

TRADE SECRET

Study Title

H-24616: Subchronic Toxicity
90-Day Gavage Study in Rats
with One-Generation Reproduction Evaluations

Volume 4 of 5

Laboratory Project ID: DuPont-4739

TEST GUIDELINES: U.S. EPA Health Effects Test Guidelines
OPPTS 870.3100 (1998)

AUTHOR: Don A. Delker, Ph.D.

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PERFORMING LABORATORY: E.I. du Pont de Nemours and Company
Haskell Laboratory for Health and Environmental Sciences
Elkton Road, P.O. Box 50
Newark, Delaware 19714-0050

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APPENDIX L (CONTINUED)

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641234	Terminal Sacrifice Killed on Day : 123 Exposure Group : RECOVERY Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.
THYROID GLAND :
ALTERATION, COLLOID, mild.
HYPERTROPHY, FOLLICULAR, minimal.

No Microscopic Abnormality Observed :
NOSE

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641237	Terminal Sacrifice Killed on Day : 92 Exposure Group : SUBCHRONIC Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.
THYROID GLAND :
ALTERATION, COLLOID, minimal.
HYPERTROPHY, FOLLICULAR, minimal.
PARATHYROID GLANDS :
NOT PRESENT.
UTERUS :
DILATATION, LUMEN, minimal.
NOSE :
DEGENERATION/NECROSIS, OLFACTORY, EPITHELIUM, minimal.

641237 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641237 Continued from previous page

Histopathology :

No Microscopic Abnormality Observed :
KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN, AORTA,
BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM,
PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH NODE,
SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS, ADRENAL
GLANDS, SCIATIC NERVE, PITUITARY GLAND, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, URINARY BLADDER, FEMUR/KNEE JOINT,
STERNUM, BONE MARROW

641256 Terminal Sacrifice
Killed on Day : 123
Exposure Group : RECOVERY
Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

641256 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641256 Continued from previous page

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.

THYROID GLAND :
ALTERATION, COLLOID, minimal.

No Microscopic Abnormality Observed :
NOSE

641259 Terminal Sacrifice
Killed on Day : 182
Exposure Group : FLUORINE ANALYSIS
Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

641259 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641259 Continued from previous page

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.
FATTY CHANGE, MEDIAN CLEFT, minimal.

No Microscopic Abnormality Observed :
THYROID GLAND, NOSE

641264 Terminal Sacrifice
Killed on Day : 182
Exposure Group : FLUORINE ANALYSIS
Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS; MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.

641264 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641264 Continued from previous page

Histopathology :

THYROID GLAND :
ALTERATION, COLLOID, mild.
HYPERTROPHY, FOLLICULAR, minimal.

No Microscopic Abnormality Observed :
NOSE

641275 Terminal Sacrifice
Killed on Day : 92
Exposure Group : SUBCHRONIC
Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
FATTY CHANGE, MEDIAN CLEFT, minimal.

641275 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641275 Continued from previous page

Histopathology :

LIVER :
 INFLAMMATION, SUBACUTE/CHRONIC, minimal.
LUNGS :
 INFLAMMATION, SUBACUTE/CHRONIC, minimal.
THYROID GLAND :
 ALTERATION, COLLOID, minimal.
PARATHYROID GLANDS :
 NOT PRESENT.
UTERUS :
 DILATATION, LUMEN, mild.
NOSE :
 DEGENERATION/NECROSIS, OLFACTORY, EPITHELIUM, minimal.

No Microscopic Abnormality Observed :
KIDNEYS, HEART, SKELETAL MUSCLE, SPLEEN, AORTA, BRAIN,
SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM, PANCREAS,
CECUM, COLON, RECTUM, MESENTERIC LYMPH NODE, SALIVARY
GLANDS, MANDIBULAR LYMPH NODE, THYMUS, ADRENAL GLANDS,
SCIATIC NERVE, PITUITARY GLAND, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, URINARY BLADDER, FEMUR/KNEE JOINT,
STERNUM, BONE MARROW

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
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641278	Terminal Sacrifice Killed on Day : 182 Exposure Group : FLUORINE ANALYSIS Animal is signed off from necropsy
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Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.
THYROID GLAND :
ALTERATION, COLLOID, minimal.

No Microscopic Abnormality Observed :
NOSE

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641279 Terminal Sacrifice
 Killed on Day : 92
 Exposure Group : SUBCHRONIC
 Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
 INFLAMMATION, SUBACUTE/CHRONIC, minimal.
THYROID GLAND :
 ALTERATION, COLLOID, mild.
 HYPERTROPHY, FOLLICULAR, mild.
PARATHYROID GLANDS :
 NOT PRESENT.

641279 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641279	Continued from previous page

Histopathology :

No Microscopic Abnormality Observed :
KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN, AORTA,
BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM,
PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH NODE,
SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS, ADRENAL
GLANDS, SCIATIC NERVE, PITUITARY GLAND, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, BONE MARROW, NOSE

641282 Terminal Sacrifice
 Killed on Day : 123
 Exposure Group : RECOVERY
 Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

641282 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641282 Continued from previous page

Histopathology :

LIVER :
 INFLAMMATION, SUBACUTE/CHRONIC, minimal.
THYROID GLAND :
 ALTERATION, COLLOID, minimal.
 HYPERTROPHY, FOLLICULAR, minimal.

No Microscopic Abnormality Observed :
 NOSE

641299 Terminal Sacrifice
 Killed on Day : 92
 Exposure Group : SUBCHRONIC
 Animal is signed off from necropsy

Gross Pathology :

SPLEEN :
 DISCOLORATION, BROWN.

No Macroscopic Abnormality Observed :
 LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, AORTA,
 BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM,
 PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH NODE,
 SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS, ADRENAL
 GLANDS, SCIATIC NERVE, PITUITARY GLAND, THYROID GLAND,
 PARATHYROID GLANDS, TRACHEA, ESOPHAGUS, PHARYNX/LARYNX,
 EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY GLAND (FEMALE),
 OVARIES, UTERUS, URINARY BLADDER, FEMUR/KNEE JOINT,
 STERNUM, NOSE

641299 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641299 Continued from previous page

Histopathology :

LIVER :

INFLAMMATION, SUBACUTE/CHRONIC, minimal.

THYROID GLAND :

HYPERTROPHY, FOLLICULAR, mild.

ALTERATION, COLLOID, minimal.

UTERUS :

DILATATION, LUMEN, minimal.

NOSE :

DEGENERATION/NECROSIS, OLFACTORY, EPITHELIUM, minimal.

