were sent to the analytical laboratory at the beginning of administration for verification of the concentrations. The samples were also used to verify the homogeneity of the low and the high concentrations (30 and 300 mg/kg bw/d). Three samples (one from the top, middle and bottom in each case) were taken from the beaker with a magnetic stirrer running.

3.6.2. Analytical methods

Details on the methods used for the analytical investigations of the test substance preparations can be found in PART III.

3.6.3. Food analyses

The food used in the study was assayed for chemical and for microbiological contaminants.

3.6.4. Drinking water analyses

The drinking water was regularly assayed for chemical contaminants both by the municipal authorities of Frankenthal and by the Environmental Analytics Water/Steam Monitoring of BASF SE as well as for bacteria by a contract laboratory.

3.6.5. Bedding and enrichment analyses

The bedding and the enrichment were regularly assayed for contaminants (chlorinated hydrocarbons and heavy metals).

3.7. EXPERIMENTAL PROCEDURE

The animals were paired by the breeder ("time-mated"); the day of evidence of mating (= detection of vaginal plug/sperm) was referred to as GD 0. The animals arrived on the same day (GD 0) at the experimental laboratory. The following day was designated as "GD 1". The animals were acclimated to the laboratory conditions between start of the study (beginning of the experimental phase) and first administration (GD 6).

The body weight of the pregnant animals on day 0 varied between 142.8 – 197.8 g.

The test substance preparations were administered to the animals once a day orally by gavage, from implantation to one day prior to the expected day of parturition (GD 6 to GD 19), always at approximately the same time in the morning. The animals of the control group were treated with the vehicle (1% CMC) in the same way. The volume administered each day was 10 ml/kg body weight. The calculation of the administration volume was based on the most recent individual body weight.

On GD 20, the females were sacrificed in a randomized order and examined macroscopically. The fetuses were removed from the uterus and investigated with the methods described in section 3.9. Examinations of the fetuses.

3.8. CLINICAL EXAMINATIONS OF THE DAMS

3.8.1. Mortality

A check was made twice a day on working days or once a day on Saturdays, Sundays or on public holidays (GD 0-20).

3.8.2. Clinical symptoms

A cage-side examination was conducted at least once daily for any signs of morbidity, pertinent behavioral changes and signs of overt toxicity. If such signs occurred, the animals were examined several times daily (GD 0-20).

3.8.3. Food consumption

The consumption of food was recorded for the intervals GD 0-1, 1-3, 3-6, 6-8, 8-10, 10-13, 13-15, 15-17, 17-19 and 19-20.

3.8.4. Body weight data

All animals were weighed on GD 0, 1, 3, 6, 8, 10, 13, 15, 17, 19 and 20. The body weight change of the animals was calculated based on the obtained results.

3.8.5. Corrected (net) body weight gain

Furthermore, the corrected body weight gain was calculated after terminal sacrifice (terminal body weight on GD 20 minus weight of the unopened uterus minus body weight on GD 6).

3.8.6. Terminal examinations of the dams

3.8.6.1. Necropsy

On GD 20, the dams were sacrificed under isoflurane anesthesia by cervical dislocation, in randomized order.

After the dams had been sacrificed, they were necropsied and assessed for gross pathology, in randomized order. The uteri and the ovaries were removed and the following data were recorded:

- Weight of the unopened uterus
- Number of corpora lutea
- Number and distribution of implantation sites classified as:
 - Live fetuses
 - Dead implantations:
 - a) Early resorptions (only decidual or placental tissues visible or according to SALEWSKI from uteri from apparently non-pregnant animals and the empty uterus horn in the case of single horn pregnancy)
 - b) Late resorptions (embryonic or fetal tissue in addition to placental tissue visible)
 - c) Dead fetuses (hypoxemic fetuses which did not breathe spontaneously after the uterus had been opened)

After the weight of the uterus had been determined, all subsequent evaluations of the dams and the gestational parameters were conducted by technicians unaware of treatment group in order to minimize bias. For this purpose animal numbers were encoded.

These data were used to calculate conception rate and pre- and postimplantation losses.

The **conception rate** (in %) was calculated according to the following formula:

$$\frac{\text{number of pregnant animals}}{\text{number of fertilized animals}} \times 100$$

The **preimplantation loss** (in %) for each individual pregnant animal which underwent scheduled sacrifice was calculated according to the following formula:

$$\frac{\text{number of corpora lutea} - \text{number of implantations}}{\text{number of corpora lutea}} \times 100$$

The **postimplantation loss** (in %) for each individual pregnant animal which underwent scheduled sacrifice was calculated according to the following formula:

$$\frac{\text{number of implantations} - \text{number of live fetuses}}{\text{number of implantations}} \times 100$$

3.9. EXAMINATIONS OF THE FETUSES

All fetal analyses were conducted by technicians unaware of the treatment group, in order to minimize bias.

3.9.1. Examinations of the fetuses after dissection from the uterus

At necropsy each fetus was weighed, sexed, and external tissues and all orifices were examined macroscopically. The sex was determined by observing the distance between the anus and the base of the genitalia.

Furthermore, the viability of the fetuses and the condition of placentae, umbilical cords, fetal membranes, and fluids were examined. The placentas were weighed and their individual weights were recorded.

Thereafter, the fetuses were sacrificed by a subcutaneous injection of pentobarbital (Narcoren®; dose: 0.1 mL/fetus).

After these examinations, approximately one half of the fetuses per dam were eviscerated, skinned and fixed in ethanol; the other half were placed in Harrison's fluid for fixation.

3.9.2. Soft tissue examination of the fetuses

The fetuses fixed in Harrison's fluid were examined for any visceral findings according to the method of BARROW and TAYLOR. After this examination these fetuses were discarded.

3.9.3. Skeletal examination of the fetuses

The skeletons of the fetuses fixed in ethanol were stained according to a modified method of KIMMEL and TRAMMELL. Thereafter, the skeletons of these fetuses were examined under a stereomicroscope. After this examination the stained fetal skeletons were archived individually.

3.9.4. Evaluation criteria for assessing the fetuses

In the present study the glossary of WISE et al. (1997) and its updated version of MAKRIS et al. (2009) was essentially used to describe findings in fetal morphology. Classification of these findings was based on the terms and definitions proposed by CHAHOUD et al. (1999) and SOLECKI et al. (2001, 2003):

Malformation

A permanent structural change that is likely to adversely affect the survival or health.

Variation

A change that also occurs in the fetuses of control animals and/or is unlikely to adversely affect the survival or health. This includes delays in growth or morphogenesis that have otherwise followed a normal pattern of development.

The term "**unclassified observation**" was used for those fetal findings, which could not be classified as malformations or variations.

All fetal findings were listed in tables according to these classifications.

3.10. STATISTICS

3.10.1. Statistics of clinical and fetal examinations

Statistical analyses were performed according to the following table

Parameter	Statistical test	Markers in the tables	References
Food consumption ^{a)} , body weight, body weight change, corrected body weight gain (net maternal body weight change), carcass weight, weight of unopened uterus, number of corpora lutea, number of implantations, number of resorptions, number of live fetuses, proportions of preimplantation loss, proportions of postimplantation loss, proportions of resorptions, proportion of live fetuses in each litter, litter mean fetal body weight, litter mean placental weight	Simultaneous comparison of all dose groups with the control group using the DUNNETT-test (two-sided) for the hypothesis of equal means	* for $p \leq 0.05$; ** for $p \leq 0.01$	DUNNETT C W (1955): A multiple comparison procedure for comparing several treatments with a control. JASA, Vol. 50, 1096-1121 DUNNETT C W (1964): New tables for multiple comparisons with a control. Biometrics, Vol. 20, 482-491
Female mortality, females pregnant at terminal sacrifice, number of litters with fetal findings	Pairwise comparison of each dose group with the control group using FISHER'S EXACT test (one-sided) for the hypothesis of equal proportions	* for $p \leq 0.05$; ** for $p \leq 0.01$	SIEGEL S (1956): Non-parametric statistics for behavioral sciences. McGraw-Hill New York
Proportions of fetuses with malformations, variations and/or unclassified observations in each litter	Pairwise comparison of each dose group with the control group using the WILCOXON-test (one-sided) for the hypothesis of equal medians	* for $p \leq 0.05$; ** for $p \leq 0.01$	NIJENHUIS A, and WILF H S (1978): Combinatorial Algorithms. Academic Press New York, 32-33 HETTMANSPERGER T P (1984): Statistical Inference based on Ranks. John Wiley & Sons New York, 132-142

^{a)} For the parameter food consumption the "mean of means" was calculated and can be found in the relevant summary tables. The "mean of means" values allow a rough estimation of the total food consumption during different time intervals (pretreatment (days 0-6), treatment (days 6-19) and entire study (days 0-20)); they are not exactly precise values, because the size of the intervals taken for calculation differs. For the "mean of means" values no statistical analysis was performed.

4. RESULTS

4.1. ANALYSES

4.1.1. Stability analysis

The stability of the test substance in 1% Carboxymethylcellulose in drinking water over a period of 4 days at room temperature and over 7 days in a refrigerator was demonstrated (see PART III, SUPPLEMENT).

4.1.2. Homogeneity analysis of the test substance preparations

The homogeneous distribution of the test substance in the vehicle (1% CMC) was demonstrated (see PART III, SUPPLEMENT).

4.1.3. Concentration control analyses of the test substance preparations

The results of the analysis of the test substance preparations in 1% Carboxymethylcellulose in drinking water confirmed the correctness of the prepared concentrations. The analytical values of the samples corresponded to the expected values within the limits of the analytical method, i.e. were always above 90% and below 110% of the nominal concentrations (see PART III, SUPPLEMENT).

4.1.4. Food analyses

On the basis of the duration of use and the analytical findings with respect to chemical and microbiological contaminants, the food was found to be suitable. The EPA Fed. Reg. of 09 May 1979 (Vol. 44, No. 91, p. 27354) served as a guideline for maximum tolerable chemical contaminants. The amount of microorganisms did not exceed $1 \cdot 10^5$ /g feed.

The individual results are found in the archives of Experimental Toxicology and Ecology, BASF SE, Ludwigshafen, Germany.

4.1.5. Drinking water analyses

On the basis of the analytical findings, the drinking water was found to be suitable. The German Drinking Water Regulation ("Trinkwasserverordnung") served as the guideline for maximum tolerable contaminants.

The individual results are found in the archives of Experimental Toxicology and Ecology, BASF SE, Ludwigshafen, Germany.

4.1.6. Bedding and enrichment analyses

On the basis of the analytical findings, the bedding and the enrichment were found to be suitable. Levels given in Lab Animal, Nov-Dec 1979, pp. 24-34, served as a guideline for maximum tolerable contaminants.

The individual results are found in the archives of Experimental Toxicology and Ecology, BASF SE, Ludwigshafen, Germany.

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4.2. EXAMINATION OF THE DAMS

Summary tables are given in Part A of PART I. Individual values are given in Part A of PART II.

4.2.1. Clinical examinations of the dams

Only pregnant dams were used for the calculations of mean maternal food consumption, body weight and body weight change. Only pregnant dams with scheduled sacrifice (GD 20) were used for the calculation of mean gravid uterine weights, corrected (net) body weight gain and summary of reproduction data.

The following female was excluded from the above-mentioned calculations:

Test group 3 (300 mg/kg bw/d):

- female No. 93 – not pregnant

4.2.1.1. Mortality

There were no test substance-related or spontaneous mortalities in any females of all test groups (0, 30, 100 or 300 mg/kg bw/d).

4.2.1.2. Clinical symptoms

(Tab. IA-001)

Seven (out of 25) high-dose females (300 mg/kg bw/d) showed transient salivation during major parts of the treatment period. Salivation persisted in the respective animals only for some minutes after daily gavage dosing (i.e. up to 20 minutes) and was initially observed on GD 10.

No clinical signs or changes of general behavior, which may be attributed to the test substance, were detected in any female of the low- and the mid-dose groups (30 or 100 mg/kg bw/d) during the entire study period.

4.2.1.3. Food consumption

(Tabs. IA-002 - IA-003)

The mean food consumption of the high-dose dams (300 mg/kg bw/d) was statistically significantly reduced at the beginning of the treatment period (GD 6-13; up to 15% below control), but recovered afterwards. If calculated for the entire treatment period (GD 6-19) or the entire study period (GD 0-20), the high-dose dams consumed 8% or 6%, respectively, less food in comparison to the concurrent control group.

The mean food consumption of the dams in test groups 1 and 2 (30 or 100 mg/kg bw/d) was generally comparable to the concurrent control throughout the entire study period.

The only exception was a slightly, but statistically significantly lower mean food consumption of the low- and high-dose dams (30 and 300 mg/kg bw/d) on GD 0-1, which was a solitary event and considered to be accidental.

4.2.1.4. Body weight data

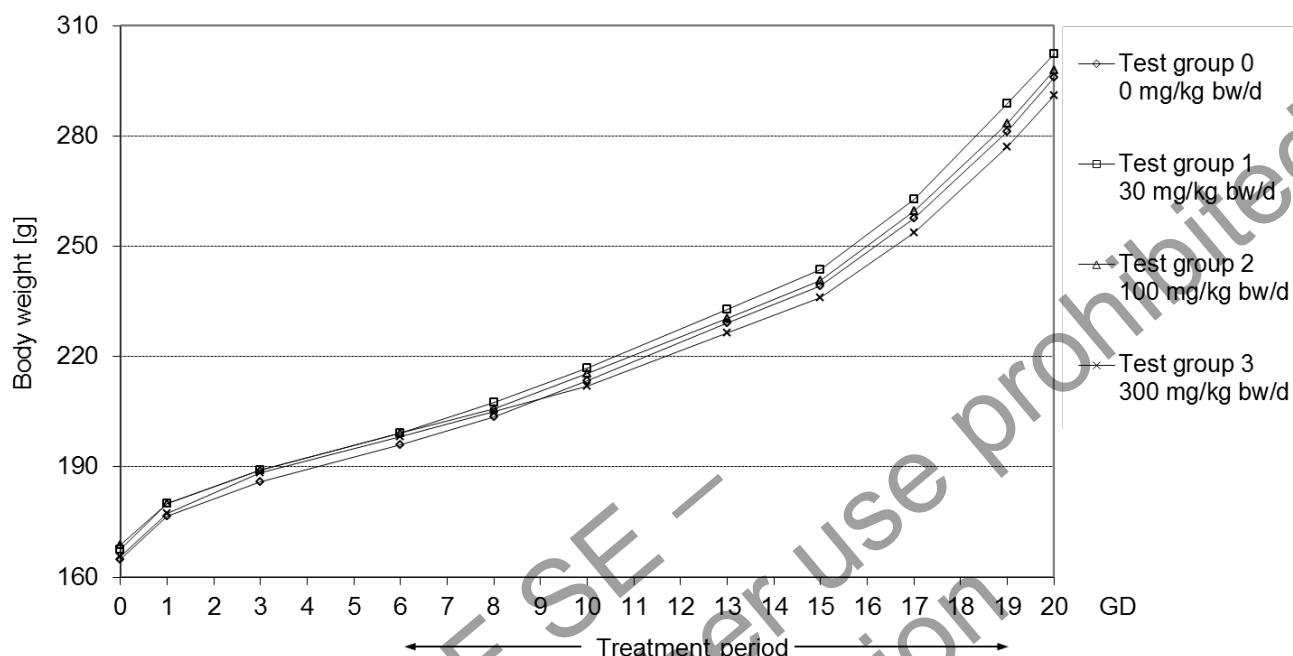
(Tabs. IA-004 - IA-006; Fig. 4.2.1.4.1.)

The mean body weights of the dams in test group 3 (300 mg/kg bw/d) were in general comparable to the controls throughout the entire study period.

The body weight change of the high-dose dams was statistically significantly reduced on GD 8-10 (approx. 29% below control) and if calculated for the entire treatment period (GD 6-19; 8% below control).

The mean body weights and the average body weight gains of the low- and mid-dose dams (30 or 100 mg/kg bw/d) were in general comparable to the controls throughout the entire study period.

Fig. 4.2.1.4.1. Mean body weight of pregnant animals



4.2.1.5. Corrected (net) body weight gain (Tab. IA-007)

The corrected body weight gain (terminal body weight on GD 20 minus weight of the unopened uterus minus body weight on GD 6) of test group 3 (300 mg/kg bw/d) was clearly lower than the concurrent control value (approx. 10% below control), but without attaining statistical significance.

The corrected body weight gain of test groups 1 and 2 (30 and 100 mg/kg bw/d) revealed no difference of any biological relevance to the corresponding control group.

Moreover, mean carcass weights of all test groups remained unaffected by the treatment.

4.2.2. Terminal examinations of the dams

4.2.2.1. Uterus weight

(Tab. IA-007)

The mean gravid uterus weights of the animals of test group 1-3 (30, 100 and 300 mg/kg bw/d) were not influenced by the test substance. The differences between these groups and the control group revealed no dose-dependency and were assessed to be without biological relevance.

4.2.2.2. Necropsy findings

(Tab. IA-008)

No necropsy findings which could be attributed to the test substance were seen in any dam (0, 30, 100 or 300 mg/kg bw/d).

A number of spontaneous findings were noted in individual females of test group 3 (300 mg/kg bw/d). These gross findings were:

- a diaphragmatic hernia in female No. 93 (this female was not pregnant),
- a dilated renal pelvis in female No. 97 (left side) and
- a hemometra in female No. 98 (right side).

4.2.2.3. Reproduction data

(Tabs. IA-009 - IA-011)

The conception rate varied between 96% in test group 3 (300 mg/kg bw/d) and 100% in test groups 0, 1 and 2 (0, 30 and 100 mg/kg bw/d). With these rates, a sufficient number of pregnant females were available for the purpose of the study (according to the test guidelines listed in chapter 2.3.).

There were no test substance-related and/or biologically relevant differences between test groups 0, 1, 2 and 3 (0, 30, 100 and 300 mg/kg bw/d) in conception rate, in the mean number of corpora lutea and implantation sites or in the values calculated for the pre- and the postimplantation losses, the number of resorptions and viable fetuses. All observed differences are considered to reflect the normal range of fluctuations for animals of this strain and age; see also PART III (SUPPLEMENT) for historical control data.

4.3. EXAMINATION OF THE FETUSES

Summary tables are given in Part B of PART I. Individual values are given in Part B of PART II.

4.3.1. Data recorded after cesarean section

4.3.1.1. Sex distribution of the fetuses

(Tab. IA-011)

The sex distribution of the fetuses in test groups 1-3 (30, 100 and 300 mg/kg bw/d) was comparable to the control fetuses. Any observable differences were without biological relevance.

4.3.1.2. Weight of the placentae

(Tab. IB-001)

The mean placental weights of the low-, mid- and high-dose groups (30, 100 and 300 mg/kg bw/d) were comparable to the corresponding control group.

4.3.1.3. Weight of the fetuses

(Tab. IB-001)

The mean fetal weights of test groups 1, 2 and 3 (30, 100 and 300 mg/kg bw/d) were not influenced by the test substance and did not show any biologically relevant differences in comparison to the control group.

4.3.2. External examination of the fetuses

(Tab. IB-002)

4.3.2.1. Fetal external malformations

(Tab. IB-003)

One mid-dose fetus with multiple external malformations was seen: male fetus No. 61-03 (100 mg/kg bw/d) had a misshapen head and an absent face (comprising anophthalmia, anotia and astomia). These external findings were associated with multiple skeletal malformations (see Tab. 4.3.2.1.1.).

The overall incidences of external malformations were comparable to those found in the historical control data (PART III, SUPPLEMENT).

Tab. 4.3.2.1.1. Individual fetal external malformations

Test group	Dam No.-Fetus No., Sex	Finding
0 (0 mg/kg bw/d)	none	
1 (30 mg/kg bw/d)	none	
2 (100 mg/kg bw/d)	61-03 M ^{a)}	multiple external malformations
3 (300 mg/kg bw/d)	none	

mg/kg bw/d = milligram per kilogram body weight per day; No. = number; M = male; F = female

^{a)} fetus with additional skeletal malformation (see Tab. 4.3.4.1.1.)

Tab. 4.3.2.1.2. Total external malformations

		Test group 0 0 mg/kg bw/d	Test group 1 30 mg/kg bw/d	Test group 2 100 mg/kg bw/d	Test group 3 300 mg/kg bw/d
Litter	N	25	25	25	24
Fetuses	N	264	265	265	243
Fetal incidence	N (%)	0.0	0.0	1 (0.4)	0.0
Litter incidence	N (%)	0.0	0.0	1 (4.0)	0.0
Affected fetuses/litter	Mean%	0.0	0.0	0.8	0.0

mg/kg bw/d = milligram per kilogram body weight per day; N = number; % = per cent

4.3.2.2. Fetal external variations

(Tab. IB-004)

No external variations were recorded.

4.3.2.3. Fetal external unclassified observations

(Tab. IB-005)

One unclassified external observation, i.e. blood coagulum around placenta, was recorded in two fetuses of the high-dose group (300 mg/kg bw/d). This finding was not considered biologically relevant.

Tab. 4.3.2.3.1. Total external unclassified observations

		Test group 0 0 mg/kg bw/d	Test group 1 30 mg/kg bw/d	Test group 2 100 mg/kg bw/d	Test group 3 300 mg/kg bw/d
Litter	N	25	25	25	24
Fetuses	N	264	265	265	243
Fetal incidence	N (%)	0.0	0.0	0.0	2 (0.8)
Litter incidence	N (%)	0.0	0.0	0.0	2 (8.3)
Affected fetuses/litter	Mean%	0.0	0.0	0.0	1.3

mg/kg bw/d = milligram per kilogram body weight per day; N = number; % = per cent

4.3.3. Soft tissue examination of the fetuses

(Tab. IB-006)

4.3.3.1. Fetal soft tissue malformations

(Tab. IB-007)

No soft tissue malformations were recorded.

4.3.3.2. Fetal soft tissue variations

(Tabs. IB-008 – IB-009)

Some soft tissue variations were detected in all test groups including the control (0, 30, 100 or 300 mg/kg bw/d), i.e. short innominate, dilated renal pelvis and dilated ureter. The incidences of these variations were neither statistically significantly different from control nor dose-dependent and therefore, not considered biologically relevant. Most of them can be found in the historical control data at comparable incidences (PART III, SUPPLEMENT).

Tab. 4.3.3.2.1. Total soft tissue variations

		Test group 0 0 mg/kg bw/d	Test group 1 30 mg/kg bw/d	Test group 2 100 mg/kg bw/d	Test group 3 300 mg/kg bw/d
Litter	N	25	25	25	24
Fetuses	N	127	128	125	116
Fetal incidence	N (%)	7 (5.5)	5 (3.9)	12 (9.6)	10 (8.6)
Litter incidence	N (%)	7 (28)	4 (16)	9 (36)	9 (38)
Affected fetuses/litter	Mean%	6.1	4.1	9.1	8.5

mg/kg bw/d = milligram per kilogram body weight per day; N = number; % = per cent

4.3.3.3. Fetal soft tissue unclassified observations

(Tab. IB-010)

No unclassified soft tissue observations were recorded.

4.3.4. Skeletal examination of the fetuses

(Tab. IB-011)

4.3.4.1. Fetal skeletal malformations

(Tabs. IB-012 – IB-014)

Skeletal malformations were detected in all test groups except group 1 (0, 100 and 300 mg/kg bw/d) affecting the skull, sternum and forelimbs (Tab. 4.3.4.1.1.). One fetus of test group 2 (No. 61-03) was multiple-malformed (malformations affected the skull, vertebral column, ribs, pelvic girdle and forelimbs) and had associated external findings. The average rate of affected fetuses per litter showing skeletal malformations was statistically significantly increased in the high-dose group (300 mg/kg bw/d). However, each of the findings leading to that increased rate (presented in Tab. 4.3.4.1.1.) is present in the historical control data and no abnormality pattern became obvious. The total incidences of skeletal malformations in the low- and mid-dose groups (30 and 100 mg/kg bw/d) were comparable to the concurrent control group (Tab. 4.3.4.1.2.).

Tab. 4.3.4.1.1. Individual fetal skeletal malformations

Test group	Dam No.-Fetus No., Sex	Finding
0 (0 mg/kg bw/d)	3-11 F	shortened humerus
1 (30 mg/kg bw/d)	none	
2 (100 mg/kg bw/d)	61-03 M ^{a)} 62-09 F 65-01 M	multiple skeletal malformations shortened humerus shortened scapula
3 (300 mg/kg bw/d)	86-08 M 87-03 F 88-04 M 89-01 F 90-10 M	malpositioned and bipartite sternebra misshapen basisphenoid malpositioned and bipartite sternebra misshapen tuberositas deltoidea shortened humerus

mg/kg bw/d = milligram per kilogram body weight per day; No. = number; M = male; F = female

^{a)} fetus with additional external malformation (Tab 4.3.2.1.1.)

Tab. 4.3.4.1.2. Total fetal skeletal malformations

		Test group 0 0 mg/kg bw/d	Test group 1 30 mg/kg bw/d	Test group 2 100 mg/kg bw/d	Test group 3 300 mg/kg bw/d
Litter	N	25	25	25	24
Fetuses	N	137	137	140	127
Fetal incidence	N (%)	1 (0.7)	0.0	3 (2.1)	5 (3.9)
Litter incidence	N (%)	1 (4.0)	0.0	3 (12)	5 (21)
Affected fetuses/litter	Mean%	0.7	0.0	2.8	4.3*

mg/kg bw/d = milligram per kilogram body weight per day; N = number; % = per cent

* = p ≤ 0.05 (Wilcoxon-test [one-sided])

4.3.4.2. Fetal skeletal variations

(Tabs. IB-015 – IB-027)

For all test groups, skeletal variations of different bone structures were observed, with or without effects on corresponding cartilages. The observed skeletal variations were related to several parts of fetal skeleton and appeared without a relation to dosing (Tab. 4.3.4.2.1.). The overall incidences of skeletal variations were comparable to the historical control data (PART III, SUPPLEMENT).

Tab. 4.3.4.2.1. Total fetal skeletal variations

		Test group 0 0 mg/kg bw/d	Test group 1 30 mg/kg bw/d	Test group 2 100 mg/kg bw/d	Test group 3 300 mg/kg bw/d
Litter	N	25	25	25	24
Fetuses	N	137	137	140	127
Fetal incidence	N (%)	136 (99)	135 (99)	139 (99)	127 (100)
Litter incidence	N (%)	25 (100)	25 (100)	25 (100)	24 (100)
Affected fetuses/litter	Mean%	99.2	98.3	98.7	100.0

mg/kg bw/d = milligram per kilogram body weight per day; N = number; % = per cent

For a better overview, all skeletal variations with statistically significant differences between the control and any treated group were compiled in the table below (Tab. 4.3.4.2.2). All incidences were expressed on a fetus per litter basis and any statistically significant differences, which were outside the historical control range were marked in bold types.

Tab. 4.3.4.2.2. Occurrence of statistically significantly increased fetal skeletal variations
(expressed as mean percentage of affected fetuses/litter)

Finding	Test group 0 0 mg/kg bw/d	Test group 1 30 mg/kg bw/d	Test group 2 100 mg/kg bw/d	Test group 3 300 mg/kg bw/d	HCD Mean % (range)
Incomplete ossification of supraoccipital; unchanged cartilage	34.1	35.2	37.6	45.2*	43.5 (10.3 – 64.3)
Dumbbell ossification of thoracic centrum; unchanged cartilage	0.7	3.0	0.0	5.6**	6.9 (0.0 – 14.5)
Unossified sternebra; unchanged cartilage	1.5	5.0	4.6	11.0**	8.2 (2.6 – 20.7)
Incomplete ossification of pubis; cartilage present	0.0	0.8	2.0*	1.7	0.3 (0.0 – 2.4)
Incomplete ossification of ischium; cartilage present	0.0	0.0	2.0*	1.7	0.2 (0.0 – 0.8)

mg/kg bw/d = milligram per kilogram body weight per day; HCD = Historical control data; % = per cent

* = $p \leq 0.05$ (Wilcoxon-test [one-sided])

** = $p \leq 0.01$ (Wilcoxon-test [one-sided])

Concerning the statistically significant findings, no dose dependency was observed and/or all values were clearly inside the historical control range, thus, an association to the test substance and a toxicological relevance is not assumed.

4.3.4.3. Fetal skeletal unclassified cartilage observations

(Tabs. IB-028 – IB-029)

Additionally, some isolated cartilage findings without impact on the respective bony structures, which were designated as unclassified cartilage observations, occurred in all test groups (Tab. 4.3.4.3.1.). The observed unclassified cartilage findings were related to the skull, the sternum and ribs and did not show any relation to dosing. The overall incidences of skeletal unclassified cartilage observations in the substance-treated groups did not differ significantly from the concurrent control group.

Tab. 4.3.4.3.1. Total unclassified cartilage observations

		Test group 0 0 mg/kg bw/d	Test group 1 30 mg/kg bw/d	Test group 2 100 mg/kg bw/d	Test group 3 300 mg/kg bw/d
Litter	N	25	25	25	24
Fetuses	N	137	137	140	127
Fetal incidence	N (%)	98 (72)	92 (67)	96 (69)	90 (71)
Litter incidence	N (%)	24 (96)	25 (100)	24 (96)	24 (100)
Affected fetuses/litter	Mean%	71.4	65.8	67.2	69.9

mg/kg bw/d = milligram per kilogram body weight per day; N = number; % = per cent

4.3.5. Assessment of all fetal external, soft tissue and skeletal observations

(Tab. IB-030)

Soft tissue malformations did not occur in any of the fetuses in this study. There were noted external (Tab. IB-003) and skeletal (Tabs. IB-012 – IB-014) malformations in all test groups.

One fetus in the mid-dose group (100 mg/kg bw/d) was multiple-malformed. Male fetus No. 61-03 had several external malformations, i.e. a misshapen head and an absent face (anophthalmia, anotia, astomia), associated with multiple skeletal malformations concerning the whole fetal body. An association of these findings to the treatment is not assumed.

Other malformations, such as misshapen basisphenoid, shortened scapula, malpositioned and bipartite sternebra, shortened humerus and misshapen tuberositas deltoidea, observed in test groups 0, 2 and 3, were scattered observations in individual fetuses of these groups, are common for this rat strain and all of them can be found in the historical control data at a comparable or higher frequency.

The number of those sporadic skeletal malformations adds up to a statistically significantly higher value for the skeletal and, subsequently, total malformation rates of affected fetuses per litter in test group 3 (300 mg/kg bw/d). However, no ontogenetic pattern is recognizable for the individual malformations nor was there any cluster of any of these individual malformations seen in the other offspring of this test group. In addition, the respective total malformation rate value for test group 3 is well within the historical control range (0.00 – 2.96%). Thus, a relationship to treatment does not exist.

Tab. 4.3.5.1. Total fetal malformations

		Test group 0 0 mg/kg bw/d	Test group 1 30 mg/kg bw/d	Test group 2 100 mg/kg bw/d	Test group 3 300 mg/kg bw/d
Litter	N	25	25	25	24
Fetuses	N	264	265	265	243
Fetal incidence	N (%)	1 (0.4)	0.0	3 (1.1)	5 (2.1)
Litter incidence	N (%)	1 (4.0)	0.0	3 (12)	5 (21)
Affected fetuses/litter	Mean%	0.4	0.0	1.6	2.3*

mg/kg bw/d = milligram per kilogram body weight per day; N = number; % = per cent

* = $p \leq 0.05$ (Wilcoxon-test [one-sided])

External variations did not occur in any of the fetuses in this study. Three soft tissue variations (Tabs. IB-008 – IB-009) and a broad range of skeletal variations (Tabs. IB-015 – IB-027) occurred in all test groups including the controls. None of the incidences showed a relation to dosing. The majority of the skeletal variations are equally distributed among the different test groups, if normal biological variation is taken into account, and can be found in the historical control data at a comparable frequency.

Tab. 4.3.5.2. Total fetal variations

		Test group 0 0 mg/kg bw/d	Test group 1 30 mg/kg bw/d	Test group 2 100 mg/kg bw/d	Test group 3 300 mg/kg bw/d
Litter	N	25	25	25	24
Fetuses	N	264	265	265	243
Fetal incidence	N (%)	143 (54)	140 (53)	151 (57)	137 (56)
Litter incidence	N (%)	25 (100)	25 (100)	25 (100)	24 (100)
Affected fetuses/litter	Mean%	54.6	52.9	56.6	56.4

mg/kg bw/d = milligram per kilogram body weight per day; N = number; % = per cent

No unclassified soft tissue observations were recorded for any of the fetuses in this study.

A spontaneous origin is assumed for the unclassified external observation (Tab. IB-005) and the unclassified skeletal cartilage observations (Tabs. IB-028 – IB-029) which were observed in several fetuses of test groups 0, 1, 2 and 3 (0, 30, 100 and 300 mg/kg bw/d). The distribution and type of these findings do not suggest any relation to treatment.

Finally, fetal examinations revealed that there is no biological meaningful effect of the compound on any morphological structures up to the highest dose tested (300 mg/kg bw/d).

5. DISCUSSION

In a prenatal developmental toxicity study the test substance **DHDPS** was administered to pregnant Wistar rats daily by gavage from implantation to one day prior to the expected day of parturition (GD 6-19) to evaluate its potential maternal and prenatal developmental toxicity.

Analyses confirmed the correctness of the prepared concentrations, the homogeneous distribution and the stability of the test substance in the vehicle.

The test substance caused no mortality nor clinical symptoms of systemic toxicity in any of the exposed groups receiving 30, 100 or 300 mg/kg bw/d **DHDPS**. Some females (7 out of 25) of the high-dose group (300 mg/kg bw/d) showed transient salivation after treatment. This salivation persisted in the respective females for a few minutes immediately after each administration. It is considered to be treatment-related, likely as a result of the bad taste of the test substance/vehicle preparation or due to local irritation of the upper digestive tract. It is not considered to be a sign of systemic toxicity.

The high-dose of the test substance (300 mg/kg bw/d) caused a significant decrease in food consumption (mainly at the beginning of treatment) and body weight gain as well as a distinct decrease in the corrected (net) body weight gain. These effects are considered to be treatment-related and adverse. No toxicologically relevant effect on food consumption and body weight gain was noted for the animals exposed to 30 or 100 mg/kg bw/d **DHDPS**.

No differences of toxicological relevance between the control and the treated groups (30, 100 or 300 mg/kg bw/d) were determined for any reproductive parameters, such as conception rate, mean number of corpora lutea, mean number of implantations, as well as pre- and postimplantation loss. Similarly, no influence of the test substance on fetal weight and sex distribution of the fetuses was noted at any dose.

Overall, there was no evidence for toxicologically relevant adverse effects of the test substance on fetal morphology at any dose.

6. CONCLUSION

Under the conditions of this prenatal developmental toxicity study, the oral administration of **DHDPS** to pregnant Wistar rats from implantation to one day prior to the expected day of parturition (GD 6-19) at a dose of **300 mg/kg bw/d** caused evidence of maternal toxicity, such as reduced food consumption and (net) body weight gain.

In conclusion, the **no observed adverse effect level (NOAEL)** for **maternal toxicity** is **100 mg/kg bw/d**.

There were no toxicologically relevant adverse fetal findings evident. Thus, the **no observed adverse effect level (NOAEL)** for **prenatal developmental toxicity** is **300 mg/kg bw/d**.

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8. APPENDIX

The following list contains abbreviations and definitions generally used in reports for this study type.

This report will not necessarily use all expressions listed below.

8.1. LIST OF ABBREVIATIONS USED IN PART IA, IB, IIA AND IIB

G	=	gram
MEAN	=	mean value
MG/KG BW/D	=	milligram per kilogram body weight per day
N/#/NO.	=	number/number of animals or litters
S.D.	=	standard deviation
%	=	per cent
P	=	significance level
TOTAL	=	total number
0	=	no such finding / observation
-	=	no animals examined

All other abbreviations used are explained in the tables.

10-APR-14	05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) SUMMARY OF MATERNAL CLINICAL OBSERVATIONS DURING GESTATION																				TABLE : 1A-	001	
		DAY OF GESTATION																						
		GROUP#	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	TOTAL
# OF FEMALES EXAMINED		0	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	1	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	2	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	3	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25
NORMAL																								
NOTHING ABNORMAL DETECTED		0	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	1	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	2	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25
	3	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25
ORAL-BUCCAL																								
SALIVATION AFTER TREATMENT		0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	3	0	0	0	0	0	0	0	0	0	0	1	0	1	3	2	3	2	3	2	3	4	0	7

10-APR-14

05R018

TABLE : IA- 002

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

MEAN MATERNAL FOOD CONSUMPTION DURING GESTATION -- GRAMS/ANIMAL/DAY

DAYS	0 TO 1	MEAN S.D. N	TEST GROUP 0 0 MG/KG BW/D			TEST GROUP 1 30 MG/KG BW/D			TEST GROUP 2 100 MG/KG BW/D			TEST GROUP 3 300 MG/KG BW/D		
			14.2 D	1.85	25	12.6*	2.15	25	13.8	1.63	25	12.0**	2.81	24
DAYS	1 TO 3	MEAN S.D. N	16.9 D	1.40	25	16.8	1.91	25	17.0	1.20	25	17.1	1.76	24
DAYS	3 TO 6	MEAN S.D. N	18.3 D	1.25	25	17.8	1.74	25	18.1	1.47	25	18.5	1.47	24
DAYS	6 TO 8	MEAN S.D. N	18.3 D	1.34	25	18.1	1.90	25	18.2	1.52	25	15.7**	1.60	24
DAYS	8 TO 10	MEAN S.D. N	19.3 D	1.70	25	19.0	2.32	25	18.7	1.71	25	16.5**	1.89	24
DAYS	10 TO 13	MEAN S.D. N	20.6 D	1.80	25	20.4	2.19	25	20.7	1.90	25	18.7**	2.02	24
DAYS	13 TO 15	MEAN S.D. N	20.5 D	1.19	25	20.5	2.07	25	20.5	1.94	25	19.5	2.18	24
DAYS	15 TO 17	MEAN S.D. N	21.9 D	1.56	25	21.6	2.17	25	22.0	2.07	25	21.2	1.93	24
DAYS	17 TO 19	MEAN S.D. N	22.2 D	1.74	25	22.6	2.23	25	22.4	1.96	25	21.7	2.44	24
DAYS	19 TO 20	MEAN S.D. N	20.8 D	2.72	25	21.8	2.62	25	21.3	2.24	25	21.0	3.16	24

Statistics: D=Dunnett-test (two-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IA- 003

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
MEAN MATERNAL FOOD CONSUMPTION DURING GESTATION -- GRAMS/ANIMAL/DAY

DAYS	0 TO 6	MEAN OF MEANS S.D. N	TEST GROUP 0 0 MG/KG BW/D			TEST GROUP 1 30 MG/KG BW/D			TEST GROUP 2 100 MG/KG BW/D			TEST GROUP 3 300 MG/KG BW/D		
			16.4	2.07	3	15.7	2.76	3	16.3	2.25	3	15.9	3.42	3
DAYS	6 TO 19	MEAN OF MEANS S.D. N	20.5	1.49	6	20.4	1.63	6	20.4	1.69	6	18.9	2.44	6
DAYS	0 TO 20	MEAN OF MEANS S.D. N	19.3	2.46	10	19.1	2.97	10	19.3	2.66	10	18.2	2.98	10

10-APR-14

05R018

TABLE : IA- 004

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

MEAN MATERNAL BODY WEIGHTS DURING GESTATION -- GRAMS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
DAY 0	MEAN	164.9 D	167.5	168.7	165.6
	S.D.	7.66	12.55	10.28	12.88
	N	25	25	25	24
DAY 1	MEAN	176.6 D	180.0	180.0	177.2
	S.D.	8.22	13.42	9.40	12.05
	N	25	25	25	24
DAY 3	MEAN	185.9 D	189.0	189.2	188.3
	S.D.	8.34	13.68	9.27	12.71
	N	25	25	25	24
DAY 6	MEAN	195.9 D	199.1	199.2	198.3
	S.D.	8.41	13.73	10.19	12.41
	N	25	25	25	24
DAY 8	MEAN	203.7 D	207.5	205.9	205.1
	S.D.	7.93	13.80	10.78	12.27
	N	25	25	25	24
DAY 10	MEAN	213.3 D	216.8	215.4	211.9
	S.D.	8.71	15.18	11.57	13.85
	N	25	25	25	24
DAY 13	MEAN	229.2 D	232.9	230.4	226.4
	S.D.	9.88	16.38	12.29	14.57
	N	25	25	25	24
DAY 15	MEAN	239.3 D	243.5	240.6	236.1
	S.D.	9.38	16.95	14.08	15.71
	N	25	25	25	24
DAY 17	MEAN	257.5 D	262.8	259.5	253.6
	S.D.	11.23	18.25	15.82	16.40
	N	25	25	25	24
DAY 19	MEAN	281.1 D	288.9	283.4	276.9
	S.D.	11.38	22.62	18.09	18.57
	N	25	25	25	24
DAY 20	MEAN	295.9 D	302.4	297.8	291.0
	S.D.	13.47	23.30	19.30	19.85
	N	25	25	25	24

Statistics: D=Dunnett-test (two-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IA- 005

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

MEAN MATERNAL BODY WEIGHT CHANGE DURING GESTATION -- GRAMS

DAYS	0 TO 1	MEAN S.D. N	TEST GROUP 0 0 MG/KG BW/D			TEST GROUP 1 30 MG/KG BW/D			TEST GROUP 2 100 MG/KG BW/D			TEST GROUP 3 300 MG/KG BW/D		
			MEAN	S.D.	N	MEAN	S.D.	N	MEAN	S.D.	N	MEAN	S.D.	N
DAYS	0 TO 1		11.7 D	3.17	25	12.5	3.60	25	11.3	3.66	25	11.5	7.52	24
DAYS	1 TO 3		9.4 D	2.29	25	9.0	2.63	25	9.2	2.64	25	11.2	4.71	24
DAYS	3 TO 6		10.0 D	2.36	25	10.1	2.38	25	10.0	2.19	25	10.0	2.44	24
DAYS	6 TO 8		7.8 D	4.27	25	8.5	3.70	25	6.8	4.00	25	6.8	3.33	24
DAYS	8 TO 10		9.6 D	3.96	25	9.3	3.42	25	9.4	2.94	25	6.8*	3.81	24
DAYS	10 TO 13		15.9 D	3.59	25	16.1	3.37	25	15.1	3.23	25	14.5	3.40	24
DAYS	13 TO 15		10.1 D	2.70	25	10.6	3.71	25	10.1	3.71	25	9.7	3.99	24
DAYS	15 TO 17		18.1 D	3.08	25	19.4	2.98	25	18.9	3.83	25	17.5	3.45	24
DAYS	17 TO 19		23.7 D	5.58	25	26.0	5.41	25	24.0	4.22	25	23.3	3.83	24
DAYS	19 TO 20		14.8 D	5.02	25	13.5	4.21	25	14.4	4.03	25	14.1	3.77	24

Statistics: D=Dunnett-test (two-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IA- 006

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

MEAN MATERNAL BODY WEIGHT CHANGE DURING GESTATION -- GRAMS

		TEST GROUP 0			TEST GROUP 1			TEST GROUP 2			TEST GROUP 3		
		0 MG/KG BW/D			30 MG/KG BW/D			100 MG/KG BW/D			300 MG/KG BW/D		
DAYS 0 TO 6	MEAN	31.0 D			31.5			30.4			32.7		
	S.D.	4.49			4.97			4.90			5.37		
	N	25			25			25			24		
DAYS 6 TO 19	MEAN	85.2 D			89.8			84.3			78.6*		
	S.D.	7.27			11.88			9.59			8.61		
	N	25			25			25			24		
DAYS 0 TO 20	MEAN	131.0 D			134.9			129.1			125.4		
	S.D.	11.53			17.21			13.74			13.37		
	N	25			25			25			24		

Statistics: D=Dunnett-test (two-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IA- 007

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
MEAN GRAVID UTERINE WEIGHTS AND NET MATERNAL BODY WEIGHT CHANGE -- GRAMS

		TEST GROUP 0	TEST GROUP 1	TEST GROUP 2	TEST GROUP 3
		0 MG/KG BW/D	30 MG/KG BW/D	100 MG/KG BW/D	300 MG/KG BW/D
GRAVID UTERUS	MEAN	59.1 D	59.6	58.7	55.8
	S.D.	7.50	9.93	9.70	8.64
	N	25	25	25	24
CARCASS	MEAN	236.8 D	242.8	239.1	235.2
	S.D.	9.32	16.56	13.68	17.38
	N	25	25	25	24
NET WEIGHT CHANGE FROM DAY 6	MEAN	40.9 D	43.7	40.0	36.9
	S.D.	5.11	8.76	7.18	7.99
	N	25	25	25	24

Statistics: D=Dunnett-test (two-sided)
* : p<=0.05 ** : p<=0.01

CARCASS WEIGHT = TERMINAL BODY WEIGHT MINUS UTERINE WEIGHT
NET WEIGHT CHANGE FROM DAY 6 = CARCASS WEIGHT MINUS DAY 6 BODY WEIGHT

10-APR-14

05R018

TABLE : 1A- 008

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF MATERNAL NECROPSY OBSERVATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
FEMALES EXAMINED	N	25	25	25	25
NOTHING ABNORMAL DETECTED	N	25	25	25	22
	%	100	100	100	88
DIAPHRAGMATIC HERNIA	N	0	0	0	1
	%	0.0	0.0	0.0	4.0
DILATED RENAL PELVIS	N	0	0	0	1
	%	0.0	0.0	0.0	4.0
HEMOMETRA	N	0	0	0	1
	%	0.0	0.0	0.0	4.0

10-APR-14

05R018

TABLE : IA- 009

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF REPRODUCTION DATA

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Females Mated	N	25	25	25	25
Pregnant	N	25	25	25	24
Conception Rate	%	100	100	100	96
Aborted	N	0	0	0	0
Premature Births	N	0	0	0	0
Dams with Viable Fetuses	N	25	25	25	24
Dams with all Resorptions	N	0	0	0	0
Female Mortality	N	0	0	0	0
	%	0.0	0.0	0.0	0.0
Pregnant at Terminal Sacrifice	N	25	25	25	24
	%	100	100	100	96
Corpora Lutea	MEAN	11.5 D	11.8	11.7	11.4
	S.D.	1.23	1.80	1.28	1.35
	TOTAL	287	295	293	274
Implantation Sites	MEAN	11.1 D	11.0	11.1	10.8
	S.D.	1.55	1.86	1.89	1.69
	TOTAL	277	276	277	259
Preimplantation Loss	MEAN%	3.6 D	6.1	5.4	5.3
	S.D.	7.60	9.43	12.33	10.94
Postimplantation Loss	MEAN%	4.7 D	3.9	3.9	6.3
	S.D.	5.95	5.82	5.71	6.47

Statistics: D=Dunnett-test (two-sided) Fi =Fisher's exact test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IA- 010

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF REPRODUCTION DATA

	TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Pregnant at Terminal Sacrifice	25	25	25	24
Resorptions: Total				
MEAN	0.5 D	0.4	0.5	0.7
S.D.	0.65	0.65	0.71	0.70
TOTAL	13	11	12	16
MEAN%	4.7 D	3.9	3.9	6.3
S.D.	5.95	5.82	5.71	6.47
Early				
MEAN	0.5 D	0.4	0.4	0.6
S.D.	0.65	0.65	0.65	0.71
TOTAL	12	11	10	15
MEAN%	4.4 D	3.9	3.3	5.7
S.D.	5.97	5.82	5.22	6.45
Late				
MEAN	0.0 D	0.0	0.1	0.0
S.D.	0.20	0.00	0.28	0.20
TOTAL	1	0	2	1
MEAN%	0.3 D	0.0	0.6	0.5
S.D.	1.67	0.00	2.22	2.55
Dead Fetuses				
N	0	0	0	0

Statistics: D=Dunnett-test (two-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IA- 011

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF REPRODUCTION DATA

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Dams with Viable Fetuses	N	25	25	25	24
Live Fetuses	MEAN	10.6 D	10.6	10.6	10.1
	S.D.	1.61	1.83	1.71	1.80
	TOTAL	264	265	265	243
	MEAN%	95.3 D	96.1	96.1	93.7
	S.D.	5.95	5.82	5.71	6.47
Females	MEAN	5.2 D	5.0	6.0	4.8
	S.D.	1.97	1.63	2.10	1.79
	TOTAL	129	125	150	115
	MEAN%	45.9 D	45.2	54.0	44.6
	S.D.	16.06	11.71	17.82	15.91
Males	MEAN	5.4 D	5.6	4.6	5.3
	S.D.	1.68	1.73	1.66	1.97
	TOTAL	135	140	115	128
	MEAN%	49.4 D	50.9	42.1	49.1
	S.D.	16.39	13.20	15.49	15.89
PER CENT LIVE FEMALES		48.9	47.2	56.6	47.3
PER CENT LIVE MALES		51.1	52.8	43.4	52.7

Statistics: D=Dunnett-test (two-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 001

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

MEAN PLACENTAL AND FETAL BODY WEIGHTS (ON A LITTER BASIS)

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
PLACENTAL WEIGHTS UNITS: GRAMS	of all Viable Fetuses	MEAN S.D. N	0.45 D 0.036 25	0.46 0.048 25	0.45 0.041 25
	of Male Fetuses	MEAN S.D. N	0.47 D 0.036 25	0.47 0.055 25	0.46 0.048 25
	of Female Fetuses	MEAN S.D. N	0.44 D 0.043 25	0.45 0.045 25	0.45 0.043 25
FETAL WEIGHTS UNITS: GRAMS	of all Viable Fetuses	MEAN S.D. N	3.6 D 0.22 25	3.6 0.15 25	3.4 0.21 24
	of Male Fetuses	MEAN S.D. N	3.6 D 0.21 25	3.7 0.16 25	3.5 0.21 24
	of Female Fetuses	MEAN S.D. N	3.5 D 0.23 25	3.5 0.19 25	3.3 0.23 24

Statistics: D=Dunnett-test (two-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 002

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF ALL CLASSIFIED FETAL EXTERNAL OBSERVATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	264	265	265	243
Live	N	264	265	265	243
Dead	N	0	0	0	0
TOTAL MALFORMATIONS					
Fetal Incidence	N	0	0	1	0
	%	0.0	0.0	0.4	0.0
Litter Incidence	N	OFI	0	1	0
	%	0.0	0.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.0W1	0.0	0.8	0.0
	S.D.	0.00	0.00	4.00	0.00
TOTAL VARIATIONS					
Fetal Incidence	N	0	0	0	0
	%	0.0	0.0	0.0	0.0
Litter Incidence	N	OFI	0	0	0
	%	0.0	0.0	0.0	0.0
Affected Fetuses/Litter	MEAN%	0.0W1	0.0	0.0	0.0
	S.D.	0.00	0.00	0.00	0.00

Statistics: F1 =Fisher's exact test (one-sided) W1 =Wilcoxon-test (one-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 003

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL EXTERNAL MALFORMATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	264	265	265	243
Live	N	264	265	265	243
Dead	N	0	0	0	0
FETUS WITH MULTIPLE EXTERNAL MALFORMATIONS					
Fetal Incidence	N	0	0	1	0
	%	0.0	0.0	0.4	0.0
Litter Incidence	N	0Fi	0	1	0
	%	0.0	0.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.0Wi	0.0	0.8	0.0
	S.D.	0.00	0.00	4.00	0.00
TOTAL FETAL EXTERNAL MALFORMATIONS					
Fetal Incidence	N	0	0	1	0
	%	0.0	0.0	0.4	0.0
Litter Incidence	N	0Fi	0	1	0
	%	0.0	0.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.0Wi	0.0	0.8	0.0
	S.D.	0.00	0.00	4.00	0.00