No Microscopic Abnormality Observed :

KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN, AORTA,
BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM,
PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH NODE,
SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS, ADRENAL
GLANDS, SCIATIC NERVE, PITUITARY GLAND, PARATHYROID GLANDS,
TRACHEA, ESOPHAGUS, PHARYNX/LARYNX,
EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY GLAND (FEMALE),
OVARIES, URINARY BLADDER, FEMUR/KNEE JOINT, STERNUM, BONE
MARROW

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641300 Terminal Sacrifice
 Killed on Day : 92
 Exposure Group : SUBCHRONIC
 Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
 INFLAMMATION, SUBACUTE/CHRONIC, minimal.
THYROID GLAND :
 ALTERATION, COLLOID, minimal.
PARATHYROID GLANDS :
 NOT PRESENT.
NOSE :
 INFLAMMATION, SUBACUTE/CHRONIC, minimal.
 DEGENERATION/NECROSIS, OLFACTORY, EPITHELIUM, mild.

641300 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641300	Continued from previous page

Histopathology :

No Microscopic Abnormality Observed :
KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN, AORTA,
BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM,
PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH NODE,
SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS, ADRENAL
GLANDS, SCIATIC NERVE, PITUITARY GLAND, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, BONE MARROW

641301 Terminal Sacrifice
 Killed on Day : 182
 Exposure Group : FLUORINE ANALYSIS
 Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

641301 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641301	Continued from previous page

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.

No Microscopic Abnormality Observed :
THYROID GLAND, NOSE

641314 Terminal Sacrifice
 Killed on Day : 92
 Exposure Group : SUBCHRONIC
 Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.

641314 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641314 Continued from previous page

Histopathology :

THYROID GLAND :

ALTERATION, COLLOID, minimal.

NOSE :

DEGENERATION/NECROSIS, OLFACTORY, EPITHELIUM, minimal.

No Microscopic Abnormality Observed :

KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN, AORTA,
BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM,
PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH NODE,
SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS, ADRENAL
GLANDS, SCIATIC NERVE, PITUITARY GLAND, PARATHYROID GLANDS,
TRACHEA, ESOPHAGUS, PHARYNX/LARYNX,
EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY GLAND (FEMALE),
OVARIES, UTERUS, URINARY BLADDER, FEMUR/KNEE JOINT,
STERNUM, BONE MARROW

641326

Terminal Sacrifice

Killed on Day : 92

Exposure Group : SUBCHRONIC

Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :

LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,

641326 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641326 Continued from previous page

Gross Pathology :

No Macroscopic Abnormality Observed :
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.
THYROID GLAND :
ALTERATION, COLLOID, minimal.
HYPERTROPHY, FOLLICULAR, minimal.
PARATHYROID GLANDS :
NOT PRESENT.

No Microscopic Abnormality Observed :
KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN, AORTA,
BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM,
PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH NODE,
SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS, ADRENAL
GLANDS, SCIATIC NERVE, PITUITARY GLAND, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, BONE MARROW, NOSE

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641327	Terminal Sacrifice Killed on Day : 92 Exposure Group : SUBCHRONIC Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.
FATTY CHANGE, MEDIAN CLEFT, minimal.
THYROID GLAND :
ALTERATION, COLLOID, minimal.
HYPERTROPHY, FOLLICULAR, minimal.
PARATHYROID GLANDS :
NOT PRESENT.
NOSE :
DEGENERATION/NECROSIS, OLFACTORY, EPITHELIUM, minimal.

641327 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641327 Continued from previous page

Histopathology :

No Microscopic Abnormality Observed :
KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN, AORTA,
BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM,
PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH NODE,
SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS, ADRENAL
GLANDS, SCIATIC NERVE, PITUITARY GLAND, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, BONE MARROW

641352 Terminal Sacrifice
Killed on Day : 123
Exposure Group : RECOVERY
Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

641352 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641352 Continued from previous page

Histopathology :

LIVER :

INFLAMMATION, SUBACUTE/CHRONIC, minimal.

No Microscopic Abnormality Observed :
THYROID GLAND, NOSE

641356

Found Dead
Duration of dosing-days: 121
Exposure Group : RECOVERY
Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :

LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

BRAIN :

MIXED GLIOMA [M].

[B]=BENIGN, [M]=MALIGNANT

641356 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641356	Continued from previous page

Histopathology :

ILEUM :
AUTOLYSIS: NECROPSY AND HISTOLOGY PERFORMED.
CECUM :
AUTOLYSIS: NECROPSY AND HISTOLOGY PERFORMED.
COLON :
AUTOLYSIS: NECROPSY AND HISTOLOGY PERFORMED.
THYROID GLAND :
AUTOLYSIS: NECROPSY AND HISTOLOGY PERFORMED.
EYE(S) WITH OPTIC NERVE :
AUTOLYSIS: NECROPSY AND HISTOLOGY PERFORMED.
SKIN :
NOT PRESENT.
MAMMARY GLAND (FEMALE) :
NOT PRESENT.

No Microscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, PANCREAS,
RECTUM, MESENTERIC LYMPH NODE, SALIVARY GLANDS, MANDIBULAR
LYMPH NODE, THYMUS, ADRENAL GLANDS, SCIATIC NERVE,
PITUITARY GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, BONE MARROW, NOSE

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641366	Terminal Sacrifice Killed on Day : 123 Exposure Group : RECOVERY Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.
NECROSIS, FOCAL, minimal.

No Microscopic Abnormality Observed :
THYROID GLAND, NOSE

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641377	Terminal Sacrifice Killed on Day : 123 Exposure Group : RECOVERY Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.
FATTY CHANGE, MEDIAN CLEFT, minimal.
THYROID GLAND :
ALTERATION, COLLOID, minimal.
HYPERTROPHY, FOLLICULAR, minimal.

No Microscopic Abnormality Observed :
NOSE

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641380	Terminal Sacrifice Killed on Day : 92 Exposure Group : SUBCHRONIC Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.
HEART :
CARDIOMYOPATHY, minimal.
SCIATIC NERVE :
NOT PRESENT.
THYROID GLAND :
ALTERATION, COLLOID, minimal.
HYPERTROPHY, FOLLICULAR, minimal.
PARATHYROID GLANDS :
NOT PRESENT.
NOSE :
DEGENERATION/NECROSIS, OLFACTORY, EPITHELIUM, mild.

641380 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641380 Continued from previous page

Histopathology :

No Microscopic Abnormality Observed :

KIDNEYS, LUNGS, SKELETAL MUSCLE, SPLEEN, AORTA, BRAIN,
SPINAL CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM, PANCREAS,
CECUM, COLON, RECTUM, MESENTERIC LYMPH NODE, SALIVARY
GLANDS, MANDIBULAR LYMPH NODE, THYMUS, ADRENAL GLANDS,
PITUITARY GLAND, TRACHEA, ESOPHAGUS, PHARYNX/LARYNX,
EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY GLAND (FEMALE),
OVARIES, UTERUS, URINARY BLADDER, FEMUR/KNEE JOINT,
STERNUM, BONE MARROW

641385

Terminal Sacrifice
Killed on Day : 123
Exposure Group : RECOVERY
Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :

LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

641385 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641385 Continued from previous page

Histopathology :

LIVER :

INFLAMMATION, SUBACUTE/CHRONIC, minimal.