Statistics: Fi =Fisher's exact test (one-sided) Wi =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 004

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL EXTERNAL VARIATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	264	265	265	243
Live	N	264	265	265	243
Dead	N	0	0	0	0
TOTAL FETAL EXTERNAL VARIATIONS					
Fetal Incidence	N	0	0	0	0
	%	0.0	0.0	0.0	0.0
Litter Incidence	N	0	0	0	0
	%	0.0	0.0	0.0	0.0
Affected Fetuses/Litter	MEAN%	0.0W1	0.0	0.0	0.0
	S.D.	0.00	0.00	0.00	0.00

Statistics: F1 =Fisher's exact test (one-sided) W1 =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 005

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL EXTERNAL UNCLASSIFIED OBSERVATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	264	265	265	243
Live	N	264	265	265	243
Dead	N	0	0	0	0
BLOOD COAGULUM AROUND PLACENTA					
Fetal Incidence	N	0	0	0	2
	%	0.0	0.0	0.0	0.8
Litter Incidence	N	OFI	0	0	2
	%	0.0	0.0	0.0	8.3
Affected Fetuses/Litter	MEAN%	0.0Wi	0.0	0.0	1.3
	S.D.	0.00	0.00	0.00	4.38
TOTAL FETAL EXTERNAL UNCLASSIFIED OBSERVATIONS					
Fetal Incidence	N	0	0	0	2
	%	0.0	0.0	0.0	0.8
Litter Incidence	N	OFI	0	0	2
	%	0.0	0.0	0.0	8.3
Affected Fetuses/Litter	MEAN%	0.0Wi	0.0	0.0	1.3
	S.D.	0.00	0.00	0.00	4.38

Statistics: Fi =Fisher's exact test (one-sided) Wi =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 006

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF ALL CLASSIFIED FETAL SOFT TISSUE OBSERVATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	127	128	125	116
Live	N	127	128	125	116
Dead	N	0	0	0	0
TOTAL MALFORMATIONS					
Fetal Incidence	N %	0 0.0	0 0.0	0 0.0	0 0.0
Litter Incidence	N %	0 0.0	0 0.0	0 0.0	0 0.0
Affected Fetuses/Litter	MEAN% S.D.	0.0W1 0.00	0.0 0.00	0.0 0.00	0.0 0.00
TOTAL VARIATIONS					
Fetal Incidence	N %	7 5.5	5 3.9	12 9.6	10 8.6
Litter Incidence	N %	7F1 28	4 16	9 36	9 38
Affected Fetuses/Litter	MEAN% S.D.	6.1W1 10.35	4.1 10.18	9.1 15.08	8.5 11.95

Statistics: F1 =Fisher's exact test (one-sided) W1 =Wilcoxon-test (one-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 007

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SOFT TISSUE MALFORMATIONS

		TEST GROUP 0	TEST GROUP 1	TEST GROUP 2	TEST GROUP 3
		0 MG/KG BW/D	30 MG/KG BW/D	100 MG/KG BW/D	300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	127	128	125	116
Live	N	127	128	125	116
Dead	N	0	0	0	0
TOTAL FETAL SOFT TISSUE MALFORMATIONS					
Fetal Incidence	N	0	0	0	0
	%	0.0	0.0	0.0	0.0
Litter Incidence	N	0	0	0	0
	%	0.0	0.0	0.0	0.0
Affected Fetuses/Litter	MEAN%	0.0	0.0	0.0	0.0
	S.D.	0.00	0.00	0.00	0.00

Statistics: F1 =Fisher's exact test (one-sided) W1 =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 008

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SOFT TISSUE VARIATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	127	128	125	116
Live	N	127	128	125	116
Dead	N	0	0	0	0
SHORT INNOMINATE Fetal Incidence	N %	1 0.8	2 1.6	2 1.6	1 0.9
Litter Incidence	N %	1Fi 4.0	2 8.0	2 8.0	1 4.2
Affected Fetuses/Litter	MEAN% S.D.	0.8Wi 4.00	1.6 5.54	1.5 5.10	1.0 5.10
DILATED RENAL PELVIS Fetal Incidence	N %	6 4.7	3 2.3	10 8.0	9 7.8
Litter Incidence	N %	6Fi 24	3 12	8 32	8 33
Affected Fetuses/Litter	MEAN% S.D.	5.3Wi 10.00	2.5 6.92	7.6 13.83	7.4 11.53
DILATED URETER Fetal Incidence	N %	1 0.8	1 0.8	1 0.8	3 2.6
Litter Incidence	N %	1Fi 4.0	1 4.0	1 4.0	3 13
Affected Fetuses/Litter	MEAN% S.D.	0.8Wi 4.00	1.0 5.00	0.7 3.33	2.4 6.41

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 009

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SOFT TISSUE VARIATIONS

	TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
TOTAL FETAL SOFT TISSUE VARIATIONS				
Fetal Incidence	N 7 5.5	5 3.9	12 9.6	10 8.6
Litter Incidence	N 7Fi 28	4 16	9 36	9 38
Affected Fetuses/Litter	MEAN% 6.1W1 S.D. 10.35	4.1 10.18	9.1 15.08	8.5 11.95

Statistics: Fi = Fisher's exact test (one-sided) W1 = Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 010

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SOFT TISSUE UNCLASSIFIED OBSERVATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	127	128	125	116
Live	N	127	128	125	116
Dead	N	0	0	0	0
TOTAL FETAL SOFT TISSUE UNCLASSIFIED OBSERVATIONS					
Fetal Incidence	N	0	0	0	0
	%	0.0	0.0	0.0	0.0
Litter Incidence	N	0	0	0	0
	%	0.0	0.0	0.0	0.0
Affected Fetuses/Litter	MEAN%	0.0W1	0.0	0.0	0.0
	S.D.	0.00	0.00	0.00	0.00

Statistics: F1 =Fisher's exact test (one-sided) W1 =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 011

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF ALL CLASSIFIED FETAL SKELETAL OBSERVATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
TOTAL MALFORMATIONS					
Fetal Incidence	N %	1 0.7	0 0.0	3 2.1	5 3.9
Litter Incidence	N %	1Fi 4.0	0 0.0	3 12	5 21
Affected Fetuses/Litter	MEAN% S.D.	0.7Wi 3.33	0.0 0.00	2.8 8.15	4.3* 8.85
TOTAL VARIATIONS					
Fetal Incidence	N %	136 99	135 99	139 99	127 100
Litter Incidence	N %	25Fi 100	25 100	25 100	24 100
Affected Fetuses/Litter	MEAN% S.D.	99.2Wi 4.00	98.3 5.89	98.7 6.67	100.0 0.00

Statistics: Fi =Fisher's exact test (one-sided) Wi =Wilcoxon-test (one-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 012

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL MALFORMATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
FETUS WITH MULTIPLE SKELETAL MALFORMATIONS					
Fetal Incidence	N	0	0	1	0
	%	0.0	0.0	0.7	0.0
Litter Incidence	N	OFI	0	1	0
	%	0.0	0.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.0WI	0.0	1.3	0.0
	S.D.	0.00	0.00	6.67	0.00
MISSHAPEN BASISPHEOID					
Fetal Incidence	N	0	0	0	1
	%	0.0	0.0	0.0	0.8
Litter Incidence	N	OFI	0	0	1
	%	0.0	0.0	0.0	4.2
Affected Fetuses/Litter	MEAN%	0.0WI	0.0	0.0	0.8
	S.D.	0.00	0.00	0.00	4.08
SHORTENED SCAPULA					
Fetal Incidence	N	0	0	1	0
	%	0.0	0.0	0.7	0.0
Litter Incidence	N	OFI	0	1	0
	%	0.0	0.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.0WI	0.0	0.7	0.0
	S.D.	0.00	0.00	3.33	0.00

Statistics: Fi =Fisher's exact test (one-sided) Wi =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 013

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL MALFORMATIONS

		TEST GROUP 0	TEST GROUP 1	TEST GROUP 2	TEST GROUP 3
		0 MG/KG BW/D	30 MG/KG BW/D	100 MG/KG BW/D	300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
MALPOSITIONED AND BIPARTITE STERNEBRA; Unchanged cartilage					
Fetal Incidence	N	0	0	0	2
	%	0.0	0.0	0.0	1.6
Litter Incidence	N	0Fi	0	0	2
	%	0.0	0.0	0.0	8.3
Affected Fetuses/Litter	MEAN%	0.0Wi	0.0	0.0	2.1
	S.D.	0.00	0.00	0.00	7.06
SHORTENED HUMERUS					
Fetal Incidence	N	1	0	1	1
	%	0.7	0.0	0.7	0.8
Litter Incidence	N	1Fi	0	1	1
	%	4.0	0.0	4.0	4.2
Affected Fetuses/Litter	MEAN%	0.7Wi	0.0	0.8	0.8
	S.D.	3.33	0.00	4.00	4.08
MISSHAPEN TUBEROSITAS DELTOIDEA					
Fetal Incidence	N	0	0	0	1
	%	0.0	0.0	0.0	0.8
Litter Incidence	N	0Fi	0	0	1
	%	0.0	0.0	0.0	4.2
Affected Fetuses/Litter	MEAN%	0.0Wi	0.0	0.0	0.6
	S.D.	0.00	0.00	0.00	2.92

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 014

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL MALFORMATIONS

	TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
TOTAL FETAL SKELETAL MALFORMATIONS				
Fetal Incidence	1 0.7	0 0.0	3 2.1	5 3.9
Litter Incidence	1Fi 4.0	0 0.0	3 12	5 21
Affected Fetuses/Litter	0.7Wi 3.33	0.0 0.00	2.8 8.15	4.3* 8.85

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)

* : $p \leq 0.05$ ** : $p \leq 0.01$

10-APR-14

05R018

TABLE : IB- 015

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
SUPRAOCCIPITAL HOLE(S)					
Fetal Incidence	N	1	0	1	0
	%	0.7	0.0	0.7	0.0
Litter Incidence	N	1Fi	0	1	0
	%	4.0	0.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.8Wi	0.0	0.7	0.0
	S.D.	4.00	0.00	3.33	0.00
INCOMPLETE OSSIFICATION OF BASISPHENOID					
Fetal Incidence	N	30	36	37	25
	%	22	26	26	20
Litter Incidence	N	16Fi	15	19	15
	%	64	60	76	63
Affected Fetuses/Litter	MEAN%	20.9Wi	25.4	26.2	18.9
	S.D.	20.87	28.54	22.84	18.70
INCOMPLETE OSSIFICATION OF PARIETAL; Unchanged cartilage					
Fetal Incidence	N	13	19	15	18
	%	9.5	14	11	14
Litter Incidence	N	9Fi	11	12	11
	%	36	44	48	46
Affected Fetuses/Litter	MEAN%	9.7Wi	14.6	11.3	13.4
	S.D.	15.51	19.11	13.40	16.13

Statistics: Fi =Fisher's exact test (one-sided) Wi =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 016

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage					
Fetal Incidence	N	48	36	37	41
	%	35	26	26	32
Litter Incidence	N	21Fi	17	18	18
	%	84	68	72	75
Affected Fetuses/Litter	MEAN%	34.3Wi	26.0	26.8	31.4
	S.D.	22.50	22.11	22.82	24.06
INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage					
Fetal Incidence	N	48	50	52	59
	%	35	36	37	46
Litter Incidence	N	20Fi	20	23	21
	%	80	80	92	88
Affected Fetuses/Litter	MEAN%	34.1Wi	35.2	37.6	45.2*
	S.D.	29.65	23.55	22.14	25.62
INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage					
Fetal Incidence	N	15	21	15	18
	%	11	15	11	14
Litter Incidence	N	8Fi	11	10	15*
	%	32	44	40	63
Affected Fetuses/Litter	MEAN%	11.3Wi	15.9	11.1	14.1
	S.D.	19.53	23.76	17.45	12.58

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 017

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
INCOMPLETE OSSIFICATION OF NASAL; Unchanged cartilage					
Fetal Incidence	N	1	1	1	0
	%	1.5	0.7	0.7	0.0
Litter Incidence	N	2Fi	1	1	0
	%	8.0	4.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	1.5Wi	1.0	0.7	0.0
	S.D.	5.10	5.00	3.33	0.00
UNOSSIFIED HYOID; Cartilage present					
Fetal Incidence	N	0	0	1	0
	%	0.0	0.0	0.7	0.0
Litter Incidence	N	0Fi	0	1	0
	%	0.0	0.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.0Wi	0.0	0.8	0.0
	S.D.	0.00	0.00	4.00	0.00
BASIOCCIPITAL HOLE (S)					
Fetal Incidence	N	0	1	0	0
	%	0.0	0.7	0.0	0.0
Litter Incidence	N	0Fi	1	0	0
	%	0.0	4.0	0.0	0.0
Affected Fetuses/Litter	MEAN%	0.0Wi	0.7	0.0	0.0
	S.D.	0.00	3.33	0.00	0.00

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 018

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
INCOMPLETE OSSIFICATION OF FRONTAL; Unchanged cartilage					
Fetal Incidence	N	0	1	1	0
	%	0.0	0.7	0.7	0.0
Litter Incidence	N	0Fi	1	1	0
	%	0.0	4.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.0Wi	0.8	0.7	0.0
	S.D.	0.00	4.00	3.33	0.00
INCOMPLETE OSSIFICATION OF HYOID; Cartilage present					
Fetal Incidence	N	1	0	1	0
	%	0.7	0.0	0.7	0.0
Litter Incidence	N	1Fi	0	1	0
	%	4.0	0.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.7Wi	0.0	0.6	0.0
	S.D.	3.33	0.00	2.86	0.00
INCOMPLETE OSSIFICATION OF CERVICAL ARCH; Cartilage present					
Fetal Incidence	N	0	1	0	0
	%	0.0	0.7	0.0	0.0
Litter Incidence	N	0Fi	1	0	0
	%	0.0	4.0	0.0	0.0
Affected Fetuses/Litter	MEAN%	0.0Wi	1.0	0.0	0.0
	S.D.	0.00	5.00	0.00	0.00

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 019

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0	TEST GROUP 1	TEST GROUP 2	TEST GROUP 3
		0 MG/KG BW/D	30 MG/KG BW/D	100 MG/KG BW/D	300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage					
Fetal Incidence	N	1	4	0	7
	%	0.7	2.9	0.0	5.5
Litter Incidence	N	1Fi	3	0	7*
	%	4.0	12	0.0	29
Affected Fetuses/Litter	MEAN%	0.7Wi	3.0	0.0	5.6**
	S.D.	3.33	8.63	0.00	8.88
DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum					
Fetal Incidence	N	3	2	2	0
	%	2.2	1.5	1.4	0.0
Litter Incidence	N	3Fi	1	2	0
	%	12	4.0	8.0	0.0
Affected Fetuses/Litter	MEAN%	2.5Wi	1.3	1.2	0.0
	S.D.	6.92	6.67	4.30	0.00
INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage					
Fetal Incidence	N	4	2	3	0
	%	2.9	1.5	2.1	0.0
Litter Incidence	N	4Fi	2	3	0
	%	16	8.0	12	0.0
Affected Fetuses/Litter	MEAN%	2.9Wi	1.7	3.1	0.0
	S.D.	6.99	5.89	8.88	0.00

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 020

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum					
Fetal Incidence	N	0	1	0	0
	%	0.0	0.7	0.0	0.0
Litter Incidence	N	0Fi	1	0	0
	%	0.0	4.0	0.0	0.0
Affected Fetuses/Litter	MEAN%	0.0Wi	0.7	0.0	0.0
	S.D.	0.00	3.33	0.00	0.00
SUPERNUMERARY THORACIC VERTEBRA					
Fetal Incidence	N	17	10	5	4
	%	12	7.3	3.6	3.1
Litter Incidence	N	11Fi	6	5	2
	%	44	24	20	8.3
Affected Fetuses/Litter	MEAN%	12.6Wi	7.1	3.7	3.2
	S.D.	17.59	15.53	7.64	12.57
INCOMPLETE OSSIFICATION OF THORACIC ARCH; Cartilage present					
Fetal Incidence	N	0	1	1	0
	%	0.0	0.7	0.7	0.0
Litter Incidence	N	0Fi	1	1	0
	%	0.0	4.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.0Wi	1.0	0.7	0.0
	S.D.	0.00	5.00	3.33	0.00

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 021

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
BIPARTITE OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum					
Fetal Incidence	N	1	0	0	0
	%	0.7	0.0	0.0	0.0
Litter Incidence	N	1Fi	0	0	0
	%	4.0	0.0	0.0	0.0
Affected Fetuses/Litter	MEAN%	0.8Wi	0.0	0.0	0.0
	S.D.	4.00	0.00	0.00	0.00
INCOMPLETE OSSIFICATION OF LUMBAR ARCH; Cartilage present					
Fetal Incidence	N	0	0	1	0
	%	0.0	0.0	0.7	0.0
Litter Incidence	N	0Fi	0	1	0
	%	0.0	0.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.0Wi	0.0	0.7	0.0
	S.D.	0.00	0.00	3.33	0.00
INCOMPLETE OSSIFICATION OF SACRAL ARCH; Cartilage present					
Fetal Incidence	N	1	2	1	1
	%	0.7	1.5	0.7	0.8
Litter Incidence	N	1Fi	2	1	1
	%	4.0	8.0	4.0	4.2
Affected Fetuses/Litter	MEAN%	0.7Wi	1.3	0.7	0.7
	S.D.	3.33	4.61	3.33	3.40

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 022

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
MISSHAPEN SACRAL VERTEBRA					
Fetal Incidence	N	9	7	2	1
	%	6.6	5.1	1.4	0.8
Litter Incidence	N	6Fi	6	2	1
	%	24	24	8.0	4.2
Affected Fetuses/Litter	MEAN%	6.8Wi	5.0	1.6	1.0
	S.D.	13.73	9.71	5.54	5.10
UNOSSIFIED STERNEBRA; Unchanged cartilage					
Fetal Incidence	N	2	7	7	13
	%	1.5	5.1	5.0	10
Litter Incidence	N	2Fi	6	5	9*
	%	8.0	24	20	38
Affected Fetuses/Litter	MEAN%	1.5Wi	5.0	4.6	11.0**
	S.D.	5.10	10.04	10.12	17.18
INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage					
Fetal Incidence	N	124	120	131	121
	%	91	88	94	95
Litter Incidence	N	25Fi	25	25	24
	%	100	100	100	100
Affected Fetuses/Litter	MEAN%	90.4Wi	86.0	93.3	94.9
	S.D.	18.02	21.13	11.65	10.48

Statistics: Fi =Fisher's exact test (one-sided) Wi =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 023

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0	TEST GROUP 1	TEST GROUP 2	TEST GROUP 3
		0 MG/KG BW/D	30 MG/KG BW/D	100 MG/KG BW/D	300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
MISSHAPEN STERNEBRA; Unchanged cartilage					
Fetal Incidence	N	56	72	68	55
	%	41	53	49	43
Litter Incidence	N	24Fi	24	23	22
	%	96	96	92	92
Affected Fetuses/Litter	MEAN%	41.8Wi	51.8	47.9	42.7
	S.D.	20.99	22.31	26.87	24.74
UNILATERAL OSSIFICATION OF STERNEBRA; Unchanged cartilage					
Fetal Incidence	N	1	0	3	1
	%	0.7	0.0	2.1	0.8
Litter Incidence	N	1Fi	0	2	1
	%	4.0	0.0	8.0	4.2
Affected Fetuses/Litter	MEAN%	0.7Wi	0.0	2.1	0.8
	S.D.	3.33	0.00	7.63	4.08
BIPARTITE OSSIFICATION OF STERNEBRA; Unchanged cartilage					
Fetal Incidence	N	0	1	1	2
	%	0.0	0.7	0.7	1.6
Litter Incidence	N	0Fi	1	1	2
	%	0.0	4.0	4.0	8.3
Affected Fetuses/Litter	MEAN%	0.0Wi	0.7	0.7	1.5
	S.D.	0.00	3.33	3.33	5.20

Statistics: Fi =Fisher's exact test (one-sided) Wi =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 024

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

	TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N 25	25	25	24
Fetuses Evaluated	N 137	137	140	127
Live	N 137	137	140	127
Dead	N 0	0	0	0
SUPERNUMERARY RIB (14TH) ; Cartilage present				
Fetal Incidence	N 12	7	8	3
	% 8.8	5.1	5.7	2.4
Litter Incidence	N 8Fi	6	8	2
	% 32	24	32	8.3
Affected Fetuses/Litter	MEAN% 9.7Wi	5.3	6.1	3.1
	S.D. 19.18	10.00	9.17	11.21
SUPERNUMERARY RIB (14TH) ; Cartilage not present				
Fetal Incidence	N 91	78	72	67
	% 66	57	51	53
Litter Incidence	N 23Fi	25	24	22
	% 92	100	96	92
Affected Fetuses/Litter	MEAN% 66.6Wi	57.6	51.7	53.8
	S.D. 28.85	25.55	26.32	30.50
CERVICAL RIB; Cartilage present				
Fetal Incidence	N 0	0	0	1
	% 0.0	0.0	0.0	0.8
Litter Incidence	N 0Fi	0	0	1
	% 0.0	0.0	0.0	4.2
Affected Fetuses/Litter	MEAN% 0.0Wi	0.0	0.0	1.0
	S.D. 0.00	0.00	0.00	5.10

Statistics: Fi =Fisher's exact test (one-sided) Wi =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 025

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
CERVICAL RIB; Cartilage not present					
Fetal Incidence	N	1	3	3	3
	%	0.7	2.2	2.1	2.4
Litter Incidence	N	1Fi	3	3	3
	%	4.0	12	12	13
Affected Fetuses/Litter	MEAN%	0.7Wi	2.5	2.8	2.7
	S.D.	3.33	6.92	8.15	7.37
WAVY RIB					
Fetal Incidence	N	16	15	12	17
	%	12	11	8.6	13
Litter Incidence	N	8Fi	10	6	12
	%	32	40	24	50
Affected Fetuses/Litter	MEAN%	12.1Wi	12.2	8.5	13.8
	S.D.	23.00	17.18	19.27	17.01
INCOMPLETE OSSIFICATION OF TUBEROSITAS DELTOIDEA; Cartilage present					
Fetal Incidence	N	1	0	0	2
	%	0.7	0.0	0.0	1.6
Litter Incidence	N	1Fi	0	0	2
	%	4.0	0.0	0.0	8.3
Affected Fetuses/Litter	MEAN%	0.7Wi	0.0	0.0	1.7
	S.D.	3.33	0.00	0.00	5.65

Statistics: Fi =Fisher's exact test (one-sided) Wi =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 026

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

		TEST GROUP 0	TEST GROUP 1	TEST GROUP 2	TEST GROUP 3
		0 MG/KG BW/D	30 MG/KG BW/D	100 MG/KG BW/D	300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
INCOMPLETE OSSIFICATION OF METACARPAL; Cartilage present					
Fetal Incidence	N	1	1	1	0
	%	0.7	0.7	0.7	0.0
Litter Incidence					
	N	1Fi	1	1	0
	%	4.0	4.0	4.0	0.0
Affected Fetuses/Litter					
	MEAN%	0.7Wi	1.0	0.7	0.0
	S.D.	3.33	5.00	3.33	0.00
INCOMPLETE OSSIFICATION OF PUBIS; Cartilage present					
Fetal Incidence	N	0	1	3	2
	%	0.0	0.7	2.1	1.6
Litter Incidence					
	N	0Fi	1	3	2
	%	0.0	4.0	12	8.3
Affected Fetuses/Litter					
	MEAN%	0.0Wi	0.8	2.0*	1.7
	S.D.	0.00	4.00	5.53	5.65
INCOMPLETE OSSIFICATION OF ISCHIUM; Cartilage present					
Fetal Incidence	N	0	0	3	2
	%	0.0	0.0	2.1	1.6
Litter Incidence					
	N	0Fi	0	3	2
	%	0.0	0.0	12	8.3
Affected Fetuses/Litter					
	MEAN%	0.0Wi	0.0	2.0*	1.7
	S.D.	0.00	0.00	5.53	5.65

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 027

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL VARIATIONS

	TEST GROUP 0 0 MG/KG BW/D	TEST GROUP 1 30 MG/KG BW/D	TEST GROUP 2 100 MG/KG BW/D	TEST GROUP 3 300 MG/KG BW/D
TOTAL FETAL SKELETAL VARIATIONS				
Fetal Incidence	N 136 99	N 135 99	N 139 99	N 127 100
Litter Incidence	N 25F1 100	N 25 100	N 25 100	N 24 100
Affected Fetuses/Litter	MEAN% 99.2W1 S.D. 4.00	MEAN% 98.3 S.D. 5.89	MEAN% 98.7 S.D. 6.67	MEAN% 100.0 S.D. 0.00

Statistics: F1 = Fisher's exact test (one-sided) W1 = Wilcoxon-test (one-sided)

* : $p \leq 0.05$ ** : $p \leq 0.01$

10-APR-14

05R018

TABLE : IB- 028

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL UNCLASS. CARTILAGE OBS.

		TEST GROUP 0	TEST GROUP 1	TEST GROUP 2	TEST GROUP 3
		0 MG/KG BW/D	30 MG/KG BW/D	100 MG/KG BW/D	300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
NOTCHED CARTILAGE BETWEEN BASIPHENOID AND BASIOCCIPITAL					
Fetal Incidence	N	1	6	5	1
	%	0.7	4.4	3.6	0.8
Litter Incidence	N	1Fi	3	2	1
	%	4.0	12	8.0	4.2
Affected Fetuses/Litter	MEAN%	0.7Wi	4.0	3.1	0.7
	S.D.	3.33	12.06	11.95	3.40
BIPARTITE PROCESSUS XIPHOIDEUS					
Fetal Incidence	N	96	89	95	88
	%	70	65	68	69
Litter Incidence	N	24Fi	25	24	24
	%	96	100	96	100
Affected Fetuses/Litter	MEAN%	70.0Wi	63.8	66.7	68.2
	S.D.	23.79	26.32	26.12	25.72
FUSED RIB CARTILAGE					
Fetal Incidence	N	0	0	1	0
	%	0.0	0.0	0.7	0.0
Litter Incidence	N	0Fi	0	1	0
	%	0.0	0.0	4.0	0.0
Affected Fetuses/Litter	MEAN%	0.0Wi	0.0	0.7	0.0
	S.D.	0.00	0.00	3.33	0.00

Statistics: Fi = Fisher's exact test (one-sided) Wi = Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 029

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF FETAL SKELETAL UNCLASS. CARTILAGE OBS.

		TEST GROUP 0	TEST GROUP 1	TEST GROUP 2	TEST GROUP 3
		0 MG/KG BW/D	30 MG/KG BW/D	100 MG/KG BW/D	300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	137	137	140	127
Live	N	137	137	140	127
Dead	N	0	0	0	0
BRANCHED RIB CARTILAGE					
Fetal Incidence	N	2	0	0	1
	%	1.5	0.0	0.0	0.8
Litter Incidence	N	2F1	0	0	1
	%	8.0	0.0	0.0	4.2
Affected Fetuses/Litter	MEAN%	1.5W1	0.0	0.0	0.7
	S.D.	5.10	0.00	0.00	3.40
CARTILAGINOUS PARTS OF RIBS DISPLACED					
Fetal Incidence	N	0	0	0	1
	%	0.0	0.0	0.0	0.8
Litter Incidence	N	0F1	0	0	1
	%	0.0	0.0	0.0	4.2
Affected Fetuses/Litter	MEAN%	0.0W1	0.0	0.0	1.0
	S.D.	0.00	0.00	0.00	5.10
TOTAL FETAL SKELETAL UNCLASS. CARTILAGE OBS.					
Fetal Incidence	N	98	92	96	90
	%	72	67	69	71
Litter Incidence	N	24F1	25	24	24
	%	96	100	96	100
Affected Fetuses/Litter	MEAN%	71.4W1	65.8	67.2	69.9
	S.D.	22.90	27.54	26.05	26.19

Statistics: F1 =Fisher's exact test (one-sided) W1 =Wilcoxon-test (one-sided)
* : p<=0.05 ** : p<=0.01

10-APR-14

05R018

TABLE : IB- 030

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

SUMMARY OF ALL CLASSIFIED FETAL EXTERNAL, SOFT TISSUE, AND SKELETAL OBSERVATIONS

		TEST GROUP 0	TEST GROUP 1	TEST GROUP 2	TEST GROUP 3
		0 MG/KG BW/D	30 MG/KG BW/D	100 MG/KG BW/D	300 MG/KG BW/D
Litters Evaluated	N	25	25	25	24
Fetuses Evaluated	N	264	265	265	243
Live	N	264	265	265	243
Dead	N	0	0	0	0
TOTAL MALFORMATIONS					
Fetal Incidence	N	1	0	3	5
	%	0.4	0.0	1.1	2.1
Litter Incidence	N	1Fi	0	3	5
	%	4.0	0.0	12	21
Affected Fetuses/Litter	MEAN%	0.4Wi	0.0	1.6	2.3*
	S.D.	1.82	0.00	4.75	4.70
TOTAL VARIATIONS					
Fetal Incidence	N	143	140	151	137
	%	54	53	57	56
Litter Incidence	N	25Fi	25	25	24
	%	100	100	100	100
Affected Fetuses/Litter	MEAN%	54.6Wi	52.9	56.6	56.4
	S.D.	6.21	6.86	7.49	6.32

Statistics: Fi =Fisher's exact test (one-sided) Wi =Wilcoxon-test (one-sided)

* : p<=0.05 ** : p<=0.01

STUDY TITLE

Report

DHDPS

Prenatal Developmental Toxicity Study
in Wistar Rats
Oral Administration (Gavage)

TEST FACILITY PROJECT IDENTIFICATION

Project No.: 30R0066/05R018

PART II OF III
(TABLES SECTION, INDIVIDUAL VALUES AND OBSERVATIONS)

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**CONTENTS OF PART II
(TABLES SECTION, INDIVIDUAL VALUES AND OBSERVATIONS)**

Title Page
Table of contents

**Part A:
EXAMINATIONS OF THE DAMS (INDIVIDUAL VALUES)**

	Tables
Individual maternal clinical observations	IIA-001 - IIA-008
Individual maternal food consumption	IIA-009 - IIA-012
Individual maternal body weights	IIA-013 - IIA-016
Individual maternal body weight change	IIA-017 - IIA-020
Individual uterine weights and net maternal body weight change	IIA-021 - IIA-024
Individual maternal necropsy observations	IIA-025 - IIA-032
Individual reproduction data	IIA-033 - IIA-036

**Part B:
EXAMINATIONS OF THE FETUSES (INDIVIDUAL VALUES)**

	Tables
Individual fetal status and uterine location	IIB-001 - IIB-004
Individual placental weights	IIB-005 - IIB-008
Individual fetal body weights	IIB-009 - IIB-012
Individual fetal external, visceral, skeletal and/or cartilage observations	IIB-013 - IIB-130

10-APR-14		05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS										TABLE : IIA-		001							
				ORAL ADMINISTRATION (GAVAGE)																			
				INDIVIDUAL MATERNAL CLINICAL OBSERVATIONS DURING GESTATION																			
TEST GROUP 0 (0 MG/KG BW/D)																							
FEMALE#	OBSERVATIONS	DAY OF																					
		GESTATION																					
		0	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	
1	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
2	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
3	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
4	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
5	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
6	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
7	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
8	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
9	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
10	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
11	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
12	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
13	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
14	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
15	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
16	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
17	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
18	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
19	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
20	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
21	NOTHING ABNORMAL DETECTED	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P	P
CODE: 1--SLIGHT 2--MODERATE 3--MARKED P--PRESENT																							

CODE: 1-SLIGHT 2-MODERATE 3-MARKED P-PRESENT

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAUGE)

INDIVIDUAL MATERNAL CLINICAL OBSERVATIONS DURING GESTATION

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	OBSERVATIONS	DAY OF GESTATION
1	0	1
1	1	1
1	1	1
1	1	1
1	1	1
2	0	1
3	1	2
4	1	3
5	2	4
6	7	5
7	8	6
8	9	7
9	0	8
0	1	9

22 NOTHING ABNORMAL DETECTED
P P

23 NOTHING ABNORMAL DETECTED P P P P P P P P P P P P P P P P

24 NOTHING ABNORMAL DETECTED P P P P P P P P P P P P P P P

25 NOTHING ABNORMAL DETECTED

CODE: 1-SLIGHT 2-MODERATE 3-MARKED P-PRESENT

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL CLINICAL OBSERVATIONS DURING GESTATION

TEST GROUP 1 (30 MG/KG BW/D)

[illegible]

CODE: 1-SLIGHT 2-MODERATE 3-MARKED P-PRESENT

05R018

TABLE : IIA-

004

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL CLINICAL OBSERVATIONS DURING GESTATION

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	OBSERVATIONS	DAY OF GESTATION
47	NOTHING ABNORMAL DETECTED	P P P P P P P P P P P P P P P P P P
48	NOTHING ABNORMAL DETECTED	P P P P P P P P P P P P P P P P P P
49	NOTHING ABNORMAL DETECTED	P P P P P P P P P P P P P P P P P P
50	NOTHING ABNORMAL DETECTED	P P P P P P P P P P P P P P P P P P

CODE: 1-SLIGHT 2-MODERATE 3-MARKED P-PRESENT

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL CLINICAL OBSERVATIONS DURING GESTATION

[illegible]

CODE: 1-SLIGHT 2-MODERATE 3-MARKED P-PRESENT

10-APR-14	05R018	TABLE : IIA-	006
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS			
ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL MATERNAL CLINICAL OBSERVATIONS DURING GESTATION			
TEST GROUP 2 (100 MG/KG BW/D)			
FEMALE#	OBSERVATIONS	DAY OF GESTATION	
72	NOTHING ABNORMAL DETECTED	0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0	1 1 1 1 1 1 1 1 1 1 2
73	NOTHING ABNORMAL DETECTED	P P	P P
74	NOTHING ABNORMAL DETECTED	P P	P P
75	NOTHING ABNORMAL DETECTED	P P	P P
CODE: 1-SLIGHT 2-MODERATE 3-MARKED P-PRESENT			

10-APR-14	05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL MATERNAL CLINICAL OBSERVATIONS DURING GESTATION	TABLE : IIA-	007
TEST GROUP 3 (300 MG/KG BW/D)				
FEMALE#	OBSERVATIONS	DAY OF GESTATION	1 1 1 1 1 1 1 1 1 2	2
76	NOTHING ABNORMAL DETECTED	0 1 2 3 4 5 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0		
77	NOTHING ABNORMAL DETECTED SALIVATION AFTER TREATMENT	P P		

CODE: 1-SLIGHT 2-MODERATE 3-MARKED P-PRESENT

05R018

TABLE : IIA-

800

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL CLINICAL OBSERVATIONS DURING GESTATION

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	OBSERVATIONS	DAY OF GESTATION	1	1	1	1	1	1	1	1	1	1	2
94	NOTHING ABNORMAL DETECTED		P	P	P	P	P	P	P	P	P	P	P
95	NOTHING ABNORMAL DETECTED SALIVATION AFTER TREATMENT		P	P	P	P	P	P	P	P	P	P	P
96	NOTHING ABNORMAL DETECTED		P	P	P	P	P	P	P	P	P	P	P
97	NOTHING ABNORMAL DETECTED		P	P	P	P	P	P	P	P	P	P	P
98	NOTHING ABNORMAL DETECTED SALIVATION AFTER TREATMENT		P	P	P	P	P	P	P	P	P	P	P
99	NOTHING ABNORMAL DETECTED		P	P	P	P	P	P	P	P	P	P	P
100	NOTHING ABNORMAL DETECTED		P	P	P	P	P	P	P	P	P	P	P

CODE: 1-SLIGHT 2-MODERATE 3-MARKED P-PRESENT

10-APR-14

05R018

009

TABLE : IIA-

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL FOOD CONSUMPTION DURING GESTATION -- GRAMS/ANIMAL/DAY

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	DAY OF GESTATION																			
	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
1	16.0	16.8	16.5	19.2	17.9	17.7	17.9	19.7	19.7	20.2	20.3	19.9	19.9	22.4	21.8	20.2	21.8	22.4	21.8	21.8
2	14.2	15.1	16.5	16.5	17.7	17.6	17.7	18.0	19.7	19.7	18.5	19.7	19.7	19.8	20.2	20.2	19.8	22.3	23.0	23.0
3	13.8	14.9	16.4	18.5	20.3	20.2	22.9	21.3	21.3	21.3	21.3	21.3	21.3	21.3	21.3	21.3	21.3	21.3	21.3	21.3
4	13.7	15.6	18.5	20.1	19.3	22.0	24.2	22.8	22.8	22.8	22.8	22.8	22.8	22.8	22.8	22.8	22.8	22.8	22.8	22.8
5	17.2	17.5	20.1	19.3	22.0	20.0	18.7	19.3	19.3	19.3	19.3	19.3	19.3	19.3	19.3	19.3	19.3	19.3	19.3	19.3
6	14.6	16.4	18.2	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1
7	14.3	15.6	19.2	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9
8	14.8	14.8	17.4	16.9	16.9	17.1	17.1	17.1	17.1	17.1	17.1	17.1	17.1	17.1	17.1	17.1	17.1	17.1	17.1	17.1
9	17.9	17.1	19.5	17.2	17.2	17.2	17.2	17.2	17.2	17.2	17.2	17.2	17.2	17.2	17.2	17.2	17.2	17.2	17.2	17.2
10	16.0	17.3	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6
11	18.1	16.9	19.3	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6	18.6
12	12.7	16.7	17.8	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6
13	15.0	16.0	17.6	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0
14	13.2	19.1	20.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5	19.5
15	12.0	16.0	17.4	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3
16	13.3	16.1	15.3	15.2	15.2	15.2	15.2	15.2	15.2	15.2	15.2	15.2	15.2	15.2	15.2	15.2	15.2	15.2	15.2	15.2
17	11.6	17.7	18.6	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9
18	12.3	16.8	18.1	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0
19	12.3	18.9	18.4	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0
20	11.1	15.1	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0	18.0
21	15.1	18.6	18.3	20.4	20.4	20.4	20.4	20.4	20.4	20.4	20.4	20.4	20.4	20.4	20.4	20.4	20.4	20.4	20.4	20.4
22	13.8	16.0	17.0	16.4	16.4	16.4	16.4	16.4	16.4	16.4	16.4	16.4	16.4	16.4	16.4	16.4	16.4	16.4	16.4	16.4
23	15.2	19.3	18.4	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9	17.9
24	13.3	19.0	20.4	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0
25	13.1	18.6	18.0	18.2	18.2	18.2	18.2	18.2	18.2	18.2	18.2	18.2	18.2	18.2	18.2	18.2	18.2	18.2	18.2	18.2
MEAN	14.2	16.9	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3	18.3
S.D.	1.85	1.40	1.25	1.25	1.34	1.34	1.34	1.34	1.34	1.34	1.34	1.34	1.34	1.34	1.34	1.34	1.34	1.34	1.34	1.34
N	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25

10-APR-14

05R018

TABLE : IIA- 010

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL FOOD CONSUMPTION DURING GESTATION -- GRAMS/ANIMAL/DAY

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	DAY OF GESTATION																			
	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
26	11.0	13.6	15.9	16.1	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3
27	10.6	15.4	18.4	17.2	20.1	20.1	20.1	20.1	20.1	20.1	20.1	20.1	20.1	20.1	20.1	20.1	20.1	20.1	20.1	20.1
28	12.6	15.8	18.2	18.3	19.1	19.1	19.1	19.1	19.1	19.1	19.1	19.1	19.1	19.1	19.1	19.1	19.1	19.1	19.1	19.1
29	14.1	16.0	16.6	15.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6
30	17.5	18.3	22.6	22.6	25.5	25.5	25.5	25.5	25.5	25.5	25.5	25.5	25.5	25.5	25.5	25.5	25.5	25.5	25.5	25.5
31	9.3	16.6	19.5	20.1	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2
32	12.4	14.5	16.8	16.7	17.0	17.0	17.0	17.0	17.0	17.0	17.0	17.0	17.0	17.0	17.0	17.0	17.0	17.0	17.0	17.0
33	10.9	13.4	14.9	15.7	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0
34	9.8	14.5	16.6	15.2	17.5	17.5	17.5	17.5	17.5	17.5	17.5	17.5	17.5	17.5	17.5	17.5	17.5	17.5	17.5	17.5
35	17.0	19.3	20.2	20.8	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7
36	10.9	14.9	16.4	17.0	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7
37	12.6	17.2	18.3	20.4	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2	21.2
38	13.4	18.3	17.9	17.8	19.7	19.7	19.7	19.7	19.7	19.7	19.7	19.7	19.7	19.7	19.7	19.7	19.7	19.7	19.7	19.7
39	10.6	15.5	16.2	16.5	16.3	16.3	16.3	16.3	16.3	16.3	16.3	16.3	16.3	16.3	16.3	16.3	16.3	16.3	16.3	16.3
40	13.6	17.8	18.7	17.8	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9	18.9
41	15.8	18.1	16.7	18.4	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1	18.1
42	13.4	17.7	17.6	18.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8
43	11.4	17.5	17.4	17.5	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8
44	11.8	17.0	18.4	18.3	18.5	18.5	18.5	18.5	18.5	18.5	18.5	18.5	18.5	18.5	18.5	18.5	18.5	18.5	18.5	18.5
45	11.1	15.0	15.3	17.4	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6	16.6
46	15.1	19.1	20.2	21.2	23.4	23.4	23.4	23.4	23.4	23.4	23.4	23.4	23.4	23.4	23.4	23.4	23.4	23.4	23.4	23.4
47	13.1	18.3	18.6	19.8	20.7	20.7	20.7	20.7	20.7	20.7	20.7	20.7	20.7	20.7	20.7	20.7	20.7	20.7	20.7	20.7
48	14.4	19.9	18.1	18.4	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3	17.3
49	11.4	20.0	19.5	19.4	19.6	19.6	19.6	19.6	19.6	19.6	19.6	19.6	19.6	19.6	19.6	19.6	19.6	19.6	19.6	19.6
50	11.2	15.6	16.5	16.3	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8	16.8
MEAN	12.6	16.8	17.8	18.1	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0
S.D.	2.15	1.91	1.74	1.90	2.32	2.32	2.32	2.32	2.32	2.32	2.32	2.32	2.32	2.32	2.32	2.32	2.32	2.32	2.32	2.32
N	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25

10-APR-14

05R018

TABLE : IIA- 011

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL FOOD CONSUMPTION DURING GESTATION -- GRAMS/ANIMAL/DAY

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	DAY OF GESTATION																			
	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
51	15.0	17.2	18.0	18.0	18.0	18.0	18.0	17.5	17.5	20.0	20.0	19.4	23.9	23.9	24.0	23.4	23.4	23.4	23.4	23.4
52	14.3	16.3	18.9	19.8	19.8	19.8	19.8	19.9	19.9	21.2	20.9	20.9	23.1	23.1	23.8	22.3	22.3	22.3	22.3	22.3
53	13.7	15.0	16.6	17.8	17.8	17.8	17.8	17.7	17.7	18.8	17.5	17.5	20.2	20.2	21.7	22.9	22.9	22.9	22.9	22.9
54	16.2	19.1	19.4	19.4	19.4	19.4	19.4	20.8	20.8	22.3	22.0	22.0	24.7	24.7	23.6	23.7	23.7	23.7	23.7	23.7
55	15.4	16.7	19.3	18.8	18.8	18.8	18.8	21.2	21.2	22.5	22.8	22.8	23.4	23.4	25.7	25.1	25.1	25.1	25.1	25.1
56	13.5	14.9	15.4	17.1	16.3	16.3	16.3	16.3	16.3	19.5	17.9	17.9	17.5	17.5	21.9	22.0	22.0	22.0	22.0	22.0
57	14.1	17.4	19.3	20.8	21.5	21.5	21.5	22.4	22.4	24.3	24.3	24.3	27.3	27.3	26.8	25.2	25.2	25.2	25.2	25.2
58	14.4	16.2	18.6	16.3	19.0	18.5	18.5	19.0	19.0	20.1	19.7	19.7	20.9	20.9	21.2	21.2	21.2	21.2	21.2	21.2
59	13.6	15.9	16.7	18.1	18.1	18.1	18.1	18.5	18.5	20.1	19.7	19.7	20.9	20.9	21.2	21.2	21.2	21.2	21.2	21.2
60	13.8	17.4	20.4	16.9	19.1	19.1	19.1	19.1	19.1	20.5	18.5	18.5	22.7	22.7	22.3	22.3	22.3	22.3	22.3	22.3
61	13.1	17.4	18.1	18.7	19.0	19.0	19.0	19.0	19.0	20.5	19.7	19.7	21.8	21.8	21.4	21.4	21.4	21.4	21.4	21.4
62	13.6	16.9	17.9	17.3	19.0	19.0	19.0	19.0	19.0	21.7	19.4	19.4	24.1	24.1	18.0	18.9	18.9	18.9	18.9	18.9
63	11.1	16.5	16.5	16.9	18.1	18.1	18.1	18.1	18.1	19.0	19.0	19.0	19.4	19.4	21.9	19.9	19.9	19.9	19.9	19.9
64	10.3	15.6	16.5	16.8	17.5	17.5	17.5	17.5	17.5	18.4	18.4	18.4	19.0	19.0	20.6	19.2	19.2	19.2	19.2	19.2
65	12.5	17.7	19.1	19.5	19.9	19.9	19.9	19.9	19.9	19.8	22.1	22.1	22.4	22.4	21.3	21.8	21.8	21.8	21.8	21.8
66	14.6	18.1	17.3	18.6	19.1	19.1	19.1	19.1	19.1	21.3	22.0	22.0	24.5	24.5	22.6	22.6	22.6	22.6	22.6	22.6
67	11.9	17.4	17.3	17.9	16.2	16.2	16.2	16.2	16.2	20.0	20.1	20.1	21.4	21.4	21.9	18.8	18.8	18.8	18.8	18.8
68	13.7	16.2	17.8	19.5	17.7	17.7	17.7	17.7	17.7	19.8	21.1	21.1	21.6	21.6	23.3	19.8	19.8	19.8	19.8	19.8
69	17.0	17.3	17.9	16.8	16.2	16.2	16.2	16.2	16.2	21.1	19.8	19.8	21.0	21.0	22.2	21.7	21.7	21.7	21.7	21.7
70	12.6	16.8	18.2	17.0	18.9	18.9	18.9	18.9	18.9	20.1	19.9	19.9	21.6	21.6	22.4	21.9	21.9	21.9	21.9	21.9
71	16.0	19.6	21.0	20.7	20.8	20.8	20.8	20.8	20.8	21.6	20.4	20.4	22.4	22.4	23.0	22.2	22.2	22.2	22.2	22.2
72	12.1	16.4	17.1	17.1	17.0	17.0	17.0	17.0	17.0	18.2	19.5	19.5	20.5	20.5	19.1	17.5	17.5	17.5	17.5	17.5
73	12.6	16.8	18.1	18.3	18.3	18.3	18.3	18.3	18.3	19.9	23.4	23.4	23.4	23.4	22.6	20.9	20.9	20.9	20.9	20.9
74	12.7	16.2	16.1	16.1	16.8	16.8	16.8	16.8	16.8	18.4	19.6	19.6	18.6	18.6	20.2	15.6	15.6	15.6	15.6	15.6
75	16.1	19.5	21.1	22.0	22.4	22.4	22.4	22.4	22.4	24.6	25.6	25.6	23.9	23.9	25.0	22.8	22.8	22.8	22.8	22.8
MEAN	13.8	17.0	18.1	18.2	18.2	18.2	18.2	18.7	18.7	20.7	20.5	20.5	22.0	22.0	22.4	21.3	21.3	21.3	21.3	21.3
S.D.	1.63	1.20	1.47	1.52	1.52	1.52	1.52	1.71	1.71	1.90	1.94	1.94	2.07	2.07	1.96	2.24	2.24	2.24	2.24	2.24
N	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25

10-APR-14

05R018

012

TABLE : IIA-

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL FOOD CONSUMPTION DURING GESTATION -- GRAMS/ANIMAL/DAY

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	DAY OF GESTATION																			
	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
76	12.8	16.5	19.1	17.3	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6
77	10.8	16.0	17.2	16.4	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6	17.6
78	13.1	16.1	18.6	14.4	14.9	15.8	15.8	15.8	15.8	15.8	15.8	15.8	15.8	15.8	15.8	15.8	15.8	15.8	15.8	15.8
79	11.3	14.5	16.4	13.0	13.4	15.7	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0	16.0
80	10.8	17.7	17.6	15.5	14.8	20.3	21.3	22.5	22.5	22.5	22.5	22.5	22.5	22.5	22.5	22.5	22.5	22.5	22.5	22.5
81	10.9	14.5	17.3	14.0	15.1	18.5	16.8	19.1	20.0	20.0	20.0	20.0	20.0	20.0	20.0	20.0	20.0	20.0	20.0	20.0
82	11.9	14.9	16.1	15.8	16.0	17.3	18.3	20.1	21.3	20.1	21.3	20.1	21.3	20.1	21.3	20.1	21.3	20.1	21.3	20.1
83	9.1	13.9	16.2	15.9	14.3	18.6	16.2	19.0	19.4	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7	18.7
84	13.8	16.6	18.3	16.9	17.5	20.0	20.6	22.6	21.8	24.2	24.2	24.2	24.2	24.2	24.2	24.2	24.2	24.2	24.2	24.2
85	9.5	18.0	18.7	16.2	18.3	20.1	22.4	23.9	24.6	23.9	24.6	23.9	24.6	23.9	24.6	23.9	24.6	23.9	24.6	23.9
86	16.0	20.1	21.6	17.7	20.5	24.4	22.5	25.1	25.8	25.9	25.9	25.9	25.9	25.9	25.9	25.9	25.9	25.9	25.9	25.9
87	14.2	17.1	18.4	15.8	17.4	19.8	20.0	22.1	20.9	22.6	22.6	22.6	22.6	22.6	22.6	22.6	22.6	22.6	22.6	22.6
88	10.6	16.4	17.6	16.7	16.7	17.5	18.0	20.6	20.8	17.4	17.4	17.4	17.4	17.4	17.4	17.4	17.4	17.4	17.4	17.4
89	13.1	17.0	18.5	14.4	12.6	14.5	16.5	17.6	19.4	16.7	16.7	16.7	16.7	16.7	16.7	16.7	16.7	16.7	16.7	16.7
90	12.8	15.9	17.5	15.4	15.7	17.5	19.5	20.4	20.4	21.4	21.4	21.4	21.4	21.4	21.4	21.4	21.4	21.4	21.4	21.4
91	15.0	18.3	18.5	10.6	16.9	18.9	20.8	21.6	22.9	17.8	17.8	17.8	17.8	17.8	17.8	17.8	17.8	17.8	17.8	17.8
92	13.6	19.3	20.5	15.6	19.0	19.3	21.9	23.8	24.5	20.6	20.6	20.6	20.6	20.6	20.6	20.6	20.6	20.6	20.6	20.6
93x	15.8	18.1	19.3	14.3	15.3	15.3	16.0	14.0	12.9	12.7	12.7	12.7	12.7	12.7	12.7	12.7	12.7	12.7	12.7	12.7
94	13.1	18.0	18.5	16.2	17.3	20.7	20.6	22.6	22.2	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7	21.7
95	14.8	17.2	19.9	16.4	18.4	18.5	20.1	20.9	20.7	20.3	20.3	20.3	20.3	20.3	20.3	20.3	20.3	20.3	20.3	20.3
96	2.7	18.8	19.8	15.9	17.3	17.7	17.8	20.2	19.4	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8	19.8
97	12.4	21.1	21.7	18.7	17.5	19.9	24.4	23.5	28.4	26.4	26.4	26.4	26.4	26.4	26.4	26.4	26.4	26.4	26.4	26.4
98	14.3	17.0	19.1	15.3	14.3	18.3	19.6	21.1	22.3	26.5	26.5	26.5	26.5	26.5	26.5	26.5	26.5	26.5	26.5	26.5
99	7.9	18.5	19.1	15.6	14.9	16.9	17.3	20.1	18.9	18.8	18.8	18.8	18.8	18.8	18.8	18.8	18.8	18.8	18.8	18.8
100	13.8	16.4	18.5	15.9	17.2	18.7	19.1	19.6	18.1	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0	19.0
MEAN	12.0	17.1	18.5	15.7	16.5	18.7	19.5	21.2	21.7	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0	21.0
S.D.	2.81	1.76	1.47	1.60	1.89	2.02	2.18	1.93	2.44	3.16	3.16	3.16	3.16	3.16	3.16	3.16	3.16	3.16	3.16	3.16
N	24	24	24	24	24	24	24	24	24	24	24	24	24	24	24	24	24	24	24	24