THYROID GLAND :

ALTERATION, COLLOID, minimal.

No Microscopic Abnormality Observed :

NOSE

641387

Terminal Sacrifice

Killed on Day : 123

Exposure Group : RECOVERY

Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :

LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

641387 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641387 Continued from previous page

Histopathology :

LIVER :
 INFLAMMATION, SUBACUTE/CHRONIC, minimal.
THYROID GLAND :
 ALTERATION, COLLOID, minimal.
 HYPERTROPHY, FOLLICULAR, minimal.

No Microscopic Abnormality Observed :
 NOSE

641390 Terminal Sacrifice
 Killed on Day : 92
 Exposure Group : SUBCHRONIC
 Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
 LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
 AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
 ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
 NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
 ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
 THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
 PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
 GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
 FEMUR/KNEE JOINT, STERNUM, NOSE

641390 Continued on the next page

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641390 Continued from previous page

Histopathology :

LIVER :

INFLAMMATION, SUBACUTE/CHRONIC, minimal.

KIDNEYS :

INFLAMMATION, SUBACUTE/CHRONIC, minimal.

THYROID GLAND :

ALTERATION, COLLOID, minimal.

HYPERTROPHY, FOLLICULAR, mild.

PARATHYROID GLANDS :

NOT PRESENT.

EYE(S) WITH OPTIC NERVE :

OPTIC NERVE NOT PRESENT.

NOSE :

DEGENERATION/NECROSIS, OLFACTORY, EPITHELIUM, minimal.

No Microscopic Abnormality Observed :

LUNGS, HEART, SKELETAL MUSCLE, SPLEEN, AORTA, BRAIN, SPINAL
CORD, STOMACH, DUODENUM, JEJUNUM, ILEUM, PANCREAS, CECUM,
COLON, RECTUM, MESENTERIC LYMPH NODE, SALIVARY GLANDS,
MANDIBULAR LYMPH NODE, THYMUS, ADRENAL GLANDS, SCIATIC
NERVE, PITUITARY GLAND, TRACHEA, ESOPHAGUS, PHARYNX/LARYNX,
EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY GLAND (FEMALE),
OVARIES, UTERUS, URINARY BLADDER, FEMUR/KNEE JOINT,
STERNUM, BONE MARROW

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref Microscopic & Macroscopic Findings

641395 Terminal Sacrifice
 Killed on Day : 182
 Exposure Group : FLUORINE ANALYSIS
 Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
 INFLAMMATION, SUBACUTE/CHRONIC, minimal.
 FATTY CHANGE, MEDIAN CLEFT, mild.
THYROID GLAND :
 ALTERATION, COLLOID, minimal.

No Microscopic Abnormality Observed :
NOSE

INDIVIDUAL ANIMAL GROSS AND MICROSCOPIC OBSERVATIONS

Dose Group : VIII - 1000 mg/kg/day Sex: Females

Animal Ref	Microscopic & Macroscopic Findings
641399	Terminal Sacrifice Killed on Day : 123 Exposure Group : RECOVERY Animal is signed off from necropsy

Gross Pathology :

No Macroscopic Abnormality Observed :
LIVER, KIDNEYS, LUNGS, HEART, SKELETAL MUSCLE, SPLEEN,
AORTA, BRAIN, SPINAL CORD, STOMACH, DUODENUM, JEJUNUM,
ILEUM, PANCREAS, CECUM, COLON, RECTUM, MESENTERIC LYMPH
NODE, SALIVARY GLANDS, MANDIBULAR LYMPH NODE, THYMUS,
ADRENAL GLANDS, SCIATIC NERVE, PITUITARY GLAND,
THYROID GLAND, PARATHYROID GLANDS, TRACHEA, ESOPHAGUS,
PHARYNX/LARYNX, EYE(S) WITH OPTIC NERVE, SKIN, MAMMARY
GLAND (FEMALE), OVARIES, UTERUS, URINARY BLADDER,
FEMUR/KNEE JOINT, STERNUM, NOSE

Histopathology :

LIVER :
INFLAMMATION, SUBACUTE/CHRONIC, minimal.
THYROID GLAND :
ALTERATION, COLLOID, moderate.
HYPERTROPHY, FOLLICULAR, mild.

No Microscopic Abnormality Observed :
NOSE

APPENDIX M
REPRODUCTIVE TOXICOLOGY EVALUATIONS
INDIVIDUAL BODY WEIGHTS OF P₁ MALE RATS

REPRODUCTIVE TOXICOLOGY EVALUATIONS
INDIVIDUAL BODY WEIGHTS OF P₁ MALE RATS

EXPLANATORY NOTES

Notes:

Test Days 71 – 85 = Cohabitation period

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ MALE RATS

GROUP: I-0

DOSE: 0 MG/KG/DAY

TEST DAYS:	71	78	85	92	100
ANIMAL NUMBER					
641114	660.5	668.9	685.4	675.2	681.6
641217	568.4	592.7	597.2	595.7	588.8
641204	500.1	508.7	513.9	520.3	522.2
641158	547.1	562.3	579.1	594.4	614.3
641100	495.4	507.2	518.6	531.5	533.5
641060	651.9	684.7	697.4	719.1	736.4
641188	501.5	510.9	520.0	528.4	532.6
641106	547.3	568.1	574.7	589.4	606.1
641175	528.5	541.7	552.7	561.5	560.1
641079	464.4	484.6	490.8	501.0	502.9
641125	638.4	655.5	671.2	696.9	716.1
641186	521.5	540.4	558.5	572.9	589.1
641052	529.2	535.1	545.6	558.8	563.0
641140	653.6	665.6	686.7	706.5	714.0
641135	494.6	500.9	512.5	535.1	536.0
641174	492.4	508.8	515.6	528.5	534.2
641107	517.3	518.4	534.1	556.7	568.0
641072	495.1	501.2	510.0	512.7	527.6
641053	569.3	584.3	597.1	600.5	633.6
641168	527.7	537.8	552.6	561.0	573.5

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ MALE RATS

GROUP: III-0

DOSE: 50 MG/KG/DAY

TEST DAYS:	71	78	85	92	100
ANIMAL NUMBER					
641087	555.4	583.1	594.2	610.7	609.3
641108	501.6	451.6	495.0	520.3	537.2
641118	559.5	567.0	577.3	602.6	604.2
641128	572.0	586.5	597.4	595.8	596.6
641091	459.2	464.5	484.3	498.3	507.7
641199	563.5	563.3	564.5	578.3	591.1
641209	482.5	508.1	513.9	507.5	518.2
641163	484.7	497.9	502.5	497.6	505.1
641129	516.8	522.4	538.4	546.1	560.3
641138	534.7	548.3	554.7	563.1	573.9
641103	622.9	616.3	627.7	647.1	651.3
641181	559.6	579.2	584.5	597.2	598.4
641171	598.6	617.5	622.4	640.6	652.5
641111	515.3	522.0	524.3	532.1	523.5
641062	524.2	536.6	539.7	551.1	560.3
641156	576.3	587.9	607.9	620.3	619.0
641173	556.4	563.9	574.3	592.7	601.9
641121	524.6	552.9	557.9	575.1	592.2
641074	582.5	600.3	610.3	569.0	600.9
641191	570.1	569.2	562.3	568.7	572.7

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ MALE RATS

GROUP: V-0

DOSE: 250 MG/KG/DAY

TEST DAYS:	71	78	85	92	100
ANIMAL NUMBER					
641170	578.9	600.0	614.0	639.4	641.1
641065	506.7	535.8	543.1	558.6	565.1
641126	495.0	474.0	483.9	501.7	522.9
641058	454.3	459.6	477.0	478.1	483.8
641131	504.0	526.7	524.3	531.9	523.5
641176	531.1	546.6	542.4	552.4	564.2
641119	581.3	605.5	601.7	617.4	612.7
641075	532.0	546.3	565.9	566.3	559.4
641180	528.0	541.5	544.2	564.8	579.2
641117	579.0	606.9	610.9	626.7	643.6
641124	594.3	601.6	618.3	636.2	652.6
641082	486.3	493.5	499.8	507.3	514.7
641151	519.8	528.2	538.5	547.3	562.9
641195	550.3	565.6	576.8	590.3	599.6
641122	538.9	551.0	558.7	553.5	565.4
641184	505.4	514.5	524.5	520.3	517.2
641142	476.4	478.6	476.4	487.5	502.6
641120	603.5	612.3	630.1	641.7	647.2
641179	583.4	604.1	605.0	620.7	648.9
641164	560.1	576.5	593.3	613.1	622.3

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ MALE RATS

GROUP: VII-0

DOSE: 1000 MG/KG/DAY

TEST DAYS:	71	78	85	92	100
ANIMAL NUMBER					
641110	558.4	567.9	556.4	565.8	592.1
641096	449.6	458.8	479.1	489.6	498.1
641212	459.6	476.0	483.5	487.6	501.5
641068	458.1	465.2	462.6	469.2	476.8
641093	445.9	455.9	470.5	491.6	501.7
641054	528.2	536.3	544.9	564.0	557.8
641208	402.4	402.5	408.4	420.3	409.0
641193	519.9	532.5	534.6	550.3	558.1
641207	549.8	557.9	571.1	589.2	592.2
641139	489.9	497.2	503.1	504.5	520.2
641080	498.2	495.9	514.8	522.7	529.8
641078	497.1	521.1	522.5	535.5	524.9
641177	589.1	621.4	639.1	642.3	646.3
641112	510.7	527.2	547.3	554.9	571.6
641210	563.7	579.0	535.5	564.1	583.7
641152	468.7	491.8	500.4	517.6	525.9
641182	447.7	451.9	459.0	474.9	480.5
641150	433.5	442.4	441.3	461.9	463.1
641211	430.4	446.8	455.2	465.8	473.6
641145	494.7	518.6	504.2	519.7	526.8

APPENDIX N

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHT GAINS OF P₁ MALE RATS

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHT GAINS OF P₁ MALE RATS

EXPLANATORY NOTES

Notes:

Test Days 71 – 85 = Cohabitation period

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ MALE RATS

GROUP: I-0

DOSE: 0 MG/KG/DAY

TEST DAYS:	71-78	78-85	85-92	92-100	71-100
ANIMAL NUMBER					
641114	8.4	16.5	-10.2	6.4	21.1
641217	24.3	4.5	-1.5	-6.9	20.4
641204	8.6	5.2	6.4	1.9	22.1
641158	15.2	16.8	15.3	19.9	67.2
641100	11.8	11.4	12.9	2.0	38.1
641060	32.8	12.7	21.7	17.3	84.5
641188	9.4	9.1	8.4	4.2	31.1
641106	20.8	6.6	14.7	16.7	58.8
641175	13.2	11.0	8.8	-1.4	31.6
641079	20.2	6.2	10.2	1.9	38.5
641125	17.1	15.7	25.7	19.2	77.7
641186	18.9	18.1	14.4	16.2	67.6
641052	5.9	10.5	13.2	4.2	33.8
641140	12.0	21.1	19.8	7.5	60.4
641135	6.3	11.6	22.6	0.9	41.4
641174	16.4	6.8	12.9	5.7	41.8
641107	1.1	15.7	22.6	11.3	50.7
641072	6.1	8.8	2.7	14.9	32.5
641053	15.0	12.8	3.4	33.1	64.3
641168	10.1	14.8	8.4	12.5	45.8

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ MALE RATS

GROUP: III-0

DOSE: 50 MG/KG/DAY

TEST DAYS:	71-78	78-85	85-92	92-100	71-100
ANIMAL NUMBER					
641087	27.7	11.1	16.5	-1.4	53.9
641108	-50.0	43.4	25.3	16.9	35.6
641118	7.5	10.3	25.3	1.6	44.7
641128	14.5	10.9	-1.6	0.8	24.6
641091	5.3	19.8	14.0	9.4	48.5
641199	-0.2	1.2	13.8	12.8	27.6
641209	25.6	5.8	-6.4	10.7	35.7
641163	13.2	4.6	-4.9	7.5	20.4
641129	5.6	16.0	7.7	14.2	43.5
641138	13.6	6.4	8.4	10.8	39.2
641103	-6.6	11.4	19.4	4.2	28.4
641181	19.6	5.3	12.7	1.2	38.8
641171	18.9	4.9	18.2	11.9	53.9
641111	6.7	2.3	7.8	-8.6	8.2
641062	12.4	3.1	11.4	9.2	36.1
641156	11.6	20.0	12.4	-1.3	42.7
641173	7.5	10.4	18.4	9.2	45.5
641121	28.3	5.0	17.2	17.1	67.6
641074	17.8	10.0	-41.3	31.9	18.4
641191	-0.9	-6.9	6.4	4.0	2.6