NP=NOT PREGNANT X=EXCLUDED FROM MEAN

10-APR-14

05R018

TABLE : IIA- 013

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL BODY WEIGHTS DURING GESTATION -- GRAMS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	DAY OF GESTATION																				
	0	1	3	6	8	10	13	15	17	19	20										
1	154.1	162.7	175.7	185.5	195.4	204.0	218.8	231.7	249.5	274.7	291.0										
2	159.6	169.8	181.1	189.7	202.5	208.9	226.2	234.2	246.9	270.4	285.7										
3	155.8	163.9	172.7	183.6	194.3	203.3	219.1	233.7	253.7	276.8	294.5										
4	152.3	163.8	175.7	189.6	205.8	209.3	231.6	241.1	258.0	284.2	305.3										
5	172.1	185.9	199.1	211.0	223.7	227.5	251.1	260.7	282.2	309.4	330.4										
6	157.6	168.3	181.3	188.7	200.0	206.0	216.3	232.4	252.5	275.3	292.7										
7	158.5	169.2	178.3	194.2	201.6	210.6	228.4	240.2	252.3	278.3	286.5										
8	164.8	175.7	182.8	191.6	203.9	206.1	219.6	231.2	245.7	265.7	282.0										
9	171.0	188.5	194.7	207.1	213.7	223.9	242.1	253.7	276.0	299.5	317.4										
10	149.9	167.4	175.5	182.0	194.6	199.6	212.7	223.4	238.2	258.7	270.8										
11	171.7	185.0	195.1	204.6	213.4	223.2	240.0	253.9	274.7	299.8	315.9										
12	169.3	177.2	183.3	195.4	203.8	211.9	228.2	239.8	254.3	283.5	297.1										
13	155.6	172.5	181.1	193.0	204.0	209.5	229.5	239.1	254.2	271.0	281.2										
14	169.5	186.6	198.0	208.3	216.3	230.7	249.2	257.2	280.3	281.4	313.9										
15	160.5	169.2	176.7	186.4	191.2	201.8	219.8	227.0	244.7	271.3	281.0										
16	166.2	178.2	186.6	193.6	196.3	208.1	223.9	229.7	246.0	272.9	285.1										
17	173.3	185.1	197.3	206.4	209.6	221.9	235.6	247.0	267.0	294.5	305.6										
18	166.7	177.8	185.3	196.1	198.5	210.4	226.6	237.2	252.7	279.7	290.8										
19	174.9	184.3	194.4	200.4	201.5	215.8	227.0	236.1	255.5	278.3	289.1										
20	168.8	173.9	182.1	192.3	201.4	214.4	231.4	238.4	257.6	284.9	296.8										
21	172.7	186.6	195.6	204.1	214.4	227.5	238.9	245.4	263.5	285.3	295.1										
22	169.4	181.0	186.6	195.4	201.0	211.2	222.7	234.4	254.3	275.2	289.3										
23	176.1	186.9	197.4	207.0	207.0	216.0	232.3	240.7	258.7	283.7	298.7										
24	169.5	179.2	189.4	202.0	204.8	222.7	235.5	242.2	262.0	290.6	303.4										
25	162.8	175.7	182.4	190.1	195.0	209.8	222.0	233.2	256.1	282.6	298.5										
MEAN	164.9	176.6	185.9	195.9	203.7	213.3	229.2	239.3	257.5	281.1	295.9										
S.D.	7.66	8.22	8.34	8.41	7.93	8.71	9.88	9.38	11.23	11.38	13.47										
N	25	25	25	25	25	25	25	25	25	25	25										

10-APR-14

05R018

TABLE : IIA- 014

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL BODY WEIGHTS DURING GESTATION -- GRAMS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	DAY OF GESTATION																				
	0	1	3	6	8	10	13	15	17	19	20										
26	154.8	163.6	172.1	181.3	189.6	198.0	212.8	224.8	240.0	258.7	271.5										
27	156.2	165.5	177.2	192.7	203.6	213.6	239.3	251.1	273.5	305.3	320.0										
28	168.1	181.7	192.0	201.9	216.4	222.4	240.3	251.2	272.4	297.7	313.9										
29	156.0	170.4	177.7	184.7	195.2	198.8	212.8	224.8	242.4	263.0	275.7										
30	171.3	192.5	200.3	213.3	228.3	240.5	257.1	275.3	297.0	327.6	348.8										
31	172.9	184.3	197.3	209.5	222.2	228.7	245.3	259.9	276.6	297.4	313.2										
32	154.6	167.6	176.6	184.6	197.0	200.3	213.9	220.5	234.4	250.6	265.1										
33	149.0	164.5	172.0	178.8	189.5	194.0	208.6	223.1	239.2	260.5	275.7										
34	142.8	150.8	157.7	169.9	176.9	184.0	198.4	211.7	232.0	248.7	263.7										
35	165.2	182.5	192.9	204.7	216.9	224.6	243.3	253.7	269.9	303.9	322.0										
36	166.5	177.2	187.9	199.2	210.4	217.5	235.2	250.2	269.8	296.7	315.9										
37	163.7	178.6	189.2	201.7	216.0	222.5	235.3	253.1	267.9	292.8	303.2										
38	173.9	186.8	196.4	206.4	210.9	226.2	240.1	250.7	270.9	304.8	319.7										
39	145.9	157.4	165.0	174.3	180.9	188.9	207.2	216.3	235.3	258.5	269.7										
40	170.2	181.3	188.8	196.8	200.9	214.6	225.2	235.0	252.2	276.4	286.1										
41	181.9	196.9	201.1	206.6	213.9	223.2	237.1	238.8	260.5	291.9	304.3										
42	165.0	178.3	187.1	196.3	204.7	215.2	229.0	236.4	255.2	279.3	286.4										
43	181.5	192.7	204.4	216.0	219.6	228.3	248.1	258.8	281.3	312.7	328.3										
44	163.6	172.9	184.2	196.2	200.5	214.7	231.6	239.8	260.7	291.9	307.1										
45	180.8	192.2	192.9	203.2	210.1	219.0	234.7	241.4	264.5	289.5	301.2										
46	177.9	197.0	206.3	216.2	223.6	234.6	253.9	262.7	285.8	317.9	325.0										
47	187.2	199.5	208.8	218.2	224.5	237.7	259.4	269.1	297.4	318.1	334.5										
48	184.5	197.9	207.6	213.3	217.1	225.6	239.4	249.2	267.4	292.9	306.6										
49	183.6	190.4	203.0	213.8	218.6	232.6	248.8	258.0	283.7	313.0	314.3										
50	171.5	177.4	186.4	197.3	201.3	214.7	225.5	231.4	251.3	271.9	288.4										
MEAN	167.5	180.0	189.0	199.1	207.5	216.8	232.9	243.5	262.8	288.9	302.4										
S.D.	12.55	13.42	13.68	13.73	13.80	15.18	16.38	16.95	18.25	22.62	23.30										
N	25	25	25	25	25	25	25	25	25	25	25										

10-APR-14

05R018

TABLE : IIA- 015

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL MATERNAL BODY WEIGHTS DURING GESTATION -- GRAMS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	DAY OF GESTATION										
	0	1	3	6	8	10	13	15	17	19	20
51	166.5	178.2	189.2	200.8	210.9	214.5	231.0	245.5	264.8	292.2	310.9
52	170.8	180.0	189.9	201.7	213.3	219.8	232.1	248.7	264.8	293.9	311.4
53	163.6	175.4	182.1	191.1	198.5	203.0	216.4	226.5	244.0	266.0	283.6
54	163.2	180.4	189.2	201.7	207.6	216.2	239.5	253.0	270.2	290.1	306.4
55	177.2	187.5	197.4	211.6	220.6	228.7	244.6	261.9	276.3	295.4	313.2
56	157.2	166.9	175.7	182.8	191.3	197.9	219.3	225.5	247.3	266.7	289.6
57	175.7	188.9	200.5	210.9	227.4	235.1	255.1	268.8	292.2	318.1	338.9
58	153.4	165.8	176.2	185.5	193.5	200.7	218.9	226.4	237.9	264.0	276.6
59	154.4	169.2	180.6	189.6	201.3	209.4	225.5	236.9	252.8	274.4	292.1
60	169.0	186.5	196.1	209.3	214.0	222.8	234.8	245.6	269.9	295.6	316.5
61	151.8	165.7	177.2	187.3	195.1	200.4	214.3	224.7	238.4	250.5	260.0
62	153.7	166.5	178.8	185.5	198.9	205.7	219.8	229.9	240.4	258.7	272.4
63	157.7	169.3	178.2	188.6	190.4	203.8	216.7	226.1	245.1	271.0	282.1
64	165.6	175.4	185.3	194.7	198.3	208.6	220.9	230.8	251.3	274.2	285.1
65	185.9	187.1	198.6	211.5	217.6	227.8	239.4	251.0	271.9	299.5	311.9
66	183.0	197.0	204.3	212.1	217.3	231.0	243.5	252.6	278.2	306.1	320.6
67	185.6	191.2	203.2	213.7	218.8	229.4	246.7	255.6	274.4	303.3	311.6
68	170.7	184.9	193.9	200.9	208.9	219.9	232.5	242.0	262.5	291.9	303.2
69	171.7	184.1	190.7	201.5	201.7	211.0	224.7	227.8	249.7	273.8	288.6
70	172.3	181.7	189.0	198.9	199.7	211.6	227.5	231.6	250.9	278.7	291.9
71	174.9	188.6	188.8	201.8	207.0	218.6	231.8	240.0	257.0	280.5	294.2
72	169.7	175.0	183.0	188.7	194.3	205.0	218.2	222.7	242.0	266.6	276.3
73	181.2	189.9	196.8	205.9	209.3	223.5	239.0	254.8	278.3	302.7	317.4
74	164.6	173.0	180.0	189.1	192.9	204.3	215.0	224.5	243.3	262.2	272.5
75	179.3	192.4	204.6	213.7	222.7	235.6	254.0	261.0	283.2	309.6	318.8
MEAN	168.7	180.0	189.2	199.2	205.9	215.4	230.4	240.6	259.5	283.4	297.8
S.D.	10.28	9.40	9.27	10.19	10.78	11.57	12.29	14.08	15.82	18.09	19.30
N	25	25	25	25	25	25	25	25	25	25	25

10-APR-14

05R018

TABLE : IIA- 016

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

TEST GROUP 3 (300 MG/KG BW/D)
INDIVIDUAL MATERNAL BODY WEIGHTS DURING GESTATION -- GRAMS

FEMALE#	DAY OF GESTATION																			
	0	1	3	6	8	10	13	15	17	19	20									
76	154.9	168.4	178.9	192.7	199.9	205.7	218.6	235.2	252.6	275.2	292.2									
77	155.2	169.8	180.6	189.7	200.1	205.7	219.9	230.3	246.6	275.4	298.8									
78	157.0	170.6	179.6	191.4	198.3	198.9	212.8	223.5	235.3	256.0	265.6									
79	150.8	165.5	173.8	183.6	187.8	193.2	205.9	212.0	229.6	247.2	260.8									
80	155.8	177.7	187.6	193.2	201.4	202.4	222.5	234.1	253.1	278.6	295.1									
81	160.9	173.2	181.2	198.0	204.9	210.0	223.2	231.9	247.4	268.2	282.8									
82	150.1	164.1	171.4	180.3	187.6	191.4	203.0	216.2	235.2	256.4	272.3									
83	154.5	166.3	173.9	183.4	193.8	197.8	214.6	224.7	236.8	259.1	269.3									
84	163.8	181.5	194.8	202.5	214.2	216.0	235.0	245.3	260.6	281.4	302.0									
85	158.1	177.3	190.9	199.9	213.8	222.5	236.2	254.7	270.4	293.6	309.9									
86	162.6	183.3	194.4	205.6	211.6	216.8	239.8	253.4	268.6	289.3	302.2									
87	174.8	195.1	204.5	213.2	223.1	226.0	243.9	255.9	267.6	291.7	306.9									
88	147.0	160.1	173.0	182.7	194.3	199.1	211.5	216.9	233.4	247.5	257.9									
89	157.6	167.1	174.9	184.5	185.5	192.2	202.7	213.8	230.8	258.4	270.2									
90	169.3	176.5	186.8	196.7	198.9	211.5	222.2	229.9	250.4	272.0	286.6									
91	177.5	189.5	197.8	209.3	210.4	224.3	238.6	251.1	269.1	297.0	307.2									
92	181.0	186.1	201.4	213.0	216.7	230.9	243.3	255.9	275.3	304.2	319.7									
93x NP	180.1	192.4	203.3	210.9	210.1	216.9	221.0	212.1	217.9	217.7	221.0									
94	181.6	190.0	201.1	209.2	215.9	224.4	243.7	251.3	271.2	292.0	299.7									
95	179.1	190.3	199.1	212.3	217.2	227.5	238.0	244.2	264.0	288.2	302.9									
96	174.3	162.7	189.8	196.9	202.6	213.6	224.9	229.4	250.5	275.2	286.2									
97	197.8	209.7	223.6	230.0	235.9	246.0	259.3	269.3	292.2	322.8	339.0									
98	172.8	183.5	188.7	199.7	204.4	210.9	224.0	228.9	241.2	266.6	275.3									
99	180.0	174.6	195.2	205.7	210.3	216.8	229.9	237.5	258.8	283.8	301.8									
100	158.5	168.9	176.6	185.8	193.1	202.2	220.5	221.4	245.7	265.9	279.1									
MEAN	165.6	177.2	188.3	198.3	205.1	211.9	226.4	236.1	253.6	276.9	291.0									
S.D.	12.88	12.05	12.71	12.41	12.27	13.85	14.57	15.71	16.40	18.57	19.85									
N	24	24	24	24	24	24	24	24	24	24	24									

NP=NOT PREGNANT x=EXCLUDED FROM MEAN

NP=NOT PREGNANT X=EXCLUDED FROM MEAN

10-APR-14

05R018

TABLE : IIA- 017

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL MATERNAL BODY WEIGHT CHANGE DURING GESTATION -- GRAMS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	DAY OF GESTATION																			
	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
1	8.6	13.0	9.8	9.9	8.6	12.8	9.9	8.6	14.8	12.9	12.9	17.8	25.2	16.3						
2	10.2	11.3	8.6	12.8	6.4	17.3	8.0	12.7	23.5	15.3										
3	8.1	8.8	10.9	10.7	9.0	15.8	14.6	20.0	23.1	17.7										
4	11.5	11.9	13.9	16.2	3.5	24.3	7.5	16.9	26.2	21.1										
5	13.8	13.2	11.9	12.7	3.8	23.6	9.6	21.5	27.7	20.5										
6	10.7	13.0	7.4	11.3	6.0	10.3	16.1	20.1	22.8	17.4										
7	10.7	9.1	15.9	7.4	8.4	18.4	11.8	12.1	26.0	8.2										
8	10.9	7.1	8.8	12.3	2.2	13.5	11.6	14.5	20.0	16.3										
9	17.5	6.2	12.4	6.6	10.2	18.2	11.6	22.3	23.5	17.9										
10	17.5	8.1	6.5	12.6	5.0	13.1	10.7	14.8	20.5	12.1										
11	13.3	10.1	9.5	8.8	9.8	13.9	13.9	20.8	25.1	16.1										
12	7.9	6.1	12.1	8.4	8.1	16.3	11.6	14.5	29.2	13.6										
13	16.9	8.6	11.9	11.0	5.5	20.0	9.6	15.1	16.8	10.2										
14	17.1	11.4	10.3	8.0	14.4	18.5	8.0	23.1	1.1	32.5										
15	8.7	7.5	9.7	4.8	10.6	18.0	7.2	17.7	26.6	9.7										
16	12.0	8.4	7.0	2.7	11.8	15.8	5.8	16.3	26.9	12.2										
17	11.8	12.2	9.1	3.2	12.3	13.7	11.4	20.0	27.5	11.1										
18	11.1	7.5	10.8	2.4	11.9	16.2	10.6	15.5	27.0	11.1										
19	9.4	10.1	6.0	1.1	14.3	11.2	9.1	19.4	22.8	10.8										
20	5.1	8.2	10.2	9.1	13.0	17.0	7.0	19.2	27.3	11.9										
21	13.9	9.0	8.5	10.3	13.1	11.4	6.5	18.1	21.8	9.8										
22	11.6	5.6	8.8	5.6	10.2	11.5	11.7	19.9	20.9	14.1										
23	10.8	10.5	9.6	0.0	9.0	16.3	8.4	18.0	25.0	15.0										
24	9.7	10.2	12.6	2.8	17.9	12.8	6.7	19.8	28.6	12.8										
25	12.9	6.7	7.7	4.9	14.8	12.2	11.2	22.9	26.5	15.9										
MEAN	11.7	9.4	10.0	7.8	9.6	15.9	10.1	18.1	23.7	14.8										
S.D.	3.17	2.29	2.36	4.27	3.96	3.59	2.70	3.08	5.58	5.02										
N	25	25	25	25	25	25	25	25	25	25										

10-APR-14

05R018

TABLE : IIA- 018

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL MATERNAL BODY WEIGHT CHANGE DURING GESTATION -- GRAMS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	DAY OF GESTATION																			
	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
26	8.8	8.5	9.2	9.2	8.3	8.4	8.4	10.0	10.0	14.8	12.0	12.0	15.2	18.7	12.8					
27	9.3	11.7	15.5	15.5	10.9	10.0	10.0	10.0	10.0	25.7	11.8	11.8	22.4	31.8	14.7					
28	13.6	10.3	9.9	9.9	14.5	6.0	6.0	6.0	6.0	17.9	10.9	10.9	21.2	25.3	16.2					
29	14.4	7.3	7.0	7.0	10.5	3.6	3.6	3.6	3.6	14.0	12.0	12.0	17.6	20.6	12.7					
30	21.2	7.8	13.0	13.0	15.0	12.3	12.3	12.3	12.3	16.5	18.2	18.2	21.7	30.6	21.2					
31	11.4	13.0	12.2	12.2	12.7	6.5	6.5	6.5	6.5	16.6	14.6	14.6	16.7	20.8	15.8					
32	13.0	9.0	8.0	8.0	12.4	3.3	3.3	3.3	3.3	13.6	6.6	6.6	13.9	16.2	14.5					
33	15.5	7.5	6.8	6.8	10.7	4.5	4.5	4.5	4.5	14.6	14.5	14.5	16.1	21.3	15.2					
34	8.0	6.9	12.2	12.2	7.0	7.1	7.1	7.1	7.1	14.4	13.3	13.3	20.3	16.7	15.0					
35	17.3	10.4	11.8	11.8	12.2	7.7	7.7	7.7	7.7	18.7	10.4	10.4	16.2	34.0	18.1					
36	10.7	10.7	11.3	11.3	11.2	7.1	7.1	7.1	7.1	17.7	15.0	15.0	19.6	26.9	19.2					
37	14.9	10.6	12.5	12.5	14.3	6.5	6.5	6.5	6.5	12.8	17.8	17.8	14.8	24.9	10.4					
38	12.9	9.6	10.0	10.0	4.5	15.3	13.9	13.9	13.9	10.6	20.2	20.2	33.9	14.9	11.2					
39	11.5	7.6	9.3	9.3	6.6	8.0	8.0	8.0	8.0	18.3	9.1	9.1	19.0	23.2	11.2					
40	11.1	7.5	8.0	8.0	4.1	13.7	10.6	10.6	10.6	10.6	9.8	9.8	17.2	24.2	9.7					
41	15.0	4.2	5.5	5.5	7.3	9.3	9.3	9.3	9.3	13.9	1.7	1.7	21.7	31.4	12.4					
42	13.3	8.8	9.2	9.2	8.4	10.5	13.8	13.8	13.8	7.4	18.8	18.8	24.1	15.6	7.1					
43	11.2	11.7	11.6	11.6	3.6	8.7	8.7	8.7	8.7	19.8	10.7	10.7	22.5	31.4	15.6					
44	9.3	11.3	12.0	12.0	4.3	14.2	16.9	16.9	16.9	8.2	20.9	20.9	31.2	15.2	15.2					
45	11.4	0.7	10.3	10.3	6.9	8.9	8.9	8.9	8.9	15.7	6.7	6.7	23.1	25.0	11.7					
46	19.1	9.3	9.9	9.9	7.4	11.0	19.3	19.3	19.3	8.8	23.1	23.1	32.1	7.1	7.1					
47	12.3	9.3	9.4	9.4	6.3	13.2	21.7	21.7	21.7	9.7	17.9	17.9	31.1	16.4	16.4					
48	13.4	9.7	5.7	5.7	3.8	8.5	13.8	13.8	13.8	9.8	18.2	18.2	25.5	13.7	13.7					
49	6.8	12.6	10.8	10.8	4.8	14.0	16.2	16.2	16.2	9.2	25.7	25.7	29.3	1.3	1.3					
50	5.9	9.0	10.9	10.9	4.0	13.4	10.8	10.8	10.8	5.9	19.9	19.9	20.6	16.5	16.5					
MEAN	12.5	9.0	10.1	10.1	8.5	9.3	16.1	16.1	16.1	10.6	19.4	19.4	26.0	13.5	13.5					
S.D.	3.60	2.63	2.38	2.38	3.70	3.42	3.37	3.37	3.37	3.71	2.98	2.98	5.41	4.21	4.21					
N	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25					

10-APR-14

05R018

TABLE : IIA- 019

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL MATERNAL BODY WEIGHT CHANGE DURING GESTATION -- GRAMS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	DAY OF GESTATION																			
	0	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
51	11.7	11.0	11.6	11.6	10.1	10.1	9.6	16.5	16.5	16.5	14.5	14.5	19.3	19.3	27.4	27.4	18.7	18.7	18.7	18.7
52	9.2	9.9	11.6	11.6	11.6	11.6	6.5	12.3	12.3	12.3	16.6	16.6	16.1	16.1	29.1	29.1	17.5	17.5	17.5	17.5
53	11.8	6.7	9.0	9.0	7.4	7.4	4.5	13.4	13.4	13.4	10.1	10.1	17.5	17.5	22.0	22.0	17.6	17.6	17.6	17.6
54	16.8	9.2	12.5	12.5	5.9	5.9	8.6	23.3	23.3	23.3	13.5	13.5	13.5	13.5	19.9	19.9	16.3	16.3	16.3	16.3
55	10.3	9.9	14.2	14.2	9.0	9.0	8.1	15.9	15.9	15.9	17.3	17.3	14.4	14.4	19.1	19.1	17.8	17.8	17.8	17.8
56	9.7	8.8	7.1	7.1	8.5	8.5	6.6	21.4	21.4	21.4	6.2	6.2	21.8	21.8	19.4	19.4	22.9	22.9	22.9	22.9
57	13.2	11.6	10.4	10.4	9.3	9.3	7.2	18.2	18.2	18.2	7.5	7.5	11.5	11.5	26.1	26.1	20.8	20.8	20.8	20.8
58	12.4	10.4	9.3	9.3	8.0	8.0	7.2	16.1	16.1	16.1	11.4	11.4	15.9	15.9	21.6	21.6	17.7	17.7	17.7	17.7
59	14.8	11.4	9.0	9.0	11.7	11.7	8.1	14.8	14.8	14.8	10.8	10.8	24.3	24.3	25.7	25.7	20.9	20.9	20.9	20.9
60	17.5	9.6	13.2	13.2	1.7	1.7	5.3	13.9	13.9	13.9	10.4	10.4	13.7	13.7	12.1	12.1	9.5	9.5	9.5	9.5
61	13.9	11.5	10.1	10.1	7.8	7.8	6.8	14.1	14.1	14.1	10.1	10.1	10.5	10.5	18.3	18.3	13.7	13.7	13.7	13.7
62	12.8	12.3	6.7	6.7	13.4	13.4	6.8	14.1	14.1	14.1	10.1	10.1	10.5	10.5	18.3	18.3	13.7	13.7	13.7	13.7
63	11.6	8.9	10.4	10.4	1.8	1.8	13.4	12.9	12.9	12.9	9.4	9.4	19.0	19.0	25.9	25.9	11.1	11.1	11.1	11.1
64	9.8	9.9	9.4	9.4	3.6	3.6	10.3	12.3	12.3	12.3	9.9	9.9	20.5	20.5	22.9	22.9	10.9	10.9	10.9	10.9
65	1.2	11.5	12.9	12.9	6.1	6.1	10.2	11.6	11.6	11.6	11.6	11.6	20.9	20.9	27.6	27.6	12.4	12.4	12.4	12.4
66	14.0	7.3	7.8	7.8	5.2	5.2	13.7	12.5	12.5	12.5	9.1	9.1	25.6	25.6	27.9	27.9	14.5	14.5	14.5	14.5
67	5.6	12.0	10.5	10.5	5.1	5.1	10.6	17.3	17.3	17.3	8.9	8.9	18.8	18.8	28.9	28.9	8.3	8.3	8.3	8.3
68	14.2	9.0	7.0	7.0	8.0	8.0	11.0	12.6	12.6	12.6	9.5	9.5	20.5	20.5	29.4	29.4	11.3	11.3	11.3	11.3
69	12.4	6.6	10.8	10.8	0.2	0.2	9.3	13.7	13.7	13.7	3.1	3.1	21.9	21.9	24.1	24.1	14.8	14.8	14.8	14.8
70	9.4	7.3	9.9	9.9	0.8	0.8	11.9	15.9	15.9	15.9	4.1	4.1	19.3	19.3	27.8	27.8	13.2	13.2	13.2	13.2
71	13.7	0.2	13.0	13.0	5.2	5.2	11.6	13.2	13.2	13.2	8.2	8.2	17.0	17.0	23.5	23.5	13.7	13.7	13.7	13.7
72	5.3	8.0	5.7	5.7	5.6	5.6	10.7	13.2	13.2	13.2	4.5	4.5	19.3	19.3	24.6	24.6	9.7	9.7	9.7	9.7
73	8.7	6.9	9.1	9.1	3.4	3.4	14.2	15.5	15.5	15.5	15.8	15.8	23.5	23.5	24.4	24.4	14.7	14.7	14.7	14.7
74	8.4	7.0	9.1	9.1	3.8	3.8	11.4	10.7	10.7	10.7	9.5	9.5	18.8	18.8	18.9	18.9	10.3	10.3	10.3	10.3
75	13.1	12.2	9.1	9.1	9.0	9.0	12.9	18.4	18.4	18.4	7.0	7.0	22.2	22.2	26.4	26.4	9.2	9.2	9.2	9.2
MEAN	11.3	9.2	10.0	10.0	6.8	6.8	9.4	15.1	15.1	15.1	10.1	10.1	18.9	18.9	24.0	24.0	14.4	14.4	14.4	14.4
S.D.	3.66	2.64	2.19	2.19	4.00	4.00	2.94	3.23	3.23	3.23	3.71	3.71	3.83	3.83	4.22	4.22	4.03	4.03	4.03	4.03
N	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25	25

10-APR-14

05R018

TABLE : IIA- 020

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL MATERNAL BODY WEIGHT CHANGE DURING GESTATION -- GRAMS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	DAY OF GESTATION																													
	0	1	1	2	3	3	4	5	6	6	7	8	8	9	10	10	11	12	13	13	14	15	15	16	17	17	18	19	20	
76	13.5	10.5	13.8	13.8	7.2	7.2	5.8	5.8	12.9	16.6	17.4	22.6	17.0																	
77	14.6	10.8	9.1	10.4	5.6	5.6	14.2	10.4	16.3	16.3	28.8	23.4																		
78	13.6	9.0	11.8	6.9	0.6	0.6	13.9	10.7	11.8	11.8	20.7	9.6																		
79	14.7	8.3	9.8	4.2	5.4	5.4	12.7	6.1	17.6	17.6	13.6	16.5																		
80	21.9	9.9	5.6	8.2	1.0	1.0	20.1	11.6	19.0	19.0	25.5	14.6																		
81	12.3	8.0	16.8	6.9	5.1	5.1	13.2	8.7	15.5	15.5	20.8	14.6																		
82	14.0	7.3	8.9	7.3	3.8	3.8	11.6	13.2	19.0	21.2	15.9	10.2																		
83	11.8	7.6	9.5	10.4	4.0	4.0	16.8	10.1	12.1	22.3	10.2	20.6																		
84	17.7	13.3	7.7	11.7	1.8	1.8	19.0	10.3	15.3	20.8	20.6																			
85	19.2	13.6	9.0	13.9	8.7	8.7	13.7	18.5	15.7	23.2	16.3																			
86	20.7	11.1	11.2	6.0	5.2	5.2	23.0	13.6	15.2	20.7	12.9																			
87	20.3	9.4	8.7	9.9	2.9	2.9	17.9	12.0	11.7	24.1	15.2																			
88	13.1	12.9	9.7	11.6	4.8	4.8	12.4	5.4	16.5	14.1	10.4																			
89	9.5	7.8	9.6	1.0	6.7	6.7	10.5	11.1	17.0	27.6	11.8																			
90	7.2	10.3	9.9	2.2	12.6	10.7	7.7	20.5	21.6	14.6																				
91	12.0	8.3	11.5	1.1	13.9	14.3	12.5	18.0	27.9	10.2																				
92	5.1	15.3	11.6	3.7	14.2	12.4	12.6	19.4	28.9	15.5																				
93x	12.3	10.9	7.6	-0.8	6.8	4.1	-8.9	5.8	-0.2	3.3																				
94	8.4	11.1	8.1	6.7	8.5	19.3	7.6	19.9	20.8	7.7																				
95	11.2	8.8	13.2	4.9	10.3	10.5	6.2	19.8	24.2	14.7																				
96	-11.6	27.1	7.1	5.7	11.0	11.3	4.5	21.1	24.7	11.0																				
97	11.9	13.9	6.4	5.9	10.1	13.3	10.0	22.9	30.6	16.2																				
98	10.7	5.2	11.0	4.7	6.5	13.1	4.9	12.3	25.4	8.7																				
99	-5.4	20.6	10.5	4.6	6.5	13.1	7.6	21.3	25.0	18.0																				
100	10.4	7.7	9.2	7.3	9.1	18.3	0.9	24.3	20.2	13.2																				
MEAN	11.5	11.2	10.0	6.8	6.8	14.5	9.7	17.5	23.3	14.1																				
S.D.	7.52	4.71	2.44	3.33	3.81	3.40	3.99	3.45	3.83	3.77																				
N	24	24	24	24	24	24	24	24	24	24																				

NP=NOT PREGNANT X=EXCLUDED FROM MEAN

10-APR-14 05R018 PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
TABLE : IIA- 021

ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL UTERINE WEIGHTS AND NET MATERNAL BODY WEIGHT CHANGE -- GRAMS

TEST GROUP 0 (0 MG/KG BW/D)	UTERUS WEIGHT	CARCASS WEIGHT	NET WEIGHT CHANGE FROM DAY 6	WEIGHT CHANGE FROM DAY 6
ANIMAL#				
1	60.5	230.5	45.0	105.5
2	58.2	227.5	37.8	96.0
3	63.0	231.5	47.9	110.9
4	66.6	238.7	49.1	115.7
5	74.9	255.5	44.5	119.4
6	65.1	227.6	38.9	104.0
7	52.3	234.2	40.0	92.3
8	56.3	225.7	34.1	90.4
9	69.3	248.1	41.0	110.3
10	43.4	227.4	45.4	88.8
11	64.9	251.0	46.4	111.3
12	55.1	242.0	46.6	101.7
13	43.5	237.7	44.7	88.2
14	57.9	256.0	47.7	105.6
15	56.0	225.0	38.6	94.6
16	56.8	228.3	34.7	91.5
17	62.5	243.1	36.7	99.2
18	64.3	226.5	30.4	94.7
19	54.5	234.6	34.2	88.7
20	62.8	234.0	41.7	104.5
21	47.7	247.4	43.3	91.0
22	55.7	233.6	38.2	93.9
23	57.0	241.7	34.7	91.7
24	62.8	240.6	38.6	101.4
25	65.6	232.9	42.8	108.4
MEAN	59.1	236.8	40.9	100.0
S.D.	7.5	9.3	5.1	9.2
N	25	25	25	25

CARCASS WEIGHT = TERMINAL BODY WEIGHT MINUS UTERINE WEIGHT
NET WEIGHT CHANGE FROM DAY 6 = CARCASS WEIGHT MINUS DAY 6 BODY WEIGHT

10-APR-14	05R018	TABLE : IIA-	022
PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS			
ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL UTERINE WEIGHTS AND NET MATERNAL BODY WEIGHT CHANGE -- GRAMS			
TEST GROUP 1 (30 MG/KG BW/D)	UTERUS WEIGHT	CARCASS WEIGHT	NET WEIGHT CHANGE FROM DAY 6
ANIMAL#			FROM DAY 6
26	50.9	220.6	39.3
27	66.4	253.6	60.9
28	58.1	255.8	53.9
29	44.8	230.9	46.2
30	79.4	269.4	56.1
31	45.3	267.9	58.4
32	45.2	219.9	35.3
33	61.9	213.8	35.0
34	48.4	215.3	45.4
35	66.7	255.3	50.6
36	62.5	253.4	54.2
37	46.7	256.5	54.8
38	71.0	248.7	42.3
39	51.4	218.3	44.0
40	54.1	232.0	35.2
41	64.8	239.5	32.9
42	52.2	234.2	37.9
43	69.9	258.4	42.4
44	67.9	239.2	43.0
45	59.0	242.2	39.0
46	69.7	255.3	39.1
47	71.4	263.1	44.9
48	65.9	240.7	27.4
49	64.8	249.5	35.7
50	51.3	237.1	39.8
MEAN	59.6	242.8	43.7
S.D.	9.9	16.6	8.8
N	25	25	25

CARCASS WEIGHT = TERMINAL BODY WEIGHT MINUS UTERINE WEIGHT
NET WEIGHT CHANGE FROM DAY 6 = CARCASS WEIGHT MINUS DAY 6 BODY WEIGHT

10-APR-14 05R018 PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS TABLE : IIA- 023

ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL UTERINE WEIGHTS AND NET MATERNAL BODY WEIGHT CHANGE -- GRAMS
TEST GROUP 2 (100 MG/KG BW/D)

ANIMAL#	UTERUS WEIGHT	CARCASS WEIGHT	NET WEIGHT CHANGE FROM DAY 6	WEIGHT CHANGE FROM DAY 6
51	69.2	241.7	40.9	110.1
52	61.6	249.8	48.1	109.7
53	54.6	229.0	37.9	92.5
54	58.6	247.8	46.1	104.7
55	50.2	263.0	51.4	101.6
56	67.1	222.5	39.7	106.8
57	67.1	271.8	60.9	128.0
58	45.1	231.5	46.0	91.1
59	60.0	232.1	42.5	102.5
60	75.1	241.4	32.1	107.2
61	27.8	232.2	44.9	72.7
62	49.5	222.9	37.4	86.9
63	56.8	225.3	36.7	93.5
64	57.1	228.0	33.3	90.4
65	62.2	249.7	38.2	100.4
66	73.0	247.6	35.5	108.5
67	62.1	249.5	35.8	97.9
68	63.9	239.3	38.4	102.3
69	56.9	231.7	30.2	87.1
70	60.6	231.3	32.4	93.0
71	56.1	238.1	36.3	92.4
72	53.4	222.9	34.2	87.6
73	67.0	250.4	44.5	111.5
74	52.5	220.0	30.9	83.4
75	60.3	258.5	44.8	105.1
MEAN	58.7	239.1	40.0	98.7
S.D.	9.7	13.7	7.2	11.5
N	25	25	25	25

CARCASS WEIGHT = TERMINAL BODY WEIGHT MINUS UTERINE WEIGHT
NET WEIGHT CHANGE FROM DAY 6 = CARCASS WEIGHT MINUS DAY 6 BODY WEIGHT

10-APR-14 05R018 PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS TABLE : IIA- 024

ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL UTERINE WEIGHTS AND NET MATERNAL BODY WEIGHT CHANGE -- GRAMS
TEST GROUP 3 (300 MG/KG BW/D)

ANIMAL#	UTERUS WEIGHT	CARCASS WEIGHT	NET WEIGHT CHANGE FROM DAY 6	WEIGHT CHANGE FROM DAY 6
76	63.2	229.0	36.3	99.5
77	66.8	232.0	42.3	109.1
78	42.1	223.5	32.1	74.2
79	50.6	210.2	26.6	77.2
80	64.6	230.5	37.3	101.9
81	53.9	228.9	30.9	84.8
82	56.2	216.1	35.8	92.0
83	57.5	211.8	28.4	85.9
84	54.8	247.2	44.7	99.5
85	57.0	252.9	53.0	110.0
86	41.9	260.3	54.7	96.6
87	55.0	251.9	38.7	93.7
88	43.2	214.7	32.0	75.2
89	66.4	203.8	19.3	85.7
90	53.1	233.5	36.8	89.9
91	66.0	241.2	31.9	97.9
92	63.2	256.5	43.5	106.7
93x NP	0.5	220.5	9.6	10.1
94	49.2	250.5	41.3	90.5
95	60.6	242.3	30.0	90.6
96	58.1	228.1	31.2	89.3
97	66.0	273.0	43.0	109.0
98	35.8	239.5	39.8	75.6
99	60.7	241.1	35.4	96.1
100	53.0	226.1	40.3	93.3
MEAN	55.8	235.2	36.9	92.7
S.D.	8.6	17.4	8.0	10.7
N	24	24	24	24

CARCASS WEIGHT = TERMINAL BODY WEIGHT MINUS UTERINE WEIGHT
NET WEIGHT CHANGE FROM DAY 6 = CARCASS WEIGHT MINUS DAY 6 BODY WEIGHT

NP=NOT PREGNANT X=EXCLUDED FROM MEAN

10-APR-14	05R018	TABLE : IIA-	025
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS			
ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL MATERNAL NECROPSY OBSERVATIONS			
TEST GROUP 0 (0 MG/KG BW/D)			
ANIMAL#	OBSERVATION	POSITION/GRADE	
1	NOTHING AENORMAL DETECTED		
2	NOTHING AENORMAL DETECTED		
3	NOTHING AENORMAL DETECTED		
4	NOTHING AENORMAL DETECTED		
5	NOTHING AENORMAL DETECTED		
6	NOTHING AENORMAL DETECTED		
7	NOTHING AENORMAL DETECTED		
8	NOTHING AENORMAL DETECTED		
9	NOTHING AENORMAL DETECTED		
10	NOTHING AENORMAL DETECTED		
11	NOTHING AENORMAL DETECTED		
12	NOTHING AENORMAL DETECTED		
13	NOTHING AENORMAL DETECTED		
14	NOTHING AENORMAL DETECTED		
15	NOTHING AENORMAL DETECTED		
16	NOTHING AENORMAL DETECTED		
17	NOTHING AENORMAL DETECTED		
18	NOTHING AENORMAL DETECTED		
19	NOTHING AENORMAL DETECTED		
20	NOTHING AENORMAL DETECTED		
POSITION/GRADE CODE: R-RIGHT, L-LEFT, B-BILATE , 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT			

10-APR-14	05R018	TABLE : IIA-	026
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS			
ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL MATERNAL NECROPSY OBSERVATIONS			
TEST GROUP 0 (0 MG/KG BW/D)			
ANIMAL#	OBSERVATION	POSITION/GRADE	
21	NOTHING ABNORMAL DETECTED		
22	NOTHING ABNORMAL DETECTED		
23	NOTHING ABNORMAL DETECTED		
24	NOTHING ABNORMAL DETECTED		
25	NOTHING ABNORMAL DETECTED		
POSITION/GRADE CODE: R-RIGHT, L-LEFT, B-BILATE, 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT			

10-APR-14	05R018	TABLE : IIA-	027
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS			
ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL MATERNAL NECROPSY OBSERVATIONS			
TEST GROUP 1 (30 MG/KG BW/D)			
ANIMAL#	OBSERVATION	POSITION/GRADE	
26	NOTHING ABNORMAL DETECTED		
27	NOTHING ABNORMAL DETECTED		
28	NOTHING ABNORMAL DETECTED		
29	NOTHING ABNORMAL DETECTED		
30	NOTHING ABNORMAL DETECTED		
31	NOTHING ABNORMAL DETECTED		
32	NOTHING ABNORMAL DETECTED		
33	NOTHING ABNORMAL DETECTED		
34	NOTHING ABNORMAL DETECTED		
35	NOTHING ABNORMAL DETECTED		
36	NOTHING ABNORMAL DETECTED		
37	NOTHING ABNORMAL DETECTED		
38	NOTHING ABNORMAL DETECTED		
39	NOTHING ABNORMAL DETECTED		
40	NOTHING ABNORMAL DETECTED		
41	NOTHING ABNORMAL DETECTED		
42	NOTHING ABNORMAL DETECTED		
43	NOTHING ABNORMAL DETECTED		
44	NOTHING ABNORMAL DETECTED		
45	NOTHING ABNORMAL DETECTED		
POSITION/GRADE CODE: R-RIGHT, L-LEFT, B-BILATE, 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT			

10-APR-14	05R018	TABLE : IIA-	028
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS			
ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL MATERNAL NECROPSY OBSERVATIONS			
TEST GROUP 1 (30 MG/KG BW/D)			
ANIMAL#	OBSERVATION	POSITION/GRADE	
46	NOTHING AENORMAL DETECTED		
47	NOTHING AENORMAL DETECTED		
48	NOTHING AENORMAL DETECTED		
49	NOTHING AENORMAL DETECTED		
50	NOTHING AENORMAL DETECTED		
POSITION/GRADE CODE: R-RIGHT, L-LEFT, B-BILATE, 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT			

10-APR-14	05R018	TABLE : IIA-	029
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS			
ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL MATERNAL NECROPSY OBSERVATIONS			
TEST GROUP 2 (100 MG/KG BW/D)			
ANIMAL#	OBSERVATION	POSITION/GRADE	
51	NOTHING ABNORMAL DETECTED		
52	NOTHING ABNORMAL DETECTED		
53	NOTHING ABNORMAL DETECTED		
54	NOTHING ABNORMAL DETECTED		
55	NOTHING ABNORMAL DETECTED		
56	NOTHING ABNORMAL DETECTED		
57	NOTHING ABNORMAL DETECTED		
58	NOTHING ABNORMAL DETECTED		
59	NOTHING ABNORMAL DETECTED		
60	NOTHING ABNORMAL DETECTED		
61	NOTHING ABNORMAL DETECTED		
62	NOTHING ABNORMAL DETECTED		
63	NOTHING ABNORMAL DETECTED		
64	NOTHING ABNORMAL DETECTED		
65	NOTHING ABNORMAL DETECTED		
66	NOTHING ABNORMAL DETECTED		
67	NOTHING ABNORMAL DETECTED		
68	NOTHING ABNORMAL DETECTED		
69	NOTHING ABNORMAL DETECTED		
70	NOTHING ABNORMAL DETECTED		
POSITION/GRADE CODE: R-RIGHT, L-LEFT, B-BILATE, 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT			

10-APR-14	05R018	TABLE : IIA-	030
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS			
ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL MATERNAL NECROPSY OBSERVATIONS			
TEST GROUP 2 (100 MG/KG BW/D)			
ANIMAL#	OBSERVATION	POSITION/GRADE	
71	NOTHING ABNORMAL DETECTED		
72	NOTHING ABNORMAL DETECTED		
73	NOTHING ABNORMAL DETECTED		
74	NOTHING ABNORMAL DETECTED		
75	NOTHING ABNORMAL DETECTED		
POSITION/GRADE CODE: R-RIGHT, L-LEFT, B-BILATE, 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT			

10-APR-14	05R018	TABLE : IIA-	031
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS			
ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL MATERNAL NECROPSY OBSERVATIONS			
TEST GROUP 3 (300 MG/KG BW/D)			
ANIMAL#	OBSERVATION	POSITION/GRADE	
76	NOTHING ABNORMAL DETECTED		
77	NOTHING ABNORMAL DETECTED		
78	NOTHING ABNORMAL DETECTED		
79	NOTHING ABNORMAL DETECTED		
80	NOTHING ABNORMAL DETECTED		
81	NOTHING ABNORMAL DETECTED		
82	NOTHING ABNORMAL DETECTED		
83	NOTHING ABNORMAL DETECTED		
84	NOTHING ABNORMAL DETECTED		
85	NOTHING ABNORMAL DETECTED		
86	NOTHING ABNORMAL DETECTED		
87	NOTHING ABNORMAL DETECTED		
88	NOTHING ABNORMAL DETECTED		
89	NOTHING ABNORMAL DETECTED		
90	NOTHING ABNORMAL DETECTED		
91	NOTHING ABNORMAL DETECTED		
92	NOTHING ABNORMAL DETECTED		
93	NOT PREGNANT		
94	DIAPHRAGMATIC HERNIA		
95	NOTHING ABNORMAL DETECTED		
POSITION/GRADE CODE: R-RIGHT, L-LEFT, B-BILATE, 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT			

10-APR-14	05R018	TABLE : IIA-	032
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS			
ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL MATERNAL NECROPSY OBSERVATIONS			
TEST GROUP 3 (300 MG/KG BW/D)			
ANIMAL#	OBSERVATION	POSITION/GRADE	
96	NOTHING AENORMAL DETECTED		
97	DILATED RENAL PELVIS	L	
98	HEMOMETRA	R	
99	NOTHING AENORMAL DETECTED		
100	NOTHING AENORMAL DETECTED		
POSITION/GRADE CODE: R-RIGHT, L-LEFT, B-BILATE, 1-SLIGHT, 2-MODERATE, 3-MARKED, P-PRESENT			

10-APR-14

05R018

TABLE : IIA- 033

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL REPRODUCTION DATA

TEST GROUP 0 (0 MG/KG BW/D)									
FEMALE#	CORPORA LUTEA	IMPLANT SITES	RESORPTIONS		FETUSES		SEX		%IMPLANTATION LOSS
			EARLY	LATE	DEAD	LIVE	MALE	FEMALE	
1	12	12	1	0	0	11	6	5	3.4
2	13	13	2	0	0	11	4	7	3.3
3	11	11	0	0	0	11	6	5	3.6
4	12	12	0	1	0	11	6	5	3.4
5	14	14	0	0	0	14	6	8	3.8
6	13	13	1	0	0	12	7	5	3.5
7	10	10	1	0	0	9	5	4	3.3
8	10	10	0	0	0	10	6	4	3.9
9	13	12	0	0	0	12	6	6	3.3
10	9	7	0	0	0	7	6	1	3.6
11	11	11	0	0	0	11	6	5	3.7
12	10	10	0	0	0	10	2	8	3.8
13	11	8	1	0	0	7	4	3	3.4
14	12	10	0	0	0	10	7	3	3.7
15	11	11	0	0	0	10	3	7	3.6
16	11	11	0	0	0	11	8	3	3.5
17	12	12	0	0	0	12	7	5	3.2
18	12	12	0	0	0	12	5	7	3.2
19	10	10	0	0	0	10	6	4	3.4
20	11	11	0	0	0	11	3	8	3.5
21	11	10	0	0	0	8	4	4	3.7
22	11	11	1	0	0	10	4	6	4.1
23	11	11	1	0	0	10	8	2	3.9
24	13	13	1	0	0	12	6	6	3.6
25	13	12	0	0	0	12	5	7	3.4
MEAN	11.5	11.1	0.5	0.0	0.0	10.6	5.4	5.2	3.6
S.D.	1.2	1.6	0.7	0.2	0.0	1.6	1.7	2.0	0.2
N	25	25	25	25	25	25	25	25	25

10-APR-14

05R018

TABLE : IIA- 034

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL REPRODUCTION DATA

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	CORPORA LUTEA	IMPLANT SITES	RESORPTIONS		FETUSES		SEX		AVERAGE FETAL BODY WEIGHT		% IMPLANTATION LOSS	
			EARLY	LATE	TOTAL	DEAD	LIVE		MALES	FEMALES	LITTER	PRE POST
26	15	9	0	0	0	0	9	5	3.6	4.0	3.8	40.0 0.0
27	13	12	0	0	0	0	12	3	4.0	3.5	3.6	7.7 0.0
28	12	12	1	0	1	0	11	4	3.4	3.2	3.3	0.0 8.3
29	10	8	0	0	0	0	8	4	3.7	3.4	3.5	20.0 0.0
30	15	15	1	0	1	0	14	7	3.9	3.5	3.7	0.0 6.7
31	9	8	0	0	0	0	8	4	3.8	3.4	3.6	11.1 0.0
32	10	10	2	0	2	0	8	3	3.7	3.3	3.5	0.0 20.0
33	11	11	0	0	0	0	11	5	3.7	3.4	3.6	0.0 0.0
34	10	9	1	0	1	0	8	4	4.1	3.9	4.0	10.0 11.1
35	13	12	0	0	0	0	12	5	3.7	3.6	3.6	7.7 0.0
36	12	12	1	0	1	0	11	7	3.8	3.6	3.8	0.0 8.3
37	9	9	1	0	1	0	8	5	3.6	3.3	3.5	0.0 11.1
38	12	12	0	0	0	0	12	3	3.8	3.6	3.7	0.0 0.0
39	10	10	1	0	1	0	9	4	3.6	3.4	3.5	0.0 10.0
40	11	10	0	0	0	0	10	5	3.6	3.3	3.5	9.1 0.0
41	13	13	1	0	1	0	12	7	3.7	3.5	3.6	0.0 7.7
42	11	9	0	0	0	0	9	6	3.7	3.4	3.6	18.2 0.0
43	14	12	0	0	0	0	12	7	3.7	3.6	3.7	14.3 0.0
44	13	12	0	0	0	0	12	6	3.7	3.4	3.6	7.7 0.0
45	11	11	0	0	0	0	11	4	3.4	3.2	3.3	0.0 0.0
46	12	12	0	0	0	0	12	2	3.5	3.3	3.5	0.0 0.0
47	13	13	0	0	0	0	13	7	3.7	3.5	3.6	0.0 0.0
48	14	14	2	0	2	0	12	5	3.6	3.4	3.5	0.0 14.3
49	13	12	0	0	0	0	12	6	3.7	3.4	3.5	7.7 0.0
50	9	9	0	0	0	0	9	5	3.6	3.4	3.5	0.0 0.0
MEAN	11.8	11.0	0.4	0.0	0.4	0.0	10.6	5.6	3.7	3.5	3.6	6.1 3.9
S.D.	1.8	1.9	0.7	0.0	0.7	0.0	1.8	1.7	0.2	0.2	0.1	9.4 5.8
N	25	25	25	25	25	25	25	25	25	25	25	25 25