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ MALE RATS

GROUP: V-0

DOSE: 250 MG/KG/DAY

TEST DAYS:	71-78	78-85	85-92	92-100	71-100
ANIMAL NUMBER					
641170	21.1	14.0	25.4	1.7	62.2
641065	29.1	7.3	15.5	6.5	58.4
641126	-21.0	9.9	17.8	21.2	27.9
641058	5.3	17.4	1.1	5.7	29.5
641131	22.7	-2.4	7.6	-8.4	19.5
641176	15.5	-4.2	10.0	11.8	33.1
641119	24.2	-3.8	15.7	-4.7	31.4
641075	14.3	19.6	0.4	-6.9	27.4
641180	13.5	2.7	20.6	14.4	51.2
641117	27.9	4.0	15.8	16.9	64.6
641124	7.3	16.7	17.9	16.4	58.3
641082	7.2	6.3	7.5	7.4	28.4
641151	8.4	10.3	8.8	15.6	43.1
641195	15.3	11.2	13.5	9.3	49.3
641122	12.1	7.7	-5.2	11.9	26.5
641184	9.1	10.0	-4.2	-3.1	11.8
641142	2.2	-2.2	11.1	15.1	26.2
641120	8.8	17.8	11.6	5.5	43.7
641179	20.7	0.9	15.7	28.2	65.5
641164	16.4	16.8	19.8	9.2	62.2

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ MALE RATS

GROUP: VII-0

DOSE: 1000 MG/KG/DAY

TEST DAYS:	71-78	78-85	85-92	92-100	71-100
ANIMAL NUMBER					
641110	9.5	-11.5	9.4	26.3	33.7
641096	9.2	20.3	10.5	8.5	48.5
641212	16.4	7.5	4.1	13.9	41.9
641068	7.1	-2.6	6.6	7.6	18.7
641093	10.0	14.6	21.1	10.1	55.8
641054	8.1	8.6	19.1	-6.2	29.6
641208	0.1	5.9	11.9	-11.3	6.6
641193	12.6	2.1	15.7	7.8	38.2
641207	8.1	13.2	18.1	3.0	42.4
641139	7.3	5.9	1.4	15.7	30.3
641080	-2.3	18.9	7.9	7.1	31.6
641078	24.0	1.4	13.0	-10.6	27.8
641177	32.3	17.7	3.2	4.0	57.2
641112	16.5	20.1	7.6	16.7	60.9
641210	15.3	-43.5	28.6	19.6	20.0
641152	23.1	8.6	17.2	8.3	57.2
641182	4.2	7.1	15.9	5.6	32.8
641150	8.9	-1.1	20.6	1.2	29.6
641211	16.4	8.4	10.6	7.8	43.2
641145	23.9	-14.4	15.5	7.1	32.1

APPENDIX O

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHTS OF P₁ FEMALE RATS DURING GESTATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHTS OF P₁ FEMALE RATS DURING GESTATION

EXPLANATORY NOTES

Footnotes

- a* Dam delivered prior to collection of day 21 body weight.
- b* No litter was delivered; data were excluded from statistical analysis.
- c* No gestation data were collected since no plug was observed; however, a litter was delivered.
- d* No gestation data were collected since no plug was observed; no litter was delivered.

Abbreviations

--- = No Data

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ FEMALE RATS DURING GESTATION

GROUP: II-0

DOSE: 0 MG/KG/DAY

GESTATION DAYS:	0	7	14	21
ANIMAL NUMBER				
641345	277.4	289.9	309.3	--- ^a
641236	294.9	330.3	366.7	461.3
641337	324.8	356.9	383.8	450.2
641328	325.8	352.1	381.6	455.0
641310	274.0	309.6	331.5	430.1
641324	288.1	318.0	350.2	438.3
641371	322.6	345.4	380.0	484.2
641332	331.2	360.4	405.4	498.0
641358 ^b	300.7	309.2	314.4	311.7
641389 ^b	297.7	309.9	317.2	314.9
641317 ^c	---	---	---	---
641255	302.8	327.2	347.7	447.4
641403	319.2	362.1	395.8	507.4
641388	354.2	376.0	425.9	--- ^a
641261	287.5	310.4	339.9	420.6
641306	289.3	329.8	356.1	450.4
641349 ^c	---	---	---	---
641346	321.8	359.8	382.0	473.5
641308	281.7	306.4	342.6	419.0
641289	300.8	330.4	366.9	441.5

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ FEMALE RATS DURING GESTATION

GROUP: IV-0

DOSE: 50 MG/KG/DAY

GESTATION DAYS:	0	7	14	21
ANIMAL NUMBER				
641378	336.0	372.8	405.4	512.7
641263	278.3	298.0	326.3	410.0
641297	311.4	347.8	378.7	483.9
641304	315.9	347.7	365.3	460.8
641274 ^b	350.5	362.8	363.2	367.2
641351	276.4	293.1	320.2	421.4
641272	290.4	317.5	346.3	437.9
641258	339.7	362.5	393.1	--- ^a
641276 ^b	376.6	393.4	411.5	401.2
641350	292.7	306.8	339.9	442.4
641376	292.3	313.6	331.8	390.8
641315	310.4	337.9	364.9	469.8
641286	335.2	350.7	386.3	471.0
641231	286.1	316.9	342.3	429.6
641338	283.6	320.5	353.0	441.2
641318	308.5	322.9	342.7	430.1
641373	330.4	366.7	405.1	517.5
641365	298.0	325.0	357.0	441.3
641348	388.0	422.7	461.3	532.9
641323	306.9	339.8	368.8	457.4

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ FEMALE RATS DURING GESTATION

GROUP: VI-0

DOSE: 250 MG/KG/DAY

GESTATION DAYS:	0	7	14	21
ANIMAL NUMBER				
641271	299.7	319.4	346.3	448.1
641280 ^b	282.6	313.4	340.2	334.2
641252 ^b	311.4	326.0	320.1	328.2
641386 ^b	288.3	315.8	338.6	317.6
641277	318.8	352.5	390.5	489.0
641233	311.3	343.5	386.4	480.7
641238	298.4	312.7	345.4	438.2
641235	253.8	282.8	304.4	362.1
641392	310.2	327.8	364.3	440.3
641360	273.2	297.8	323.2	408.3
641381 ^b	334.3	351.5	361.2	363.3
641333 ^d	---	---	---	---
641239	333.1	353.3	386.0	487.2
641228 ^b	334.9	361.6	389.4	387.4
641334	285.0	314.6	342.4	431.7
641347	315.2	351.3	394.3	490.2
641367 ^b	350.5	340.1	350.5	349.5
641303	286.2	308.0	329.0	417.4
641339	321.0	349.0	378.0	472.4
641316	293.1	319.2	349.3	443.2

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ FEMALE RATS DURING GESTATION

GROUP: VIII-0

DOSE: 1000 MG/KG/DAY

GESTATION DAYS:	0	7	14	21
ANIMAL NUMBER				
641265	298.6	319.8	354.7	451.3
641393	291.3	312.0	349.2	443.6
641402	267.6	281.4	302.5	368.9
641246	264.9	279.3	311.0	390.5
641320	321.2	333.2	374.7	471.9
641353	256.8	292.7	328.7	406.9
641401	281.2	309.5	336.6	437.4
641307	274.4	297.6	317.2	405.7
641368	311.5	330.6	362.1	476.1
641343	348.6	369.1	411.8	522.0
641254	301.9	340.9	373.2	472.8
641242 ^b	343.9	336.9	346.8	346.9
641284	304.4	336.5	365.3	485.1
641230	279.6	297.2	325.9	408.0
641273 ^b	286.7	318.6	329.4	320.0
641296	279.5	295.3	329.9	421.7
641248	346.4	373.1	409.0	519.2
641370	310.9	337.5	370.3	444.3
641243	312.0	335.4	372.3	470.0
641329	264.4	281.6	313.0	409.1

APPENDIX P

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHT GAINS OF P₁ FEMALE RATS DURING GESTATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHT GAINS OF P₁ FEMALE RATS DURING GESTATION

EXPLANATORY NOTES

Footnotes

- a* Dam delivered prior to collection of day 21 body weight.
- b* No litter was delivered; data were excluded from statistical analysis.
- c* No gestation data were collected since no plug was observed; however, a litter was delivered.
- d* No gestation data were collected since no plug was observed; no litter was delivered.

Abbreviations

--- = No Data

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ FEMALE RATS DURING GESTATION

GROUP: II-0

DOSE: 0 MG/KG/DAY

GESTATION DAYS:	0-7	7-14	14-21	0-21
ANIMAL NUMBER				
641345	12.5	19.4	--- ^a	--- ^a
641236	35.4	36.4	94.6	166.4
641337	32.1	26.9	66.4	125.4
641328	26.3	29.5	73.4	129.2
641310	35.6	21.9	98.6	156.1
641324	29.9	32.2	88.1	150.2
641371	22.8	34.6	104.2	161.6
641332	29.2	45.0	92.6	166.8
641358 ^b	8.5	5.2	-2.7	11.0
641389 ^b	12.2	7.3	-2.3	17.2
641317 ^c	---	---	---	---
641255	24.4	20.5	99.7	144.6
641403	42.9	33.7	111.6	188.2
641388	21.8	49.9	--- ^a	--- ^a
641261	22.9	29.5	80.7	133.1
641306	40.5	26.3	94.3	161.1
641349 ^c	---	---	---	---
641346	38.0	22.2	91.5	151.7
641308	24.7	36.2	76.4	137.3
641289	29.6	36.5	74.6	140.7

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ FEMALE RATS DURING GESTATION

GROUP: IV-0

DOSE: 50 MG/KG/DAY

GESTATION DAYS:	0-7	7-14	14-21	0-21
ANIMAL NUMBER				
641378	36.8	32.6	107.3	176.7
641263	19.7	28.3	83.7	131.7
641297	36.4	30.9	105.2	172.5
641304	31.8	17.6	95.5	144.9
641274 ^b	12.3	0.4	4.0	16.7
641351	16.7	27.1	101.2	145.0
641272	27.1	28.8	91.6	147.5
641258	22.8	30.6	--- ^a	--- ^a
641276 ^b	16.8	18.1	-10.3	24.6
641350	14.1	33.1	102.5	149.7
641376	21.3	18.2	59.0	98.5
641315	27.5	27.0	104.9	159.4
641286	15.5	35.6	84.7	135.8
641231	30.8	25.4	87.3	143.5
641338	36.9	32.5	88.2	157.6
641318	14.4	19.8	87.4	121.6
641373	36.3	38.4	112.4	187.1
641365	27.0	32.0	84.3	143.3
641348	34.7	38.6	71.6	144.9
641323	32.9	29.0	88.6	150.5

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ FEMALE RATS DURING GESTATION

GROUP: VI-0

DOSE: 250 MG/KG/DAY

GESTATION DAYS:	0-7	7-14	14-21	0-21
ANIMAL NUMBER				
641271	19.7	26.9	101.8	148.4
641280 ^b	30.8	26.8	-6.0	51.6
641252 ^b	14.6	-5.9	8.1	16.8
641386 ^b	27.5	22.8	-21.0	29.3
641277	33.7	38.0	98.5	170.2
641233	32.2	42.9	94.3	169.4
641238	14.3	32.7	92.8	139.8
641235	29.0	21.6	57.7	108.3
641392	17.6	36.5	76.0	130.1
641360	24.6	25.4	85.1	135.1
641381 ^b	17.2	9.7	2.1	29.0
641333 ^d	---	---	---	---
641239	20.2	32.7	101.2	154.1
641228 ^b	26.7	27.8	-2.0	52.5
641334	29.6	27.8	89.3	146.7
641347	36.1	43.0	95.9	175.0
641367 ^b	-10.4	10.4	-1.0	-1.0
641303	21.8	21.0	88.4	131.2
641339	28.0	29.0	94.4	151.4
641316	26.1	30.1	93.9	150.1

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ FEMALE RATS DURING GESTATION

GROUP: VIII-0

DOSE: 1000 MG/KG/DAY

GESTATION DAYS:	0-7	7-14	14-21	0-21
ANIMAL NUMBER				
641265	21.2	34.9	96.6	152.7
641393	20.7	37.2	94.4	152.3
641402	13.8	21.1	66.4	101.3
641246	14.4	31.7	79.5	125.6
641320	12.0	41.5	97.2	150.7
641353	35.9	36.0	78.2	150.1
641401	28.3	27.1	100.8	156.2
641307	23.2	19.6	88.5	131.3
641368	19.1	31.5	114.0	164.6
641343	20.5	42.7	110.2	173.4
641254	39.0	32.3	99.6	170.9
641242 ^b	-7.0	9.9	0.1	3.0
641284	32.1	28.8	119.8	180.7
641230	17.6	28.7	82.1	128.4
641273 ^b	31.9	10.8	-9.4	33.3
641296	15.8	34.6	91.8	142.2
641248	26.7	35.9	110.2	172.8
641370	26.6	32.8	74.0	133.4
641243	23.4	36.9	97.7	158.0
641329	17.2	31.4	96.1	144.7

APPENDIX Q

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHTS OF P₁ FEMALE RATS DURING LACTATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHTS OF P₁ FEMALE RATS DURING LACTATION

EXPLANATORY NOTES

Footnotes

- a* No litter was delivered; therefore, no lactation data were collected.
- b* Weighing error. Data excluded from statistical analysis.
- c* Animal was found dead on lactation day 13. All pups were cannibalized before lactation day 7. Animal was excluded from statistical analysis.