10-APR-14

05R018

TABLE : IIA- 035

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL REPRODUCTION DATA

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	CORPORA LUTEA	IMPLANT SITES	RESORPTIONS		FETUSES		SEX		AVERAGE FETAL BODY WEIGHT		% IMPLANTATION LOSS	
			EARLY	LATE	TOTAL	DEAD	LIVE	MALE	MALES	FEMALES	LITTER	PRE POST
51	12	12	0	0	0	0	12	5	3.8	3.4	3.6	0.0 0.0
52	12	11	0	0	0	0	11	5	3.5	3.3	3.4	8.3 0.0
53	11	11	0	0	1	0	10	4	3.5	3.4	3.4	0.0 9.1
54	11	11	0	0	1	0	10	4	3.8	3.6	3.7	0.0 9.1
55	12	10	1	0	1	0	9	4	3.6	3.5	3.5	16.7 10.0
56	14	13	1	0	1	0	12	4	3.7	3.4	3.5	7.1 7.7
57	12	12	0	0	0	0	12	2	3.6	3.5	3.5	0.0 0.0
58	9	8	0	0	0	0	8	1	3.8	3.3	3.4	11.1 0.0
59	13	13	2	0	2	0	11	9	3.3	3.2	3.3	0.0 15.4
60	14	13	0	0	0	0	13	4	3.9	3.7	3.8	7.1 0.0
61	12	5	0	0	0	0	5	4	3.2	3.6	3.3	58.3 0.0
62	11	9	0	0	0	0	9	3	3.6	3.3	3.4	18.2 0.0
63	12	12	1	0	1	0	11	5	3.3	3.3	3.3	0.0 8.3
64	11	11	0	0	0	0	11	5	3.3	3.3	3.3	0.0 0.0
65	11	11	0	0	0	0	11	6	3.5	3.4	3.5	0.0 0.0
66	13	13	0	0	0	0	13	5	3.5	3.4	3.4	0.0 0.0
67	11	11	0	0	0	0	11	3	3.7	3.5	3.6	0.0 0.0
68	12	12	0	0	0	0	12	4	3.2	3.0	3.1	0.0 0.0
69	10	10	0	0	0	0	10	4	3.7	3.6	3.7	0.0 0.0
70	13	13	1	1	2	0	11	7	3.5	3.2	3.4	0.0 15.4
71	13	13	2	0	2	0	11	6	3.4	3.3	3.3	0.0 15.4
72	12	12	0	1	1	0	11	4	3.2	2.7	3.3	8.3 0.0
73	12	12	0	0	0	0	12	6	3.6	3.5	3.5	0.0 0.0
74	9	9	0	0	0	0	9	6	3.8	3.5	3.7	0.0 0.0
75	11	10	0	0	0	0	10	3	3.6	3.8	3.8	9.1 0.0
MEAN	11.7	11.1	0.4	0.1	0.5	0.0	10.6	4.6	3.5	3.4	3.4	5.4 3.9
S.D.	1.3	1.9	0.6	0.3	0.7	0.0	1.7	1.7	0.2	0.2	0.2	12.3 5.7
N	25	25	25	25	25	25	25	25	25	25	25	25 25

10-APR-14

05R018

TABLE : IIA- 036

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL REPRODUCTION DATA

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	CORPORA LUTEA	IMPLANT SITES	RESORPTIONS		FETUSES		SEX		AVERAGE FETAL BODY WEIGHT		% IMPLANTATION LOSS	
			EARLY	LATE	TOTAL	DEAD	MALE	FEMALE	MALES	FEMALES	PRE	POST
76	13	11	0	0	0	0	4	7	4.0	3.7	15.4	0.0
77	12	12	0	0	0	0	9	3	3.4	3.2	0.0	0.0
78	9	9	1	0	1	0	4	4	3.3	3.1	0.0	11.1
79	10	10	1	0	1	0	4	5	3.8	3.4	0.0	10.0
80	12	12	0	0	0	0	5	7	3.6	3.3	0.0	0.0
81	11	11	2	0	2	0	6	3	3.9	3.5	0.0	18.2
82	12	10	0	0	0	0	3	7	3.6	3.4	16.7	0.0
83	12	12	1	0	1	0	11	2	3.1	3.2	0.0	8.3
84	11	11	1	0	1	0	10	5	3.5	3.0	0.0	9.1
85	11	11	1	0	1	0	5	5	3.7	3.6	0.0	9.1
86	9	8	0	1	1	0	7	5	3.8	3.6	11.1	12.5
87	11	11	1	0	1	0	5	5	3.4	3.4	0.0	9.1
88	10	10	2	0	2	0	5	3	3.3	3.2	0.0	20.0
89	14	14	0	0	0	0	6	8	3.0	2.8	0.0	0.0
90	10	10	1	0	1	0	6	3	3.6	3.5	0.0	10.0
91	13	12	0	0	0	0	6	6	3.6	3.4	7.7	0.0
92	12	11	0	0	0	0	8	3	3.6	3.6	8.3	0.0
93	NP											
94	10	10	1	0	1	0	1	8	3.5	3.3	0.0	10.0
95	12	12	1	0	1	0	6	5	3.4	3.3	0.0	8.3
96	12	11	0	0	0	0	5	6	3.4	3.3	8.3	0.0
97	14	14	2	0	2	0	7	5	3.5	3.3	0.0	14.3
98	12	6	0	0	0	0	4	2	3.7	3.3	50.0	0.0
99	11	11	0	0	0	0	8	3	3.5	3.4	0.0	0.0
100	11	10	0	0	0	0	5	5	3.4	3.2	9.1	0.0
MEAN	11.4	10.8	0.6	0.0	0.7	0.0	5.3	4.8	3.5	3.3	5.3	6.3
S.D.	1.3	1.7	0.7	0.2	0.7	0.0	2.0	1.8	0.2	0.2	10.9	6.5
N	24	24	24	24	24	24	24	24	24	24	24	24

NP=NOT PREGNANT

10-APR-14 05R018 PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL FETAL STATUS AND UTERINE LOCATION

TABLE : IIB- 001

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE# IMPLANT # 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23

1	FA	MA	MA	MA	FA	FA	FA	MA	FA	FA	MA	MA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA
2	FA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
3	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
4	MA	MA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA	FA
5	MA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
6	FA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
7	MA	FA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
8	MA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
9	MA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
10	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
11	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
12	FA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
13	FA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
14	MA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
15	FA	FA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
16	FA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
17	MA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
18	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
19	MA	FA	MA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
20	FA	FA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
21	MA	E	MA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
22	FA	E	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
23	MA	MA	FA	FA	E	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
24	FA	FA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
25	FA	MA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA

M MALE F FEMALE U UNCERTAIN / DENOTES POSITION OF CERVIX
A ALIVE E EARLY RESORPTION L LATE RESORPTION D DEAD FETUS B ABORTED FETUS P PREMATURE DELIVERY

10-APR-14

05R018

002

TABLE : IIB-

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL STATUS AND UTERINE LOCATION

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE# IMPLANT # 1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23

26	MA	FA	FA	FA	MA	MA	MA	FA	FA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
27	FA	FA	FA	MA	MA	FA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
28	FA	FA	MA	MA	MA	FA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
29	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
30	MA	MA	MA	MA	E	MA	FA	FA	FA	FA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA
31	MA	MA	MA	MA	FA	FA	FA	FA	FA	FA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA
32	FA	FA	MA	E	MA	E	MA	FA	FA	FA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA
33	FA	MA	MA	MA	FA	FA	MA	FA	FA	FA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA
34	FA	FA	E	MA	MA	MA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
35	FA	FA	MA	MA	FA	FA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
36	MA	FA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
37	MA	MA	MA	FA	E	MA	MA	FA	FA	FA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA
38	FA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
39	FA	MA	E	FA	FA	FA	MA	E	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
40	MA	FA	FA	FA	FA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
41	MA	MA	E	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
42	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
43	MA	MA	FA	MA	MA	MA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
44	MA	MA	FA	FA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
45	FA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
46	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
47	FA	FA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
48	FA	FA	MA	FA	MA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
49	FA	FA	MA	FA	MA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA
50	FA	MA	FA	MA	MA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA	MA

M MALE F FEMALE U UNCERTAIN / DENOTES POSITION OF CERVIX
A ALIVE E EARLY RESORPTION L LATE RESORPTION D DEAD FETUS B ABORTED FETUS P PREMATURE DELIVERY

10-APR-14 05R018 PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL FETAL STATUS AND UTERINE LOCATION
TABLE : IIB- 003

TEST GROUP 2 (100 MG/KG BW/D)		INDIVIDUAL FETAL STATUS AND UTERINE LOCATION																						
FEMALE#	IMPLANT #	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23
51	FA	MA	FA	FA	FA	MA	FA	MA	MA	FA	FA	MA	MA											
52	MA	MA	FA	FA	FA	FA	MA	FA	MA	FA	MA	FA	FA											
53	MA	FA	FA	FA	FA	MA	MA	FA	E	FA	FA	MA	MA											
54	FA	MA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	E											
55	FA	FA	MA	FA	E	FA	MA	MA	MA	MA	MA	MA	MA											
56	FA	E	FA	MA	FA	FA	MA	FA	FA	FA	MA	FA	FA	MA										
57	FA	FA	MA	FA	FA	FA	FA	FA	FA	FA	FA	FA	MA											
58	FA	FA	MA	FA	FA	FA	FA	FA	FA	FA	FA	FA	MA											
59	FA	MA	E	MA	E	MA	MA	MA	FA	MA	MA	MA	MA	MA										
60	MA	FA	FA	MA	MA	MA	FA	FA	FA	MA	FA	FA	FA	FA										
61	MA	MA	MA	MA	FA	FA	FA	FA	FA	MA	FA	FA	FA	FA										
62	FA	FA	MA	MA	MA	FA	FA	MA	FA	FA	MA	FA	E											
63	MA	FA	FA	MA	MA	MA	MA	FA	FA	FA	MA	FA	MA											
64	FA	MA	FA	MA	MA	MA	MA	MA	FA	FA	FA	MA	MA											
65	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	FA	MA	MA	FA										
66	FA	MA	MA	FA	FA	FA	FA	FA	FA	FA	MA	MA	FA	FA										
67	FA	MA	FA	FA	FA	FA	MA	MA	FA	FA	MA	MA	FA	FA										
68	FA	MA	MA	FA	FA	FA	FA	MA	FA	MA	FA	MA	FA	FA										
69	MA	FA	MA	FA	FA	FA	FA	MA	FA	MA	FA	MA	FA	FA										
70	MA	MA	MA	MA	E	MA	FA	MA	MA	MA	MA	MA	FA	FA										
71	E	E	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA	FA	FA										
72	MA	FA	FA	MA	MA	FA	FA	FA	MA	MA	L	MA	FA	FA										
73	MA	MA	MA	FA	MA	FA	MA	FA	FA	FA	MA	MA	FA	FA										
74	MA	MA	MA	MA	FA	MA	MA	MA	FA	MA	MA	MA	FA	FA										
75	MA	FA	MA	MA	FA	FA	FA	FA	FA	MA	MA	FA	FA											

M MALE F FEMALE U UNCERTAIN / DENOTES POSITION OF CERVIX
A ALIVE E EARLY RESORPTION L LATE RESORPTION D DEAD FETUS B ABORTED FETUS P PREMATURE DELIVERY

10-APR-14

05R018

TABLE : IIB- 004

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL STATUS AND UTERINE LOCATION

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	IMPLANT #	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23
76	FA	FA	MA	MA	MA	FA	FA	MA	MA	FA	FA	FA	FA	MA										
77	MA	FA	MA	MA	MA	MA	MA	MA	MA	FA	MA	FA	MA	MA										
78	MA	FA	FA	MA	MA	MA	MA	MA	FA	E														
79	FA	FA	MA	MA	FA	FA	FA	FA	FA	MA	MA	MA	MA	FA										
80	FA	MA	MA	MA	FA	FA	FA	FA	FA	MA	MA	MA	MA	FA										
81	FA	FA	E	MA	E	MA	MA	MA	MA	MA	MA	MA	MA	MA										
82	FA	MA	MA	MA	FA	FA	FA	FA	FA	FA	FA	FA	FA	MA										
83	MA	MA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	E	MA									
84	FA	MA	MA	MA	MA	FA	FA	MA	MA	E	MA	FA	FA	FA	FA									
85	MA	MA	E	FA	FA	FA	MA	MA	MA	FA	MA	MA	MA	FA	FA									
86	L	FA	FA	FA	FA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA									
87	MA	FA	FA	MA	MA	FA	FA	E	MA	MA	FA	MA	MA	FA	FA									
88	FA	E	MA	FA	FA	E	MA	FA	MA	MA	MA	MA	MA	FA	FA									
89	FA	MA	FA	FA	FA	MA	MA	MA	MA	FA	MA	MA	MA	FA	FA									
90	MA	FA	MA	MA	MA	E	FA	MA	MA	MA	MA	MA	MA	MA	FA									
91	MA	FA	FA	MA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	FA									
92	MA	MA	MA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	FA	FA									
93																								
94	FA	FA	FA	MA	MA	MA	FA	FA	FA	E	FA	FA	FA	FA	FA									
95	FA	FA	MA	MA	MA	MA	FA	FA	FA	MA	MA	MA	FA	E										
96	MA	FA	MA	MA	MA	FA	MA	FA	FA	MA	FA	FA	FA	FA	FA									
97	FA	MA	MA	E	MA	MA	FA	FA	FA	MA	FA	E	MA	FA	MA									
98	MA	MA	MA	FA	MA	MA	FA	MA	MA	MA	MA	MA	MA	MA	MA									
99	FA	MA	MA	MA	MA	FA	FA	MA	MA	MA	MA	MA	MA	MA	MA									
100	FA	MA	FA	MA	FA	MA	MA	FA	MA	FA	MA	MA	MA	MA	MA									

NOT PREGNANT

M MALE F FEMALE U UNCERTAIN / DENOTES POSITION OF CERVIX
A ALIVE E EARLY RESORPTION L LATE RESORPTION D DEAD FETUS B ABORTED FETUS P PREMATURE DELIVERY

10-APR-14 05R018 PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS TABLE : IIB- 005

ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL PLACENTAL WEIGHTS -- GRAMS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	MEAN	FETUS#	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23
1	0.42	0.36	0.50	0.40	0.41	0.45	0.42	0.45	0.36	0.41	0.35	E	0.48												
2	0.43	0.45	0.43	0.41	0.36	0.49	0.41	0.45	0.46	0.45	0.41	E	0.45												
3	0.50	0.38	0.54	0.50	0.51	0.55	0.54	0.49	0.43	0.42	0.53	0.61													
4	0.46	0.45	0.45	0.50	0.48	0.44	1	0.50	0.41	0.51	0.45	0.41	0.41												
5	0.46	0.39	0.42	0.41	0.42	0.50	0.39	0.47	0.46	0.48	0.41	0.46	0.49	0.44	0.40										
6	0.47	0.47	0.51	0.40	0.42	0.46	0.46	0.47	0.52	0.47	0.48	0.47	E	0.48											
7	0.45	0.45	0.43	0.41	0.46	0.49	E	0.48	0.45	0.47	0.40														
8	0.48	0.54	0.42	0.52	0.51	0.50	0.46	0.40	0.48	0.42	0.55														
9	0.40	0.39	0.31	0.44	0.38	0.46	0.55	0.38	0.36	0.34	0.36	0.36	0.45												
10	0.57	0.50	0.56	0.61	0.57	0.59	0.53	0.61																	
11	0.45	0.41	0.33	0.50	0.46	0.43	0.41	0.54	0.41	0.44	0.51	0.55													
12	0.45	0.44	0.43	0.44	0.39	0.44	0.51	0.47	0.55	0.43	0.37														
13	0.49	0.44	0.56	0.47	0.52	0.47	0.47	0.50	E																
14	0.46	0.57	0.39	0.48	0.45	0.39	0.46	0.46	0.43	0.47	0.47	0.47	0.37												
15	0.44	0.46	0.41	0.51	0.43	0.38	0.44	0.50	0.45	0.46	E	0.37													
16	0.41	0.34	0.43	0.40	0.43	0.39	0.46	0.38	0.45	0.39	0.39	0.42													
17	0.41	0.39	0.43	0.42	0.37	0.37	0.40	0.44	0.37	0.38	0.42	0.45	0.53												
18	0.42	0.39	0.41	0.42	0.67	0.43	0.44	0.37	0.41	0.37	0.44	0.41	0.33												
19	0.42	0.45	0.38	0.39	0.41	0.45	0.42	0.43	0.45	0.44	0.38														
20	0.44	0.43	0.49	0.43	0.46	0.37	0.49	0.49	0.38	0.45	0.42	0.41													
21	0.50	0.52	E	0.49	0.57	0.50	0.41	0.58	0.48	E	0.48														
22	0.47	0.44	E	0.45	0.42	0.52	0.49	0.54	0.44	0.54	0.48	0.42													
23	0.47	0.47	0.61	0.37	0.46	E	0.48	0.45	0.46	0.49	0.46	0.45													
24	0.44	0.39	0.46	0.44	0.39	0.40	0.41	0.48	0.51	0.63	E	0.41	0.40	0.40											
25	0.46	0.43	0.42	0.51	0.46	0.40	0.58	0.44	0.45	0.43	0.45	0.48	0.45												

MEAN 0.45
S.D. 0.04
N 25

E-EARLY RESORPTION L-LATE RESORPTION D-DEAD FETUS /-DENOTES POSITION OF CERVIX

10-APR-14		05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS														TABLE : IIB-		006					
				ORAL ADMINISTRATION (GAVAGE)																					
				INDIVIDUAL PLACENTAL WEIGHTS -- GRAMS																					
TEST GROUP 1 (30 MG/KG BW/D)																									
FEMALE#	MEAN	FETUS#	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23
26	0.42	0.43	0.50/0.40	0.38	0.44	0.42	0.30	0.49	0.45																
27	0.53	0.46	0.51	0.55	0.50/0.56	0.52	0.53	0.56	0.59	0.52	0.56	0.55													
28	0.45	0.36	0.46	0.46	0.44	0.51	0.42	0.46	0.51/0.46	0.45	E	0.44													
29	0.48	0.48	0.43	0.41	0.43/0.46	0.47	0.53	0.62																	
30	0.50	0.51	0.52	0.52	0.51	E	0.47	0.46	0.59/0.50	0.47	0.39	0.50	0.45	0.49	0.56										
31	0.47	0.43	0.42	0.55	0.52	0.48/0.41	0.50	0.48																	
32	0.46	0.43	0.49	0.47	E	0.51	E	0.54	0.49	0.44	0.33														
33	0.46	0.42	0.43	0.44	0.42	0.49/0.49	0.47	0.46	0.47	0.51	0.48														
34	0.51	0.47	0.61	E	0.53	0.56/0.48	0.51	0.43	0.51																
35	0.44	0.42	0.43	0.40	0.42	0.36	0.44	0.34	0.60	0.50/0.44	0.47	0.49													
36	0.51	0.60	0.50	0.47	0.61	0.41	0.52	0.54	0.54/0.51	0.50	0.46	E													
37	0.52	0.53	0.43	0.54	0.55	E	0.60/0.56	0.51	0.48																
38	0.43	0.47	0.42	0.43	0.43	0.38/0.45	0.50	0.42	0.39	0.38	0.45	0.46													
39	0.52	0.44	0.45	0.56	0.57	0.61/0.62	0.44	0.54	E	0.41															
40	0.39	0.38	0.40	0.38	0.39	0.38/0.40	0.38	0.39	0.43	0.34															
41	0.39	0.40	0.39	E	0.40	0.39	0.40	0.39/0.35	0.37	0.42	0.40	0.39	0.35												
42	0.47	0.48	0.48	0.35/0.46	0.54	0.53	0.44	0.48	0.47																
43	0.49	0.44	0.52	0.51	0.51	0.54/0.49	0.44	0.45	0.48	0.49	0.51	0.44													
44	0.40	0.36	0.34	0.34	0.34	0.36	0.48	0.43/0.54	0.43	0.37	0.41	0.36													
45	0.41	0.39	0.43	0.48	0.40	0.40	0.40	0.45/0.38	0.41	0.36	0.41														
46	0.55	0.45	0.43	0.52	0.58	0.43	0.48	0.54/1.01	0.60	0.45	0.57	0.49													
47	0.43	0.43	0.41	0.34	0.37	0.42	0.44	0.51/0.46	0.41	0.51	0.44	0.43	0.40												
48	0.47	0.47	0.39	0.58	0.42	0.40	0.53	0.48	0.41	E	0.55	0.59	0.42	0.44	E										
49	0.38	0.36	0.40	0.42	0.33	0.39	0.38	0.42	0.38/0.38	0.34	0.36	0.36													
50	0.44	0.40	0.42	0.44	0.46/0.51	0.50	0.50	0.46	0.39	0.41															
MEAN																									
S.D.	0.46																								
N	0.05																								
	25																								
				E-EARLY RESORPTION														L-LATE RESORPTION		D-DEAD FETUS		/-DENOTES POSITION OF CERVIX			

E-EARLY RESORPTION L-LATE RESORPTION D-DEAD FETUS /-DENOTES POSITION OF CERVIX

10-APR-14

05R018

TABLE : IIB-

007

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL PLACENTAL WEIGHTS -- GRAMS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	MEAN	FETUS#	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23
51	0.48	0.45	0.53	0.53	0.38	0.50	0.47	0.45	0.48	0.45	0.44	0.56	0.46												
52	0.50	0.44	0.59	0.57	0.51	0.49	0.44	0.56	0.60	0.43	0.46	0.44													
53	0.43	0.42	0.40	0.42	0.40	0.40	0.51	0.41/	E	0.39	0.49	0.50													
54	0.51	0.48	0.52	0.47	0.44	0.51	0.58	0.48	0.60	0.56	0.51	E													
55	0.46	0.42	0.41	0.46	0.47	E	0.59	0.50	0.49	0.36	0.45														
56	0.49	0.45	E	0.56	0.50	0.48	0.44	0.52	0.50	0.50	0.46	0.54	0.45	0.46											
57	0.46	0.50	0.39	0.45	0.53	0.47	0.51	0.51	0.44	0.47	0.43	0.42	0.40												
58	0.45	0.39	0.53	0.44	0.41	0.37	0.36	0.56	0.54																
59	0.48	0.41	0.52	E	0.54	E	0.45	0.48	0.48	0.51	0.46	0.49	0.50	0.45											
60	0.49	0.43	0.38	0.40	0.44	0.52	0.52	0.56	0.59	0.59	0.45	0.53	0.56	0.44											
61	0.45	0.44	0.46	0.42	0.45	0.47																			
62	0.44	0.42	0.51	0.48	0.48	0.40	0.38	0.43	0.38	0.46															
63	0.37	0.24	0.34	0.37	0.40	0.32	0.42	0.39	0.39	0.40	0.40	0.38	E												
64	0.39	0.32	0.37	0.39	0.45	0.42	0.37	0.40	0.40	0.42	0.37	0.40													
65	0.48	0.48	0.43	0.43	0.43	0.53	0.42	0.41	0.67	0.53	0.48	0.45													
66	0.46	0.49	0.37	0.45	0.45	0.40	0.47	0.38	0.43	0.49	0.50	0.41	0.57	0.54											
67	0.42	0.42	0.46	0.39	0.44	0.43	0.46	0.40	0.50	0.41	0.37	0.39													
68	0.50	0.44	0.55	0.43	0.52	0.53	0.49	0.56	0.46	0.61	0.50	0.46	0.49												
69	0.47	0.41	0.47	0.45	0.47	0.51	0.47	0.56	0.44	0.45	0.48														
70	0.44	0.47	0.38	0.48	E	0.47	0.36	0.46/	L	0.48	0.45	0.42	0.41	0.43											
71	0.38	E	0.35	0.41	0.43	0.39	0.47	0.36	0.33	0.41	0.34	0.38	0.32												
72	0.40	0.41	0.36	0.35	0.42	0.31	0.56	0.43	0.44	L	0.38	0.37													
73	0.42	0.45	0.36	0.42	0.53	0.43	0.38	0.48	0.43	0.44	0.40	0.39	0.39												
74	0.41	0.38	0.41	0.45	0.35	0.43	0.35	0.48	0.41	0.45															
75	0.51	0.58	0.46	0.53	0.49	0.54	0.44	0.48	0.46	0.58	0.55														

MEAN 0.45
S.D. 0.04
N 25

E-EARLY RESORPTION L-LATE RESORPTION D-DEAD FETUS /-DENOTES POSITION OF CERVIX

10-APR-14		05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS														TABLE : IIB-		008					
				ORAL ADMINISTRATION (GAVAGE)																					
				INDIVIDUAL PLACENTAL WEIGHTS -- GRAMS																					
TEST GROUP 3 (300 MG/KG BW/D)																									
FEMALE#	MEAN	FETUS#	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19	20	21	22	23
76	0.42	0.40	0.37	0.46	0.44	0.48	0.49	0.34	0.44	0.41	0.40	0.41													
77	0.50	0.53	0.48	0.50	0.53	0.53	0.48	0.55	0.46	0.50	0.53	0.58	0.39												
78	0.51	0.44	0.53	0.46	0.55	0.47	0.56	0.54	0.53	E															
79	0.45	0.44	0.40	0.50	0.44	0.39	0.40	0.46	0.49	0.53	E														
80	0.46	0.39	0.45	0.50	0.47	0.51	0.51	0.47	0.42	0.39	0.51	0.44	0.45												
81	0.49	0.41	0.43	E	0.54	E	0.48	0.59	0.43	0.53	0.58	0.46													
82	0.47	0.44	0.45	0.54	0.63	0.45	0.46	0.40	0.45	0.43	0.46														
83	0.53	0.58	0.58	0.41	0.56	0.49	0.59	0.53	0.48	0.52	0.57	E	0.57												
84	0.45	0.43	0.52	0.51	0.40	0.42	0.41	0.49	0.46	E	0.39	0.42													
85	0.42	0.30	0.45	E	0.47	0.34	0.46	0.48	0.39	0.46	0.40	0.44													
86	0.52	L	0.46	0.57	0.45	0.62	0.47	0.48	0.58																
87	0.50	0.55	0.57	0.45	0.60	0.54	0.54	E	0.39	0.50	0.39	0.47													
88	0.49	0.42	E	0.60	0.51	0.47	E	0.43	0.45	0.50	0.55														
89	0.38	0.36	0.48	0.31	0.40	0.41	0.37	0.35	0.41	0.37	0.41	0.37	0.34	0.36	0.38										
90	0.55	0.54	0.56	0.55	0.62	E	0.48	0.64	0.55	0.52	0.50														
91	0.38	0.32	0.37	0.38	0.36	0.41	0.46	0.34	0.43	0.36	0.39	0.37	0.39												
92	0.50	0.47	0.52	0.47	0.47	0.44	0.50	0.55	0.56	0.46	0.51	0.54													
93x	NP																								
94	0.49	0.47	0.50	0.45	0.43	0.49	0.46	0.57	0.54	E	0.46														
95	0.45	0.35	0.36	0.53	0.50	0.52	0.51	0.46	0.45	0.39	0.43	0.49	E												
96	0.46	0.43	0.39	0.49	0.48	0.46	0.52	0.46	0.46	0.39	0.45	0.49													
97	0.43	0.35	0.47	0.49	E	0.46	0.37	0.39	0.40	0.43	0.44	E	0.44	0.43	0.46										
98	0.52	0.39	1.07	0.42	0.36	0.45	0.41																		
99	0.42	0.41	0.36	0.44	0.38	0.42	0.47	0.42	0.44	0.41	0.51	0.39													
100	0.46	0.44	0.47	0.47	0.49	0.45	0.49	0.45	0.46	0.46	0.46	0.46													
MEAN	0.47																								
S.D.	0.05																								
N	24																								

E-EARLY RESORPTION L-LATE RESORPTION D-DEAD FETUS /-DENOTES POSITION OF CERVIX

NP=NOT PREGNANT x=EXCLUDED FROM MEAN

10-APR-14

05R018

TABLE : IIB-

009

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL BODY WEIGHTS -- GRAMS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	MEAN	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
1	3.4	3.1	3.3	3.7	3.4	3.7	3.5	3.1	3.5	3.6	3.0	E	3.7							
2	3.3	3.3	3.2	3.2	3.5	3.0	3.2	3.2	3.8	3.6	3.3	E	E	3.2						
3	3.6	3.2	3.4	4.0	3.8	4.0	3.5	3.8	3.5	3.4	3.7	3.5								
4	3.8	3.7	3.7	3.8	3.8	3.8	3.5	4.2	3.7	4.1	3.7	3.8	3.8							
5	3.5	3.1	3.4	3.3	2.8	3.8	3.5	3.6	3.6	3.6	3.7	3.6	3.8							
6	3.3	2.8	3.3	3.5	3.2	3.3	3.5	3.3	3.6	3.3	3.4	3.4	E	3.5	3.1					
7	3.9	3.2	4.1	4.3	4.0	4.0	E	4.2	4.1	4.0	3.5									
8	3.5	3.4	3.8	3.3	3.7	3.7	3.5	3.2	3.3	3.4	3.6									
9	3.6	3.6	3.3	4.3	3.7	3.8	4.0	3.3	3.5	3.1	3.6	3.5	4.0							
10	3.7	3.8	3.2	3.9	3.7	3.3	4.2	3.9	3.7	3.9	3.8	3.5								
11	3.8	3.8	4.0	3.9	3.7	3.9	3.5	4.1	3.7	3.6	3.4									
12	3.5	3.5	3.5	3.5	3.1	3.5	3.5	3.5	3.7	3.6	3.4									
13	3.8	3.8	4.2	3.6	3.6	3.7	3.9	3.7	3.6	3.8	3.5									
14	3.7	3.7	3.4	3.6	3.8	3.7	3.7	4.0	3.6	3.8	3.5									
15	3.6	3.4	3.5	3.6	3.6	3.2	3.7	4.1	3.6	3.6	E	3.7								
16	3.2	3.0	3.1	3.0	3.5	3.4	3.3	3.3	3.4	2.5	3.4	3.2								
17	3.2	3.5	3.2	3.4	3.2	3.1	3.5	3.4	3.5	3.2	3.4	3.3	2.1							
18	3.5	3.3	3.6	3.8	3.4	3.5	3.5	3.5	3.6	3.3	3.5	3.2	3.3							
19	3.4	3.4	3.4	3.5	3.5	3.6	3.7	3.4	3.4	3.3	3.4									
20	3.7	3.4	3.5	3.3	3.5	3.8	3.8	4.1	3.6	3.7	3.7	3.8								
21	4.0	4.0	E	4.1	4.0	3.7	4.0	4.2	4.0	E	4.0									
22	3.6	3.7	E	3.3	3.7	3.6	3.5	3.6	3.6	3.7	3.7	3.8								
23	3.6	3.6	3.6	3.3	3.7	E	3.7	4.0	3.3	3.5	4.0	3.6								
24	3.2	3.3	2.5	3.4	3.1	3.1	3.5	3.3	3.4	3.3	E	3.4	3.2	3.0						
25	3.4	3.2	3.4	3.6	3.5	3.3	3.5	3.4	3.3	3.4	3.5	3.8	3.3							

MEAN 3.6
S.D. 0.2
N 25

E-EARLY RESORPTION L-LATE RESORPTION D-DEAD FETUS /-DENOTES POSITION OF CERVIX

10-APR-14

05R018

TABLE : IIB-

010

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL BODY WEIGHTS -- GRAMS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	MEAN	FETUS#	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
26	3.8	3.8	4.2/	4.1	3.8	4.1	3.7	2.8	3.7	3.9											
27	3.6	3.3	3.6/	3.5	3.6/	4.1	3.7	3.5	3.5	4.1	3.4	3.9	3.3								
28	3.3	3.2	3.1	3.5	3.3	3.5	3.3	3.1	3.4/	3.4	2.8	E	3.5								
29	3.5	3.6	3.4	3.5	3.4/	3.8	3.6	3.4	3.6/	3.7	3.5	3.3	3.4								
30	3.7	3.8	4.1	4.2	4.0	E	3.8	3.3	3.5/	3.7	3.5	3.3	3.4								
31	3.6	3.8	3.5	4.2	3.7	3.7/	3.8	2.7	3.7	3.7											
32	3.5	3.4	3.4	3.9	E	3.6	E/	3.7	3.4	3.6	3.0										
33	3.6	3.1	4.1	3.7	3.7	3.3/	3.5	3.8	3.5	3.4	3.5	3.6									
34	4.0	3.8	3.9	E	4.2	4.1/	4.1	4.0	4.0	3.8											
35	3.6	3.6	3.6	3.9	3.7	3.3	3.8	3.4	3.6	3.3/	3.8	3.7	3.9								
36	3.8	4.0	3.7	3.7	3.6	3.8	4.0	3.8	3.8/	3.8	3.5	3.6	E								
37	3.5	3.4	3.7	3.7	3.3	E	3.8/	3.7	3.5	3.1	3.9	3.5	3.8								
38	3.7	3.6	3.8	3.7	3.9	3.7/	3.7	3.7	3.7	3.8	3.9	3.3									
39	3.5	3.2	3.4	3.7	3.4	3.4	3.8	3.5	3.5	3.7	E	3.3									
40	3.5	3.6	3.3	3.2	3.1	3.8/	3.6	3.4	3.8	3.8	3.3	3.6									
41	3.6	3.6	3.5	E	3.6	3.3	3.6	3.6/	3.5	3.4	3.6	3.9	4.0								
42	3.6	3.6	4.0	3.5/	3.5	3.4	3.9	3.4	3.6	3.4											
43	3.7	3.8	3.7	3.4	3.7	3.8/	3.8	3.5	3.6	3.8	3.8	3.7	3.5								
44	3.6	3.7	3.7	3.2	3.6	4.0	3.5	3.5/	3.8	3.8	3.5	3.5	3.4								
45	3.3	3.2	3.4	3.6	3.3	3.0	3.4	3.5/	3.3	3.0	3.4	3.2	3.4								
46	3.5	3.7	3.5	3.5	3.7	2.9	3.7	3.4/	3.8	3.4	3.3	3.6	3.3								
47	3.6	3.6	3.3	3.3	3.3	3.8	3.6	3.9/	3.9	3.7	3.9	3.5	3.6								
48	3.5	3.6	3.6	3.2	3.6	3.5	4.2	3.3	3.1	E/	3.8	3.3	3.3								
49	3.5	3.0	3.4	3.5	3.3	3.5	3.6	4.1	3.7/	3.7	3.4	3.8	3.4								
50	3.5	3.1	3.8	3.6	3.5/	4.0	3.2	3.7	3.6	3.2											

MEAN
S.D.
N

3.6
0.1
25

E-EARLY RESORPTION L-LATE RESORPTION D-DEAD FETUS /-DENOTES POSITION OF CERVIX

10-APR-14

05R018

TABLE : IIB-

011

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS

ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL BODY WEIGHTS -- GRAMS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	MEAN	FETUS#	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
51	3.6	3.5	3.7	3.5	2.7	3.5	3.7	3.5	3.8	3.8	3.8	3.4	3.9	3.9							
52	3.4	3.5	3.3	3.4	3.3	3.4	3.5	3.5	3.5	3.5	3.4	3.5	3.1								
53	3.4	3.6	3.4	3.6	3.3	3.5	3.3	2.9	E	3.5	3.5	3.5	3.6								
54	3.7	3.3	3.7	3.5	4.0	3.7	3.8	3.5	3.9	3.4	4.0	4.0	E								
55	3.5	3.4	3.5	3.8	3.6	E	3.6	3.4	3.4	3.5	3.6	3.6									
56	3.5	3.3	E	3.5	3.6	3.3	3.7	3.4	3.2	3.3	3.8	3.7	3.5	3.7							
57	3.5	3.6	3.6	3.5	3.5	3.5	3.8	3.4	3.5	3.4	3.3	3.3	3.7	3.5							
58	3.4	3.3	3.4	3.8	3.3	3.3	3.1	3.3	3.1	3.4	3.4	3.4	3.5	3.4	3.2						
59	3.3	3.2	3.3	E	3.3	E	3.1	3.3	3.9	4.0	3.9	4.0	3.7	3.5	3.3						
60	3.8	3.9	3.9	3.7	3.9	3.9	3.8	3.9	4.0												
61	3.3	3.9	3.5	1.8	3.5	3.6	3.6	3.3	3.6	3.3	3.2	3.5	3.2								
62	3.4	3.4	3.3	3.8	3.5	3.3	3.3	3.3	3.3	3.3	3.3	3.5	3.2	E							
63	3.3	2.5	3.2	3.5	3.3	3.4	3.7	3.1	3.3	3.3	3.3	3.3	3.3								
64	3.3	3.3	3.2	3.3	3.5	3.3	3.4	3.4	3.2	3.4	3.4	3.3	3.3								
65	3.5	3.5	3.2	3.8	3.6	3.5	3.5	3.4	3.6	3.7	3.4	3.7	3.4								
66	3.4	3.2	3.2	3.6	3.5	3.2	3.7	3.0	3.3	3.5	3.6	3.6	3.4	3.6	3.4						
67	3.6	3.6	3.6	3.1	3.5	3.6	3.9	3.7	3.7	3.0	3.7	3.7	3.7	3.2							
68	3.1	3.1	3.2	3.0	2.9	3.1	3.3	3.4	3.0	3.0	3.0	3.3	2.6	3.0							
69	3.7	3.4	3.4	3.9	3.8	3.9	3.8	3.7	3.5	3.8	3.3	3.3	3.7	3.0	3.3						
70	3.4	3.4	3.5	3.4	E	3.4	3.2	3.6	1	3.8	3.7	3.7	3.2	3.0	3.3						
71	3.3	E	3.3	3.5	3.5	3.5	3.5	3.5	3.6	3.2	2.9	3.3	3.4	3.2	3.3						
72	2.9	3.0	2.8	2.7	2.9	3.3	2.7	1.9	3.0	3.2	3.2	3.3	3.4	3.2	3.3						
73	3.5	3.3	3.4	3.7	3.0	3.7	3.9	3.8	3.7	3.8	3.7	3.3	3.7	3.5							
74	3.7	3.8	4.0	3.9	3.5	3.8	3.5	4.0	3.6	3.6	3.6	3.3	3.7	3.5							
75	3.8	3.8	3.7	3.3	3.6	4.0	3.6	3.7	4.2	3.9	3.9	3.9									

MEAN
S.D.
N

E-EARLY RESORPTION L-LATE RESORPTION D-DEAD FETUS /-DENOTES POSITION OF CERVIX

10-APR-14		TABLE : IIB-																	012		
05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS																			
		ORAL ADMINISTRATION (GAVAGE)																			
		INDIVIDUAL FETAL BODY WEIGHTS -- GRAMS																			
TEST GROUP 3 (300 MG/KG BW/D)																					
FEMALE#	MEAN	FETUS#	1	2	3	4	5	6	7	8	9	10	11	12	13	14	15	16	17	18	19
76	3.8	3.7	3.6	4.1	3.7	4.0	3.6	3.9	4.1	3.9	3.6	3.5									
77	3.3	3.0	3.2	3.2	3.4	3.3	3.3	3.7	3.1	3.4	3.3	3.2	3.4								
78	3.2	3.4	3.2	3.0	2.9	3.3	3.6	3.5	3.3	3.1	E										
79	3.6	3.5	3.0	3.9	3.7	3.6	3.5	3.5	3.5	4.0	3.7	E									
80	3.4	3.1	3.8	3.9	3.5	3.4	3.3	3.3	3.5	3.0	3.5	3.6	3.3								
81	3.7	3.3	3.6	E	3.8	E	4.3	3.9	3.6	3.8	4.0	3.3									
82	3.5	3.6	3.6	3.6	3.3	3.5	3.5	3.3	3.6	3.3	3.4	3.4	E	3.3							
83	3.1	3.0	2.9	2.5	3.1	3.3	3.3	3.3	3.4	3.2	3.4	2.9									
84	3.3	2.3	3.7	3.6	3.1	3.4	3.2	3.5	3.4	E	3.4	2.9	3.5								
85	3.6	3.0	3.9	E	3.6	3.4	4.0	3.8	3.5	3.9	3.7	3.5									
86	3.7	L	3.5	3.7	3.7	3.7	3.8	3.9	3.6												
87	3.4	3.3	3.7	3.0	3.8	3.5	3.4	E	3.3	3.2	3.5	3.1									
88	3.3	3.0	E	3.3	3.3	3.2	E	3.3	3.6	3.5	3.2										
89	2.9	2.8	3.3	2.8	2.7	3.0	3.1	2.7	3.1	2.9	3.0	2.6	3.0	2.7	2.8						
90	3.6	3.6	3.4	3.5	3.5	E	3.4	3.9	3.7	3.7	3.6										
91	3.5	3.4	3.3	3.3	3.6	3.5	3.5	3.4	3.8	3.7	3.8	3.8	3.0								
92	3.6	3.5	3.6	3.7	3.4	3.4	3.6	3.8	3.3	3.7	3.7	3.8									
93x	NP																				
94	3.4	3.6	3.4	3.0	3.5	3.5	3.4	3.3	3.0	E	3.4										
95	3.3	3.1	3.2	3.5	3.2	3.0	3.7	3.5	3.6	3.7	3.3	3.1	E								
96	3.4	3.3	2.9	3.5	3.5	3.4	3.4	3.4	3.1	3.5	3.5	3.4									
97	3.4	3.5	3.6	3.8	E	3.6	3.4	3.2	3.5	3.8	3.3	E	3.5	3.0	3.1						
98	3.6	3.5	4.0	3.7	3.3	3.6	3.4														
99	3.5	3.0	3.4	3.3	3.3	3.7	3.5	3.5	3.5	3.9	3.5	3.4									
100	3.3	3.3	3.1	3.4	3.4	3.2	3.5	3.2	3.3	3.0	3.3										
MEAN	3.4																				
S.D.	0.2																				
N	24																				

E-EARLY RESORPTION L-LATE RESORPTION D-DEAD FETUS /-DENOTES POSITION OF CERVIX

NP=NOT PREGNANT x=EXCLUDED FROM MEAN

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS					TABLE : IIB-	013
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 0 (0 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
1	1 F	NO OBSERVED GROSS FINDINGS	3.1			
		V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY THORACIC VERTEBRA- 14th				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
	2 M	NO OBSERVED GROSS FINDINGS	3.3			
		NO OBSERVED VISCERAL FINDINGS				
	3 M	NO OBSERVED GROSS FINDINGS	3.7			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY THORACIC VERTEBRA- 14th				
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	4 M	NO OBSERVED GROSS FINDINGS	3.4			
		NO OBSERVED VISCERAL FINDINGS				
	5 F	NO OBSERVED GROSS FINDINGS	3.7			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present				
		RUDIMENTARY				
		V MISHPAPEN SACRAL VERTEBRA				
		1ST SACRAL ARCH - RIGHT SIDE, CARTILAGE PRESENT				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	6 M	NO OBSERVED GROSS FINDINGS	3.5			
		NO OBSERVED VISCERAL FINDINGS				
	7 F	NO OBSERVED GROSS FINDINGS	3.1			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY THORACIC VERTEBRA- 14th				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 014

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
1	(CONTINUED)		
	8 F	V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.5
	9 M	V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	3.6
	10 F	V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.0
	12 M	V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY	3.7
	1 F	V MISHPAPEN SACRAL VERTEBRA 1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.3
	2 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.2
	3 F	NO OBSERVED GROSS FINDINGS V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.2
	4 F	V INCOMPLETE OSSIFICATION OF BASISPHENOID NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.5

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

015

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
2	(CONTINUED)		
	5 M	NO OBSERVED GROSS FINDINGS	3.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	
		V INCOMPLETE OSSIFICATION OF SACRAL ARCH; BILATERAL; Cartilage present- SEVERAL 3RD, 4TH	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage WITH BILATERAL HOLES	
	6 F	NO OBSERVED GROSS FINDINGS	3.2
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	3.2
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
	8 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	3.8
		NO OBSERVED GROSS FINDINGS	
	9 M	NO OBSERVED VISCERAL FINDINGS	3.6
		NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	
	10 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	13 F	NO OBSERVED GROSS FINDINGS	3.2
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V MISSHAPEN SACRAL VERTEBRA	
		1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT	
3	1 F	NO OBSERVED GROSS FINDINGS	3.2
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	2 M	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	3 M	NO OBSERVED GROSS FINDINGS	4.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

016

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
3	(CONTINUED)		
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	4 M	NO OBSERVED GROSS FINDINGS	3.8
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	4.0
		NO OBSERVED VISCERAL FINDINGS	
	6 F	V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	3.5
		NO OBSERVED GROSS FINDINGS	
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	3.8
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	8 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	
	9 F	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	3.4
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	10 M	NO OBSERVED GROSS FINDINGS	3.7
		NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	
	11 F	V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V WAVY RIB; BILATERAL- SEVERAL	
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
		M SHORTENED HUMERUS; BILATERAL	
4	1 M	NO OBSERVED GROSS FINDINGS	3.7

OBSERVATION CODES: M=Malformation V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB- 017

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
4	(CONTINUED)		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V WAVY RIB; RIGHT- SEVERAL	
		8TH, 9TH, 10TH	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
2 M		NO OBSERVED GROSS FINDINGS	3.7
		NO OBSERVED VISCERAL FINDINGS	
3 F		NO OBSERVED GROSS FINDINGS	3.8
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
4 F		NO OBSERVED GROSS FINDINGS	3.8
		NO OBSERVED VISCERAL FINDINGS	
5 F		NO OBSERVED GROSS FINDINGS	3.8
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V MISHAPEN SACRAL VERTEBRA	
		1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
7 M		NO OBSERVED GROSS FINDINGS	4.2
		NO OBSERVED VISCERAL FINDINGS	
8 M		NO OBSERVED GROSS FINDINGS	3.7
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL	
		2ND, 6TH	
		V MISHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		3RD, 4TH, 5TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
9 M		NO OBSERVED GROSS FINDINGS	4.1

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		018
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 0 (0 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
4 (CONTINUED)						
	10 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS C BRANCHED RIB CARTILAGE; RIGHT- 8th DISTAL V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 12th V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V MISSHAPEN SACRAL VERTEBRA 1ST SACRAL ARCH - RIGHT SIDE, CARTILAGE PRESENT V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.7			
	11 F		3.8			
	12 M		3.8			
5						
	1 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY THORACIC VERTEBRA- 14th V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 11th V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present SMALL V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY	3.1			
	2 M		3.4			
	3 F		3.3			
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 019

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
5	(CONTINUED)		
	4 M	V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	2.8
	5 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	3.8
	6 F	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS	3.5
	7 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.6
	8 F	V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS	3.6
	9 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY	3.6
	10 M	NO OBSERVED GROSS FINDINGS	3.7
	11 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.6
	12 M	NO OBSERVED GROSS FINDINGS	3.8
	13 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.7
	14 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.1

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

05R018

TABLE : IIB-

020

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAUVE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

[illegible]

OBSERVATION CODES: V=Variation C=Cartilage

SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-021

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
6 (CONTINUED)			
	8 M	NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
	9 F	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	10 M	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	11 M	NO OBSERVED GROSS FINDINGS	3.4
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF TUBEROSITAS DELTOIDEA; LEFT; Cartilage present	
		V DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum- 13th	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		NASAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
	13 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	7		3.2
	1 M	NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V UNOSIFIED STERNEBRA; Unchanged cartilage- 5th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
		WITH BILATERAL HOLES	
		V INCOMPLETE OSSIFICATION OF NASAL; BILATERAL; Unchanged cartilage	
	2 F	NO OBSERVED GROSS FINDINGS	4.1
		NO OBSERVED VISCERAL FINDINGS	
	3 M	NO OBSERVED GROSS FINDINGS	4.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL	
		5TH, 6TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
	4 F	NO OBSERVED GROSS FINDINGS	4.0
		NO OBSERVED VISCERAL FINDINGS	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-022

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
7	(CONTINUED)		
	5 M	NO OBSERVED GROSS FINDINGS	4.0
		V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BRANCHED RIB CARTILAGE; RIGHT- 8th DISTAL	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present	
		V INCOMPLETE OSSIFICATION OF BASISPHEOID	
	7 M	NO OBSERVED GROSS FINDINGS	4.2
		NO OBSERVED VISCERAL FINDINGS	
	8 M	NO OBSERVED GROSS FINDINGS	4.1
		V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V INCOMPLETE OSSIFICATION OF BASISPHEOID	
	9 F	NO OBSERVED GROSS FINDINGS	4.0
		NO OBSERVED VISCERAL FINDINGS	
	10 F	NO OBSERVED GROSS FINDINGS	3.5
		V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		NO OBSERVED GROSS FINDINGS	
	1 M	NO OBSERVED GROSS FINDINGS	3.4
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHEOID	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	2 M	NO OBSERVED GROSS FINDINGS	3.8
		NO OBSERVED VISCERAL FINDINGS	
	3 F	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	4 M	NO OBSERVED GROSS FINDINGS	3.7
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	3.7

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		023
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 0 (0 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
8 (CONTINUED)						
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	6 M	NO OBSERVED GROSS FINDINGS	3.5			
		NO OBSERVED VISCERAL FINDINGS				
	7 F	NO OBSERVED GROSS FINDINGS	3.2			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		4TH, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	8 F	NO OBSERVED GROSS FINDINGS	3.3			
		NO OBSERVED VISCERAL FINDINGS				
	9 F	NO OBSERVED GROSS FINDINGS	3.4			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 4th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	10 M	NO OBSERVED GROSS FINDINGS	3.6			
		V DILATED RENAL PELVIS; left				
		V DILATED URETER; left				
		NO OBSERVED GROSS FINDINGS				
	1 M	NO OBSERVED GROSS FINDINGS	3.6			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL				
		2ND, 6TH				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	2 F	NO OBSERVED GROSS FINDINGS	3.3			
		NO OBSERVED VISCERAL FINDINGS				
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 024

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
9	(CONTINUED)		
	3 F	NO OBSERVED GROSS FINDINGS	
		V MISHPAPEN STERNEBRA; Unchanged cartilage- 6th	4.3
		ADDITIONALLY INCOMPLETELY OSSIFIED	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); LEFT: Cartilage not present	
		RUDIMENTARY	
	4 F	NO OBSERVED GROSS FINDINGS	3.7
		V DILATED RENAL PELVIS; right	
	5 M	NO OBSERVED GROSS FINDINGS	3.8
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF BASISPHEOID	
	6 M	NO OBSERVED GROSS FINDINGS	4.0
		NO OBSERVED VISCERAL FINDINGS	
	7 F	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	8 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	9 F	NO OBSERVED GROSS FINDINGS	3.1
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL	
		5TH, 6TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); RIGHT: Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF BASISPHEOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
		WITH BILATERAL HOLES	
	10 F	NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
	11 F	NO OBSERVED GROSS FINDINGS	3.5
		V UNILATERAL OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
	12 F	V INCOMPLETE OSSIFICATION OF BASISPHEOID	4.0
		NO OBSERVED GROSS FINDINGS	
		NO OBSERVED VISCERAL FINDINGS	
10	1 M	NO OBSERVED GROSS FINDINGS	3.8