Abbreviations

--- = No Data

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ FEMALE RATS DURING LACTATION

GROUP: II-0

DOSE: 0 MG/KG/DAY

LACTATION DAYS:	0	7	14	21
ANIMAL NUMBER				
641345	295.3	292.2	315.7	293.4
641236	356.4	361.0	372.3	353.9
641337	394.4	362.3	361.6	354.4
641328	369.9	367.0	380.7	374.2
641310	313.9	299.1	350.9	317.2
641324	314.5	335.9	348.8	334.7
641371	367.4	354.4	344.9	334.8
641332	388.7	369.3	386.8	360.2
641358 ^a	---	---	---	---
641389 ^a	---	---	---	---
641317	357.2	358.5	359.2	360.7
641255	342.1	342.9	359.3	346.8
641403	372.3	378.8	420.8	389.8
641388	382.5	374.0	395.3	384.5
641261	333.1	339.9	346.3	335.2
641306	335.4	335.3	356.5	340.0
641349	354.6	353.6	393.3	361.9
641346	358.3	362.2	365.3	357.6
641308	308.0	320.6	346.2	340.5
641289	348.3	349.9	362.3	352.4

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ FEMALE RATS DURING LACTATION

GROUP: IV-0

DOSE: 50 MG/KG/DAY

LACTATION DAYS:	0	7	14	21
ANIMAL NUMBER				
641378	373.6	376.7	403.9	373.5
641263	321.8	328.5	345.7	323.1
641297	376.7	365.8	381.7	350.7
641304	386.1	341.7	355.4	339.6
641274 ^a	---	---	---	---
641351	317.2	312.9	338.5	329.3
641272	312.4	332.4	328.9	308.0
641258	340.4	354.7	392.3	376.5
641276 ^a	---	---	---	---
641350	335.1	328.1	356.8	335.5
641376	312.4	331.3	345.1	342.6
641315	--- ^b	360.5	376.0	366.1
641286	372.7	343.0	360.4	340.7
641231	328.6	306.0	326.0	318.0
641338	337.8	337.3	342.1	322.7
641318	328.8	321.4	340.5	340.5
641373	378.5	367.6	410.8	396.7
641365	340.4	335.1	351.1	349.3
641348	442.6	419.9	404.5	397.1
641323	354.1	336.5	357.4	346.2

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ FEMALE RATS DURING LACTATION

GROUP: VI-0

DOSE: 250 MG/KG/DAY

LACTATION DAYS:	0	7	14	21
ANIMAL NUMBER				
641271	329.0	348.4	355.5	343.1
641280 ^a	---	---	---	---
641252 ^a	---	---	---	---
641386 ^a	---	---	---	---
641277	392.8	377.8	379.7	355.4
641233	361.1	351.8	374.8	356.9
641238	333.4	318.1	342.6	324.0
641235	301.4	302.0	322.6	319.1
641392 ^c	342.6	329.3	---	---
641360	299.7	322.8	338.3	326.7
641381 ^a	---	---	---	---
641333 ^a	---	---	---	---
641239	368.5	369.8	378.4	388.2
641228 ^a	---	---	---	---
641334	301.0	321.9	343.5	324.0
641347	360.8	374.2	385.6	361.0
641367 ^a	---	---	---	---
641303	315.7	321.0	325.6	318.0
641339	353.4	381.3	389.0	367.5
641316	331.1	348.1	341.2	353.8

INDIVIDUAL BODY WEIGHTS (grams) OF P₁ FEMALE RATS DURING LACTATION

GROUP: VIII-0

DOSE: 1000 MG/KG/DAY

LACTATION DAYS:	0	7	14	21
ANIMAL NUMBER				
641265	339.5	351.5	378.1	360.9
641393	328.3	331.3	328.1	335.0
641402	297.5	300.2	312.8	300.0
641246	297.1	311.2	331.7	310.7
641320	373.2	362.6	384.8	367.4
641353	290.3	284.7	335.4	302.3
641401	326.2	286.6	337.2	320.9
641307	300.4	310.9	334.9	318.3
641368	376.0	364.0	379.6	349.5
641343	393.6	393.6	408.0	393.3
641254	343.5	347.8	375.2	355.5
641242 ^a	---	---	---	---
641284	358.9	358.9	386.0	357.6
641230	320.7	313.8	321.0	318.8
641273 ^a	---	---	---	---
641296	307.9	324.3	328.9	313.1
641248	401.4	390.2	391.9	355.3
641370	352.8	365.0	386.3	357.2
641243	363.0	353.8	363.0	374.3
641329	292.6	322.7	330.2	315.3

APPENDIX R

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHT GAINS OF P₁ FEMALE RATS DURING LACTATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHT GAINS OF P₁ FEMALE RATS DURING LACTATION

EXPLANATORY NOTES

Footnotes

- a* No litter was delivered; therefore, no lactation data were collected.
- b* Weighing error. Data excluded from statistical analysis.
- c* Animal was found dead on lactation day 13. All pups were cannibalized before lactation day 7. Animal was excluded from statistical analysis.

Abbreviations

--- = No Data

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ FEMALE RATS DURING LACTATION

GROUP: II-0

DOSE: 0 MG/KG/DAY

LACTATION DAYS:	0-7	7-14	14-21	0-21
ANIMAL NUMBER				
641345	-3.1	23.5	-22.3	-1.9
641236	4.6	11.3	-18.4	-2.5
641337	-32.1	-0.7	-7.2	-40.0
641328	-2.9	13.7	-6.5	4.3
641310	-14.8	51.8	-33.7	3.3
641324	21.4	12.9	-14.1	20.2
641371	-13.0	-9.5	-10.1	-32.6
641332	-19.4	17.5	-26.6	-28.5
641358 ^a	---	---	---	---
641389 ^a	---	---	---	---
641317	1.3	0.7	1.5	3.5
641255	0.8	16.4	-12.5	4.7
641403	6.5	42.0	-31.0	17.5
641388	-8.5	21.3	-10.8	2.0
641261	6.8	6.4	-11.1	2.1
641306	-0.1	21.2	-16.5	4.6
641349	-1.0	39.7	-31.4	7.3
641346	3.9	3.1	-7.7	-0.7
641308	12.6	25.6	-5.7	32.5
641289	1.6	12.4	-9.9	4.1

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ FEMALE RATS DURING LACTATION

GROUP: IV-0

DOSE: 50 MG/KG/DAY

LACTATION DAYS:	0-7	7-14	14-21	0-21
ANIMAL NUMBER				
641378	3.1	27.2	-30.4	-0.1
641263	6.7	17.2	-22.6	1.3
641297	-10.9	15.9	-31.0	-26.0
641304	-44.4	13.7	-15.8	-46.5
641274 ^a	---	---	---	---
641351	-4.3	25.6	-9.2	12.1
641272	20.0	-3.5	-20.9	-4.4
641258	14.3	37.6	-15.8	36.1
641276 ^a	---	---	---	---
641350	-7.0	28.7	-21.3	0.4
641376	18.9	13.8	-2.5	30.2
641315	--- ^b	15.5	-9.9	--- ^b
641286	-29.7	17.4	-19.7	-32.0
641231	-22.6	20.0	-8.0	-10.6
641338	-0.5	4.8	-19.4	-15.1
641318	-7.4	19.1	0.0	11.7
641373	-10.9	43.2	-14.1	18.2
641365	-5.3	16.0	-1.8	8.9
641348	-22.7	-15.4	-7.4	-45.5
641323	-17.6	20.9	-11.2	-7.9