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS		TABLE : IIB-		025	
				ORAL ADMINISTRATION (GAVAGE)					
				INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS					
TEST GROUP 0 (0 MG/KG BW/D)									
FEMALE#		FETUS# (SEX)		OBSERVATION				WEIGHT (G)	
10		(CONTINUED)							
				V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th					
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
				C BIPARTITE PROCESSUS XIPHOIDEUS					
				V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present					
				RUDIMENTARY					
				V WAVY RIB; RIGHT- SEVERAL					
				10TH, 11TH					
				V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage					
				V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage					
		2 M		NO OBSERVED GROSS FINDINGS				3.2	
				NO OBSERVED VISCERAL FINDINGS					
		3 M		NO OBSERVED GROSS FINDINGS				3.9	
				V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th					
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
				C BIPARTITE PROCESSUS XIPHOIDEUS					
				V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present					
				RUDIMENTARY					
				V WAVY RIB- SEVERAL					
				4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH, 13TH					
				V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage					
				FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES					
		4 M		NO OBSERVED GROSS FINDINGS				3.7	
				V DILATED RENAL PELVIS; left					
				NO OBSERVED GROSS FINDINGS					
		5 F		NO OBSERVED GROSS FINDINGS				3.3	
				V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th					
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
				V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present					
				RUDIMENTARY					
		6 M		NO OBSERVED GROSS FINDINGS				4.2	
				NO OBSERVED VISCERAL FINDINGS					
		7 M		NO OBSERVED GROSS FINDINGS				3.9	
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
				C BIPARTITE PROCESSUS XIPHOIDEUS					
				V SUPERNUMERARY THORACIC VERTEBRA- 14th					
				V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present					
				SMALL					
				V WAVY RIB; RIGHT- SEVERAL					
				5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH					
				V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage					
				V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage					
				OBSERVATION CODES: V=Variation C=Cartilage					
				SEX CODE: M=MALE, F=FEMALE					

10-APR-14

05R018

TABLE : IIB-

026

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
10	(CONTINUED)		
11	1 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.8
		NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present	
		SMALL	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		NO OBSERVED GROSS FINDINGS	
	2 F	NO OBSERVED VISCERAL FINDINGS	4.0
	3 F	NO OBSERVED GROSS FINDINGS	3.9
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V WAVY RIB; RIGHT- SEVERAL	
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	4 M	NO OBSERVED GROSS FINDINGS	3.7
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	3.9
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	6 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	4.1
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V WAVY RIB; BILATERAL- SEVERAL	
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	8 F	NO OBSERVED GROSS FINDINGS	3.7
		NO OBSERVED VISCERAL FINDINGS	
	9 M	NO OBSERVED GROSS FINDINGS	3.9
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

027

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
11	(CONTINUED)		
		V WAVY RIB; BILATERAL- SEVERAL	
		5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	10 M	NO OBSERVED GROSS FINDINGS	3.8
	11 M	NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V WAVY RIB; BILATERAL- SEVERAL	
		5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH	
		V DUMBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 12th	
		C NOTCHED CARTILAGE BETWEEN BASISPHEOID AND BASIOCCIPITAL	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
12	1 F	NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present	
	2 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.5
		NO OBSERVED GROSS FINDINGS	
	3 F	V SHORT INNOMINATE	3.5
		NO OBSERVED GROSS FINDINGS	
		V MISHPAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		3RD, 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	4 F	NO OBSERVED GROSS FINDINGS	3.1
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	3.5
		V MISHPAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		3RD, 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V WAVY RIB; RIGHT- SEVERAL	
		5TH, 6TH, 7TH, 8TH	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14	05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)	TABLE : IIB-	028
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				
TEST GROUP 0 (0 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION		WEIGHT (G)
12	(CONTINUED)			
		V MISSHAPEN SACRAL VERTEBRA		
		1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
	6 F	NO OBSERVED GROSS FINDINGS		3.5
	7 F	NO OBSERVED VISCERAL FINDINGS		
		NO OBSERVED GROSS FINDINGS		3.5
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		
	8 M	NO OBSERVED GROSS FINDINGS		3.7
		NO OBSERVED VISCERAL FINDINGS		
	9 F	NO OBSERVED GROSS FINDINGS		3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		
	10 F	NO OBSERVED GROSS FINDINGS		3.4
		NO OBSERVED VISCERAL FINDINGS		
13	1 F	NO OBSERVED GROSS FINDINGS		3.8
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL		
		4TH, 5TH		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage		
		BASISPHEOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES		
	2 M	NO OBSERVED GROSS FINDINGS		4.2
		NO OBSERVED VISCERAL FINDINGS		
	3 F	NO OBSERVED GROSS FINDINGS		3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY THORACIC VERTEBRA- 14th		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
	4 M	NO OBSERVED GROSS FINDINGS		3.6
		NO OBSERVED VISCERAL FINDINGS		
	5 M	NO OBSERVED GROSS FINDINGS		3.7
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
OBSERVATION CODES: V=Variation C=Cartilage				
SEX CODE: M=MALE, F=FEMALE				

10-APR-14

05R018

TABLE : IIB-

029

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
13	(CONTINUED)		
	6 M	C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage present V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present SMALL	3.9
	7 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present V DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum- 13th V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	3.7
14	1 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.7
	2 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.4
	3 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.6
	4 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.8
	5 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH	3.7
		C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	
	6 M	V SUPRAOCCIPITAL HOLE(S); RIGHT NO OBSERVED GROSS FINDINGS	3.7
	7 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS C BIPARTITE PROCESSUS XIPHOIDEUS	4.0

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14	05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)	TABLE : IIB-	030
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				
TEST GROUP 0 (0 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION		WEIGHT (G)
14	(CONTINUED)			
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present		
		SMALL		
	8 F	V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present		3.6
		NO OBSERVED GROSS FINDINGS		
	9 M	NO OBSERVED VISCERAL FINDINGS		3.8
		NO OBSERVED GROSS FINDINGS		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
	10 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		3.5
		NO OBSERVED GROSS FINDINGS		
		NO OBSERVED VISCERAL FINDINGS		
	15	1 F		3.4
		NO OBSERVED GROSS FINDINGS		
		V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		3.5
	2 F	NO OBSERVED GROSS FINDINGS		
		NO OBSERVED VISCERAL FINDINGS		
	3 M	NO OBSERVED GROSS FINDINGS		3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL		
		5TH, 6TH		
		V INCOMPLETE OSSIFICATION OF BASISPHENOID		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
	4 F	NO OBSERVED GROSS FINDINGS		3.6
		NO OBSERVED VISCERAL FINDINGS		
	5 F	NO OBSERVED GROSS FINDINGS		3.2
		V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF BASISPHENOID		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
	6 M	NO OBSERVED GROSS FINDINGS		3.7
		NO OBSERVED VISCERAL FINDINGS		
	7 M	NO OBSERVED GROSS FINDINGS		4.1
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
OBSERVATION CODES: V=Variation C=Cartilage				
SEX CODE: M=MALE, F=FEMALE				

10-APR-14

05R018

TABLE : IIB- 031

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
15	(CONTINUED)		
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	8 F	NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
	9 F	NO OBSERVED GROSS FINDINGS	3.6
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	11 F	NO OBSERVED GROSS FINDINGS	3.7
		NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	
16	1 F	NO OBSERVED GROSS FINDINGS	3.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF NASAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	2 M	NO OBSERVED GROSS FINDINGS	3.1
		NO OBSERVED VISCERAL FINDINGS	
	3 F	NO OBSERVED GROSS FINDINGS	3.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V WAVY RIB; RIGHT- 11th	
		V INCOMPLETE OSSIFICATION OF PARIETAL; LEFT; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	4 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	3.4
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	6 M	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		032
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 0 (0 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			

16	(CONTINUED)					
	7 M	NO OBSERVED GROSS FINDINGS	3.3			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 6TH				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V WAVY RIB; RIGHT- SEVERAL 6TH, 7TH, 8TH, 9TH, 10TH, 11TH				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		NEARLY ALL SKULL BONES				
	8 M	NO OBSERVED GROSS FINDINGS	3.4			
		NO OBSERVED VISCERAL FINDINGS				
	9 M	NO OBSERVED GROSS FINDINGS	2.5			
		V UNOBSIFIED STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 5TH, 6TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 4th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V WAVY RIB; BILATERAL- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH RIGHT SIDE 10TH, 11TH, 12TH LEFT SIDE				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		NEARLY ALL SKULL BONES				
	10 F	NO OBSERVED GROSS FINDINGS	3.4			
		NO OBSERVED VISCERAL FINDINGS				
	11 M	NO OBSERVED GROSS FINDINGS	3.2			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V WAVY RIB; RIGHT- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		BASISPHENOID, TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES				
	1 M	NO OBSERVED GROSS FINDINGS	3.5			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	2 M	NO OBSERVED GROSS FINDINGS	3.2			
		NO OBSERVED VISCERAL FINDINGS				

OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 033

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
17	(CONTINUED)		
	3 F	NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	3.4
		C BIPARTITE PROCESSUS XIPHOIDEUS	
	4 M	V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		NO OBSERVED GROSS FINDINGS	3.2
	5 F	NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	3.1
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	6 M	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.5
		NO OBSERVED GROSS FINDINGS	
	7 F	NO OBSERVED VISCERAL FINDINGS	3.4
		NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	8 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	9 M	NO OBSERVED GROSS FINDINGS	3.2
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		BASISPHENOID, TEMPORAL, PARIETAL, INTERPARIETAL AND	
		SUPRAOCCIPITAL BONES	
	10 M	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	11 M	NO OBSERVED GROSS FINDINGS	3.3
		V MISAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 11th	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	12 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	2.1
		NO OBSERVED GROSS FINDINGS	
		NO OBSERVED VISCERAL FINDINGS	
18	1 F	NO OBSERVED GROSS FINDINGS	3.3

OBSERVATION CODES: V=Variation C=Cartilage

SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS		TABLE : IIB-	034
TEST GROUP 0 (0 MG/KG BW/D)							
FEMALE#	FETUS# (SEX)	OBSERVATION				WEIGHT (G)	
18	(CONTINUED)						
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH					
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
		C BIPARTITE PROCESSUS XIPHOIDEUS					
		V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present RUDIMENTARY					
	2 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				3.6	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage					
		NO OBSERVED GROSS FINDINGS					
	3 M	NO OBSERVED VISCERAL FINDINGS				3.8	
		NO OBSERVED GROSS FINDINGS					
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH					
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present RUDIMENTARY					
	4 M	V INCOMPLETE OSSIFICATION OF BASISPHENOID				3.4	
		NO OBSERVED GROSS FINDINGS					
		NO OBSERVED VISCERAL FINDINGS					
	5 M	NO OBSERVED GROSS FINDINGS				3.5	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th					
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
		C BIPARTITE PROCESSUS XIPHOIDEUS					
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage					
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage					
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage					
	6 F	NO OBSERVED GROSS FINDINGS				3.5	
		NO OBSERVED VISCERAL FINDINGS					
	7 F	NO OBSERVED GROSS FINDINGS				3.5	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
		C BIPARTITE PROCESSUS XIPHOIDEUS					
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage					
		WITH BILATERAL HOLES					
		NO OBSERVED GROSS FINDINGS					
	8 F	NO OBSERVED VISCERAL FINDINGS				3.6	
		NO OBSERVED GROSS FINDINGS					
	9 F	NO OBSERVED GROSS FINDINGS				3.3	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH					
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present RUDIMENTARY					
OBSERVATION CODES: V=Variation C=Cartilage							
SEX CODE: M=MALE, F=FFEMALE							

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		035	
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS							
ORAL ADMINISTRATION (GAVAGE)							
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS							
TEST GROUP 0 (0 MG/KG BW/D)							
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)				
18 (CONTINUED)							
		V INCOMPLETE OSSIFICATION OF BASISPHENOID					
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage					
	10 M	NO OBSERVED GROSS FINDINGS				3.5	
		NO OBSERVED VISCERAL FINDINGS					
	11 F	NO OBSERVED GROSS FINDINGS				3.2	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
		C BIPARTITE PROCESSUS XIPHOIDEUS					
	12 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				3.3	
		NO OBSERVED GROSS FINDINGS					
		NO OBSERVED VISCERAL FINDINGS					
19							
	1 M	NO OBSERVED GROSS FINDINGS				3.4	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
		C BIPARTITE PROCESSUS XIPHOIDEUS					
	2 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				3.4	
		NO OBSERVED GROSS FINDINGS					
	3 M	NO OBSERVED VISCERAL FINDINGS				3.5	
		NO OBSERVED GROSS FINDINGS					
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
	4 F	V INCOMPLETE OSSIFICATION OF BASISPHENOID				3.5	
		NO OBSERVED GROSS FINDINGS					
	5 F	NO OBSERVED VISCERAL FINDINGS				3.6	
		NO OBSERVED GROSS FINDINGS					
	6 M	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				3.7	
		NO OBSERVED GROSS FINDINGS					
	7 M	NO OBSERVED VISCERAL FINDINGS				3.4	
		NO OBSERVED GROSS FINDINGS					
	8 M	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				3.4	
		NO OBSERVED VISCERAL FINDINGS					
	9 F	NO OBSERVED GROSS FINDINGS				3.3	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
		C BIPARTITE PROCESSUS XIPHOIDEUS					
	10 M	V MISHPHEN STERNEBRA; Unchanged cartilage- 5th				3.4	
		NO OBSERVED GROSS FINDINGS					
		NO OBSERVED VISCERAL FINDINGS					
20							
	1 F	NO OBSERVED GROSS FINDINGS				3.4	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
		C BIPARTITE PROCESSUS XIPHOIDEUS					
OBSERVATION CODES: V=Variation C=Cartilage							
SEX CODE: M=MALE, F=FEMALE							

10-APR-14

05R018

TABLE : IIB- 036

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 0 (0 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
20	(CONTINUED)		
	2 F	V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	3.5
	3 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	3.3
	4 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.5
	5 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	3.8
	6 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.8
	7 M	NO OBSERVED GROSS FINDINGS C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	4.1
	8 F	V CERVICAL RIB; LEFT; Cartilage not present NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.6
	9 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	3.7
	10 F	V MISHAPEN SACRAL VERTEBRA 1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT NO OBSERVED GROSS FINDINGS	3.7
	11 F	V DILATED RENAL PELVIS; right NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.8
		V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		TABLE : IIB-		037
05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)		
TEST GROUP 0 (0 MG/KG BW/D)		INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS		
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)	
21	1 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V WAVY RIB; BILATERAL- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage TEMPORAL, FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- SEVERAL 10TH, 12TH V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V MISSHAPEN SACRAL VERTEBRA 1ST SACRAL ARCH - RIGHT SIDE, CARTILAGE PRESENT V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	4.0	
	3 M	NO OBSERVED GROSS FINDINGS	4.1	
	4 M	NO OBSERVED GROSS FINDINGS	4.0	
	5 F	NO OBSERVED GROSS FINDINGS	3.7	
	6 F	NO OBSERVED GROSS FINDINGS	4.0	
	7 M	NO OBSERVED GROSS FINDINGS	4.2	
	8 F	NO OBSERVED GROSS FINDINGS	4.0	
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE				

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14	05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)	TABLE : IIB-	038
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				
TEST GROUP 0 (0 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION		WEIGHT (G)
21	(CONTINUED)			
	10 F	V MISHPEN SACRAL VERTEBRA 1ST SACRAL ARCH - RIGHT SIDE; CARTILAGE PRESENT NO OBSERVED GROSS FINDINGS V DILATED RENAL PELVIS; left		4.0
22	1 F	NO OBSERVED GROSS FINDINGS V MISHPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY THORACIC VERTEBRA- 14th V BIPARTITE OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum- 12th V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present SMALL V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY THORACIC VERTEBRA- 14th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V DUMBELL OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum- 11th V SUPERNUMERARY THORACIC VERTEBRA- 14th V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage present V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present SMALL NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS		3.7
	3 F			3.3
	4 F			3.7
	5 M			3.6
	6 M			3.5
	7 F			3.6
	8 F			3.6
	9 M			3.7
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE				

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS				TABLE : IIB-	039
ORAL ADMINISTRATION (GAVAGE)					
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS					
TEST GROUP 0 (0 MG/KG BW/D)					
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)		

22	(CONTINUED)				
	10 F	NO OBSERVED GROSS FINDINGS	3.7		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present			
		RUDIMENTARY			
	11 M	NO OBSERVED GROSS FINDINGS	3.8		
		NO OBSERVED VISCERAL FINDINGS			
23	1 M	NO OBSERVED GROSS FINDINGS	3.6		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present			
		RUDIMENTARY			
		V INCOMPLETE OSSIFICATION OF PARIETAL; RIGHT; Unchanged cartilage			
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.6		
	2 M	NO OBSERVED GROSS FINDINGS			
		NO OBSERVED VISCERAL FINDINGS			
	3 F	NO OBSERVED GROSS FINDINGS	3.3		
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL			
		4TH, 5TH			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
	4 F	NO OBSERVED GROSS FINDINGS	3.7		
		NO OBSERVED VISCERAL FINDINGS			
	6 M	NO OBSERVED GROSS FINDINGS	3.7		
		NO OBSERVED SKELETAL FINDINGS			
	7 M	NO OBSERVED GROSS FINDINGS	4.0		
		NO OBSERVED VISCERAL FINDINGS			
	8 M	NO OBSERVED GROSS FINDINGS	3.3		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present			
		RUDIMENTARY			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			
		WITH BILATERAL HOLES			
	9 M	NO OBSERVED GROSS FINDINGS	3.5		
		NO OBSERVED VISCERAL FINDINGS			
	10 M	NO OBSERVED GROSS FINDINGS	4.0		
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th			

OBSERVATION CODES: V=Variation C=Cartilage					
SEX CODE: M=MALE, F=FEMALE					

TEST GROUP 0 (0 MG/KG BW/D)	INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS	ORAL ADMINISTRATION (GAVAGE)	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
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TABLE : IIB-

040

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
23	(CONTINUED)		
	11 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.6
24	1 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY THORACIC VERTEBRA- 14th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage BASISPHENOID, TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage WITH BILATERAL HOLES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage NEARLY ALL SKULL BONES	3.3 2.5 3.4
	4 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.1
	5 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.1
	6 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.5
	7 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage NEARLY ALL SKULL BONES	3.3

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS				TABLE : IIB-	041
ORAL ADMINISTRATION (GAVAGE)					
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS					
TEST GROUP 0 (0 MG/KG BW/D)					
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)		
24	(CONTINUED)				
	8 M	NO OBSERVED GROSS FINDINGS	3.4		
		NO OBSERVED VISCERAL FINDINGS			
	9 M	NO OBSERVED GROSS FINDINGS	3.3		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present			
		RUDIMENTARY			
		V INCOMPLETE OSSIFICATION OF BASISPHENOID			
	11 M	NO OBSERVED GROSS FINDINGS	3.4		
		NO OBSERVED VISCERAL FINDINGS			
	12 F	NO OBSERVED GROSS FINDINGS	3.2		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present			
		SMALL			
		V INCOMPLETE OSSIFICATION OF BASISPHENOID			
	13 F	NO OBSERVED GROSS FINDINGS	3.0		
		NO OBSERVED VISCERAL FINDINGS			
25	1 F	NO OBSERVED GROSS FINDINGS	3.2		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present			
		RUDIMENTARY			
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage			
		V INCOMPLETE OSSIFICATION OF HYOID; Cartilage present			
	2 M	NO OBSERVED GROSS FINDINGS	3.4		
		NO OBSERVED VISCERAL FINDINGS			
	3 M	NO OBSERVED GROSS FINDINGS	3.6		
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL			
		2ND, 3RD, 4TH, 5TH			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present			
		RUDIMENTARY			
	4 M	NO OBSERVED GROSS FINDINGS	3.5		
		NO OBSERVED VISCERAL FINDINGS			
	5 F	NO OBSERVED GROSS FINDINGS	3.3		
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
OBSERVATION CODES: V=Variation C=Cartilage					
SEX CODE: M=MALE, F=FEMALE					

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14	05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)	TABLE : IIB-	042
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				
TEST GROUP 0 (0 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION		WEIGHT (g)
25	(CONTINUED)			
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF BASISPHENOID		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		3.5
	6 F	NO OBSERVED GROSS FINDINGS		
		NO OBSERVED VISCERAL FINDINGS		
	7 M	NO OBSERVED GROSS FINDINGS		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		3.4
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		SMALL		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
	8 F	NO OBSERVED GROSS FINDINGS		3.3
		V DILATED RENAL PELVIS; right		
	9 F	NO OBSERVED GROSS FINDINGS		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		3.4
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
	10 F	NO OBSERVED GROSS FINDINGS		3.5
		NO OBSERVED VISCERAL FINDINGS		
	11 M	NO OBSERVED GROSS FINDINGS		
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL		3.8
		2ND,3RD,4TH,5TH		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
	12 F	NO OBSERVED GROSS FINDINGS		3.3
		NO OBSERVED VISCERAL FINDINGS		
OBSERVATION CODES: V=Variation C=Cartilage				
SEX CODE: M=MALE, F=FEMALE				

10-APR-14		05R018		TABLE : IIB-		043
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
26	1 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	3.8			
	2 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	4.2			
	3 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	4.1			
	4 F	V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY	3.8			
	5 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	4.1			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH				
	6 M	C BIPARTITE PROCESSUS XIPHOIDEUS NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.7			
	7 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 1ST, 2ND, 3RD, 4TH, 5TH	2.8			
		V UNOSIFIED STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY THORACIC VERTEBRA- 14th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present V INCOMPLETE OSSIFICATION OF PUBIS; BILATERAL; Cartilage present V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage WITH BILATERAL HOLES				
	8 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.7			
	9 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	3.9			
27	1 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS	3.3			
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE						

10-APR-14		05R018		TABLE : IIB-		044
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			

27	(CONTINUED)					
		V SUPERNUMERARY THORACIC VERTEBRA- 14th				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		BASISPHEOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES				
	2 F	NO OBSERVED GROSS FINDINGS	3.6			
		NO OBSERVED VISCERAL FINDINGS				
	3 F	NO OBSERVED GROSS FINDINGS	3.5			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY THORACIC VERTEBRA- 14th				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
		V INCOMPLETE OSSIFICATION OF BASISPHEOID				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	3.6			
	4 F	NO OBSERVED GROSS FINDINGS				
		NO OBSERVED VISCERAL FINDINGS				
	5 M	NO OBSERVED GROSS FINDINGS	4.1			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		4TH, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY THORACIC VERTEBRA- 14th				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
	6 F	NO OBSERVED GROSS FINDINGS	3.7			
		NO OBSERVED VISCERAL FINDINGS				
	7 F	NO OBSERVED GROSS FINDINGS	3.5			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		BASISPHEOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES				
	8 F	NO OBSERVED GROSS FINDINGS	3.5			
		NO OBSERVED VISCERAL FINDINGS				
	9 M	NO OBSERVED GROSS FINDINGS	4.1			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		4TH, 5TH				

OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14		05R018		TABLE : IIB-		045
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			

27	(CONTINUED)					
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	10 F	NO OBSERVED GROSS FINDINGS	3.4			
		NO OBSERVED VISCERAL FINDINGS				
	11 M	NO OBSERVED GROSS FINDINGS	3.9			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY THORACIC VERTEBRA- 14th				
		V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	3.3			
	12 F	NO OBSERVED GROSS FINDINGS				
		NO OBSERVED VISCERAL FINDINGS				
28		NO OBSERVED GROSS FINDINGS	3.2			
	1 F	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	2 F	NO OBSERVED GROSS FINDINGS	3.1			
		NO OBSERVED VISCERAL FINDINGS				
	3 M	NO OBSERVED GROSS FINDINGS	3.5			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
	4 M	NO OBSERVED GROSS FINDINGS	3.3			
		NO OBSERVED VISCERAL FINDINGS				
	5 M	NO OBSERVED GROSS FINDINGS	3.5			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL				
		3RD, 6TH				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V WAVY RIB; BILATERAL- SEVERAL				
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH				
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14		05R018		TABLE : IIB-		046
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT(G)			

28	(CONTINUED)	V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage NASAL, BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES				
	6 F	NO OBSERVED GROSS FINDINGS	3.3			
	7 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.1			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 2nd				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
	8 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS	3.4			
		V SHORT INNOMINATE				
	9 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH	3.4			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum 12th				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
	10 F	V INCOMPLETE OSSIFICATION OF PARIETAL; LEFT; Unchanged cartilage	2.8			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
		NO OBSERVED GROSS FINDINGS				
		NO OBSERVED VISCERAL FINDINGS				
	12 M	NO OBSERVED GROSS FINDINGS	3.5			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
29	1 M	NO OBSERVED GROSS FINDINGS	3.6			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V WAVY RIB; BILATERAL- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES				
	2 F	NO OBSERVED GROSS FINDINGS	3.4			
		NO OBSERVED VISCERAL FINDINGS				
	3 F	NO OBSERVED GROSS FINDINGS	3.5			
		NO OBSERVED SKELETAL FINDINGS				
	4 F	NO OBSERVED GROSS FINDINGS	3.4			

OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 047

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
29	(CONTINUED)		
	5 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.8
	6 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.6
	7 F	NO OBSERVED GROSS FINDINGS V CERVICAL RIB; LEFT; Cartilage not present V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.4
	8 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.6
30	1 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF BASISPHENOID NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	3.8
	2 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	4.1
	3 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	4.2
	4 M	V INCOMPLETE OSSIFICATION OF BASISPHENOID NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	4.0
	6 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th C BIPARTITE PROCESSUS XIPHOIDEUS	3.8
	7 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.3
	8 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY	3.5

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		048
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			

30	(CONTINUED)					
	9 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS	3.7			
	10 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present RUDIMENTARY	3.5			
	11 F	V MISSHAPEN SACRAL VERTEBRA 1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT V INCOMPLETE OSSIFICATION OF BASISPHENOID	3.3			
	12 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH	3.4			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH				
	13 F	C BIPARTITE PROCESSUS XIPHOIDEUS NO OBSERVED GROSS FINDINGS	3.5			
	14 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present RUDIMENTARY	3.4			
	15 M	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.7			
31	1 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present RUDIMENTARY	3.8			
	2 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.5			
	3 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 6th ADDITIONALLY INCOMPLETELY OSSIFIED	4.2			
		V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 11th V INCOMPLETE OSSIFICATION OF BASISPHENOID				
	4 F	NO OBSERVED GROSS FINDINGS	3.7			

OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 049

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
31	(CONTINUED)		
	5 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS C BIPARTITE PROCESSUS XIPHOIDEUS V DUMEBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 13th	3.7
	6 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.8
	7 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 1ST, 5TH, 6TH C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage NEARLY ALL SKULL BONES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	2.7
	8 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.7
32	1 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.4
	2 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.4
	3 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.9
	5 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.6
	7 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.7

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				TABLE : IIB-	050
TEST GROUP 1 (30 MG/KG BW/D)					
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)		
(CONTINUED)					
32		V INCOMPLETE OSSIFICATION OF BASISPHENOID			
	8 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS	3.4		
	9 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.6		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		V SUPERNUMERARY RIB (14TH) ; RIGHT; Cartilage not present SMALL			
		V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage present			
		V MISSHAPEN SACRAL VERTEBRA			
		1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT			
		V INCOMPLETE OSSIFICATION OF BASISPHENOID			
10 F		NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.0		
33		NO OBSERVED GROSS FINDINGS			
	1 F	V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	3.1		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage			
		BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES			
2 M		NO OBSERVED GROSS FINDINGS	4.1		
		V DILATED RENAL PELVIS; left			
3 M		NO OBSERVED GROSS FINDINGS	3.7		
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH, 5TH			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V INCOMPLETE OSSIFICATION OF BASISPHENOID			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			
4 M		NO OBSERVED GROSS FINDINGS	3.7		
		NO OBSERVED VISCERAL FINDINGS			
5 F		NO OBSERVED GROSS FINDINGS	3.3		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th			
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V WAVY RIB; BILATERAL- SEVERAL 4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH			
		V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present RUDIMENTARY			
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE					

10-APR-14		05R018		TABLE : IIB-		051
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			

33	(CONTINUED)					
	6 M	V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	3.5			
		NEARLY ALL SKULL BONES				
		NO OBSERVED GROSS FINDINGS				
	7 F	NO OBSERVED VISCERAL FINDINGS	3.8			
		NO OBSERVED GROSS FINDINGS				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		2ND, 3RD, 4TH, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
	8 M	NO OBSERVED GROSS FINDINGS	3.5			
		NO OBSERVED VISCERAL FINDINGS				
		NO OBSERVED GROSS FINDINGS				
	9 F	NO OBSERVED GROSS FINDINGS	3.4			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		4TH, 5TH				
		5TH ADDITIONALLY INCOMPLETELY OSSIFIED				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	10 F	NO OBSERVED GROSS FINDINGS	3.5			
		V SHORT INNOMINATE				
		NO OBSERVED GROSS FINDINGS				
	11 F	NO OBSERVED GROSS FINDINGS	3.6			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V WAVY RIB; RIGHT- SEVERAL				
		7TH, 8TH, 10TH, 11TH, 12TH				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF PARIETAL; Bilateral; Unchanged cartilage				
34	1 F	NO OBSERVED GROSS FINDINGS	3.8			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); Bilateral; Cartilage not present				
		RUDIMENTARY				
		V WAVY RIB; Bilateral- SEVERAL				
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH				
	2 F	NO OBSERVED GROSS FINDINGS	3.9			

OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

05R018

TABLE : IIB-

052

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
34	(CONTINUED)		
	4 M	NO OBSERVED VISCERAL FINDINGS	4.2
		NO OBSERVED GROSS FINDINGS	
		V MISSAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
	5 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	4.1
		NO OBSERVED GROSS FINDINGS	
	6 M	NO OBSERVED VISCERAL FINDINGS	4.1
		NO OBSERVED GROSS FINDINGS	
		V MISSAPEN STERNEBRA; Unchanged cartilage- 5th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V WAVY RIB; RIGHT- SEVERAL	
		5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH	
		V INCOMPLETE OSSIFICATION OF PARIETAL; RIGHT; Unchanged cartilage	
	7 F	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	4.0
		NO OBSERVED GROSS FINDINGS	
	8 F	NO OBSERVED VISCERAL FINDINGS	4.0
		NO OBSERVED GROSS FINDINGS	
		V MISSAPEN STERNEBRA; Unchanged cartilage- 5th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	9 M	NO OBSERVED GROSS FINDINGS	3.8
		NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	
	1 F	V MISSAPEN STERNEBRA; Unchanged cartilage- SEVERAL	3.6
		4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V MISSAPEN SACRAL VERTEBRA	
		1ST SACRAL ARCH - RIGHT SIDE, CARTILAGE PRESENT	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	2 F	NO OBSERVED GROSS FINDINGS	3.6

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		053
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			

35	(CONTINUED)					
	3 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present SMALL	3.9			
	4 F	NO OBSERVED GROSS FINDINGS V DILATED RENAL PELVIS; right	3.7			
	5 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY THORACIC VERTEBRA- 14th V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage present V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.3			
	6 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.8			
	7 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH V SUPERNUMERARY RIB (14TH) ; RIGHT; Cartilage not present SMALL V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage present V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.4			
	8 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.6			
	9 F	NO OBSERVED GROSS FINDINGS V UNOSSIFIED STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present SMALL V DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum- 12th V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.3			
	10 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.8			

OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE						

10-APR-14		05R018		TABLE : IIB-		054
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			

35	(CONTINUED)					
	11 F	NO OBSERVED GROSS FINDINGS	3.7			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V DUMBELL OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum- 12th				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		NEARLY ALL SKULL BONES				
	12 M	NO OBSERVED GROSS FINDINGS	3.9			
		NO OBSERVED VISCERAL FINDINGS				
36	1 M	NO OBSERVED GROSS FINDINGS	4.0			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES				
	2 F	NO OBSERVED GROSS FINDINGS	3.7			
		NO OBSERVED VISCERAL FINDINGS				
	3 M	NO OBSERVED GROSS FINDINGS	3.7			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
		V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 12th				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES				
	4 M	NO OBSERVED GROSS FINDINGS	3.6			
		NO OBSERVED VISCERAL FINDINGS				
	5 F	NO OBSERVED GROSS FINDINGS	3.8			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES				
	6 M	NO OBSERVED GROSS FINDINGS	4.0			
		NO OBSERVED VISCERAL FINDINGS				
	7 M	NO OBSERVED GROSS FINDINGS	3.8			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present				
		RUDIMENTARY				

OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 055

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
36	(CONTINUED)		
	8 M	V INCOMPLETE OSSIFICATION OF SACRAL ARCH; BILATERAL; Cartilage present- 4th	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
		NO OBSERVED GROSS FINDINGS	3.8
	9 M	NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	3.8
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL	
		5TH, 6TH	
	10 F	C BIPARTITE PROCESSUS XIPHOIDEUS	
		NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	11 F	NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
37	1 M	NO OBSERVED GROSS FINDINGS	3.4
		V INCOMPLETE OSSIFICATION OF METACARPAL; BILATERAL; Cartilage present- SEVERAL	
		2ND RIGHT SIDE	
		2ND, 4TH LEFT SIDE	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V WAVY RIB; BILATERAL- SEVERAL	
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH, 13TH	
		V INCOMPLETE OSSIFICATION OF THORACIC ARCH; LEFT; Cartilage present- 1st	
		12TH LEFT SIDE	
		13TH BILATERAL	
		V INCOMPLETE OSSIFICATION OF CERVICAL ARCH; LEFT; Cartilage present- 1st	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		NEARLY ALL SKULL BONES	
	2 M	NO OBSERVED GROSS FINDINGS	3.7
		V DILATED RENAL PELVIS; left	
		V DILATED URETER; left	
	3 M	NO OBSERVED GROSS FINDINGS	3.7
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V WAVY RIB; BILATERAL- SEVERAL	
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH, 13TH	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

056

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
37	(CONTINUED)		
	4 F	V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage NEARLY ALL SKULL BONES	3.3
	6 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.8
	7 M	V INCOMPLETE OSSIFICATION OF NASAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.7
	8 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY	3.5
	9 F	V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage NEARLY ALL SKULL BONES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.1
38	1 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage WITH BILATERAL HOLES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.6
	2 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.8
	3 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.7

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		057
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
38	(CONTINUED)					
	4 M	C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL	3.9			
		NO OBSERVED GROSS FINDINGS				
	5 M	NO OBSERVED VISCERAL FINDINGS	3.7			
		NO OBSERVED GROSS FINDINGS				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
	6 M	C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL	3.7			
		NO OBSERVED GROSS FINDINGS				
	7 M	NO OBSERVED VISCERAL FINDINGS	3.7			
		NO OBSERVED GROSS FINDINGS				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	8 M	NO OBSERVED GROSS FINDINGS	3.8			
		NO OBSERVED VISCERAL FINDINGS				
	9 M	NO OBSERVED GROSS FINDINGS	3.9			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V MISSHAPEN SACRAL VERTEBRA				
		1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT				
	10 M	NO OBSERVED GROSS FINDINGS	3.9			
		NO OBSERVED VISCERAL FINDINGS				
	11 F	NO OBSERVED GROSS FINDINGS	3.5			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		4TH, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	12 M	NO OBSERVED GROSS FINDINGS	3.8			
		NO OBSERVED VISCERAL FINDINGS				
39	1 F	NO OBSERVED GROSS FINDINGS	3.2			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 058

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
39	(CONTINUED)		
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	2 M	V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	3.4
		NO OBSERVED GROSS FINDINGS	
	3 F	NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	3.7
		RUDIMENTARY	
		V WAVY RIB; RIGHT- SEVERAL	
		7TH, 8TH, 9TH, 10TH, 11TH, 12TH	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		TEMPORAL, FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL	
		BONES	
	4 F	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	5 F	NO OBSERVED GROSS FINDINGS	3.4
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	6 M	NO OBSERVED GROSS FINDINGS	3.8
		NO OBSERVED VISCERAL FINDINGS	
	7 F	NO OBSERVED GROSS FINDINGS	3.5
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	8 M	NO OBSERVED GROSS FINDINGS	3.7
		NO OBSERVED VISCERAL FINDINGS	
	10 M	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
40	1 M	NO OBSERVED GROSS FINDINGS	3.6
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	2 F	NO OBSERVED GROSS FINDINGS	3.3

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		059
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
(CONTINUED)						
40	3 F	NO OBSERVED VISCERAL FINDINGS	3.2			
		NO OBSERVED GROSS FINDINGS				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 4th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present				
	4 F	RUDIMENTARY	3.1			
		NO OBSERVED GROSS FINDINGS				
		NO OBSERVED VISCERAL FINDINGS				
	5 M	NO OBSERVED GROSS FINDINGS	3.8			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.6			
	6 F	NO OBSERVED GROSS FINDINGS				
		NO OBSERVED VISCERAL FINDINGS				
	7 M	NO OBSERVED GROSS FINDINGS	3.4			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V SUPERNUMERARY RIB (14TH) ; RIGHT; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	8 M	NO OBSERVED GROSS FINDINGS	3.8			
		NO OBSERVED VISCERAL FINDINGS				
	9 M	NO OBSERVED GROSS FINDINGS	3.8			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present				
		RUDIMENTARY				
	10 F	NO OBSERVED GROSS FINDINGS	3.3			
		NO OBSERVED VISCERAL FINDINGS				
41	1 M	NO OBSERVED GROSS FINDINGS	3.6			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14	05R018	TABLE : IIB-	060
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)			
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS			
TEST GROUP 1 (30 MG/KG BW/D)			
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
41	(CONTINUED)		
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		C NOTCHED CARTILAGE BETWEEN BASISPHEOID AND BASIOCCIPITAL	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	2 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	4 F	NO OBSERVED GROSS FINDINGS	3.6
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	5 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	6 M	NO OBSERVED GROSS FINDINGS	3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	7 F	NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
	8 F	NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
	9 F	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	10 M	NO OBSERVED GROSS FINDINGS	3.6
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
OBSERVATION CODES: V=Variation C=Cartilage			
SEX CODE: M=MALE, F=FEMALE			

10-APR-14

05R018

TABLE : IIB- 061

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
41	(CONTINUED)		
	11 M	NO OBSERVED GROSS FINDINGS	3.9
		NO OBSERVED VISCERAL FINDINGS	
	12 M	NO OBSERVED GROSS FINDINGS	4.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V WAVY RIB; RIGHT- SEVERAL	
		7TH, 8TH, 9TH, 10TH, 11TH	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	13 M	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	
	1 M	NO OBSERVED GROSS FINDINGS	3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V WAVY RIB; BILATERAL- SEVERAL	
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH	
		V INCOMPLETE OSSIFICATION OF FRONTAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	2 M	NO OBSERVED GROSS FINDINGS	4.0
		NO OBSERVED VISCERAL FINDINGS	
	3 M	NO OBSERVED GROSS FINDINGS	3.5
		MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	4 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	3.4
		MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		2ND, 3RD, 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				TABLE : IIB-	062
TEST GROUP 1 (30 MG/KG BW/D)					
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)		
42 (CONTINUED)					
	6 M	V INCOMPLETE OSSIFICATION OF PARIETAL; LEFT; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.9		
	7 F	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY THORACIC VERTEBRA- 14th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL	3.4		
		V WAVY RIB; BILATERAL- SEVERAL 4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH			
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES			
	8 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.6		
	9 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	3.4		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage WITH BILATERAL HOLES			
43	1 M	NO OBSERVED GROSS FINDINGS NO OBSERVED SKELETAL FINDINGS	3.8		
	2 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.7		
	3 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH	3.4		
		V BIPARTITE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY			
	4 M	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.7		
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE					

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS				TABLE : IIB-	063
ORAL ADMINISTRATION (GAVAGE)					
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS					
TEST GROUP 1 (30 MG/KG BW/D)					
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)		
43 (CONTINUED)					
	5 M	NO OBSERVED GROSS FINDINGS	3.8		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present			
		RUDIMENTARY			
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage			
	6 M	NO OBSERVED GROSS FINDINGS	3.8		
		NO OBSERVED VISCERAL FINDINGS			
	7 F	NO OBSERVED GROSS FINDINGS	3.5		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL			
		5TH, 6TH			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			
		WITH BILATERAL HOLES			
	8 F	NO OBSERVED GROSS FINDINGS	3.6		
		NO OBSERVED VISCERAL FINDINGS			
	9 F	NO OBSERVED GROSS FINDINGS	3.8		
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL			
		3RD, 4TH, 5TH			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V INCOMPLETE OSSIFICATION OF BASISPHENOID			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			
		V BASIOCCIPITAL HOLE(S)			
	10 F	NO OBSERVED GROSS FINDINGS	3.8		
		NO OBSERVED VISCERAL FINDINGS			
	11 M	NO OBSERVED GROSS FINDINGS	3.7		
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL			
		2ND, 3RD, 4TH, 5TH			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present			
		RUDIMENTARY			
		V INCOMPLETE OSSIFICATION OF BASISPHENOID			
	12 M	NO OBSERVED GROSS FINDINGS	3.5		
		NO OBSERVED VISCERAL FINDINGS			
	44	1 M	3.7		
		NO OBSERVED GROSS FINDINGS			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL			
		3RD, 4TH, 5TH			
OBSERVATION CODES: V=Variation C=Cartilage					
SEX CODE: M=MALE, F=FEMALE					

10-APR-14		05R018		TABLE : IIB-		064
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#		FETUS# (SEX)		OBSERVATION		WEIGHT (G)

44		(CONTINUED)		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
	2 M			V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present		3.7
	3 F			NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS		3.2
				V UNOSSIFIED STERNEBRA; Unchanged cartilage- 5th		
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
				C BIPARTITE PROCESSUS XIPHOIDEUS		
				V INCOMPLETE OSSIFICATION OF BASISPHENOID		
				V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
	4 F			V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		3.6
				NO OBSERVED GROSS FINDINGS		
				NO OBSERVED VISCERAL FINDINGS		
	5 M			NO OBSERVED GROSS FINDINGS		4.0
				V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th		
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
				C BIPARTITE PROCESSUS XIPHOIDEUS		
				V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
				RUDIMENTARY		
	6 F			NO OBSERVED GROSS FINDINGS		3.5
				NO OBSERVED VISCERAL FINDINGS		
	7 F			NO OBSERVED GROSS FINDINGS		3.5
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH		
				V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present		
				RUDIMENTARY		
	8 M			NO OBSERVED GROSS FINDINGS		3.8
				NO OBSERVED VISCERAL FINDINGS		
	9 M			NO OBSERVED GROSS FINDINGS		3.8
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
				C BIPARTITE PROCESSUS XIPHOIDEUS		
	10 M			NO OBSERVED GROSS FINDINGS		3.5
				NO OBSERVED VISCERAL FINDINGS		
	11 F			NO OBSERVED GROSS FINDINGS		3.5
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH		
				V CERVICAL RIB; LEFT; Cartilage not present		
	12 F			NO OBSERVED GROSS FINDINGS		3.4
				NO OBSERVED VISCERAL FINDINGS		
	1 F			NO OBSERVED GROSS FINDINGS		3.2

OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 065

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
45	(CONTINUED)		
		V MISAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
		NO OBSERVED GROSS FINDINGS	3.4
	2 M	NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	3.6
	3 M	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V WAVY RIB; BILATERAL- SEVERAL	
		5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH	
		V INCOMPLETE OSSIFICATION OF BASISPHEMION	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	4 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	5 F	NO OBSERVED GROSS FINDINGS	3.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL	
		5TH, 6TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SACRAL ARCH; BILATERAL; Cartilage present- SEVERAL	
		3RD LEFT SIDE	
		4TH BILATERAL	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
	6 M	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		TEMPORAL, FRONTAL, PARIETAL AND SUPRAOCCIPITAL BONES	
	8 M	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018	TABLE : IIB-		066
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS					
TEST GROUP 1 (30 MG/KG BW/D)					
FEMALE#	FETUS# (SEX)	OBSERVATION			WEIGHT (G)

45	(CONTINUED)				
	9 M	NO OBSERVED GROSS FINDINGS			3.0
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
	10 M	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			3.4
		NO OBSERVED GROSS FINDINGS			
	11 F	NO OBSERVED VISCERAL FINDINGS			
		NO OBSERVED GROSS FINDINGS			3.2
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 4th			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH			
		V WAVY RIB; BILATERAL- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH			
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage			
		NEARLY ALL SKULL BONES			
46	1 M	NO OBSERVED GROSS FINDINGS			3.7
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH, 5TH			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present			
		RUDIMENTARY			
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			3.5
	2 M	NO OBSERVED GROSS FINDINGS			
		NO OBSERVED VISCERAL FINDINGS			
	3 M	NO OBSERVED GROSS FINDINGS			3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present			
		RUDIMENTARY			
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			3.7
	4 M	NO OBSERVED GROSS FINDINGS			
		NO OBSERVED VISCERAL FINDINGS			
	5 M	NO OBSERVED GROSS FINDINGS			2.9
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH			

OBSERVATION CODES: V=Variation C=Cartilage					
SEX CODE: M=MALE, F=FEMALE					

10-APR-14		05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS		TABLE : IIB-		067	
				ORAL ADMINISTRATION (GAVAGE)					
TEST GROUP 1 (30 MG/KG BW/D)				INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS					
FEMALE#		FETUS# (SEX)		OBSERVATION				WEIGHT (G)	
46		(CONTINUED)							
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
				C BIPARTITE PROCESSUS XIPHOIDEUS					
				V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present					
				SMALL					
		6 M		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				3.7	
				NO OBSERVED GROSS FINDINGS					
				NO OBSERVED VISCERAL FINDINGS					
		7 M		NO OBSERVED GROSS FINDINGS				3.4	
				V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL					
				2ND, 3RD, 4TH, 5TH					
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
				C BIPARTITE PROCESSUS XIPHOIDEUS				3.8	
		8 M		NO OBSERVED GROSS FINDINGS				3.4	
				NO OBSERVED VISCERAL FINDINGS					
		9 M		NO OBSERVED GROSS FINDINGS					
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
				C BIPARTITE PROCESSUS XIPHOIDEUS				3.3	
		10 F		NO OBSERVED GROSS FINDINGS				3.6	
				NO OBSERVED VISCERAL FINDINGS					
		11 M		NO OBSERVED GROSS FINDINGS					
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
				C BIPARTITE PROCESSUS XIPHOIDEUS					
				V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present					
				RUDIMENTARY					
				V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				3.3	
		12 F		NO OBSERVED GROSS FINDINGS					
				NO OBSERVED VISCERAL FINDINGS					
		1 F		NO OBSERVED GROSS FINDINGS				3.6	
				V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL					
				3RD, 4TH, 5TH					
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
				C BIPARTITE PROCESSUS XIPHOIDEUS					
				V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				3.6	
		2 F		NO OBSERVED GROSS FINDINGS					
				NO OBSERVED VISCERAL FINDINGS					
		3 M		NO OBSERVED GROSS FINDINGS				3.3	
				V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL					
				3RD, 4TH, 5TH					
				V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th					
				OBSERVATION CODES: V=Variation C=Cartilage					
				SEX CODE: M=MALE, F=FEMALE					

10-APR-14

05R018

TABLE : IIB-

068

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
47	(CONTINUED)		
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.3
	4 F	NO OBSERVED GROSS FINDINGS	
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	3.8
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	6 M	NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	3.9
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	8 F	NO OBSERVED GROSS FINDINGS	3.9
		NO OBSERVED VISCERAL FINDINGS	
	9 M	NO OBSERVED GROSS FINDINGS	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	3.7
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
	10 M	NO OBSERVED GROSS FINDINGS	3.9
		NO OBSERVED VISCERAL FINDINGS	
	11 F	NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	3.5
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	12 M	NO OBSERVED GROSS FINDINGS	3.6

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS		TABLE : IIB-	069
			ORAL ADMINISTRATION (GAVAGE)			
TEST GROUP 1 (30 MG/KG BW/D)			INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS			
FEMALE#	FETUS# (SEX)	OBSERVATION				WEIGHT (G)
47	(CONTINUED)					
	13 F	NO OBSERVED VISCERAL FINDINGS				
		NO OBSERVED GROSS FINDINGS				
		V UNOBSERVED STERNEBRA; Unchanged cartilage- 6th				2.9
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
48	1 F	NO OBSERVED GROSS FINDINGS				3.6
		V MISSAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				3.6
	2 F	NO OBSERVED GROSS FINDINGS				
		NO OBSERVED VISCERAL FINDINGS				
	3 M	NO OBSERVED GROSS FINDINGS				3.2
		V MISSAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		4TH, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL				
	4 F	NO OBSERVED GROSS FINDINGS				3.6
		NO OBSERVED VISCERAL FINDINGS				
	5 M	NO OBSERVED GROSS FINDINGS				3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present				
		SMALL				
		V DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- SEVERAL				
		4TH, 10TH				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
		C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL				
	6 M	NO OBSERVED GROSS FINDINGS				4.2
		NO OBSERVED VISCERAL FINDINGS				
		NO OBSERVED GROSS FINDINGS				
	7 F	NO OBSERVED GROSS FINDINGS				3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14		05R018		TABLE : IIB-		070
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 1 (30 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION				WEIGHT (G)
48 (CONTINUED)						
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V DUMBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 10th				
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				3.1
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				3.8
	8 F	NO OBSERVED GROSS FINDINGS				
		NO OBSERVED VISCERAL FINDINGS				
	10 M	NO OBSERVED GROSS FINDINGS				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present				
		RUDIMENTARY				
	11 M	NO OBSERVED GROSS FINDINGS				3.3
		NO OBSERVED VISCERAL FINDINGS				
	12 F	NO OBSERVED GROSS FINDINGS				3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
	13 F	NO OBSERVED GROSS FINDINGS				3.4
		NO OBSERVED VISCERAL FINDINGS				
49						
	1 F	NO OBSERVED GROSS FINDINGS				3.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
	2 F	NO OBSERVED GROSS FINDINGS				3.4
		NO OBSERVED VISCERAL FINDINGS				
	3 M	NO OBSERVED GROSS FINDINGS				3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V DUMBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 13th				
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present				
		RUDIMENTARY				
	4 F	NO OBSERVED GROSS FINDINGS				3.3
		NO OBSERVED VISCERAL FINDINGS				
	5 M	NO OBSERVED GROSS FINDINGS				3.5
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		2ND, 3RD, 4TH, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB-071

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
49	(CONTINUED)		
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V MISSHAPEN SACRAL VERTEBRA	
		1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT	
	6 F	NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	4.1
		NO OBSERVED VISCERAL FINDINGS	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage present	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		SMALL	
		V MISSHAPEN SACRAL VERTEBRA	
		1ST SACRAL ARCH - RIGHT SIDE, CARTILAGE PRESENT	
	8 F	NO OBSERVED GROSS FINDINGS	3.7
		NO OBSERVED VISCERAL FINDINGS	
	9 M	NO OBSERVED GROSS FINDINGS	3.7
		NO OBSERVED VISCERAL FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	10 M	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	11 M	NO OBSERVED GROSS FINDINGS	3.8
		NO OBSERVED VISCERAL FINDINGS	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	12 F	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	50	1 F	3.1
		NO OBSERVED GROSS FINDINGS	
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V WAVY RIB; RIGHT- SEVERAL	
		6TH, 7TH, 8TH, 9TH, 10TH	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB- 072

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 1 (30 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
50	(CONTINUED)		
	2 M	NO OBSERVED GROSS FINDINGS	3.8
		NO OBSERVED VISCERAL FINDINGS	
	3 F	NO OBSERVED GROSS FINDINGS	3.6
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	4 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	4.0
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V MISSHAPEN SACRAL VERTEBRA	
		1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	6 M	NO OBSERVED GROSS FINDINGS	3.2
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	3.7
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL	
		5TH, 6TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V CERVICAL RIB; BILATERAL; Cartilage not present	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	8 F	NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
	9 F	NO OBSERVED GROSS FINDINGS	3.2
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		4TH, 6TH	
		6TH ADDITIONALLY INCOMPLETELY OSSIFIED	
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- 5th	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	

OBSERVATION CODES: V=Variation C=Cartilage

SEX CODE: M=MALE, F=FEMALE

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS				TABLE : IIB-073
ORAL ADMINISTRATION (GAVAGE)				
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				
TEST GROUP 2 (100 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)	
51	1 F	NO OBSERVED GROSS FINDINGS	3.5	
		V MISSHPAPEN STERNEBRA; Unchanged cartilage- 5th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
	2 M	V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	3.7	
		SMALL		
		NO OBSERVED GROSS FINDINGS		
		NO OBSERVED VISCERAL FINDINGS		
	3 F	NO OBSERVED GROSS FINDINGS	3.5	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		NO OBSERVED GROSS FINDINGS		
4 F	NO OBSERVED VISCERAL FINDINGS	2.7		
	NO OBSERVED GROSS FINDINGS			
	NO OBSERVED GROSS FINDINGS			
	NO OBSERVED VISCERAL FINDINGS			
5 M	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	3.5		
	C BIPARTITE PROCESSUS XIPHOIDEUS			
	NO OBSERVED GROSS FINDINGS			
	NO OBSERVED VISCERAL FINDINGS			
6 F	NO OBSERVED GROSS FINDINGS	3.7		
	NO OBSERVED VISCERAL FINDINGS			
	NO OBSERVED GROSS FINDINGS			
	NO OBSERVED VISCERAL FINDINGS			
7 F	NO OBSERVED GROSS FINDINGS	3.5		
	V MISSHPAPEN STERNEBRA; Unchanged cartilage- SEVERAL			
	4TH, 5TH			
	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
8	8 M	V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	3.8	
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
		NO OBSERVED GROSS FINDINGS		
	9 F	NO OBSERVED VISCERAL FINDINGS	3.8	
		NO OBSERVED GROSS FINDINGS		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		NO OBSERVED GROSS FINDINGS		
	10 F	NO OBSERVED VISCERAL FINDINGS	3.4	
		NO OBSERVED GROSS FINDINGS		
		V MISSHPAPEN STERNEBRA; Unchanged cartilage- 5th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
11 M	C BIPARTITE PROCESSUS XIPHOIDEUS	3.9		
	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage			
	NO OBSERVED GROSS FINDINGS			
	V DILATED RENAL PELVIS; right			
12 M	V DILATED URETER; right	3.9		
OBSERVATION CODES: V=Variation C=Cartilage				
SEX CODE: M=MALE, F=FEMALE				

10-APR-14		05R018		TABLE : IIB-		074
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 2 (100 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
52	1 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY THORACIC VERTEBRA- 14th V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage present V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present SMALL V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage NEARLY ALL SKULL BONES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISAPEN STERNEBRA; Unchanged cartilage- 4th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V UNILATERAL OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th ADDITIONALLY REDUCED IN SIZE C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF NASAL; BILATERAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V UNILATERAL OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID	3.5			
	2 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.3			
	3 F	NO OBSERVED GROSS FINDINGS V MISAPEN STERNEBRA; Unchanged cartilage- 4th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V UNILATERAL OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th ADDITIONALLY REDUCED IN SIZE C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V UNILATERAL OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID	3.4			
	4 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.3			
	5 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V UNILATERAL OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID	3.4			
	6 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID	3.5			
	7 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID	3.5			
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 075

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
52	(CONTINUED)		
	8 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.5
	9 F	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.4
	10 M	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.5
	11 F	V UNOSSIFIED STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.1
53	1 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 11th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS V DILATED RENAL PELVIS; left NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present SMALL	3.6
	2 F		3.4
	3 F		3.6

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		076
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 2 (100 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
53	(CONTINUED)					
	4 F	V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present	3.3			
		NO OBSERVED GROSS FINDINGS				
	5 M	NO OBSERVED VISCERAL FINDINGS	3.5			
		NO OBSERVED GROSS FINDINGS				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH,5TH				
	6 M	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	3.3			
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		NO OBSERVED GROSS FINDINGS				
	7 F	NO OBSERVED VISCERAL FINDINGS	2.9			
		NO OBSERVED GROSS FINDINGS				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH,6TH				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
		V MISSHAPEN SACRAL VERTEBRA				
		1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT				
	9 F	V UNOSSIFIED HYOID; Cartilage present	3.5			
		NO OBSERVED GROSS FINDINGS				
		NO OBSERVED VISCERAL FINDINGS				
	10 F	NO OBSERVED GROSS FINDINGS	3.5			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V CERVICAL RIB; RIGHT; Cartilage not present				
		V WAVY RIB; RIGHT- SEVERAL				
		5TH,6TH,7TH,8TH,9TH,10TH				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
		C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	11 M	NO OBSERVED GROSS FINDINGS	3.6			
		NO OBSERVED VISCERAL FINDINGS				
54	1 F	NO OBSERVED GROSS FINDINGS	3.3			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH,5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE						

10-APR-14		05R018		TABLE : IIB-		077
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 2 (100 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
54 (CONTINUED)						
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
	2 M	NO OBSERVED GROSS FINDINGS				3.7
	3 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS				3.5
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present				
		RUDIMENTARY				
	4 M	V INCOMPLETE OSSIFICATION OF BASISPHENOID				4.0
		NO OBSERVED GROSS FINDINGS				
	5 F	NO OBSERVED VISCERAL FINDINGS				3.7
		NO OBSERVED GROSS FINDINGS				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
	6 M	NO OBSERVED GROSS FINDINGS				3.8
		NO OBSERVED VISCERAL FINDINGS				
	7 M	NO OBSERVED GROSS FINDINGS				3.5
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
	8 M	NO OBSERVED GROSS FINDINGS				3.9
		NO OBSERVED VISCERAL FINDINGS				
	9 M	NO OBSERVED GROSS FINDINGS				4.0
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
	10 F	NO OBSERVED GROSS FINDINGS				4.0
		NO OBSERVED VISCERAL FINDINGS				
		NO OBSERVED VISCERAL FINDINGS				
	1 F	NO OBSERVED GROSS FINDINGS				3.4
55						
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB-

078

PROJECT NO 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE# FETUS# (SEX) OBSERVATION WEIGHT (G)

55 (CONTINUED)

	V	INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
	C	BIPARTITE PROCESSUS XIPHOIDEUS	
	V	INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.5
2 F		NO OBSERVED GROSS FINDINGS	
		NO OBSERVED VISCERAL FINDINGS	
3 M		NO OBSERVED GROSS FINDINGS	3.8
	V	MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
	V	INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
	C	BIPARTITE PROCESSUS XIPHOIDEUS	
	V	INCOMPLETE OSSIFICATION OF BASISPHEOID	
	V	INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.6
4 F		NO OBSERVED GROSS FINDINGS	
		NO OBSERVED VISCERAL FINDINGS	
6 F		NO OBSERVED GROSS FINDINGS	3.6
	V	UNILATERAL OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th	
	V	INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
	C	BIPARTITE PROCESSUS XIPHOIDEUS	
	V	INCOMPLETE OSSIFICATION OF BASISPHEOID	
7 M		NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
8 M		NO OBSERVED GROSS FINDINGS	3.4
	V	MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
	V	INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
	C	BIPARTITE PROCESSUS XIPHOIDEUS	
	V	INCOMPLETE OSSIFICATION OF BASISPHEOID	
	V	INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.5
9 F		NO OBSERVED GROSS FINDINGS	
		NO OBSERVED VISCERAL FINDINGS	
10 M		NO OBSERVED GROSS FINDINGS	3.6
	V	MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
	V	INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
	C	BIPARTITE PROCESSUS XIPHOIDEUS	
	V	SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	

56	1 F	NO OBSERVED GROSS FINDINGS	3.3
	V	MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
	V	INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
	V	SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB- 079

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
56	(CONTINUED)		
	3 F	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	3.5
		NO OBSERVED GROSS FINDINGS	
	4 M	NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	3.6
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	5 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	6 M	NO OBSERVED GROSS FINDINGS	
		NO OBSERVED VISCERAL FINDINGS	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	3.7
		1ST, 2ND, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
	7 F	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	8 F	NO OBSERVED GROSS FINDINGS	3.2
		NO OBSERVED VISCERAL FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage present	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		SMALL	
	9 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	10 M	NO OBSERVED GROSS FINDINGS	3.8
		NO OBSERVED VISCERAL FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	11 F	NO OBSERVED GROSS FINDINGS	3.7
		NO OBSERVED VISCERAL FINDINGS	
	12 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
		V BIPARTITE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th	
		ADDITIONALLY INCOMPLETELY OSSIFIED	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V CERVICAL RIB; BILATERAL; Cartilage not present	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-080

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
56	(CONTINUED) 13 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.7
57	1 F	NO OBSERVED GROSS FINDINGS V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C FUSED RIB CARTILAGE; LEFT PROXIMAL V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF PARIETAL; RIGHT; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V MISHPAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	3.6
	2 F		3.6
	3 M		3.7
	4 F		3.5
	5 F		3.5
	6 F		3.8
	7 F		3.4
	8 F		3.5
	9 F		3.4

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB- 081

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
57	(CONTINUED)		
	10 F	NO OBSERVED GROSS FINDINGS	3.3
		V DILATED RENAL PELVIS; left	
	11 F	NO OBSERVED GROSS FINDINGS	3.7
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V MISHPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	12 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	
	1 F	NO OBSERVED GROSS FINDINGS	3.3
		V MISHPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	2 F	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	3 M	NO OBSERVED GROSS FINDINGS	3.8
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	4 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	5 F	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present	
		RUDIMENTARY	
	6 F	NO OBSERVED GROSS FINDINGS	3.1
		NO OBSERVED VISCERAL FINDINGS	
	7 F	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- SEVERAL 11TH, 12TH	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-082

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
58	(CONTINUED) 8 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.3
59	1 F	NO OBSERVED GROSS FINDINGS V MISHPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF BASISPHENOID NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V MISHPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V WAVY RIB; BILATERAL- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH RIGHT SIDE 7TH, 8TH, 9TH LEFT SIDE V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	3.2
	2 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.3
	4 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF BASISPHENOID NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V MISHPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V WAVY RIB; BILATERAL- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH RIGHT SIDE 7TH, 8TH, 9TH LEFT SIDE V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	3.3
	6 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.1
	7 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V MISHPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V WAVY RIB; BILATERAL- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH RIGHT SIDE 7TH, 8TH, 9TH LEFT SIDE V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	3.3
	8 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.1
	9 M	NO OBSERVED GROSS FINDINGS V MISHPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V WAVY RIB; BILATERAL- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH RIGHT SIDE 7TH, 8TH, 9TH LEFT SIDE V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	3.4
	10 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.4
	11 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	3.5

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB- 083

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
59	(CONTINUED)		
	12 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.4
	13 M	V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.2
60	1 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH 6TH ADDITIONALLY INCOMPLETELY OSSIFIED V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.9
	2 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.9
	3 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS	3.7
	4 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.9
	5 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY	3.9
	6 F	V INCOMPLETE OSSIFICATION OF BASISPHENOID NO OBSERVED GROSS FINDINGS	3.8
	7 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH 6TH ADDITIONALLY INCOMPLETELY OSSIFIED V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY	3.9
	8 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	4.0

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

084

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
60	(CONTINUED)		
	9 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 1ST, 2ND, 3RD, 4TH, 5TH	3.9
	10 F	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th NO OBSERVED GROSS FINDINGS	4.0
	11 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.7
	12 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.5
	13 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH, 5TH, 6TH 6TH ADDITIONALLY INCOMPLETELY OSSIFIED V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY	3.3
61	1 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 6TH V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage NEARLY ALL SKULL BONES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS M FETUS WITH MULTIPLE EXTERNAL MALFORMATIONS - HEAD MISSHAPEN - FACE ABSENT (ANOPHTHALMIA, ANOTIA, ASTOMIA) M FETUS WITH MULTIPLE SKELETAL MALFORMATIONS IN CORRELATION TO THE NOTED EXTERNAL MALFORMATIONS THE FOLLOWING FINDINGS WERE OBSERVED: - MOST SKULL BONES ARE MISSHAPEN, FUSED, ABSENT AND/OR INCOMPLETELY OSSIFIED ADDITIONAL DISTINCT FINDINGS IN THE AREA OF THE VERTEBRAL COLUMN, RIBS, PELVIC GIRDLE AND FORELIMBS WERE NOTED: - SHORTENED SCAPULA AND HUMERUS, LEFT - BENT RADIUS, LEFT	3.9
	2 M		3.5
	3 M		1.8

OBSERVATION CODES: M=Malformation V=Variation
SEX CODE: M=MALE, F=FEMALE

10-APR-14		TABLE : IIB-		085
05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)		
TEST GROUP 2 (100 MG/KG BW/D)		INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS		
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)	
(CONTINUED)				
61		- THORACIC VERTEBRA CENTERS MISSHAPEN AND FUSED WITH EACH OTHER - RIBS ARE BENT, WAVY AND/OR INCOMPLETELY OSSIFIED - LUMBAR AND SACRAL VERTEBRAE INCOMPLETELY OSSIFIED - PUBIS AND ISCHIUM INCOMPLETELY OSSIFIED, BILATERAL BECAUSE OF THE COMPLEXITY OF THESE SKELETAL FINDINGS, A SPECIFICATION OF EACH AND EVERY DETAIL IS INPRACTICAL. NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 12th V CERVICAL RIB; LEFT; Cartilage not present V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.5	3.6
	4 M			
	5 F			
62	1 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present RUDIMENTARY V MISSHAPEN SACRAL VERTEBRA 1ST SACRAL ARCH - RIGHT SIDE, CARTILAGE PRESENT V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 6TH C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH) ; RIGHT; Cartilage not present RUDIMENTARY V WAVY RIB; BILATERAL- SEVERAL 4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage NEARLY ALL SKULL BONES	3.4	
	2 F			3.3
	3 M			3.8
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE				

10-APR-14

05R018

TABLE : IIB- 086

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
62	(CONTINUED)		
	4 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	5 F	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
	6 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	3.6
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	8 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	9 F	NO OBSERVED GROSS FINDINGS	3.2
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present	
		V WAVY RIB: BILATERAL- SEVERAL 3RD, 4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
		M SHORTENED HUMERUS; RIGHT	
63	1 M	NO OBSERVED GROSS FINDINGS	2.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 1ST, 5TH	
		V UNOSIFIED STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHEOID	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
		WITH BILATERAL HOLES	
	2 F	NO OBSERVED GROSS FINDINGS	3.2
		NO OBSERVED VISCERAL FINDINGS	
	3 F	NO OBSERVED GROSS FINDINGS	3.5

OBSERVATION CODES: M=Malformation V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		087
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 2 (100 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
63 (CONTINUED)						
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	4 M	NO OBSERVED GROSS FINDINGS				3.3
		NO OBSERVED VISCERAL FINDINGS				
	5 M	NO OBSERVED GROSS FINDINGS				3.4
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
	6 M	NO OBSERVED GROSS FINDINGS				3.7
		NO OBSERVED VISCERAL FINDINGS				
	7 F	NO OBSERVED GROSS FINDINGS				3.1
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	8 F	NO OBSERVED GROSS FINDINGS				3.3
		NO OBSERVED VISCERAL FINDINGS				
	9 F	NO OBSERVED GROSS FINDINGS				3.3
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
	10 M	NO OBSERVED GROSS FINDINGS				3.5
		NO OBSERVED VISCERAL FINDINGS				
	11 F	NO OBSERVED GROSS FINDINGS				3.2
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		4TH, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		4TH, 5TH				
	64	1 F				3.3
		NO OBSERVED GROSS FINDINGS				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		2ND, 3RD, 4TH, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS				TABLE : IIB-	088
ORAL ADMINISTRATION (GAVAGE)					
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS					
TEST GROUP 2 (100 MG/KG BW/D)					
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)		
64 (CONTINUED)					
	2 M	V INCOMPLETE OSSIFICATION OF BASISPHENOID			
		NO OBSERVED GROSS FINDINGS			3.2
	3 F	NO OBSERVED VISCERAL FINDINGS			
		NO OBSERVED GROSS FINDINGS			3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
	4 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage			3.5
		NO OBSERVED GROSS FINDINGS			
	5 M	NO OBSERVED VISCERAL FINDINGS			3.3
		NO OBSERVED GROSS FINDINGS			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH) ; RIGHT; Cartilage not present			
		RUDIMENTARY			
	6 F	NO OBSERVED GROSS FINDINGS			3.4
		NO OBSERVED VISCERAL FINDINGS			
	7 M	NO OBSERVED GROSS FINDINGS			3.4
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			
	8 F	NO OBSERVED GROSS FINDINGS			3.2
		NO OBSERVED VISCERAL FINDINGS			
	9 F	NO OBSERVED GROSS FINDINGS			3.4
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present			
		RUDIMENTARY			
	10 F	NO OBSERVED GROSS FINDINGS			3.3
		NO OBSERVED VISCERAL FINDINGS			
	11 M	NO OBSERVED GROSS FINDINGS			3.3
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL			
		4TH,5TH			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			
		WITH BILATERAL HOLES			
65	1 M	NO OBSERVED GROSS FINDINGS			3.5
OBSERVATION CODES: V=Variation C=Cartilage					
SEX CODE: M=MALE, F=FEMALE					

10-APR-14	05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)	TABLE : IIB-	089
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				
TEST GROUP 2 (100 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION		WEIGHT (G)
65	(CONTINUED)			
		V INCOMPLETE OSSIFICATION OF METACARPAL; BILATERAL; Cartilage present- SEVERAL		
		2ND, 4TH		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL		
		2ND, 3RD, 4TH, 6TH		
		V MISHPEN STERNEBRA; Unchanged cartilage- 5th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V WAVY RIB; BILATERAL- SEVERAL		
		3RD, 4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH, 13TH		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		
		M SHORTENED SCAPULA; RIGHT		
		V INCOMPLETE OSSIFICATION OF THORACIC ARCH; BILATERAL; Cartilage present- SEVERAL		
		12TH, 13TH		
		V INCOMPLETE OSSIFICATION OF LUMBAR ARCH; RIGHT; Cartilage present- 5th		
		V INCOMPLETE OSSIFICATION OF SACRAL ARCH; BILATERAL; Cartilage present- SEVERAL		
		3RD, 4TH		
		V INCOMPLETE OSSIFICATION OF PUBIS; BILATERAL; Cartilage present		
		V INCOMPLETE OSSIFICATION OF ISCHIUM; RIGHT; Cartilage present		
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage		
		NEARLY ALL SKULL BONES		
2 F		NO OBSERVED GROSS FINDINGS		3.2
		NO OBSERVED VISCERAL FINDINGS		
3 F		NO OBSERVED GROSS FINDINGS		3.8
		V MISHPEN STERNEBRA; Unchanged cartilage- SEVERAL		
		3RD, 4TH, 5TH		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present		
		SMALL		
		V WAVY RIB; BILATERAL- SEVERAL		
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH		
		V INCOMPLETE OSSIFICATION OF FRONTAL; BILATERAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
4 M		NO OBSERVED GROSS FINDINGS		3.6
		NO OBSERVED VISCERAL FINDINGS		
		NO OBSERVED GROSS FINDINGS		
5 M		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		3.5
		V WAVY RIB; BILATERAL- SEVERAL		
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH		
OBSERVATION CODES: M=Malformation V=Variation C=Cartilage				
SEX CODE: M=MALE, F=FEMALE				

10-APR-14

05R018

TABLE : IIB-

090

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
65	(CONTINUED)		
	6 F	V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage TEMPORAL, FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	3.3
	7 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY	3.4
	8 M	V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.6
	9 F	NO OBSERVED GROSS FINDINGS NO OBSERVED STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.7
	10 F	V WAVY RIB; RIGHT- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage TEMPORAL, FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	3.4
	11 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V WAVY RIB; BILATERAL- SEVERAL 4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage TEMPORAL, FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	3.4
66	1 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	3.2

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

091

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
66	(CONTINUED)		
	2 M	C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF BASISPHENOID C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 4th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH V UNOSIFIED STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	3.2 3.6 3.5 3.2 3.7 3.0 3.3 3.5 3.6 3.2

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-092

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
66	(CONTINUED)		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF HYOID; Cartilage present	
		V INCOMPLETE OSSIFICATION OF BASISPHEOID	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	12 F	NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
	13 F	NO OBSERVED GROSS FINDINGS	3.4
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum- 13th	
		C NOTCHED CARTILAGE BETWEEN BASISPHEOID AND BASIOCCIPITAL	
		V INCOMPLETE OSSIFICATION OF BASISPHEOID	
67	1 F	NO OBSERVED GROSS FINDINGS	3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	2 M	NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
	3 F	NO OBSERVED GROSS FINDINGS	3.1
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		NO OBSERVED GROSS FINDINGS	
	4 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	5 F	NO OBSERVED GROSS FINDINGS	3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
	6 M	NO OBSERVED GROSS FINDINGS	3.9
		NO OBSERVED VISCERAL FINDINGS	
	7 F	NO OBSERVED GROSS FINDINGS	3.7
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
	8 F	NO OBSERVED GROSS FINDINGS	3.0

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		093
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 2 (100 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
67 (CONTINUED)						
	9 F	NO OBSERVED VISCERAL FINDINGS	3.7			
		NO OBSERVED GROSS FINDINGS				
		V MISHPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
	10 F	NO OBSERVED GROSS FINDINGS	3.7			
		NO OBSERVED VISCERAL FINDINGS				
	11 M	NO OBSERVED GROSS FINDINGS	3.7			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present				
		SMALL				
68	1 F	NO OBSERVED GROSS FINDINGS	3.1			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		SMALL				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES				
	2 M	NO OBSERVED GROSS FINDINGS	3.2			
		NO OBSERVED VISCERAL FINDINGS				
	3 M	NO OBSERVED GROSS FINDINGS	3.0			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
	4 F	NO OBSERVED GROSS FINDINGS	2.9			
		NO OBSERVED VISCERAL FINDINGS				
	5 F	NO OBSERVED GROSS FINDINGS	3.1			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL				
		5TH, 6TH				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V WAVY RIB; BILATERAL- SEVERAL				
		5TH, 6TH, 7TH, 8TH, 9TH, 10TH				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		BASISPHENOID, FRONTAL, PARIETAL, INTERPARIETAL AND				
		SUPRAOCCIPITAL BONES				
	6 F	NO OBSERVED GROSS FINDINGS	3.3			
		V DILATED RENAL PELVIS; right				
	7 M	NO OBSERVED GROSS FINDINGS	3.4			
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14		TABLE : IIB-		094
05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)		
TEST GROUP 2 (100 MG/KG BW/D)		INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS		
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)	
(CONTINUED)				
68		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage		
		FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES		
	8 F	NO OBSERVED GROSS FINDINGS	3.0	
		NO OBSERVED VISCERAL FINDINGS		
	9 M	NO OBSERVED GROSS FINDINGS	3.0	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL		
		2ND, 3RD, 4TH		
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- SEVERAL		
		5TH, 6TH		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
69		V WAVY RIB; BILATERAL- SEVERAL		
		5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH		
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
	10 F	NO OBSERVED GROSS FINDINGS	3.3	
		NO OBSERVED VISCERAL FINDINGS		
	11 F	NO OBSERVED GROSS FINDINGS	2.6	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 4th		
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- SEVERAL		
		5TH, 6TH		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		
12 F		V INCOMPLETE OSSIFICATION OF PUBIS; BILATERAL; Cartilage present		
		V INCOMPLETE OSSIFICATION OF ISCHIUM; BILATERAL; Cartilage present		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
		WITH BILATERAL HOLES		
		NO OBSERVED GROSS FINDINGS	3.0	
		NO OBSERVED VISCERAL FINDINGS		
		NO OBSERVED GROSS FINDINGS		
	1 M	V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	3.4	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
		OBSERVATION CODES: V=Variation C=Cartilage		
		SEX CODE: M=MALE, F=FEMALE		

10-APR-14

05R018

TABLE : IIB- 095

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
69	(CONTINUED)		
	2 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.5
		NO OBSERVED GROSS FINDINGS	
	3 M	V DILATED RENAL PELVIS; bilateral	3.9
		NO OBSERVED GROSS FINDINGS	
		V MISHPHEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	4 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.8
		NO OBSERVED GROSS FINDINGS	
	5 F	NO OBSERVED VISCERAL FINDINGS	3.9
		NO OBSERVED GROSS FINDINGS	
		V MISHPHEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	6 F	NO OBSERVED GROSS FINDINGS	3.8
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	3.7
		V MISHPHEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	8 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	9 M	NO OBSERVED GROSS FINDINGS	3.8
		V MISHPHEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
	10 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS				TABLE : IIB-	096
ORAL ADMINISTRATION (GAVAGE)					
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS					
TEST GROUP 2 (100 MG/KG BW/D)					
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)		
70	1 M	NO OBSERVED GROSS FINDINGS	3.4		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present			
	2 M	RUDIMENTARY	3.5		
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage			
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			
	3 M	NO OBSERVED GROSS FINDINGS	3.4		
		NO OBSERVED VISCERAL FINDINGS			
		NO OBSERVED GROSS FINDINGS			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
		C BIPARTITE PROCESSUS XIPHOIDEUS			
		V SUPERNUMERARY THORACIC VERTEBRA- 14th			
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present			
		SMALL			
5 M	V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	3.4			
	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	NO OBSERVED GROSS FINDINGS				
6 F	NO OBSERVED VISCERAL FINDINGS	3.2			
	NO OBSERVED GROSS FINDINGS				
	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
	V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
7 M	RUDIMENTARY	3.6			
	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	NO OBSERVED GROSS FINDINGS				
	V DILATED RENAL PELVIS; left				
9 M	NO OBSERVED GROSS FINDINGS	3.8			
	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
	C BIPARTITE PROCESSUS XIPHOIDEUS				
	V INCOMPLETE OSSIFICATION OF BASISPHENOID				
10 M	NO OBSERVED GROSS FINDINGS	3.7			
	NO OBSERVED VISCERAL FINDINGS				
	NO OBSERVED GROSS FINDINGS				
	V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
11 F	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	3.2			
	V INCOMPLETE OSSIFICATION OF BASISPHENOID				
	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
OBSERVATION CODES: V=Variation C=Cartilage					
SEX CODE: M=MALE, F=FEMALE					

10-APR-14

05R018

TABLE : IIB- 097

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
70	(CONTINUED)		
	12 F	NO OBSERVED GROSS FINDINGS	3.0
		V SHORT INNOMINATE	
	13 F	NO OBSERVED GROSS FINDINGS	3.3
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
71	3 M	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage present	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	4 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	6 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	8 M	NO OBSERVED GROSS FINDINGS	3.2
		NO OBSERVED VISCERAL FINDINGS	
	9 F	NO OBSERVED GROSS FINDINGS	2.9
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
	10 F	NO OBSERVED GROSS FINDINGS	3.3
		V DILATED RENAL PELVIS; left	
	11 F	NO OBSERVED GROSS FINDINGS	3.4

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

098

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
71	(CONTINUED)		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Dumbbell-shaped cartilage of centrum- 11th	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	12 M	NO OBSERVED GROSS FINDINGS	3.2
		NO OBSERVED VISCERAL FINDINGS	
	13 F	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
72	1 M	NO OBSERVED GROSS FINDINGS	3.0
		V MISHPAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	2 F	NO OBSERVED GROSS FINDINGS	2.8
		NO OBSERVED VISCERAL FINDINGS	
	3 F	NO OBSERVED GROSS FINDINGS	2.7
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage WITH BILATERAL HOLES	
	4 F	NO OBSERVED GROSS FINDINGS	2.9
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	6 F	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	2.7
		NO OBSERVED GROSS FINDINGS	
		NO OBSERVED VISCERAL FINDINGS	
	7 F	NO OBSERVED GROSS FINDINGS	1.9
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 1st	
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH, 5TH, 6TH	

OBSERVATION CODES: V=Variation C=Cartilage

SEX CODE: M=MALE, F=FEMALE

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB- 099

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
72	(CONTINUED)		
		V INCOMPLETE OSSIFICATION OF PUBIS; BILATERAL; Cartilage present	
		V INCOMPLETE OSSIFICATION OF ISCHIIUM; BILATERAL; Cartilage present	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		NEARLY ALL SKULL BONES	
	8 F	NO OBSERVED GROSS FINDINGS	3.0
		NO OBSERVED VISCERAL FINDINGS	
	9 M	NO OBSERVED GROSS FINDINGS	3.2
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	11 M	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	12 F	NO OBSERVED GROSS FINDINGS	2.9
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
		WITH BILATERAL HOLES	
73	1 M	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL	
		2ND, 6TH	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		4TH, 5TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
	2 M	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	3 M	NO OBSERVED GROSS FINDINGS	3.7
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	4 F	NO OBSERVED GROSS FINDINGS	3.0

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		100
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 2 (100 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
73 (CONTINUED)						
	5 M	V SHORT INNOMINATE NO OBSERVED GROSS FINDINGS	3.7			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
	6 F	C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF BASISPHEOID	3.9			
	7 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.8			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
	8 F	C BIPARTITE PROCESSUS XIPHOIDEUS V SUPRAOCCIPITAL HOLE(S) ; BILATERAL	3.7			
	9 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.8			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage WITH BILATERAL HOLES	3.3			
	10 F	NO OBSERVED GROSS FINDINGS				
	11 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.7			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
	12 F	V SUPERNUMERARY RIB (14TH) ; LEFT; Cartilage not present RUDIMENTARY	3.5			
		NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS				
74	1 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	3.8			
		6TH ADDITIONALLY INCOMPLETELY OSSIFIED				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
	2 M	NO OBSERVED GROSS FINDINGS	4.0			
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB-

101

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 2 (100 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
74	(CONTINUED)		
	3 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V MISCHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISCHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V WAVY RIB; BILATERAL- SEVERAL 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH RIGHT SIDE 11TH, 12TH LEFT SIDE V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS V MISCHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES NO OBSERVED GROSS FINDINGS V DILATED RENAL PELVIS; right NO OBSERVED GROSS FINDINGS	3.9 3.5 3.8 3.5 4.0 3.6 3.6 3.8
75	1 M	NO OBSERVED GROSS FINDINGS V MISCHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES NO OBSERVED GROSS FINDINGS V DILATED RENAL PELVIS; right NO OBSERVED GROSS FINDINGS	3.7 3.3

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		TABLE : IIB-		102
05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS		
TEST GROUP 2 (100 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)	
75 (CONTINUED)				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		
	4 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		3.6
		NO OBSERVED GROSS FINDINGS		
	5 F	NO OBSERVED VISCERAL FINDINGS		4.0
		NO OBSERVED GROSS FINDINGS		
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage		
	6 F	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		3.6
		NO OBSERVED GROSS FINDINGS		
	7 F	NO OBSERVED VISCERAL FINDINGS		3.7
		NO OBSERVED GROSS FINDINGS		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		V INCOMPLETE OSSIFICATION OF BASISPHENOID		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
	8 F	NO OBSERVED GROSS FINDINGS		4.2
		V DILATED RENAL PELVIS; left		
	9 M	NO OBSERVED GROSS FINDINGS		3.9
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL		
		3RD, 4TH, 5TH		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present		
	10 F	NO OBSERVED GROSS FINDINGS		3.9
		V DILATED RENAL PELVIS; left		
OBSERVATION CODES: V=Variation C=Cartilage				
SEX CODE: M=MALE, F=FEMALE				

05R018		TABLE : IIB-		103
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)				
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				
TEST GROUP 3 (300 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)	
76	1 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH C BIPARTITE PROCESSUS XIPHOIDEUS V WAVY RIB; BILATERAL- SEVERAL 4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH RIGHT SIDE 8TH, 9TH, 10TH, 11TH LEFT SIDE V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF BASISPHENOID NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present RUDIMENTARY	3.7	
	2 F			3.6
	3 M			4.1
	4 M			3.7
	5 F			4.0
	6 F			3.6
	7 M			3.9
	8 M			4.1
	9 F			3.9
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE				

10-APR-14		05R018		TABLE : IIB-		104
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS						
ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 3 (300 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			

76	(CONTINUED)					
	10 F	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	3.6			
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
		NO OBSERVED GROSS FINDINGS				
	11 F	NO OBSERVED VISCERAL FINDINGS	3.5			
		NO OBSERVED GROSS FINDINGS				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V WAVY RIB; BILATERAL- SEVERAL				
		6TH, 7TH, 8TH, 9TH, 11TH RIGHT SIDE				
		10TH, 11TH LEFT SIDE				
		V INCOMPLETE OSSIFICATION OF SACRAL ARCH; BILATERAL; Cartilage present- SEVERAL 3RD LEFT SIDE				
		4TH BILATERAL				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		NEARLY ALL SKULL BONES				
77	1 M	NO OBSERVED GROSS FINDINGS	3.0			
		V MISHPAPEN STERNEBRA; Unchanged cartilage- 3rd				
		V BIPARTITE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH				
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- 2nd				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
	2 F	NO OBSERVED GROSS FINDINGS	3.2			
	3 M	NO OBSERVED VISCERAL FINDINGS	3.2			
		NO OBSERVED GROSS FINDINGS				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
	4 M	NO OBSERVED GROSS FINDINGS	3.4			
		V DILATED RENAL PELVIS; left				
		V DILATED URETER; left				
	5 M	NO OBSERVED GROSS FINDINGS	3.3			

OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14	05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)	TABLE : IIB-	105
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				
TEST GROUP 3 (300 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION		WEIGHT (G)
77	(CONTINUED)			
	6 M	V MISHPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY		3.3
	7 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISHPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY		3.7
	8 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS		3.1
	9 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V MISHPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS C BRANCHED RIB CARTILAGE; LEFT- 8th DISTAL		3.4
	10 M	V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY		3.3
	11 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present SMALL		3.2
	12 M	V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage BASISPHEOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS		3.4
78	1 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY		3.4
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE				

10-APR-14

05R018

TABLE : IIB- 106

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
78	(CONTINUED)		
	2 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.2
		NO OBSERVED GROSS FINDINGS	
	3 F	NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- 5th	3.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V CERVICAL RIB; RIGHT; Cartilage present	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
		WITH BILATERAL HOLES	
	4 M	NO OBSERVED GROSS FINDINGS	2.9
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	3.3
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		4TH 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V WAVY RIB; RIGHT- SEVERAL	
		5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
	6 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	7 F	NO OBSERVED GROSS FINDINGS	3.3
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V CERVICAL RIB; LEFT; Cartilage not present	
	8 F	NO OBSERVED GROSS FINDINGS	3.1
		NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	
	1 F	NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	2 F	NO OBSERVED GROSS FINDINGS	3.0
		NO OBSERVED VISCERAL FINDINGS	
	3 M	NO OBSERVED GROSS FINDINGS	3.9
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		107
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 3 (300 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
79 (CONTINUED)						
	4 M	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.7			
		NO OBSERVED GROSS FINDINGS				
	5 F	NO OBSERVED VISCERAL FINDINGS	3.6			
		NO OBSERVED GROSS FINDINGS				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.5			
	6 F	NO OBSERVED GROSS FINDINGS				
		V DILATED RENAL PELVIS; left				
	7 F	NO OBSERVED GROSS FINDINGS	3.5			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	8 M	NO OBSERVED GROSS FINDINGS	4.0			
		NO OBSERVED VISCERAL FINDINGS				
	9 M	NO OBSERVED GROSS FINDINGS	3.7			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.1			
80						
	1 F	NO OBSERVED GROSS FINDINGS				
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.8			
	2 M	NO OBSERVED GROSS FINDINGS				
		NO OBSERVED VISCERAL FINDINGS				
	3 M	NO OBSERVED GROSS FINDINGS	3.9			
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL				
		4TH, 5TH				
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB- 108

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
80	(CONTINUED)		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
	4 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	5 F	NO OBSERVED GROSS FINDINGS	3.4
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present	
		RUDIMENTARY	
	6 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	7 F	NO OBSERVED GROSS FINDINGS	3.3
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present	
		RUDIMENTARY	
		C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL	
	8 M	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.5
		NO OBSERVED GROSS FINDINGS	
		NO OBSERVED VISCERAL FINDINGS	
	9 M	NO OBSERVED GROSS FINDINGS	3.0
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		3RD, 4TH, 5TH	
		5TH ADDITIONALLY INCOMPLETELY OSSIFIED	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
		WITH BILATERAL HOLES	
	10 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	11 M	NO OBSERVED GROSS FINDINGS	3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	12 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
81	1 F	NO OBSERVED GROSS FINDINGS	3.3

OBSERVATION CODES: V=Variation C=Cartilage

SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		109
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)						
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 3 (300 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
81 (CONTINUED)						
		V MISHPAPEN STERNEBRA; Unchanged cartilage- 4th				
		V BIPARTITE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th				
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	2 F	NO OBSERVED GROSS FINDINGS	3.6			
		NO OBSERVED VISCERAL FINDINGS				
	4 M	NO OBSERVED GROSS FINDINGS	3.8			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage				
		TEMPORAL, NASAL, PARIETAL AND INTERPARIETAL BONES				
	6 M	NO OBSERVED GROSS FINDINGS	4.3			
		NO OBSERVED VISCERAL FINDINGS				
	7 M	NO OBSERVED GROSS FINDINGS	3.9			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
	8 F	NO OBSERVED GROSS FINDINGS	3.6			
		NO OBSERVED VISCERAL FINDINGS				
	9 M	NO OBSERVED GROSS FINDINGS	3.8			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th				
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V SUPERNUMERARY RIB (14TH) ; BILATERAL; Cartilage not present				
		RUDIMENTARY				
		V WAVY RIB; BILATERAL- SEVERAL				
		5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH RIGHT SIDE				
		7TH, 8TH, 11TH, 12TH LEFT SIDE				
		V DUMBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 12th				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage				
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage				
	10 M	NO OBSERVED GROSS FINDINGS	4.0			
		NO OBSERVED VISCERAL FINDINGS				
		NO OBSERVED GROSS FINDINGS				
	11 M	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	3.3			
		C BIPARTITE PROCESSUS XIPHOIDEUS				
		V INCOMPLETE OSSIFICATION OF BASISPHENOID				
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage				
OBSERVATION CODES: V=Variation C=Cartilage						
SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB-

110

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
81	(CONTINUED)		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
82	1 F	NO OBSERVED GROSS FINDINGS	3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	2 M	NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
	3 M	NO OBSERVED GROSS FINDINGS	3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V WAVY RIB; RIGHT- SEVERAL	
		6TH, 7TH, 8TH, 11TH, 12TH	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	4 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY THORACIC VERTEBRA- 14th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	6 F	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	7 F	NO OBSERVED GROSS FINDINGS	3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL	
		5TH, 6TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
	8 F	NO OBSERVED GROSS FINDINGS	3.6

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

111

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
82	(CONTINUED)		
	9 F	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY THORACIC VERTEBRA- 14th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.3 3.4
83	1 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V WAVY RIB; BILATERAL- SEVERAL 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH RIGHT SIDE 11TH, 12TH LEFT SIDE V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage BASISPHENOID, TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS 3 M V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS 4 F 5 F NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS 6 M 7 M NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH	3.0 3.1 3.3

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14	05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)	TABLE : IIB-	112
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				
TEST GROUP 3 (300 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION		WEIGHT (G)
83	(CONTINUED)			
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF BASISPHENOID		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
	8 M	NO OBSERVED GROSS FINDINGS		3.4
		NO OBSERVED VISCERAL FINDINGS		
	9 M	NO OBSERVED GROSS FINDINGS		3.2
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF BASISPHENOID		
	10 M	NO OBSERVED GROSS FINDINGS		3.3
		NO OBSERVED VISCERAL FINDINGS		
	12 M	NO OBSERVED GROSS FINDINGS		3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage		
		BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES		
84	1 F	NO OBSERVED GROSS FINDINGS		2.3
		V INCOMPLETE OSSIFICATION OF TUBEROSITAS DELTOIDEA; BILATERAL; Cartilage present		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL		
		1ST,2ND		
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- SEVERAL		
		5TH,6TH		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V INCOMPLETE OSSIFICATION OF PUBIS; BILATERAL; Cartilage present		
		V INCOMPLETE OSSIFICATION OF ISCHIUM; BILATERAL; Cartilage present		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
		WITH BILATERAL HOLES		
	2 M	NO OBSERVED GROSS FINDINGS		3.7
		NO OBSERVED VISCERAL FINDINGS		
	3 M	NO OBSERVED GROSS FINDINGS		3.6
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL		
		4TH,5TH		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
OBSERVATION CODES: V=Variation C=Cartilage				
SEX CODE: M=MALE, F=FEMALE				

10-APR-14

05R018

TABLE : IIB- 113

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
84	(CONTINUED)		
	4 M	NO OBSERVED GROSS FINDINGS	3.1
		V DILATED RENAL PELVIS; left	
		V DILATED URETER; left	
	5 F	NO OBSERVED GROSS FINDINGS	3.4
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	6 F	NO OBSERVED GROSS FINDINGS	3.2
		NO OBSERVED VISCERAL FINDINGS	
	7 M	NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V DUMEBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 10th	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	8 F	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	10 M	NO OBSERVED GROSS FINDINGS	3.4
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
	11 F	NO OBSERVED GROSS FINDINGS	2.9
		NO OBSERVED VISCERAL FINDINGS	
	1 M	NO OBSERVED GROSS FINDINGS	3.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 1ST, 2ND	
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14	05R018	PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)	TABLE : IIB-	114
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS				
TEST GROUP 3 (300 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION		WEIGHT (G)
85	(CONTINUED)			
		V MISHPEN STERNEBRA; Unchanged cartilage- 4th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V INCOMPLETE OSSIFICATION OF TUBEROSITAS DELTOIDEA; BILATERAL; Cartilage present		
		V DUMBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- SEVERAL		
		8TH, 10TH		
		V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present		
		RUDIMENTARY		
		V INCOMPLETE OSSIFICATION OF PUBIS; BILATERAL; Cartilage present		
		V INCOMPLETE OSSIFICATION OF ISCHIUM; BILATERAL; Cartilage present		
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage		
		TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES		
	2 M	NO OBSERVED GROSS FINDINGS		3.9
		NO OBSERVED VISCERAL FINDINGS		
	4 F	NO OBSERVED GROSS FINDINGS		3.6
		V MISHPEN STERNEBRA; Unchanged cartilage- 5th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
	5 F	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		3.4
		NO OBSERVED GROSS FINDINGS		
		NO OBSERVED VISCERAL FINDINGS		
	6 M	NO OBSERVED GROSS FINDINGS		4.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		
		V WAVY RIB; RIGHT- SEVERAL		
		5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH		
		V INCOMPLETE OSSIFICATION OF BASISPHENOID		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
	7 M	NO OBSERVED GROSS FINDINGS		3.8
		NO OBSERVED VISCERAL FINDINGS		
	8 F	NO OBSERVED GROSS FINDINGS		3.5
		V MISHPEN STERNEBRA; Unchanged cartilage- 5th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V INCOMPLETE OSSIFICATION OF BASISPHENOID		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
	9 F	NO OBSERVED GROSS FINDINGS		3.9
		NO OBSERVED VISCERAL FINDINGS		
	10 M	NO OBSERVED GROSS FINDINGS		3.7
		V MISHPEN STERNEBRA; Unchanged cartilage- 5th		
OBSERVATION CODES: V=Variation C=Cartilage				
SEX CODE: M=MALE, F=FEMALE				

10-APR-14

05R018

TABLE : IIB-

115

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
85	(CONTINUED)		
	11 F	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS 3.5	3.5
86	2 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V WAVY RIB; BILATERAL- SEVERAL 7TH, 8TH, 9TH, 10TH, 11TH V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage BASISPHENOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present SMALL V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS U BLOOD COAGULUM AROUND PLACENTA V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH 3.6	3.5 3.5
	3 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present SMALL V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS U BLOOD COAGULUM AROUND PLACENTA V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH 3.7	3.7
	4 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present SMALL V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS U BLOOD COAGULUM AROUND PLACENTA V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH 3.7	3.7
	5 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present SMALL V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS U BLOOD COAGULUM AROUND PLACENTA V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH 3.8	3.8
	6 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present SMALL V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS U BLOOD COAGULUM AROUND PLACENTA V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH 3.9	3.9
	7 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present SMALL V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS U BLOOD COAGULUM AROUND PLACENTA V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH 3.6	3.6
	8 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present SMALL V SUPERNUMERARY RIB (14TH); LEFT; Cartilage present NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS U BLOOD COAGULUM AROUND PLACENTA V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH 3.6	3.6

OBSERVATION CODES: V=Variation C=Cartilage U=Unclassified
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

116

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
86	(CONTINUED)		
87	1 M	M MALPOSITIONED AND BIPARTITE STERNEBRA; Unchanged cartilage- 5th	3.3
		V NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
	2 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.7
		V NO OBSERVED GROSS FINDINGS	
		V NO OBSERVED VISCERAL FINDINGS	
	3 F	V NO OBSERVED GROSS FINDINGS	3.0
		V MISSAPEN STERNEBRA; Unchanged cartilage- 4th	
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V CERVICAL RIB; LEFT; Cartilage not present	
		V DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 2nd	
	4 M	M MISSAPEN BASISPHENOID	3.8
		V NO OBSERVED GROSS FINDINGS	
		V DILATED RENAL PELVIS; left	
	5 F	V NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	6 F	V NO OBSERVED GROSS FINDINGS	3.4
		V NO OBSERVED VISCERAL FINDINGS	
		V NO OBSERVED GROSS FINDINGS	
	8 M	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	3.3
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	9 M	V NO OBSERVED GROSS FINDINGS	3.2
		V NO OBSERVED VISCERAL FINDINGS	
	10 F	V NO OBSERVED GROSS FINDINGS	3.5
		V MISSAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
	11 M	V NO OBSERVED GROSS FINDINGS	3.1
		V NO OBSERVED VISCERAL FINDINGS	

OBSERVATION CODES: M=Malformation V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
88	1 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH C BIPARTITE PROCESSUS XIPHOIDEUS V WAVY RIB; BILATERAL- SEVERAL 8TH, 9TH, 10TH, 11TH, RIGHT SIDE 10TH, 11TH LEFT SIDE V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present V UNOSSIFIED STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 6TH V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 1ST, 3RD M MALPOSITIONED AND BIPARTITE STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH C CARTILAGINOUS PARTS OF RIBS DISPLACED; RIGHT- SEVERAL 1ST, 2ND, 3RD, 4TH, 5TH NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage present V MISSHAPEN SACRAL VERTEBRA 1ST SACRAL ARCH - LEFT SIDE, CARTILAGE PRESENT V INCOMPLETE OSSIFICATION OF BASISPHENOID NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.0 3.3 3.3
	3 M		3.3
	4 M		3.3
	5 F		3.2
	7 M		3.3
	8 F		3.6
	9 M		3.5
	10 M		3.2

OBSERVATION CODES: M=Malformation V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

05R018

TABLE : IIB-

118

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
89	1 F	NO OBSERVED GROSS FINDINGS M MISSAPEN TUBEROSITAS DELTOIDEA, LEFT V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH 5TH, 6TH	2.8
	2 M	C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage BASISPHEOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	3.3
	3 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	2.8
	4 F	C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF BASISPHEOID V INCOMPLETE OSSIFICATION OF PARIETAL, BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	2.7
	5 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH, 5TH	3.0
	6 M	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF PARIETAL, BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.1
	7 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 4th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	2.7
	8 F	C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF BASISPHEOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage WITH BILATERAL HOLES NO OBSERVED GROSS FINDINGS	3.1

OBSERVATION CODES: M=Malformation V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS				TABLE : IIB-	119
ORAL ADMINISTRATION (GAVAGE)					
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS					
TEST GROUP 3 (300 MG/KG BW/D)					
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)		

89	(CONTINUED)				
	9 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND,3RD,4TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH,6TH C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	2.9		
	10 M		3.0		
	11 F		2.6		
	12 F		3.0		
	13 F		2.7		
	14 F		2.8		
90	1 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- SEVERAL 10TH,13TH V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY V WAVY RIB; BILATERAL- SEVERAL 10TH,11TH V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.6		
OBSERVATION CODES: V=Variation C=Cartilage					
SEX CODE: M=MALE, F=FEMALE					

10-APR-14

05R018

TABLE : IIB-120

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
90	(CONTINUED)		
	2 F	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	3 M	NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V WAVY RIB; BILATERAL- SEVERAL	
		5TH,6TH,7TH,8TH,9TH,10TH,11TH,12TH	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	4 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	6 F	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
	7 M	NO OBSERVED GROSS FINDINGS	3.9
		NO OBSERVED VISCERAL FINDINGS	
	8 F	NO OBSERVED GROSS FINDINGS	3.7
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
	9 M	NO OBSERVED GROSS FINDINGS	3.7
		V SHORT INNOMINATE	
	10 M	NO OBSERVED GROSS FINDINGS	3.6
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		4TH,5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V WAVY RIB; BILATERAL- SEVERAL	
		4TH,5TH,6TH,7TH,8TH,9TH,10TH,11TH,12TH,13TH	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

121

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
90	(CONTINUED)		
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage NEARLY ALL SKULL BONES M SHORTENED HUMERUS	
91	1 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY	3.4
	2 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.3
	3 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY	3.3
	4 M	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.6
	5 M	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present RUDIMENTARY	3.5
	6 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.5
	7 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.4
	8 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.8
	9 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	3.7

OBSERVATION CODES: M=Malformation V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-122

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
91	(CONTINUED)		
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		SMALL	
		V INCOMPLETE OSSIFICATION OF BASISPHENOID	
10 F		NO OBSERVED GROSS FINDINGS	3.8
11 M		NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	3.8
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
12 F		NO OBSERVED GROSS FINDINGS	3.0
		NO OBSERVED VISCERAL FINDINGS	
92			
		NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V DUMBLELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 12th	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
2 M		NO OBSERVED GROSS FINDINGS	3.6
		DILATED RENAL PELVIS; left	
3 M		NO OBSERVED GROSS FINDINGS	3.7
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
4 M		NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
5 F		NO OBSERVED GROSS FINDINGS	3.4
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
6 F		NO OBSERVED GROSS FINDINGS	3.6
		NO OBSERVED VISCERAL FINDINGS	
7 M		NO OBSERVED GROSS FINDINGS	3.8
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
8 M		NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
9 M		NO OBSERVED GROSS FINDINGS	3.7

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		TABLE : IIB-		123
05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE)		
TEST GROUP 3 (300 MG/KG BW/D)		INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS		
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)	

92	(CONTINUED)	V MISSHPEN STERNEBRA; Unchanged cartilage- 4th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
		NO OBSERVED GROSS FINDINGS		
		NO OBSERVED VISCERAL FINDINGS		
		NO OBSERVED GROSS FINDINGS		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
10 M			3.7	
11 F				
	NO OBSERVED GROSS FINDINGS			
	NO OBSERVED VISCERAL FINDINGS			
	NO OBSERVED GROSS FINDINGS			
	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th			
	C BIPARTITE PROCESSUS XIPHOIDEUS			
	V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage			
	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage			
	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage			
	NO OBSERVED GROSS FINDINGS			
	NO OBSERVED VISCERAL FINDINGS			
	NO OBSERVED GROSS FINDINGS			
	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 5TH, 6TH			
	C BIPARTITE PROCESSUS XIPHOIDEUS			
V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present				
RUDIMENTARY				
94	1 F		3.6	
		NO OBSERVED GROSS FINDINGS		
		V MISSHPEN STERNEBRA; Unchanged cartilage- 4th		
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- 5th		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
	2 F	NO OBSERVED GROSS FINDINGS		3.4
		NO OBSERVED VISCERAL FINDINGS		
	3 F	NO OBSERVED GROSS FINDINGS		3.0
		V MISSHPEN STERNEBRA; Unchanged cartilage- 4th		
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present		
		RUDIMENTARY		
		V CERVICAL RIB; LEFT; Cartilage not present		
	4 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		3.5
		NO OBSERVED GROSS FINDINGS		
		NO OBSERVED VISCERAL FINDINGS		
	5 M	NO OBSERVED GROSS FINDINGS		3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 5TH, 6TH		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		RUDIMENTARY		

OBSERVATION CODES: V=Variation C=Cartilage				
SEX CODE: M=MALE, F=FEMALE				

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS				TABLE : IIB-	124
ORAL ADMINISTRATION (GAVAGE)					
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS					
TEST GROUP 3 (300 MG/KG BW/D)					
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)		
94 (CONTINUED)					
	6 F	V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	3.4		
	7 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.3		
	8 F	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS	3.0		
	10 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH	3.4		
95					
	1 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	3.1		
	2 F	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.2		
	3 M	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS	3.5		
	4 M	V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.2		
	5 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V UNOSSIFIED STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS	3.0		
	6 F	V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.7		
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE					

10-APR-14		TABLE : IIB-		125
05R018		PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS		
TEST GROUP 3 (300 MG/KG BW/D)				
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)	

95	(CONTINUED)			
	7 F	NO OBSERVED GROSS FINDINGS		3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
	8 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage		3.6
		NO OBSERVED GROSS FINDINGS		
	9 M	NO OBSERVED VISCERAL FINDINGS		3.7
		NO OBSERVED GROSS FINDINGS		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present		
		RUDIMENTARY		
	10 M	NO OBSERVED GROSS FINDINGS		3.3
		NO OBSERVED VISCERAL FINDINGS		
	11 F	NO OBSERVED GROSS FINDINGS		3.1
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL		
		4TH, 5TH		
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- 6th		
		C BIPARTITE PROCESSUS XIPHOIDEUS		
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage		
		HYOID, TEMPORAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL		
		BONES		

96	1 M	NO OBSERVED GROSS FINDINGS		3.3
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		V WAVY RIB; BILATERAL- SEVERAL		
		3RD, 4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
		TEMPORAL, FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL		
		BONES		
	2 F	NO OBSERVED GROSS FINDINGS		2.9
		NO OBSERVED VISCERAL FINDINGS		
	3 M	NO OBSERVED GROSS FINDINGS		3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th		
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present		
		SMALL		
		V WAVY RIB; BILATERAL- SEVERAL		
		5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH RIGHT SIDE		
		5TH, 6TH, 7TH, 8TH LEFT SIDE		
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage		
		WITH BILATERAL HOLES		

OBSERVATION CODES: V=Variation C=Cartilage				
SEX CODE: M=MALE, F=FEMALE				

10-APR-14

05R018

TABLE : IIB-126

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS
TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
96	(CONTINUED)		
	4 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	5 F	NO OBSERVED GROSS FINDINGS	3.4
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
	6 M	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.4
		NO OBSERVED GROSS FINDINGS	
	7 F	NO OBSERVED VISCERAL FINDINGS	3.4
		NO OBSERVED GROSS FINDINGS	
	8 F	V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	3.1
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		NO OBSERVED GROSS FINDINGS	
	9 M	NO OBSERVED VISCERAL FINDINGS	3.5
		NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
	10 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.5
		NO OBSERVED GROSS FINDINGS	
	11 F	NO OBSERVED VISCERAL FINDINGS	3.4
		NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V WAVY RIB; RIGHT- SEVERAL	
		6TH, 7TH, 8TH, 9TH, 10TH	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
97	1 F	NO OBSERVED GROSS FINDINGS	3.5
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
	2 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	3.6
		NO OBSERVED GROSS FINDINGS	
	3 M	NO OBSERVED VISCERAL FINDINGS	3.8
		NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V DUMBBELL OSSIFICATION OF THORACIC CENTRUM; Unchanged cartilage- 10th	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-

127

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)
INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
97	(CONTINUED)		
	5 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS	3.6
	6 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 4TH, 5TH	3.4
	7 F	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY	3.2
	8 F	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS	3.5
	9 M	V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS	3.8
	10 F	V DILATED RENAL PELVIS; bilateral NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present RUDIMENTARY	3.3
	12 M	V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.5
	13 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V SUPERNUMERARY RIB (14TH); RIGHT; Cartilage not present RUDIMENTARY	3.0
	14 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.1
98	1 M	U BLOOD COAGULUM AROUND PLACENTA	3.5

OBSERVATION CODES: V=Variation C=Cartilage U=Unclassified
SEX CODE: M=MALE, F=FEMALE

10-APR-14

05R018

TABLE : IIB-128

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
98	(CONTINUED)		
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); LEFT; Cartilage not present	
		RUDIMENTARY	
	2 M	V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	4.0
		NO OBSERVED GROSS FINDINGS	
	3 M	NO OBSERVED VISCERAL FINDINGS	3.7
		NO OBSERVED GROSS FINDINGS	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	4 F	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	5 M	NO OBSERVED GROSS FINDINGS	3.6
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
	6 F	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
99	1 F	NO OBSERVED GROSS FINDINGS	3.0
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL	
		5TH, 6TH	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	2 M	NO OBSERVED GROSS FINDINGS	3.4
		NO OBSERVED VISCERAL FINDINGS	
	3 M	NO OBSERVED GROSS FINDINGS	3.3
		V MISSHAPEN STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	4 M	NO OBSERVED GROSS FINDINGS	3.3
		V DILATED RENAL PELVIS; left	
	5 F	NO OBSERVED GROSS FINDINGS	3.7
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL	
		5TH, 6TH	
		V SUPERNUMERARY RIB (14TH); BILATERAL; Cartilage not present	
		RUDIMENTARY	

OBSERVATION CODES: V=Variation C=Cartilage
SEX CODE: M=MALE, F=FEMALE

10-APR-14		05R018		TABLE : IIB-		129
PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS ORAL ADMINISTRATION (GAVAGE) INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS						
TEST GROUP 3 (300 MG/KG BW/D)						
FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)			
99 (CONTINUED)						
	6 F	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS	3.5			
	7 M	NO OBSERVED VISCERAL FINDINGS NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF BASISPHENOID V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS	3.5			
	8 M	V DILATED RENAL PELVIS; left NO OBSERVED GROSS FINDINGS	3.5			
	9 M	V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 2ND, 3RD, 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.9			
	10 M	V MISSHAPEN STERNEBRA; Unchanged cartilage- 2nd V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th V WAVY RIB; BILATERAL- SEVERAL 6TH, 7TH, 8TH, 9TH	3.5			
	11 M	V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage HYOID, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	3.4			
100						
	1 F	NO OBSERVED GROSS FINDINGS V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- SEVERAL 5TH, 6TH	3.3			
	2 M	NO OBSERVED GROSS FINDINGS NO OBSERVED VISCERAL FINDINGS	3.3			
	3 F	NO OBSERVED GROSS FINDINGS V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL 3RD, 4TH, 5TH V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th C BIPARTITE PROCESSUS XIPHOIDEUS V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	3.1			
OBSERVATION CODES: V=Variation C=Cartilage SEX CODE: M=MALE, F=FEMALE						

10-APR-14

05R018

TABLE : IIB-130

PROJECT NO. 30R0066/05R018: PRENATAL TOXICITY STUDY IN RATS
ORAL ADMINISTRATION (GAVAGE)

INDIVIDUAL FETAL EXTERNAL, VISCERAL, SKELETAL AND/OR CARTILAGE OBSERVATIONS

TEST GROUP 3 (300 MG/KG BW/D)

FEMALE#	FETUS# (SEX)	OBSERVATION	WEIGHT (G)
100	(CONTINUED)		
	4 M	V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	3.4
		NO OBSERVED GROSS FINDINGS	
	5 F	NO OBSERVED VISCERAL FINDINGS	
		NO OBSERVED GROSS FINDINGS	3.2
		V UNILATERAL OSSIFICATION OF STERNEBRA; Unchanged cartilage- 5th	
		ADDITIONALLY INCOMPLETELY OSSIFIED	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF PARIETAL; BILATERAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF INTERPARIETAL; Unchanged cartilage	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
	6 M	NO OBSERVED GROSS FINDINGS	3.5
		NO OBSERVED VISCERAL FINDINGS	
	7 F	NO OBSERVED GROSS FINDINGS	3.2
		V UNOSSIFIED STERNEBRA; Unchanged cartilage- 5th	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V WAVY RIB; BILATERAL- SEVERAL	
		4TH, 5TH, 6TH, 7TH, 8TH, 9TH, 10TH, 11TH, 12TH	
		V INCOMPLETE OSSIFICATION OF SKULL; Unchanged cartilage	
		FRONTAL, PARIETAL, INTERPARIETAL AND SUPRAOCCIPITAL BONES	
	8 M	NO OBSERVED GROSS FINDINGS	3.3
		NO OBSERVED VISCERAL FINDINGS	
	9 F	NO OBSERVED GROSS FINDINGS	3.0
		V MISSHAPEN STERNEBRA; Unchanged cartilage- SEVERAL	
		3RD, 4TH	
		V INCOMPLETE OSSIFICATION OF STERNEBRA; Unchanged cartilage- 6th	
		C BIPARTITE PROCESSUS XIPHOIDEUS	
		V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL; Unchanged cartilage	
		WITH BILATERAL HOLES	
	10 M	NO OBSERVED GROSS FINDINGS	3.3
		V DILATED RENAL PELVIS; left	
		V DILATED URETER; left	

OBSERVATION CODES: V=Variation C=Cartilage

SEX CODE: M=MALE, F=FEMALE

STUDY TITLE

Report

DHDPS

Prenatal Developmental Toxicity Study
in Wistar Rats
Oral Administration (Gavage)

TEST FACILITY PROJECT IDENTIFICATION

Project No.: 30R0066/05R018

PART III OF III
(SUPPLEMENT)

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CONTENTS OF PART III (SUPPLEMENT)

Title Page
Table of contents

1. Analysis of the test substance

- Final Report: Characterization of “4,4’-Dihydroxydiphenylsulfon”

2. Analyses of the test substance preparations

- DHDPS: Stability Analysis in 1% Carboxymethylcellulose in Drinking Water
- Homogeneity and Concentration Control Analysis of DHDPS in 1% Carboxymethylcellulose in Drinking water

3. Historical control data

- Tables

1. Analysis of the test substance

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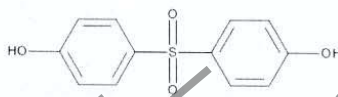
Competence Center Analytics

Final Report

Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)



Page 1 of 11

Test item	4,4'-Dihydroxydiphenylsulfon
Chemical identity	4,4'-Sulfonyldiphenol (DHDPS)
Chemical structure	
Batch identification	69611767J0
Date of production (test item)	Nov 28, 2011
Origin of test item	[REDACTED]
PSN	05/0066-4
CAS no.	80-09-1
Sponsor	[REDACTED]
Date of receipt of order	Dec 07, 2011
Date of receipt of test item	Dec 15, 2011
Testing facility	Competence Center Analytics, BASF SE, D-67056 Ludwigshafen
Study director	[REDACTED]
Storage cond. test item	Room temperature, moisture protection
Test period	Jan 13, 2012 – Feb 28, 2012
Storage of records	GLP archives, Competence Center Analytics
Storage of sample of test item	Archives, Competence Center Analytics

Competence Center Analytics

Final Report
Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)

Page 2 of 11

Summary of results

The test item was characterized by spectroscopic and chromatographic methods.

¹H-NMR spectroscopy confirmed the structure of the test item. Furthermore, the test item was imaged by its HPLC fingerprint by reversed-phase chromatography. Two peaks with area fractions >0.1% were present. The main component with 99.3% and 99.5% and a by-product with 0.30% and 0.34% at a wavelength of 230 and 250 nm, respectively.

Quantitative ¹H-NMR spectroscopy using the internal standard method yielded a mean purity of the test item of 99.4 g/100 g.

1 Appearance and Homogeneity

Method	Visual Inspection
Result	The test item was a fine white powder. It was obviously homogeneous.
Date of test	Jan 13, 2012
Head of laboratory	[REDACTED]

2 Identity via ¹H-NMR-Spectroscopy

Method	¹ H-NMR spectroscopy
Apparatus	Bruker DPX 401
Reagents	Solvent: DMSO-d ₆ (Euriso-top) Reference standard: Tetramethylsilane TMS (Cambridge Isotope)
Sample preparation	An adequate mass of the test item was filled into a sample tube and dissolved in DMSO-d ₆ containing TMS.
Test parameters	Measuring frequency = 400 MHz, measuring temperature = 30 °C; further parameters see ¹ H-NMR spectrum page 6
Result	The ¹ H-NMR spectrum shows the expected signals for the given structure (see page 6).
Date of test	Feb 23, 2012
Head of laboratory	[REDACTED]

Competence Center Analytics

Final Report
Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)

Page 3 of 11

3 Content of Main Component and By-Products

Method ¹H-NMR spectroscopy with internal standard

Apparatus Bruker DPX 401

Reagents
Solvent: DMSO-d₆ (Euriso-top)
Reference standard: Tetramethylsilane TMS (Cambridge Isotope)
Internal standard: 1,3,5-Trimethoxybenzene (Sigma Aldrich)
Purity: 100 % (for calc.)
Mol. weight: 168.19 g/mol

Sample preparation About 11-15 mg test item and about 10-15 mg of internal standard were weighed to the nearest 0.01 mg into sample tubes and dissolved in DMSO-d₆ containing TMS as reference standard.

Test parameters Measuring frequency = 400 MHz, measuring temperature = 30 °C; further parameters see at ¹H-NMR spectrum page 7

Result For quantitation, triplicate measurements were carried out. Evaluation was performed by using three protons/molecule of the internal standard (at ~ 6.1 ppm) and 4 protons/molecule of 4,4'-dihydroxydiphenylsulfone (at ~ 7.7 ppm). Exemplary spectrum see page 7.

Content of 4,4'-dihydroxydiphenylsulfone

Det.	test item		internal standard		Resulting mass fraction [g/100g]
	weight [mg]	peak intensity [area units]	weight [mg]	peak intensity [area units]	
1	11.12	400.00	10.40	420.14	99.4
2	11.70	400.00	15.09	578.85	99.5
3	14.40	400.00	10.29	320.86	99.4
mean					99.4

By-products were not detected.

Date of test Feb 28, 2012

Head of laboratory

Competence Center Analytics

Final Report
Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)

Page 4 of 11

4 HPLC Fingerprint

Method Reversed-phase high pressure liquid chromatography with UV detection at two wavelengths and area percent evaluation

Apparatus Automated HPLC system, equipped with an autosampler and a UV/Vis detector, connected to an electronical data processing system

Reagents Acetonitrile (Fluka)
Demineralized water obtained from a Milli-Q system

Sample preparation Approximately 30 mg of test item were weighed into a 25 mL volumetric flask and filled up to the mark with acetonitrile. Afterwards, a 2/25 dilution was carried out with a mixture of acetonitrile/water (1/1 v/v).

Test parameters

Column: LiChrospher 100, RP-18 ec, 5 µm, 125 x 3 mm (Macherey Nagel)

Mobile phase A: Acetonitrile/water = 10/90 (V/V)

Mobile phase B: Acetonitrile/water = 90/10 (V/V)

Injection volume: 20 µL

Flow rate: 0.7 mL/min

Oven temperature: 25 °C

UV detection: λ = 230 nm, λ = 250 nm

Gradient elution:

time [min]	0	25	30	31	46
A [%]	85	30	30	85	new
B [%]	15	70	70	15	injection

Result

The HPLC fingerprint shows two peaks with area fractions >0.1%. The peak area of the main component corresponds to 99.3 % at 230 nm and 99.5 % at 250 nm. A by-product with 0.30% and 0.34% was detected at 230 and 250 nm, respectively. These values are the means of two determinations. For details see pages 9-10.

For representative chromatograms of blank run and test item at both wavelengths see pages 8-10.

Date of test Jan 18, 2012 – Jan 19, 2012

Head of laboratory

May 21, 2012
Date

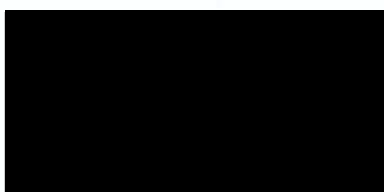
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Final Report
Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)

Page 5 of 11

GLP Compliance Statement

This study was conducted in accordance with the OECD Principles of Good Laboratory Practice and the GLP Principles of the German "Chemikaliengesetz" (Chemicals Act).



May 21, 2012
Date

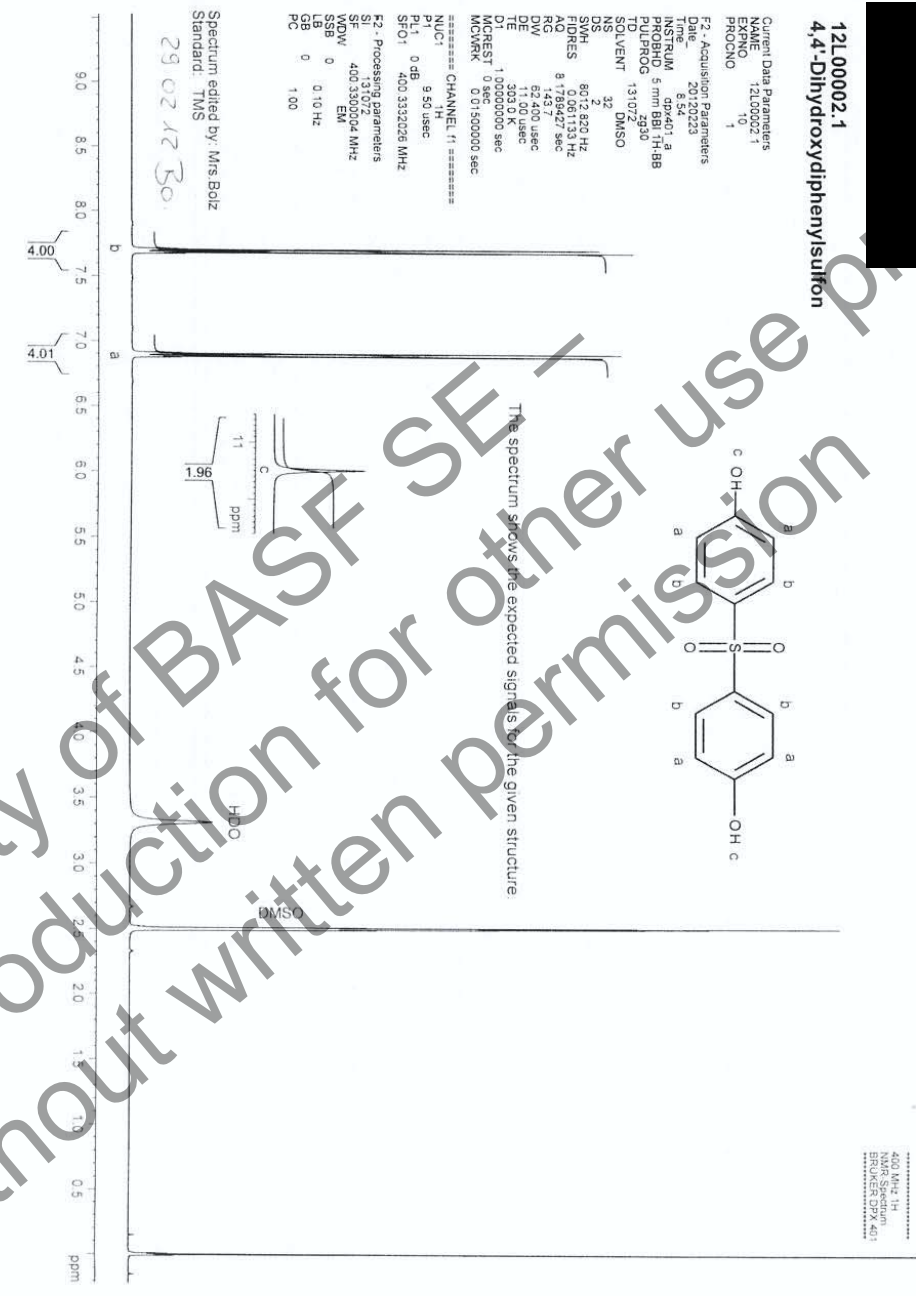
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Final Report
Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)

Page 6 of 11

Identity of test item by ¹H-NMR spectroscopy

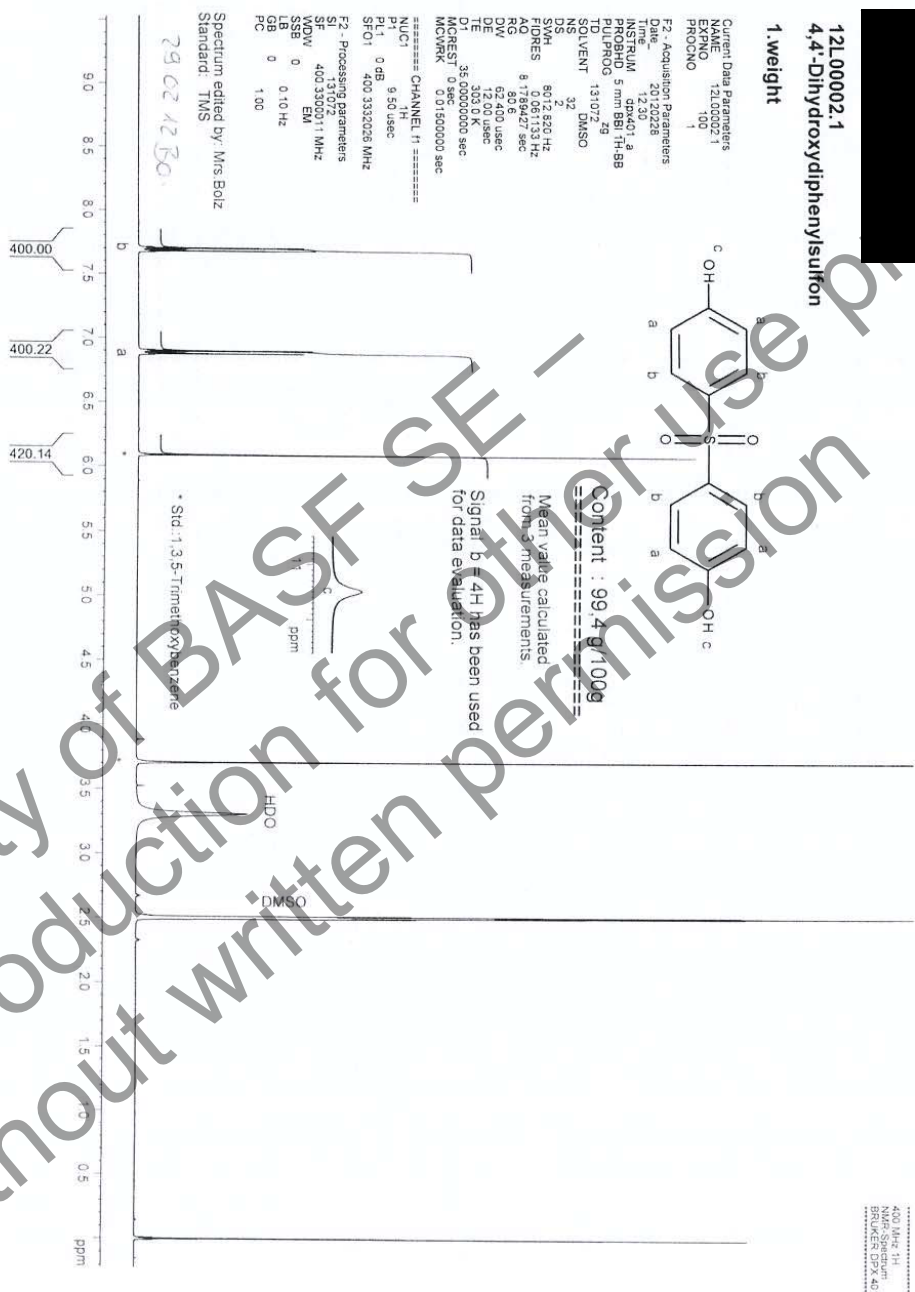


Competence Center Analytics

Final Report
Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)

Page 7 of 11

Quantitative ¹H-NMR spectrum of the test item



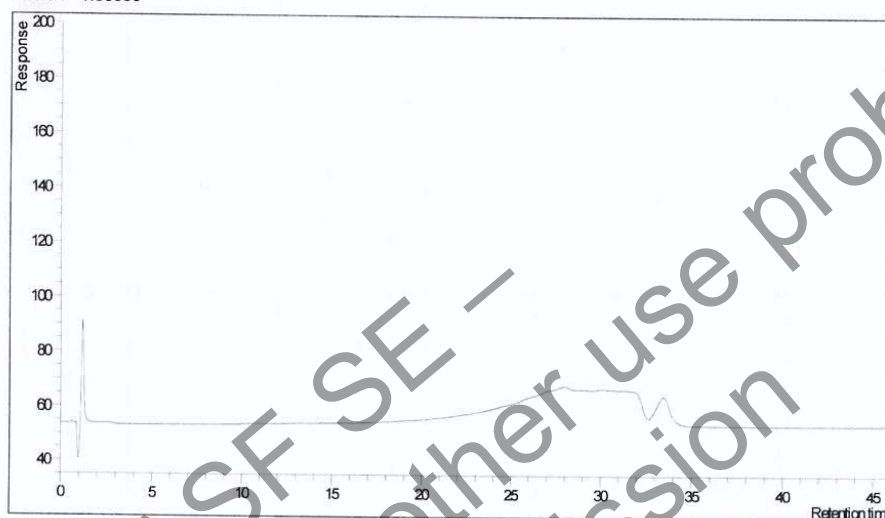
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Final Report
Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)

Page 8 of 11

Blank run of HPLC-fingerprint at $\lambda = 230$ nm

Sample : Blindlauf (Acetonitril - 230 nm)
Conc : 1.00000



Blank run of HPLC-fingerprint at 250 nm

Sample : Blindlauf (Acetonitril - 250 nm)
Conc : 1.00000



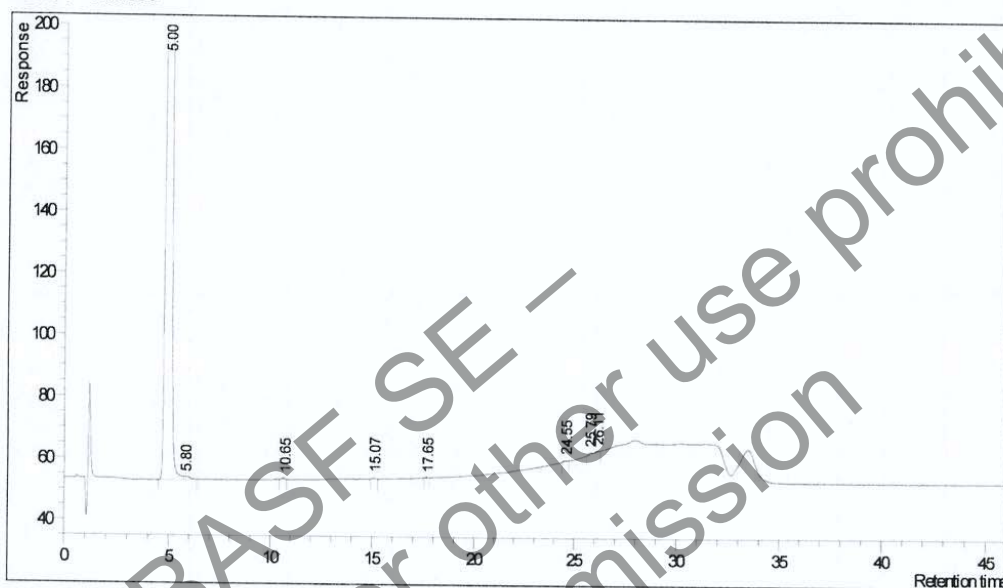
Competence Center Analytics

Final Report
Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)

Page 9 of 11

HPLC-fingerprint of test item at 230 nm

Sample : 12L00002 (230 nm) - A
 Conc : 1.00000



Named Peaks					
RT [min]	Height [mV]	Area [mVs]	Area%	Peak Name	
5.000	416.364	6773.860	99.363		BL
5.800	0.953	18.976	0.278		BVU
10.653	0.521	5.218	0.077		BBU
15.073	0.367	3.285	0.048		BBU
17.653	0.235	1.862	0.027		BBU
24.547	0.366	3.884	0.057		BBU
25.793	0.557	6.629	0.097		BVU
26.113	0.332	3.556	0.052		BVU
Sum	419.695	6817.270	99.999		

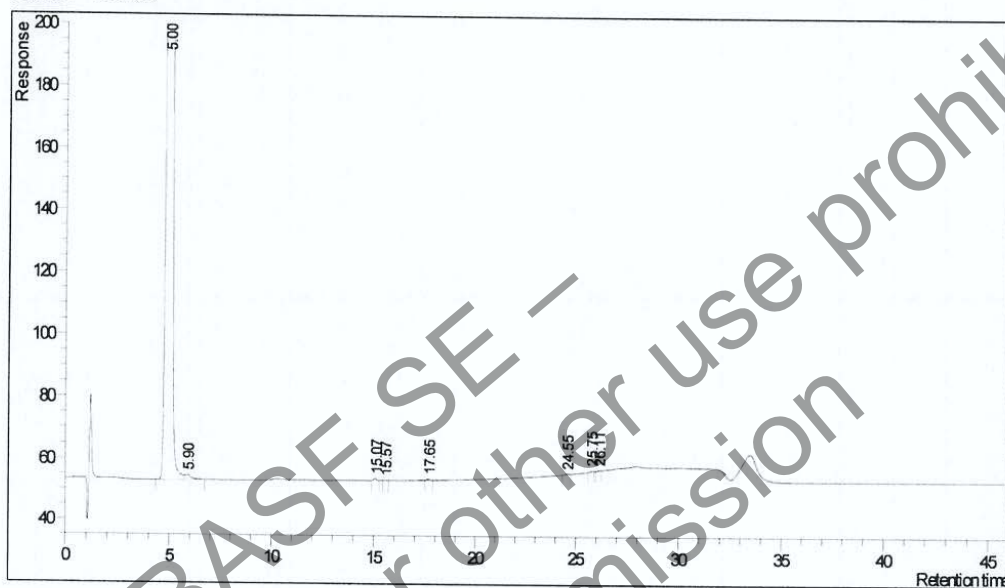
Competence Center Analytics

Final Report
Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)

Page 10 of 11

HPLC-fingerprint of test item at 250 nm

Sample : 12L00002 (250 nm) - A
 Conc : 1.00000



Named Peaks					
RT [min]	Height [mV]	Area [mVs]	Area%	Peak Name	BL
5.000	572.847	9341.446	99.478		BL
5.900	1.531	34.598	0.368		VBV
15.067	0.593	5.153	0.055		BBU
15.573	0.084	0.676	0.007		BBU
17.653	0.348	3.203	0.034		BBU
24.553	0.230	2.925	0.031		BBU
25.747	0.150	1.594	0.017		BBU
26.107	0.106	0.841	0.009		BBU
Sum	575.889	9390.436	99.999		BBU

Competence Center Analytics

Final Report
Characterization of "4,4'-Dihydroxydiphenylsulfon"
Study No. 12L00002 (confidential)

Page 11 of 11

Statement of the Quality Assurance Unit

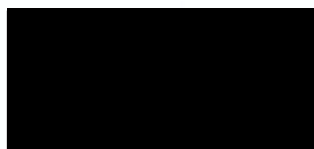
The Quality Assurance Unit inspects the laboratories of the department Competence Center Analytics in regular intervals. Besides these general inspections we inspected the following items of this study in accordance with the OECD Principles of Good Laboratory Practice and the GLP Principles of the German "Chemikaliengesetz" (Chemicals Act). Findings are reported to study director and to management.

Verification of study plan: Jan 10, 2012

Inspection of	Date of inspection	Reported to study director and to management
Conduct of study: HPLC fingerprint	Jan 18, 2012	Jan 18, 2012
Raw data:	May 16, 2012	May 21, 2012
Final report:	May 16, 2012	May 21, 2012

The final report reflects the raw data.

Ludwigshafen



May 21, 2012
Date

2. Analyses of the test substance preparations

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STUDY TITLE

ANALYTICAL REPORT

DHDPS

Stability Analysis in

1% Carboxymethylcellulose in Drinking Water

AUTHOR(S)**STUDY COMPLETION DATE**

29 August 2013

TEST FACILITYBASF SE
Experimental Toxicology and Ecology
67056 Ludwigshafen, Germany**TEST FACILITY PROJECT IDENTIFICATION**

Project No.: 01Y0066/05Y009

SPONSORBASF SE
67056 Ludwigshafen, Germany

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GLP COMPLIANCE STATEMENT

This study was conducted in accordance with the OECD Principles of Good Laboratory Practice and the GLP Principles of the German "Chemikaliengesetz" (Chemicals Act) which meet the United States Environmental Protection Agency Good Laboratory Practice Standards [40 CFR Part 160 (FIFRA) and Part 792 (TSCA)], with the exception that recognized differences exist between the GLP Principles/Standards of OECD and the Principles/Standards of FIFRA and TSCA.

Study Director

Typed name of Study Director:

Typed name of Laboratory:

BASF SE
Experimental Toxicology and Ecology
67056 Ludwigshafen
Germany

Date: 29 Aug. 2013

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SIGNATURE PAGE

Study Director:

[Redacted Signature]

29 Aug. 2013

Management:

[Redacted Signature]

27 Aug 2013

STATEMENT OF THE QUALITY ASSURANCE UNIT

The Quality Assurance Unit (QAU) inspected the study and reported any inspection results to the Study Director and to Management.

The final report reflects the raw data.

Phase of study	Date of inspection (mm-dd-yyyy)	Reported to Study Director and to Management (mm-dd-yyyy)
Study Plan:	10-09-2012	10-09-2012
Conduct of study:	10-12-2012	10-12-2012
Report:	08-08-2013	08-08-2013

Ludwigshafen, 29 August 2013

GLP CERTIFICATE (FROM THE COMPETENT AUTHORITY)



Gute Laborpraxis / Good Laboratory Practice

GLP-Bescheinigung / Statement of GLP Compliance
(gem. / according to § 19 Abs. 1 Chemikaliengesetz)

Eine GLP-Inspektion zur Überwachung und der Einhaltung der GLP-Grundsätze gemäß Chemikaliengesetz bzw. Richtlinie 2004/9/EG wurde durchgeführt in:

Assessment of conformity with GLP according to Chemikaliengesetz and Directive 2004/9/EC at::

Prüfeinrichtung / Test facility

BASF SE
Experimentelle Toxikologie und Ökologie
67056 Ludwigshafen

BASF SE
Experimental Toxicology and Ecology
67056 Ludwigshafen, Germany

Prüfung nach Kategorien / Areas of Expertise
(gem. / according ChemVwV-GLP Nr. 53/OECD guidance)
1,2,3,4,5,8,9

Kat. 9 – Biochemische und pathologische Untersuchungen zu Wirkmechanismen /
Biochemical and pathological examinations concerning mode of action

Datum der Inspektion / Date of Inspection
(Tag/Monat/Jahr / day.month.year)
19.05.2009 & 06. bis 08.07.2009

Die genannte Prüfeinrichtung befindet sich im nationalen GLP-Überwachungsverfahren und wird regelmäßig auf Einhaltung der GLP-Grundsätze überwacht.

The above mentioned test facility is included in the national GLP Compliance Programme and is inspected on a regular basis.

Auf der Grundlage des Inspektionsberichtes wird hiermit bestätigt, dass in dieser Prüfeinrichtung die oben genannten Prüfungen unter Einhaltung der GLP-Grundsätze durchgeführt werden können. Eine erneute behördliche Überprüfung der Einhaltung der GLP-Grundsätze durch die Prüfeinrichtung ist so rechtzeitig zu beantragen, dass die Folgeinspektion spätestens vier Jahre nach dem Beginn der o.g. Inspektion stattfinden kann. Ohne diesen Antrag wird die Prüfeinrichtung nach Ablauf der Frist aus dem deutschen GLP-Überwachungsprogramm genommen und diese GLP-Bescheinigung verliert ihre Gültigkeit.

Based on the inspection report it can be confirmed, that the test facility is able to conduct the aforementioned studies in compliance with the Principles of GLP. Verification of the compliance of the test facility with the Principles of the GLP has to be applied for in time to allow for a follow-up inspection to take place within four years after commencing the above mentioned inspection. Elapsing this term, the test facility will be taken out of the German GLP-Monitoring Programme and this GLP Certificate becomes invalid.

Unterschrift, Datum / Signature, Date



Dr. Pia Hirsch - stellv. Präsidentin -
(Name und Funktion der verantwortlichen Person / name and function of responsible person)



Siegel

Landesamt für Umwelt, Wasserwirtschaft und Gewerbeaufsicht
Kaiser-Friedrich-Straße 7
55116 Mainz

(Name und Adresse der GLP-Überwachungsbehörde /
Name and address of the GLP Monitoring Authority)

Landesamt für
Umwelt, Wasserwirtschaft
und Gewerbeaufsicht

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CONTENTS

TITLE PAGE

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GLP COMPLIANCE STATEMENT

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SIGNATURE PAGE

STATEMENT OF THE QUALITY ASSURANCE UNIT

GLP CERTIFICATE (FROM THE COMPETENT AUTHORITY)

CONTENTS

1. INTRODUCTION
2. RETENTION OF RECORDS
3. TIME SCHEDULE
4. MATERIAL AND METHODS
 - 4.1. TEST ITEM
 - 4.2. SAMPLE DATA
 - 4.3. TEST SUBSTANCE PREPARATION
 - 4.4. SAMPLE PREPARATION AND ANALYSIS
 - 4.5. LIST OF DEVIATIONS
 - 4.5.1. LIST OF DEVIATIONS FROM THE CONTROL PROCEDURE
5. RESULTS AND DISCUSSION
 - 5.1. ANALYSIS OF STABILITY
 - 5.2. DISCUSSION
- FIGURES
6. APPENDIX
 - 6.1. CONTROL PROCEDURE 05/0066_01-01

1. INTRODUCTION

In the context of toxicological studies the stability of the test substance DHDPS in the vehicle 1% carboxymethylcellulose in drinking water has to be verified. The results of these analyses are reported and discussed.

2. RETENTION OF RECORDS

GLP-relevant records and materials are stored at BASF SE for at least the period of time specified in the GLP principles. Details concerning responsibilities or locations of archiving can be seen from the respective SOPs and from the raw data.

3. TIME SCHEDULE

Study initiation date:	09 October 2012
Experimental starting date:	12 October 2012
Experimental completion date:	19 October 2012

4. MATERIAL AND METHODS

4.1. TEST ITEM

The analyses of the test item (= test substance) were carried out at the Competence Center Analytics of BASF SE, Ludwigshafen, Germany.

Name of test substance:	DHDPS
Test substance No.:	05/0066-4
Batch identification:	69611767J0
CAS No.:	80-09-1
Purity:	(1) 99.3 and 99.5 %, (HPLC) (2) 99.4 g/100 g, (1H-NMR) (according to the project number 12L00002)
Homogeneity:	Given
Storage stability:	stable until: 28 May 2013 The stability of the test substance under storage conditions over the test period was guaranteed by the sponsor, and the sponsor holds this responsibility.
Additional Test Substance Information	
Date of production:	28 Nov 2011
Physical state/ Appearance:	Solid / white
Storage conditions	Room temperature

4.2. SAMPLE DATA

Sponsor: Dr. Buesen; Ms. Pabst
Vehicle: 1% carboxymethylcellulose in drinking water
Target concentration: 0.05 g/100 mL
Duration of the stability test period: 7 days
Storage conditions of the samples during the stability period: Refrigerator

4.3. TEST SUBSTANCE PREPARATION

51.2 mg of the test substance were dissolved in 5 mL acetone. 0.5 mL of this solution were transferred into 100 mL volumetric flasks. After acetone evaporation at room temperature, 10 mL 1% carboxymethylcellulose in drinking water were added. For each time point a sample was prepared. The final nominal concentration was 0.0512 g / 100 mL.

4.4. SAMPLE PREPARATION AND ANALYSIS

The sample preparation and analysis of the test substance was carried out according to the valid control procedure 05/0066_01-01.

A detailed description of the control procedure is given in the appendix of this report.

4.5. LIST OF DEVIATIONS

4.5.1. LIST OF DEVIATIONS FROM THE CONTROL PROCEDURE

There were no deviations from the described control procedure 05/0066_01-01.

5. RESULTS AND DISCUSSION

5.1. ANALYSIS OF STABILITY

The results obtained for the stability of the test substance in 1% carboxymethylcellulose in drinking water are summarized in the following table.

All calculated values in the table are rounded. Calculations were performed with a full set of decimal places.

Nominal concentration [g/100 mL]	Time after starting	Concentration found [g/100 mL]	Nominal concentration (%)
0.0512	0h	0.055	107.3
0.0512	4h	0.053	104.9
0.0512	4d	0.055	108.5

The stability samples from 0 hours until 4 days were stored at room temperature

Nominal concentration [g/100 mL]	Time after starting	Concentration found [g/100 mL]	Nominal concentration (%)
0.0512	7d	0.054	106.6

The stability sample for the duration of 7 days was stored in the refrigerator.

5.2. DISCUSSION

Based on the analytical results it is concluded, that DHDPS is stable in 1% carboxymethylcellulose in drinking water over a period of 4 days at room temperature and 7 days in the refrigerator.

All determined concentrations were in the range of 90 % - 110 % of the nominal concentration.

FIGURES

Figure 1: Chromatogram of matrix solution (measured on 19 Oct 2012)

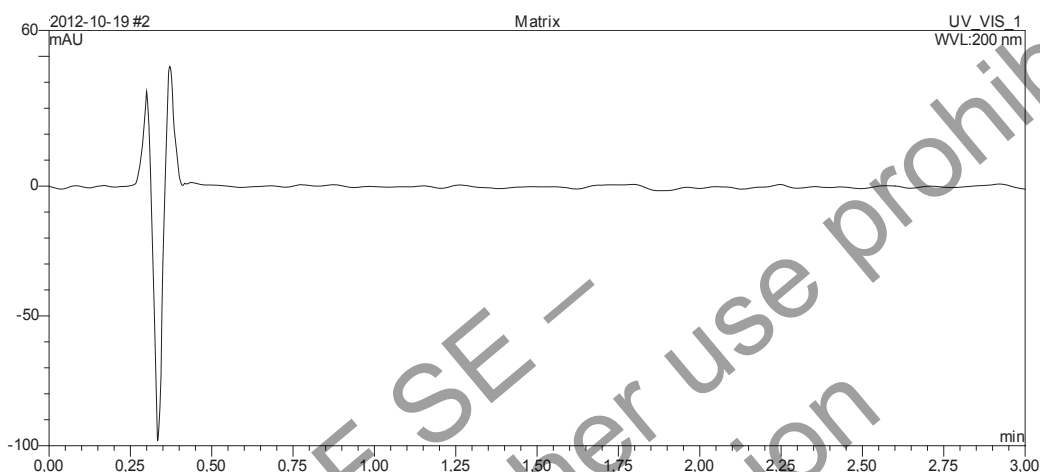


Figure 2: Chromatogram of calibration solution 1 (2.044 mg/100 mL, measured on 19 Oct 2012)

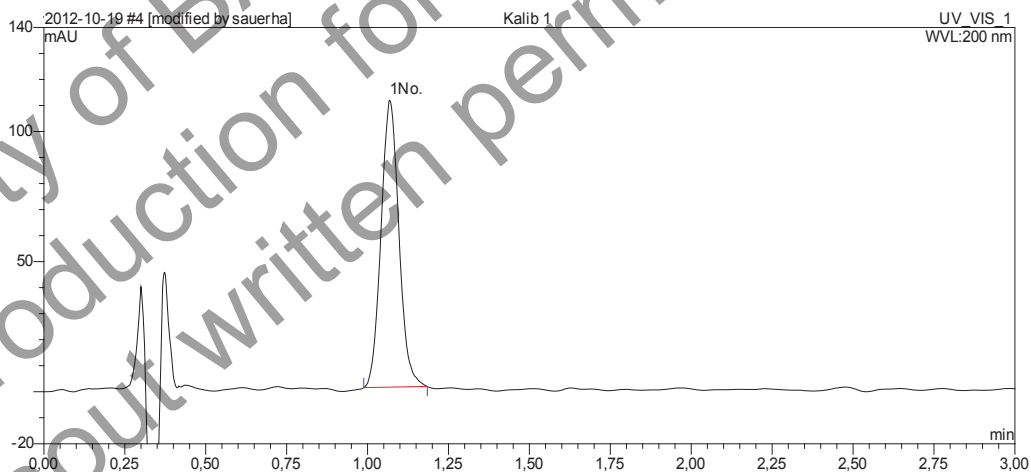


Figure 3: Chromatogram sample solution day 7 (measured on 19 Oct 2012)

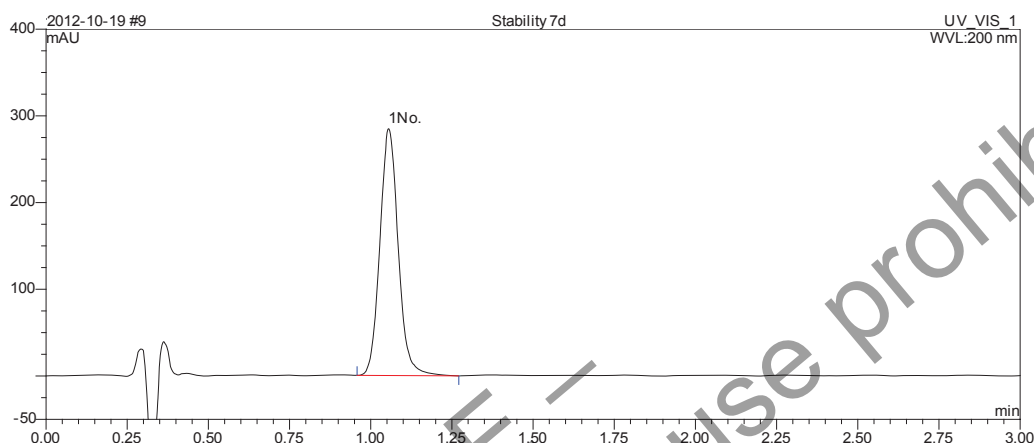
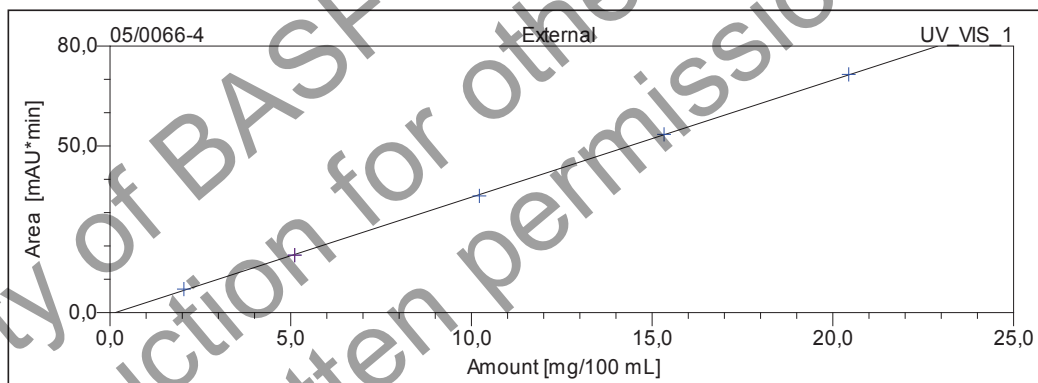


Figure 4: Calibration curve (19 Oct 2012, Concentration range 2.044 – 20.44 mg/100 mL)



6. APPENDIX

6.1. CONTROL PROCEDURE 05/0066_01-01

BASF SE
Test Facility
Experimental Toxicology and Ecology / Analytical Chemistry

**CONTROL TEST**

Test substance number: 05/0066	No.: 05/0066_01-01
Name of test substance: DHDPs	Effective from: 09 Oct 2012
Control procedure: Content (LC) / Carboxymethylcellulose(CMC) in drinking water	Page 1 of 5

Technique	HPLC
System:	Waters alliance 2487 with auto sampler, Dionex Chromeleon-Software (Dionex), or equivalent system
Column:	Length: 100 mm Inner diameter: 4.6 mm
Stationary Phase:	Chromolith Performance RP 18e, Merck or equivalent
Mobile Phase A:	1000 mL acetonitrile are mixed with 1 mL formic acid (HCOOH)
Mobile Phase B:	1000 mL water are mixed with 1 mL formic acid (HCOOH)
Isocratic:	
Mobile Phase A 20 %	Mobile Phase B 80 %
Injection volume:	10 µL
Flow rate:	5 mL/min
Detection:	200 nm
Column temperature:	Ambient
Run time:	Approx. 3 min

BASF SE
Test Facility
Experimental Toxicology and Ecology / Analytical Chemistry



CONTROL TEST

Test substance number: 05/0066	No.: 05/0066_01-01
Name of test substance: DHDPs	Effective from: 09 Oct 2012
Control procedure: Content (LC) / Carboxymethylcellulose(CMC) in drinking water	Page 2 of 5

Sample solution: Samples are diluted completely with methanol using appropriate volumetric flasks to obtain sample solutions with test substance concentrations that match the calibration range.
If required, all dilutions are sonicated for 5 minutes to ensure a complete dissolution of the test substance.

The samples are filtered (cellulose filter, 0.2 µm) prior HPLC analysis.

Annotation: If the amount of test substance in the sample solution is outside the calibration range (calibration solutions 1 – 5), an adequate dilution step with matrix solution has to be performed to match the described concentration range.

Matrix solution: The preparation of the matrix solution has to be performed according to the procedure described for sample solution preparation

Stock solution: Approx. 50 mg test substance are dissolved to a final volume of 100 mL with methanol (50 mg/100 mL)

Calibration solution 1: 1.0 mL stock solution are diluted with matrix solution to 25 mL (2 mg/100 mL)

Calibration solution 2: 1.0 mL stock solution are diluted with matrix solution to 10 mL (5 mg/100 mL)

Calibration solution 3: 1.0 mL stock solution are diluted with matrix solution to 5 mL (10 mg/100 mL)

Calibration solution 4: 1.5 mL stock solution are diluted with matrix solution to 5 mL (15 mg/100 mL)

Calibration solution 5: 2.0 mL stock solution are diluted with matrix solution to 5 mL (20 mg/100 mL)

System-suitability solution: System-suitability solution is prepared with a second independent weighing according to calibration solution 3 (10 mg/100 mL)

Procedure After conditioning the HPLC system, sample solutions, matrix solution, calibration solutions and system-suitability solution are injected according to the sequence described in the raw data. All solutions are injected at least once.

BASF SE
Test Facility
Experimental Toxicology and Ecology / Analytical Chemistry


The Chemical Company

CONTROL TEST

Test substance number: 05/0066	No.: 05/0066_01-01
Name of test substance: DHDPs	Effective from: 09 Oct 2012
Control procedure: Content (LC) / Carboxymethylcellulose(CMC) in drinking water	Page 3 of 5

Retention time:

Test substance : 4,4'-Dihydroxydiphenylsulfon:
Approx. 1 min

System suitability:

The calculated content of the system-suitability solution has to be in the range from 95 % to 105 %.

The coefficient of determination (R^2) has to be ≥ 0.990 . If the correlation coefficient (R) is used, this value has to be ≥ 0.995 .

Calculation:

The concentration control measurements are based on external calibration (calibration solutions 1 – 5).

The calculation of the content is performed electronically. (e.g. Dionex Chromeleon – Software, Microsoft Excel). Basic formulas for calculations are described below (e.g. Dionex Chromeleon – Software)

Formulas:

Calibration curve

$$Y = a \cdot x + b$$

a = slope of calibration curve
b = intercept

Analysed concentration (C_A)

$$C_A = \frac{(Y-b) \cdot V \cdot d}{a \cdot w}$$

or

$$C_A = \frac{(Y-b)}{a} \cdot \frac{V \cdot d}{v}$$

w = weight sample
V = final sample volume
d = dilution factor

v = volume sample
V = final sample volume
d = dilution factor

Analysed concentration (C_A)

BASF SE
Test Facility
Experimental Toxicology and Ecology / Analytical Chemistry

CONTROL TEST

Test substance number: 05/0066

No.: 05/0066_01-01

Name of test substance: DHDPs

Effective from: 09 Oct 2012

Control procedure: Content (LC) / Carboxymethylcellulose(CMC) in drinking water

Page 4 of 5

Figure 1.1: Example chromatogram matrix solution (08 Oct 2012, Project no.: 01Y0066/05Y009) for illustration

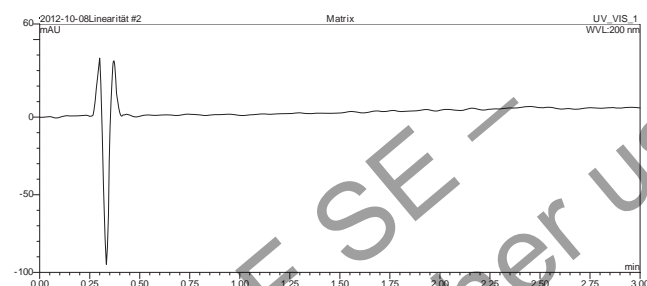
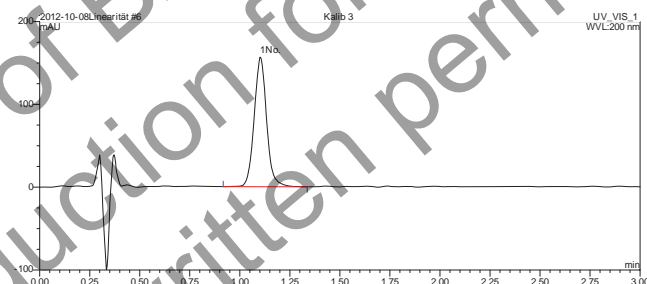


Figure 1.2: Example chromatogram calibration solution (08 Oct 2012, Project no.: 01Y0066/05Y009) for illustration



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CONTROL TEST

Test substance number: 05/0066

No.: 05/0066_01-01

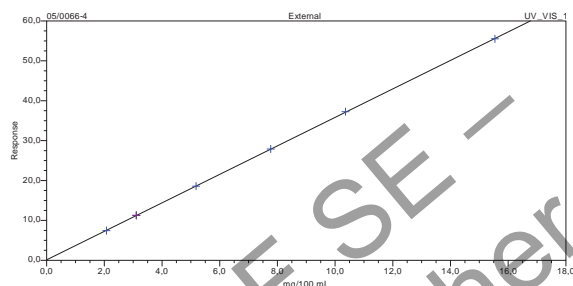
Name of test substance: DHDPs

Effective from: 09 Oct 2012

Control procedure: Content (LC) / Carboxymethylcellulose(CMC) in drinking water

Page 5 of 5

Figure 1.3 Example calibration curve (08 Oct 2012, Project no.: 01Y0066/05Y009) for illustration



Homogeneity and Concentration Control Analysis of DHDPS in 1% Carboxymethylcellulose in Drinking water

1. PROJECT AND TEST SUBSTANCE INFORMATION

Project No.: 30R0066/05R018

Test item (= test substance): DHDPS

Batch No.: 69611767J0

2. SAMPLE DATA

2.1. HOMOGENEITY AND CONCENTRATION CONTROL ANALYSIS

Vehicle: 1% carboxymethylcellulose in drinking water

Storage conditions of the
samples until analysis: Freezer

3. MATERIAL AND METHODS

3.1. SAMPLE PREPARATION AND ANALYSIS

The sample preparation and analysis of the test substance was carried out according to the valid control procedure 05/0066_01-01.

3.2. LIST OF DEVIATIONS

3.2.1. List of deviations from the control procedure

There was no deviation from the described control procedure 05/0066_01-01.

4. RESULTS AND DISCUSSION

4.1. HOMOGENEITY AND CONCENTRATION CONTROL ANALYSIS

The results obtained for the homogeneity and concentration control analysis of DHDPS in 1% carboxymethylcellulose in drinking water are summarized in the following table:

All calculated values in the table are rounded. Calculations were performed with a full set of decimal places

Date of sample preparation: 16 Sep 2013
Date of sampling: 16 Sep 2013
Date of receipt of sample in analytical laboratory: 16 Sep 2013
Starting date of analytical determination: 16 Sep 2013

Name	Amount	Nominal Conc	Nominal Conc	Mean	RSD
	mg/100 mL	mg/100 mL	%	%	%
Sample 03	284.591	300	94.9%		
Sample 04	279.672	300	93.2%		
Sample 05	284.346	300	94.8%	94.3%	1.0%
Sample 06	941.714	1000	94.2%		
Sample 07	2938.028	3000	97.9%		
Sample 08	3023.634	3000	100.8%		
Sample 09	2912.861	3000	97.1%	98.6%	2.0%

Considering the low relative standard deviation in the homogeneity analysis, it can be concluded that DHDPS was distributed homogeneously in 1% carboxymethylcellulose in drinking water.

The mean values (samples 03 – 05 and samples 07 – 09) and single value (sample 06) of DHDPS in 1% carboxymethylcellulose in drinking water were found to be in the range of 90 % – 110 % of the nominal concentrations.

These results demonstrated the correctness of the concentrations of DHDPS in 1% carboxymethylcellulose in drinking water.

Figures of the calibration curve and examples of chromatograms will follow within this report.

Figure 1: Chromatogram of matrix solution (measured on 16 Sep 2013)

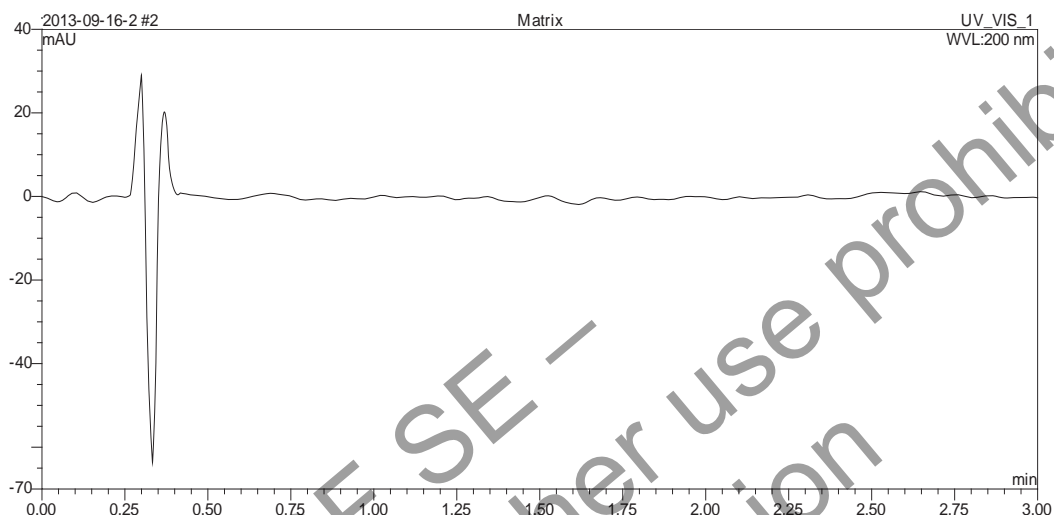


Figure 2: Chromatogram of calibration solution 1 (2.04 mg/100 mL, measured on 16 Sep 2013)

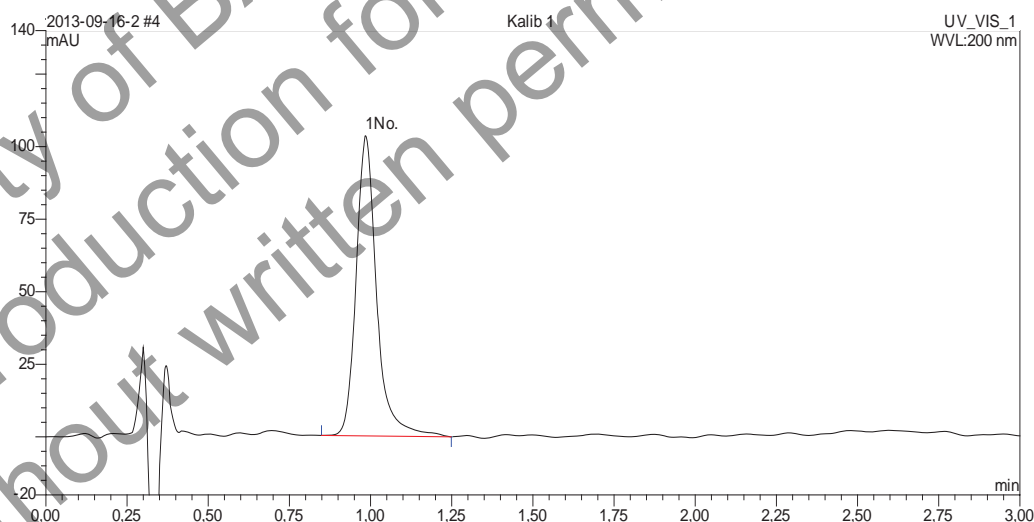


Figure 3: Chromatogram of sample 07 (measured on 16 Sep 2013)

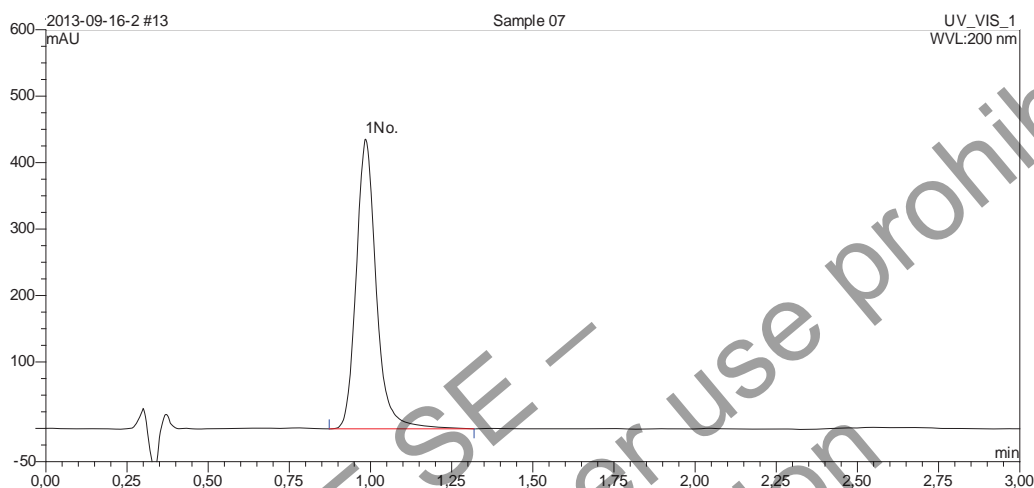
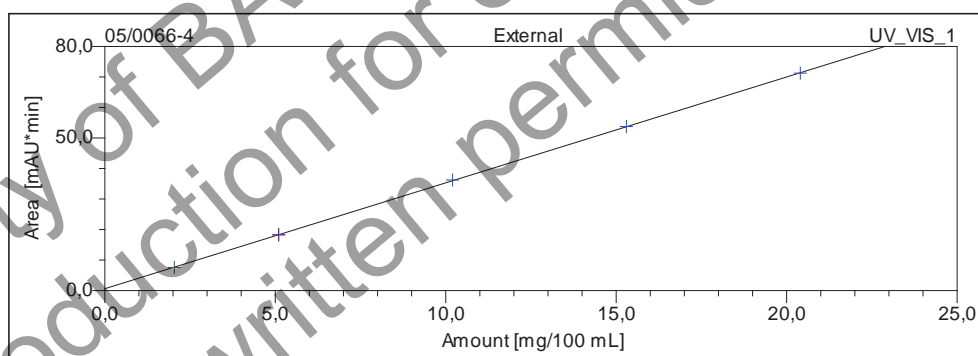


Figure 4: Calibration curve (measured on 16 Sep 2013, Concentration range 2.04 – 20.4 mg/100 mL)



5. APPENDIX

5.1. CONTROL PROCEDURE 05/0066_01-01

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The Chemical Company

CONTROL TEST

Test substance number: 05/0066

No.: 05/0066_01-01

Name of test substance: DHDPs

Effective from: 09 Oct 2012

Control procedure: Content (LC) / Carboxymethylcellulose(CMC) in drinking water

Page 1 of 5

Technique	HPLC
System:	Waters alliance 2487 with auto sampler, Dionex Chromeleon-Software (Dionex), or equivalent system
Column:	Length: 100 mm Inner diameter: 4.6 mm
Stationary Phase:	Chromolith Performance RP 18e, Merck or equivalent
Mobile Phase A:	1000 mL acetonitrile are mixed with 1 mL formic acid (HCOOH)
Mobile Phase B:	1000 mL water are mixed with 1 mL formic acid (HCOOH)
Isocratic:	
Mobile Phase A 20 %	Mobile Phase B 80 %
Injection volume:	10 µL
Flow rate:	5 mL/min
Detection:	200 nm
Column temperature:	Ambient
Run time:	Approx. 3 min

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Experimental Toxicology and Ecology / Analytical Chemistry

 **BASF**
The Chemical Company

CONTROL TEST

Test substance number: 05/0066	No.: 05/0066_01-01
Name of test substance: DHDPs	Effective from: 09 Oct 2012
Control procedure: Content (LC) / Carboxymethylcellulose(CMC) in drinking water	Page 2 of 5

Sample solution: Samples are diluted completely with methanol using appropriate volumetric flasks to obtain sample solutions with test substance concentrations that match the calibration range. If required, all dilutions are sonicated for 5 minutes to ensure a complete dissolution of the test substance.

The samples are filtered (cellulose filter, 0.2 µm) prior HPLC analysis.

Annotation: If the amount of test substance in the sample solution is outside the calibration range (calibration solutions 1 – 5), an adequate dilution step with matrix solution has to be performed to match the described concentration range.

Matrix solution: The preparation of the matrix solution has to be performed according to the procedure described for sample solution preparation

Stock solution: Approx. 50 mg test substance are dissolved to a final volume of 100 mL with methanol (50 mg/100 mL)

Calibration solution 1: 1.0 mL stock solution are diluted with matrix solution to 25 mL (2 mg/100 mL)

Calibration solution 2: 1.0 mL stock solution are diluted with matrix solution to 10 mL (5 mg/100 mL)

Calibration solution 3: 1.0 mL stock solution are diluted with matrix solution to 5 mL (10 mg/100 mL)

Calibration solution 4: 1.5 mL stock solution are diluted with matrix solution to 5 mL (15 mg/100 mL)

Calibration solution 5: 2.0 mL stock solution are diluted with matrix solution to 5 mL (20 mg/100 mL)

System-suitability solution: System-suitability solution is prepared with a second independent weighing according to calibration solution 3 (10 mg/100 mL)

Procedure After conditioning the HPLC system, sample solutions, matrix solution, calibration solutions and system-suitability solution are injected according to the sequence described in the raw data. All solutions are injected at least once.

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Experimental Toxicology and Ecology / Analytical Chemistry

 **BASF**
The Chemical Company

CONTROL TEST

Test substance number: 05/0066

No.: 05/0066_01-01

Name of test substance: DHDPs

Effective from: 09 Oct 2012

Control procedure: Content (LC) / Carboxymethylcellulose(CMC) in drinking water

Page 3 of 5

Retention time:

Test substance : 4,4'-Dihydroxydiphenylsulfon:
Approx. 1 min

System suitability:

The calculated content of the system-suitability solution has to be in the range from 95 % to 105 %.

The coefficient of determination (R^2) has to be ≥ 0.990 . If the correlation coefficient (R) is used, this value has to be ≥ 0.995 .

Calculation:

The concentration control measurements are based on external calibration (calibration solutions 1 – 5).

The calculation of the content is performed electronically. (e.g. Dionex Chromeleon – Software, Microsoft Excel). Basic formulas for calculations are described below (e.g. Dionex Chromeleon – Software)

Formulae:

Calibration curve

$$Y = a \cdot x + b$$

a = slope of calibration curve

b = intercept

Analysed concentration (C_A)

$$C_A = \frac{(Y - b) \cdot V \cdot d}{a \cdot w}$$

or

$$C_A = \frac{(Y - b) \cdot V \cdot d}{a \cdot v}$$

w = weight sample

V = final sample volume

d = dilution factor

v = volume sample

V = final sample volume

d = dilution factor

Analysed concentration (C_A)

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BASF
The Chemical Company

CONTROL TEST

Test substance number: 05/0066

No.: 05/0066_01-01

Name of test substance: DHDPS

Effective from: 09 Oct 2012

Control procedure: Content (LC) / Carboxymethylcellulose(CMC) in drinking water

Page 4 of 5

Figure 1.1: Example chromatogram matrix solution (08 Oct 2012, Project no.: 01Y0066/05Y009) for illustration

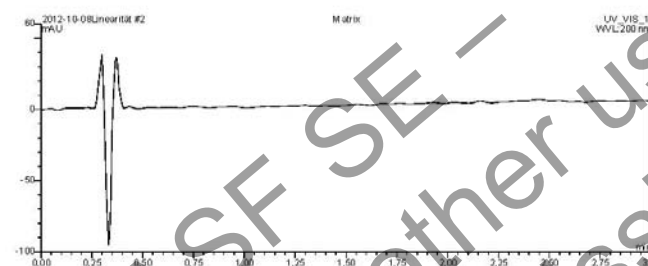
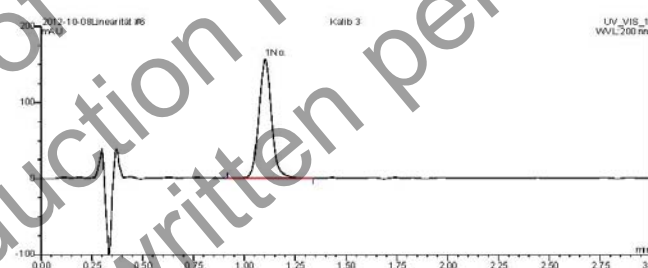


Figure 1.2: Example chromatogram calibration solution (08 Oct 2012, Project no.: 01Y0066/05Y009) for illustration



BASF SE
Test Facility
Experimental Toxicology and Ecology / Analytical Chemistry

BASF
The Chemical Company

CONTROL TEST

Test substance number: 05/0066

No.: 05/0066_01-01

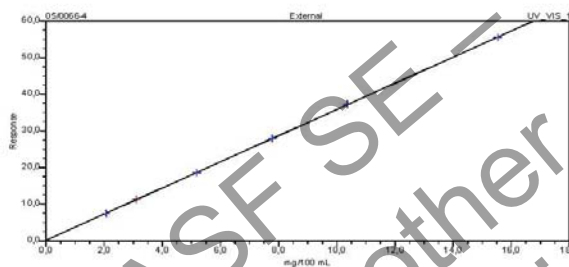
Name of test substance: DHDPs

Effective from: 09 Oct 2012

Control procedure: Content (LC) / Carboxymethylcellulose(CMC) in drinking water

Page 5 of 5

Figure 1.3 Example calibration curve (08 Oct 2012, Project no.: 01Y0066/05Y009) for illustration



3. Historical control data

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15-APR-14

TABLE :
01-JAN-09 to 31-MAR-14

HISTORICAL CONTROL DATA

SPECIES: RAT
STRAIN: WISTAR
SUPPLIER: CHARLES RIVER
FILENAME: 05R018

Study No./Control Group	Start Date	End Date	Route of Administration
04R038 1	9-JUL-13		GAVAGE
06I005 1	19-OCT-10		INHALATION
06R010 1	22-MAR-11		GAVAGE
06I07 1	15-APR-09		INHALATION
07R048 1	29-JAN-13		GAVAGE
08018 1	3-MAR-09		GAVAGE
08099 1	19-JAN-10		GAVAGE
09R118 1	6-MAR-13		GAVAGE
09R128 1	3-SEP-13		GAVAGE
10R134 1	23-NOV-11		GAVAGE
10R157 1	2-OCT-12		GAVAGE
11R069 1	7-AUG-12		GAVAGE
11R087 1	8-NOV-11		GAVAGE
11R159 1	3-MAY-12		GAVAGE
11R199 1	13-NOV-12		GAVAGE
11R223 1	25-JUN-13		GAVAGE
12R041 1	4-SEP-12		GAVAGE
12R079 1	19-FEB-13		GAVAGE
13R015 1	27-AUG-13		GAVAGE
99R009 1	15-FEB-11		GAVAGE

15-APR-14

SPECIES: RAT
STRAIN: WISTAR
SUPPLIER: CHARLES RIVER

TABLE :
01-JAN-09 to 31-MAR-14

HISTORICAL CONTROL DATA
MEAN MATERNAL BODY WEIGHTS DURING GESTATION GRAMS

	NO. OF ANIMALS	MEAN	RANGE OF ACTUAL VALUES	95 % 2.5%	SPREAD 97.5%
DAY 0	465	165.6	137.6 192.3	143.0	188.1
DAY 1	489	177.8	150.6 213.4	153.7	201.9
DAY 3	489	187.2	159.1 238.4	161.5	212.8
DAY 6	489	198.2	159.1 257.9	170.7	225.8
DAY 8	489	205.2	168.2 264.1	177.0	233.4
DAY 10	489	214.3	179.1 266.2	184.8	243.9
DAY 13	489	229.7	184.0 286.5	197.9	261.6
DAY 15	489	240.5	182.5 294.1	207.0	274.0
DAY 17	489	257.5	184.2 311.9	220.4	294.5
DAY 19	489	279.0	184.4 341.1	236.9	321.2
DAY 20	489	291.1	184.3 351.9	244.8	337.4

15-APR-14

TABLE :

SPECIES RAT
STRAIN WISTAR
SUPPLIER CHARLES RIVER

HISTORICAL CONTROL DATA
REPRODUCTION DATA

DATES: 01-JAN-09 - 31-MAR-14

RANGE (per study)

Females Mated	N	500			
Pregnant	N	489			
	%	98			
Aborted	N	0			
Premature Births	N	0			
Dams with Viable Fetuses	N	486			
Dams with all Resorptions	N	3			
Female Mortality	N	0			
	%	0.0			
Pregnant at C-section	N	489			
	%	98			
Corpora Lutea	N	5575	10.7	12.5	
	MEAN	11.4			
	S.D.	1.56			
Implantation Sites	N	5231	9.6	11.7	
	MEAN	10.7			
	S.D.	1.94			
Preimplantation Loss	%	6.2	1.4	15.8	
	S.D.	12.39			
Postimplantation Loss	%	7.1	4.7	12.0	
	S.D.	11.80			
Dams with Resorptions	N	228			
	%	47			
Resorptions	N	489			
	MEAN	0.7	0.5	1.0	
	S.D.	0.95			
	MEAN %	7.1	4.7	12.0	
	S.D.	11.80			
Dead Fetuses	N	0			
Viable Fetuses	N	4888			
Litter Size	MEAN	10.0	8.7	10.9	
	S.D.	2.09			
Viable Male Fetuses	N	2449			
	%	50			

15-APR-14

TABLE :

SPECIES RAT
STRAIN WISTAR
SUPPLIER CHARLES RIVER

HISTORICAL CONTROL DATA
PLACENTA WEIGHTS -- GRAMS

DATES: 01-JAN-09 - 31-MAR-14

	NO. OF LITTERS	MEAN	RANGE OF ACTUAL VALUES	95 % 2.5 %	SPREAD 97.5 %
MALES	482	0.48	0.34 1.09	0.34	0.62
FEMALES	483	0.46	0.31 0.85	0.32	0.60
BOTH	486	0.47	0.35 0.99	0.33	0.61

15-APR-14

SPECIES		RAT		HISTORICAL CONTROL DATA		TABLE :		DATES: 01-JAN-09 - 31-MAR-14	
STRAIN		WISTAR		MEAN FETAL WEIGHTS		-- GRAMS			
SUPPLIER		CHARLES RIVER							
				NO. OF LITTERS		MEAN		RANGE OF ACTUAL VALUES	
								95 % SPREAD	
								2.5% 97.5%	
MALES				482		3.6		2.6 5.3	
FEMALES				483		3.4		2.5 4.9	
BOTH				486		3.5		2.5 5.1	

15-APR-14

TABLE :

SPECIES		HISTORICAL CONTROL DATA									
RAT		FETAL EXTERNAL MALFORMATIONS									
STRAIN WISTAR		SUPPLIER CHARLES RIVER									
DATES: 01-JAN-09 - 31-MAR-14											
		FETUSES		%RANGE (per study)		LITTERS		%RANGE (per study)		AFFECTED FETUSES /LITTER	
		N	%	%	%	N	%	%	%	MEAN	%
NUMBER EVALUATED		4888				486					
Live		4888									
Dead		0									
M FETUS WITH MULTIPLE EXTERNAL MALFORMATIONS		1	0.02	0.0	0.4	1	0.2	0.0	4.0	0.0	0.4
M ANASARCA		1	0.02	0.0	0.4	1	0.2	0.0	4.2	0.0	0.4
M UMBILICAL HERNIA		1	0.02	0.0	0.4	1	0.2	0.0	4.2	0.0	0.3
M MANDIBULAR MICROGNATHIA		1	0.02	0.0	0.4	1	0.2	0.0	4.0	0.0	0.5
M AGLOSSIA		1	0.02	0.0	0.4	1	0.2	0.0	4.0	0.0	0.5
M MALROTATED LIMB		1	0.02	0.0	0.4	1	0.2	0.0	4.2	0.0	0.4
M SHORT TAIL		1	0.02	0.0	0.4	1	0.2	0.0	4.0	0.0	0.4
TOTAL FETAL EXTERNAL MALFORMATIONS		6	0.1	0.0	0.8	6	1.2	0.0	8.3	0.1	0.7

OBSERVATION CODE: M=Malformation

15-APR-14

TABLE :

SPECIES		RAT		HISTORICAL CONTROL DATA		FETAL EXTERNAL VARIATIONS		FETUSES		LITTERS		%RANGE (per study)		%RANGE (per study)		%MEAN (per study)		AFFECTED FETUSES /LITTER	
STRAIN		WISTAR																	
SUPPLIER		CHARLES RIVER																	
DATES:		01-JAN-09 - 31-MAR-14																	
NUMBER EVALUATED		4888		4888		0		1		0.2		0.0		0.0		4.0		0.0	
Live		4888		4888		0		1		0.2		0.0		0.0		4.0		0.0	
Dead		0		0		0		1		0.2		0.0		0.0		4.0		0.0	
V LIMB HYPERFLEXION		1		0.02		0.0		0.4		0.2		0.0		0.0		4.0		0.0	
V LIMB HYPEREXTENSION		3		0.06		0.0		0.8		0.6		0.0		0.1		8.0		0.0	
TOTAL FETAL EXTERNAL VARIATIONS		4		0.08		0.0		0.8		0.8		0.0		0.1		8.0		0.0	

OBSERVATION CODE: V=Variation

15-APR-14

TABLE :

SPECIES		HISTORICAL CONTROL DATA									
STRAIN		FETAL EXTERNAL UNCLASSIFIED FINDINGS									
SUPPLIER		CHARLES RIVER									
DATES:		01-JAN-09 - 31-MAR-14									
		FETUSES		LITTERS		%RANGE (per study)		%RANGE (per study)		AFFECTED FETUSES /LITTER	
		N	%	N	%	%	%	%	%	MEAN	RANGE (per study)
NUMBER EVALUATED		4888		486							
Live		4888									
Dead		0									
U BLOOD COAGULUM AROUND PLACENTA		1	0.02	1	0.2	0.0	0.4	0.0	4.8	0.0	0.0 0.6
U PLACENTAE FUSED		2	0.04	2	0.4	0.0	0.4	0.0	4.2	0.0	0.0 0.4
U AMNIOTIC FLUID DISCOLORED		3	0.06	1	0.2	0.0	1.2	0.0	4.2	0.2	0.0 4.2
TOTAL FETAL EXTERNAL UNCLASSIFIED FIND		6	0.1	4	0.8	0.0	1.2	0.0	4.8	0.3	0.0 4.2

OBSERVATION CODE: U=Unclassified

15-APR-14

TABLE :

SPECIES RAT
STRAIN WISTAR
SUPPLIER CHARLES RIVER

DATES: 01-JAN-09 - 31-MAR-14

HISTORICAL CONTROL DATA		FETUSES		FETUSES (per study)		LITTERS		%RANGE (per study)		AFFECTED FETUSES /LITTER	
		N	%	%RANGE	%	N	%	%RANGE	%	%MEAN	%
NUMBER EVALUATED		2329				485					
Live		2329									
Dead		0									
M SITUS INVERSUS		1	0.04	0.0	0.0	1	0.2	0.0	0.0	0.0	0.8
M FETUS WITH MULTIPLE VISCERAL MALFORMATIONS		2	0.09	0.0	0.0	2	0.4	0.0	0.1	0.0	0.8
M INCOMPLETE CLOSURE OF PALATUM MOLLE		1	0.04	0.0	0.0	1	0.2	0.0	0.0	0.0	0.8
M ANOPHTHALMIA		2	0.09	0.0	1.7	2	0.4	0.0	0.1	0.0	2.8
M RIGHT-SIDED AORTIC ARCH		1	0.04	0.0	0.8	1	0.2	0.0	0.0	0.0	0.8
M ABNORMAL LUNG LOBATION		1	0.04	0.0	0.8	1	0.2	0.0	0.0	0.0	0.8
M DIAHRAGMATIC HERNIA		1	0.04	0.0	0.9	1	0.2	0.0	0.0	0.0	0.8
M HYDRONEPHROSIS		1	0.04	0.0	0.8	1	0.2	0.0	0.0	0.0	0.8
M HYDROURETER		1	0.04	0.0	0.8	1	0.2	0.0	0.0	0.0	0.8
TOTAL FETAL SOFT TISSUE MALFORMATIONS		10	0.4	0.0	3.3	10	2.1	0.0	0.5	0.0	4.4

OBSERVATION CODE: M=Malformation

TABLE :

HISTORICAL CONTROL DATA

FETAL SOFT TISSUE VARIATIONS

DATES: 01-JAN-09 - 31-MAR-14

NUMBER EVALUATED		FETUSES		%RANGE (per study)		LITTERS		%RANGE (per study)		AFFECTED FETUSES / LITTER	
		N	%	%	%	N	%	%	%	MEAN	RANGE
										%	%
Live	2329					485					
Dead	2329										
	0										
V SHORT INNOMINATE	5	0.2	0.0	1.7	5	1.0	0.0	8.3	0.2	0.0	1.3
V MALPOSITIONED CAROTID ORIGIN	1	0.04	0.0	0.9	1	0.2	0.0	4.0	0.0	0.0	0.8
V MALPOSITIONED SUBCLAVIAN ORIGIN	1	0.04	0.0	0.8	1	0.2	0.0	4.0	0.0	0.0	0.8
V ABSENT LUNG LOBE (L. INFERIOR MEDIALIS)	1	0.04	0.0	0.8	1	0.2	0.0	4.2	0.0	0.0	0.8
V DILATED RENAL PELVIS	52	2.2	0.0	6.1	51	10.5	0.0	28.0	2.2	0.0	6.3
V DILATED URETER	15	0.6	0.0	2.7	15	3.1	0.0	12.5	0.6	0.0	3.1
TOTAL FETAL SOFT TISSUE VARIATIONS	61	2.6	0.0	7.0	60	12.4	0.0	32.0	2.6	0.0	6.9

OBSERVATION CODE: V=Variation

15-APR-14

TABLE :

SPECIES		HISTORICAL CONTROL DATA									
STRAIN WISTAR		FETAL SOFT TISSUE UNCLASSIFIED FINDINGS									
SUPPLIER CHARLES RIVER											
		FETUSES		LITTERS		%RANGE (per study)		%RANGE (per study)		%MEAN (per study)	
		N	%	N	%	%	%	%	%	%	%
NUMBER EVALUATED		2329		485							
Live		2329									
Dead		0									
U DISCOLORED VITREOUS HUMOR		1	0.04	1	0.2	0.0	0.0	4.2	0.0	0.0	0.7
TOTAL FETAL SOFT TISSUE UNCLASSIFIED FIND		1	0.04	1	0.2	0.0	0.0	4.2	0.0	0.0	0.7

OBSERVATION CODE: U=Unclassified

15-APR-14

TABLE :

SPECIES RAT
STRAIN WISTAR
SUPPLIER CHARLES RIVER

HISTORICAL CONTROL DATA
FETAL SKELETAL MALFORMATIONS

DATES: 01-JAN-09 - 31-MAR-14

	FETUSES		LITTERS		%RANGE (per study)		%RANGE (per study)		AFFECTED FETUSES /LITTER	
	N	%	N	%	%	%	%	%	MEAN %	RANGE (per study)
NUMBER EVALUATED	2559		486							
Live	2559									
Dead	0									
M SEVERELY MALFORMED SKULL BONES	2	0.08	2	0.4	0.0	0.0	4.0	0.1	0.0	1.0
M MISSHAPEN BASISPHENOID	2	0.08	2	0.4	0.0	0.0	4.0	0.1	0.0	0.7
M BIPARTITE BASISPHENOID	1	0.04	1	0.2	0.0	0.0	4.2	0.0	0.0	0.8
M MISSHAPEN BASIOCCIPITAL	3	0.1	3	0.6	0.0	0.0	12.0	0.1	0.0	2.3
M SEVERELY MALFORMED VERTEBRAL COLUMN	1	0.04	1	0.2	0.0	0.0	4.0	0.0	0.0	0.8
M MISSHAPEN THORACIC VERTEBRA	1	0.04	1	0.2	0.0	0.0	4.8	0.0	0.0	0.7
M SHORTENED SCAPULA	1	0.04	1	0.2	0.0	0.0	4.0	0.0	0.0	0.7
M CLEFT STERNUM	2	0.08	2	0.4	0.0	0.0	4.0	0.1	0.0	1.0
M MALPOSITIONED AND BIPARTITE STERNEBRA Unchanged cartilage	1	0.04	1	0.2	0.0	0.0	4.0	0.0	0.0	0.6
M SEVERELY MALFORMED STERNUM	3	0.1	3	0.6	0.0	0.0	4.2	0.1	0.0	0.7
M MISSHAPEN TUBEROSITAS DELTOIDEA	5	0.2	5	1.0	0.0	0.0	8.0	0.2	0.0	1.5
M SHORTENED HUMERUS	1	0.04	1	0.2	0.0	0.0	4.0	0.0	0.0	0.7
M BENT RADIUS	1	0.04	1	0.2	0.0	0.0	4.0	0.0	0.0	0.7
TOTAL FETAL SKELETAL MALFORMATIONS	22	0.9	21	4.3	0.0	0.0	12.0	0.8	0.0	3.0

OBSERVATION CODE: M=Malformation

TABLE :

SPECIES RAT
 STRAIN WISTAR
 HISTORICAL CONTROL DATA
 FETAL SKELETAL VARIATIONS
 DATES: 01-JAN-09 - 31-MAR-14

NUMBER EVALUATED		FETUSES		%RANGE (per study)		LITTERS		%RANGE (per study)		AFFECTED FETUSES / LITTER	
Live	Dead	N	%	%	%	N	%	%	%	MEAN	%
2559	0	486									
V SUPRAOCCIPITAL HOLE (S)											
2559	279	10.9	2.3	31.5	176	36.2	12.5	84.0	10.7	2.6	30.1
V INCOMPLETE OSSIFICATION OF BASISPHENOID											
2559	418	16.3	7.9	26.9	236	48.6	28.0	76.0	16.1	8.7	27.1
V INCOMPLETE OSSIFICATION OF PARIETAL Unchanged cartilage											
2559	299	11.7	6.9	19.2	200	41.2	28.0	66.7	11.7	6.4	21.5
V INCOMPLETE OSSIFICATION OF INTERPARIETAL Unchanged cartilage											
2559	662	25.9	17.7	34.3	334	68.7	50.0	84.0	25.7	16.7	34.7
V INCOMPLETE OSSIFICATION OF SUPRAOCCIPITAL Unchanged cartilage											
2559	1106	43.2	10.6	63.7	401	82.5	48.0	96.0	43.5	10.3	64.3
V INCOMPLETE OSSIFICATION OF SKULL Unchanged cartilage											
2559	220	8.6	0.8	19.4	137	28.2	4.0	56.0	8.6	0.8	21.4
V INCOMPLETE OSSIFICATION OF BASIOCCIPITAL											
2559	1	0.04	0.0	0.8	1	0.2	0.0	4.0	0.1	0.0	1.0
V INCOMPLETE OSSIFICATION OF NASAL Unchanged cartilage											
2559	40	1.6	0.0	10.8	33	6.8	0.0	36.0	1.5	0.0	10.1
V BASIOCCIPITAL HOLE (S)											
2559	11	0.4	0.0	2.1	11	2.3	0.0	12.0	0.4	0.0	2.1
V INCOMPLETE OSSIFICATION OF HYOID Cartilage present											
2559	15	0.6	0.0	2.2	14	2.9	0.0	12.0	0.5	0.0	2.0
V INCOMPLETE OSSIFICATION OF PALATINE BONE											
2559	2	0.08	0.0	1.6	2	0.4	0.0	8.3	0.1	0.0	1.7
V INCOMPLETE OSSIFICATION OF FRONTAL Unchanged cartilage											
2559	3	0.1	0.0	1.0	3	0.6	0.0	4.8	0.2	0.0	2.4

OBSERVATION CODE: V=Variation

TABLE :

DATE: 01-JAN-09 - 31-MAR-14

SPECIES		RAT		HISTORICAL CONTROL DATA				FETAL SKELETAL VARIATIONS				DATES: 01-JAN-09 - 31-MAR-14			
SUPPAIN		WISTAR		FETAL SKELETAL VARIATIONS				FETAL SKELETAL VARIATIONS				FETAL SKELETAL VARIATIONS			
SUPPLIER		CHARLES RIVER		FETAL SKELETAL VARIATIONS				FETAL SKELETAL VARIATIONS				FETAL SKELETAL VARIATIONS			
NUMBER EVALUATED		NUMBER EVALUATED		NUMBER EVALUATED				NUMBER EVALUATED				NUMBER EVALUATED			
Live		Live		Live				Live				Live			
Dead		Dead		Dead				Dead				Dead			
V BIPARTITE OSSIFICATION OF SUPRAOCCIPITAL Unchanged cartilage		V BIPARTITE OSSIFICATION OF SUPRAOCCIPITAL Unchanged cartilage		V BIPARTITE OSSIFICATION OF SUPRAOCCIPITAL Unchanged cartilage				V BIPARTITE OSSIFICATION OF SUPRAOCCIPITAL Unchanged cartilage				V BIPARTITE OSSIFICATION OF SUPRAOCCIPITAL Unchanged cartilage			
V UNOSSIFIED HYOID Cartilage present		V UNOSSIFIED HYOID Cartilage present		V UNOSSIFIED HYOID Cartilage present				V UNOSSIFIED HYOID Cartilage present				V UNOSSIFIED HYOID Cartilage present			
V INCOMPLETE OSSIFICATION OF TEMPORAL		V INCOMPLETE OSSIFICATION OF TEMPORAL		V INCOMPLETE OSSIFICATION OF TEMPORAL				V INCOMPLETE OSSIFICATION OF TEMPORAL				V INCOMPLETE OSSIFICATION OF TEMPORAL			
V INCOMPLETE OSSIFICATION OF CERVICAL ARCH Cartilage present		V INCOMPLETE OSSIFICATION OF CERVICAL ARCH Cartilage present		V INCOMPLETE OSSIFICATION OF CERVICAL ARCH Cartilage present				V INCOMPLETE OSSIFICATION OF CERVICAL ARCH Cartilage present				V INCOMPLETE OSSIFICATION OF CERVICAL ARCH Cartilage present			
V DUMBELL OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage		V DUMBELL OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage		V DUMBELL OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage				V DUMBELL OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage				V DUMBELL OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage			
V DUMBELL OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum		V DUMBELL OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum		V DUMBELL OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum				V DUMBELL OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum				V DUMBELL OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum			
V SUPERNUMERARY THORACIC VERTEBRA		V SUPERNUMERARY THORACIC VERTEBRA		V SUPERNUMERARY THORACIC VERTEBRA				V SUPERNUMERARY THORACIC VERTEBRA				V SUPERNUMERARY THORACIC VERTEBRA			
V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage		V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage		V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage				V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage				V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage			
V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum		V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum		V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum				V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum				V INCOMPLETE OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum			
V BIPARTITE OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage		V BIPARTITE OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage		V BIPARTITE OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage				V BIPARTITE OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage				V BIPARTITE OSSIFICATION OF THORACIC CENTRUM Unchanged cartilage			
V BIPARTITE OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum		V BIPARTITE OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum		V BIPARTITE OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum				V BIPARTITE OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum				V BIPARTITE OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum			
V INCOMPLETE OSSIFICATION OF THORACIC ARCH Cartilage present		V INCOMPLETE OSSIFICATION OF THORACIC ARCH Cartilage present		V INCOMPLETE OSSIFICATION OF THORACIC ARCH Cartilage present				V INCOMPLETE OSSIFICATION OF THORACIC ARCH Cartilage present				V INCOMPLETE OSSIFICATION OF THORACIC ARCH Cartilage present			

OBSERVATION CODE: V=Variation

15-APR-14

TABLE :

SPECIES RAT
STRAIN WISTAR
SUPPLIER CHARLES RIVER

HISTORICAL CONTROL DATA
FETAL SKELETAL VARIATIONS

DATES: 01-JAN-09 - 31-MAR-14

	NUMBER EVALUATED	LIVE	DEAD	FETUSES N	% FETUSES	%RANGE (per study) %	LITTERS N	% LITTERS	%RANGE (per study) %	%MEAN (per study) %	AFFECTED FETUSES /LITTER RANGE (per study) %
V UNOSSIFIED THORACIC CENTRUM Unchanged cartilage	2559	2559	0	2	0.08	0.0	2	0.4	0.0	4.2	0.1 0.0 0.8
V UNILATERAL OSSIFICATION OF THORACIC CENTRUM Dumbbell-shaped cartilage of centrum	1	0.04	0.0	1	0.8	0.0	1	0.2	0.0	4.2	0.0 0.0 0.8
V BIPARTITE OSSIFICATION OF LUMBAR CENTRUM Dumbbell-shaped cartilage of centrum	2	0.08	0.0	2	0.8	0.0	2	0.4	0.0	4.0	0.1 0.0 0.8
V SUPERNUMERARY LUMBAR VERTEBRA	3	0.1	0.0	3	1.5	0.0	3	0.6	0.0	8.0	0.1 0.0 1.2
V DUMBELL OSSIFICATION OF LUMBAR CENTRUM Unchanged cartilage	13	0.5	0.0	12	1.6	0.0	12	2.5	0.0	8.0	0.5 0.0 1.7
V DUMBELL OSSIFICATION OF LUMBAR CENTRUM Dumbbell-shaped cartilage of centrum	2	0.08	0.0	2	0.8	0.0	2	0.4	0.0	4.0	0.1 0.0 1.0
V INCOMPLETE OSSIFICATION OF LUMBAR ARCH Cartilage present	1	0.04	0.0	1	0.7	0.0	1	0.2	0.0	4.0	0.0 0.0 0.8
V BIPARTITE OSSIFICATION OF LUMBAR CENTRUM Dumbbell-shaped cartilage of centrum	1	0.04	0.0	1	0.8	0.0	1	0.2	0.0	4.2	0.0 0.0 0.8
V INCOMPLETE OSSIFICATION OF LUMBAR CENTRUM Unchanged cartilage	1	0.04	0.0	1	1.0	0.0	1	0.2	0.0	4.8	0.0 0.0 0.8
V UNILATERAL OSSIFICATION OF LUMBAR CENTRUM Dumbbell-shaped cartilage of centrum	1	0.04	0.0	1	0.8	0.0	1	0.2	0.0	4.2	0.0 0.0 0.8
V MISSHAPEN SACRAL VERTEBRA	84	3.3	0.8	72	7.1	4.0	14.8	32.0	3.2	1.0	6.9
V INCOMPLETE OSSIFICATION OF SACRAL ARCH Cartilage present	40	1.6	0.0	32	7.7	0.0	6.6	24.0	1.6	0.0	8.1

OBSERVATION CODE: V=Variation

TABLE :

DATE: 01-JAN-09 - 31-MAR-14

SPECIES			RAT			HISTORICAL CONTROL DATA					DATES: 01-JAN-09 - 31-MAR-14				
STRAIN			WISTAR			FETAL SKELETAL VARIATIONS									
SUPPLIER			CHARLES RIVER												

NUMBER EVALUATED															
Live															
Dead															
V FUSED SACRAL CENTRUM Fused centrum cartilage			1	0.04	0.0	0.7	1	0.2	0.0	4.0	0.0	0.0	0.8		
V UNOSSIFIED STERNEBRA Unchanged cartilage			208	8.1	2.8	20.9	125	25.7	8.0	52.0	8.2	2.6	20.7		
V INCOMPLETE OSSIFICATION OF STERNEBRA Unchanged cartilage			2166	84.6	69.7	94.5	479	98.6	95.2	100.0	84.2	69.3	94.7		
V MISSHAPEN STERNEBRA Unchanged cartilage			1351	52.8	29.6	66.4	466	95.9	72.0	100.0	53.3	29.5	66.8		
V BIPARITE OSSIFICATION OF STERNEBRA Unchanged cartilage			6	0.2	0.0	0.8	6	1.2	0.0	4.2	0.3	0.0	1.4		
V UNILATERAL OSSIFICATION OF STERNEBRA Unchanged cartilage			23	0.9	0.0	3.8	23	4.7	0.0	20.0	1.0	0.0	3.7		
V EXTRA STERNEBRAL OSSIFICATION SITE Unchanged cartilage			1	0.04	0.0	0.8	1	0.2	0.0	4.8	0.0	0.0	1.0		
V SUPERNUMERARY RIB (14TH) Cartilage present			148	5.8	1.5	11.6	110	22.6	8.0	41.7	5.6	1.4	11.7		
V SUPERNUMERARY RIB (14TH) Cartilage not present			1396	54.6	40.8	64.6	435	89.5	76.0	100.0	54.4	41.5	66.8		
V CERVICAL RIB Cartilage present			3	0.1	0.0	0.8	3	0.6	0.0	4.2	0.1	0.0	0.8		
V CERVICAL RIB Cartilage not present			50	2.0	0.0	8.5	41	8.4	0.0	29.2	2.0	0.0	8.5		
V WAVY RIB			113	4.4	0.0	11.0	79	16.3	0.0	33.3	4.6	0.0	10.5		
V INCOMPLETE OSSIFICATION OF TUBEROSITAS DELTOIDEA Cartilage present			1	0.04	0.0	0.8	1	0.2	0.0	4.0	0.0	0.0	0.7		

OBSERVATION CODE: V=Variation

15-APR-14

TABLE :

SPECIES RAT
STRAIN WISTAR
SUPPLIER CHARLES RIVER

DATES: 01-JAN-09 - 31-MAR-14

HISTORICAL CONTROL DATA		FETAL SKELETAL VARIATIONS		FETUSES		LITTERS		%RANGE (per study)		%RANGE (per study)		%MEAN		AFFECTED FETUSES /LITTER	
				N	%	N	%								
NUMBER EVALUATED				2559		486									
Live				2559											
Dead				0											
V HOLE IN TUBEROSITAS DELTOIDEA				4	0.2	4	0.8	0.0	0.0	12.0	0.2	0.0	0.0	2.3	
V INCOMPLETE OSSIFICATION OF PUBIS				7	0.3	5	1.0	0.0	0.0	8.0	0.3	0.0	0.0	2.4	
Cartilage present															
V INCOMPLETE OSSIFICATION OF ISCHIUM				4	0.2	4	0.8	0.0	0.0	4.8	0.2	0.0	0.0	0.8	
Cartilage present															
TOTAL FETAL SKELETAL VARIATIONS				2533	99.0	486	100.0	95.8	100.0	100.0	99.0	95.7	100.0		

OBSERVATION CODE: V=Variation

DATE: 01-JAN-09 - 31-MAR-14

SPECIES	RAT	HISTORICAL CONTROL DATA
WISTAR STRAIN		FEMAL SKELETAL CARTILAGE

NUMBER EVALUATED	FETUSES		%RANGE (per study)		LITTERS		%RANGE (per study)		AFFECTED FETUSES / LITTER		
	N	%	%	%	N	%	%	%	%MEAN	%	
Live	2559				486						
Dead	2559										
	0										
C NOTCHED CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL	62	2.4	0.0	10.0	50	10.3	0.0	36.0	2.4	0.0	10.0
C SPLIT CARTILAGE BETWEEN BASISPHENOID AND BASIOCCIPITAL	2	0.08	0.0	0.8	2	0.4	0.0	4.2	0.1	0.0	0.8
C DUMBELL-SHAPED CARTILAGE OF CERVICAL CENTRUM	1	0.04	0.0	0.7	1	0.2	0.0	4.0	0.0	0.0	0.8
C SPLIT CARTILAGE OF CERVICAL CENTRUM	1	0.04	0.0	0.7	1	0.2	0.0	4.0	0.0	0.0	0.7
C C SACRAL ARCH CARTILAGE NOT CONNECTED	4	0.2	0.0	2.3	4	0.8	0.0	12.0	0.2	0.0	2.3
C BIPARTITE PROCESSUS XIPHOIDEUS	1722	67.3	47.5	81.4	464	95.5	83.3	100.0	67.3	46.6	82.4
C C BRANCHED RIB CARTILAGE	29	1.1	0.0	6.1	26	5.3	0.0	24.0	1.2	0.0	6.0
C C FUSED RIB CARTILAGE	2	0.08	0.0	0.8	2	0.4	0.0	4.0	0.1	0.0	0.8
C C STERNUM: EXTRA CARTILAGINOUS PART	1	0.04	0.0	0.7	1	0.2	0.0	4.0	0.0	0.0	0.8
C NOTCHED MANUBRIUM	17	0.7	0.0	4.6	14	2.9	0.0	16.0	0.7	0.0	4.3
C CARTILAGINOUS PART OF RIBS NOT CONNECTED WITH STERNUM	1	0.04	0.0	0.8	1	0.2	0.0	4.0	0.0	0.0	0.8
C C BIPARTITE RIB CARTILAGE	2	0.08	0.0	0.8	2	0.4	0.0	4.0	0.1	0.0	0.8
C CARTILAGINOUS PARTS OF RIBS DISPLACED	1	0.04	0.0	0.8	1	0.2	0.0	4.8	0.0	0.0	1.0
TOTAL FETAL SKELETAL CARTILAGE	1759	68.7	48.5	82.9	468	96.3	83.3	100.0	68.7	47.6	83.8

OBSERVATION CODE: C=Cartilage

15-APR-14

TABLE :
HISTORICAL CONTROL DATA FOR TOTAL MALFORMATIONS AND TOTAL VARIATIONS
01-JAN-09 to 31-MAR-14

SPECIES: RAT
STRAIN: WISTAR
SUPPLIER: CHARLES RIVER

	FETUSES				LITTERS				AFFECTED FETUSES/LITTER			
	POOLED N	%	LO%	HI%	POOLED N	%	LO%	HI%	POOLED MEAN%	LO%	HI%	BY STUDY LO% HI%
NUMBER EVALUATED	4888				486							
TOTAL FETAL MALFORMATIONS	35	0.72	0.00	2.02	33	6.79	0.00	20.83	0.74	0.00	2.96	
TOTAL FETAL VARIATIONS	2595	53.09	50.37	56.20	486	100.00	100.00	100.0	53.22	50.44	56.61	