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ FEMALE RATS DURING LACTATION

GROUP: VI-0

DOSE: 250 MG/KG/DAY

LACTATION DAYS:	0-7	7-14	14-21	0-21
ANIMAL NUMBER				
641271	19.4	7.1	-12.4	14.1
641280 ^a	---	---	---	---
641252 ^a	---	---	---	---
641386 ^a	---	---	---	---
641277	-15.0	1.9	-24.3	-37.4
641233	-9.3	23.0	-17.9	-4.2
641238	-15.3	24.5	-18.6	-9.4
641235	0.6	20.6	-3.5	17.7
641392 ^c	-13.3	---	---	---
641360	23.1	15.5	-11.6	27.0
641381 ^a	---	---	---	---
641333 ^a	---	---	---	---
641239	1.3	8.6	9.8	19.7
641228 ^a	---	---	---	---
641334	20.9	21.6	-19.5	23.0
641347	13.4	11.4	-24.6	0.2
641367 ^a	---	---	---	---
641303	5.3	4.6	-7.6	2.3
641339	27.9	7.7	-21.5	14.1
641316	17.0	-6.9	12.6	22.7

INDIVIDUAL BODY WEIGHT GAINS (grams) OF P₁ FEMALE RATS DURING LACTATION

GROUP: VIII-0

DOSE: 1000 MG/KG/DAY

LACTATION DAYS:	0-7	7-14	14-21	0-21
ANIMAL NUMBER				
641265	12.0	26.6	-17.2	21.4
641393	3.0	-3.2	6.9	6.7
641402	2.7	12.6	-12.8	2.5
641246	14.1	20.5	-21.0	13.6
641320	-10.6	22.2	-17.4	-5.8
641353	-5.6	50.7	-33.1	12.0
641401	-39.6	50.6	-16.3	-5.3
641307	10.5	24.0	-16.6	17.9
641368	-12.0	15.6	-30.1	-26.5
641343	0.0	14.4	-14.7	-0.3
641254	4.3	27.4	-19.7	12.0
641242 ^a	---	---	---	---
641284	0.0	27.1	-28.4	-1.3
641230	-6.9	7.2	-2.3	-2.0
641273 ^a	---	---	---	---
641296	16.4	4.6	-15.8	5.2
641248	-11.2	1.7	-36.6	-46.1
641370	12.2	21.3	-29.1	4.4
641243	-9.2	9.2	11.3	11.3
641329	30.1	7.5	-14.9	22.7

APPENDIX S

REPRODUCTIVE TOXICOLOGY EVALUATIONS

**INDIVIDUAL DAILY FOOD CONSUMPTION BY P₁ FEMALE RATS
DURING GESTATION**

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL DAILY FOOD CONSUMPTION BY P₁ FEMALE RATS DURING GESTATION

EXPLANATORY NOTES

Footnotes

- a* No litter was delivered; data were excluded from statistical analysis.
- b* No gestation data were collected since no plug was observed; however, a litter was delivered.
- c* No gestation data were collected since no plug was observed; no litter was delivered.
- d* Weighing error; data were excluded from statistical analysis.

Abbreviations

--- = No data

INDIVIDUAL DAILY FOOD CONSUMPTION (grams) BY P₁ FEMALE RATS
DURING GESTATION

GROUP: II-0

DOSE: 0 MG/KG/DAY

GESTATION DAYS:	0-7	7-14	0-14
ANIMAL NUMBER			
641345	23.0	22.7	22.9
641236	26.1	29.6	27.8
641337	25.6	29.9	27.8
641328	24.5	29.5	27.0
641310	23.0	24.3	23.7
641324	23.3	27.2	25.3
641371	24.0	27.1	25.6
641332	30.1	33.2	31.7
641358 ^a	19.7	20.3	20.0
641389 ^a	21.1	20.1	20.6
641317 ^b	---	---	---
641255	24.2	25.7	24.9
641403	24.7	29.8	27.3
641388	25.7	27.9	26.8
641261	22.9	25.4	24.1
641306	24.8	25.0	24.9
641349 ^b	---	---	---
641346	27.5	27.9	27.7
641308	25.3	26.9	26.1
641289	24.0	26.4	25.2

INDIVIDUAL DAILY FOOD CONSUMPTION (grams) BY P₁ FEMALE RATS
DURING GESTATION

GROUP: IV-0

DOSE: 50 MG/KG/DAY

GESTATION DAYS:	0-7	7-14	0-14
ANIMAL NUMBER			
641378	28.2	30.1	29.2
641263	23.9	26.6	25.2
641297	24.1	28.2	26.1
641304	26.6	26.1	26.4
641274 ^a	27.2	24.0	25.6
641351	20.1	22.0	21.1
641272	25.7	26.4	26.0
641258	24.8	29.6	27.2
641276 ^a	26.4	28.7	27.6
641350	19.7	24.0	21.9
641376	22.0	22.4	22.2
641315	26.5	25.8	26.1
641286	24.9	28.8	26.8
641231	22.3	24.5	23.4
641338	26.4	28.1	27.3
641318	22.0	22.4	22.2
641373	31.1	33.7	32.4
641365	24.0	23.6	23.8
641348	31.1	33.5	32.3
641323	27.0	28.1	27.6

INDIVIDUAL DAILY FOOD CONSUMPTION (grams) BY P₁ FEMALE RATS
DURING GESTATION

GROUP: VI-0

DOSE: 250 MG/KG/DAY

GESTATION DAYS:	0-7	7-14	0-14
ANIMAL NUMBER			
641271	21.1	23.2	22.2
641280 ^a	24.3	26.3	25.3
641252 ^a	23.9	22.8	23.4
641386 ^a	21.0	23.8	22.4
641277	24.9	29.1	27.0
641233	25.5	29.4	27.4
641238	22.0	26.8	24.4
641235	21.0	21.3	21.1
641392	23.4	25.4	24.4
641360	20.4	20.9	20.6
641381 ^a	27.0	25.7	26.4
641333 ^c	---	---	---
641239	24.7	26.8	25.8
641228 ^a	23.7	29.4	26.6
641334	23.0	--- ^d	--- ^d
641347	27.7	29.6	28.7
641367 ^a	20.2	31.0	25.6
641303	24.1	24.2	24.2
641339	26.3	28.1	27.2
641316	25.2	25.9	25.6

INDIVIDUAL DAILY FOOD CONSUMPTION (grams) BY P₁ FEMALE RATS
DURING GESTATION

GROUP: VIII-0

DOSE: 1000 MG/KG/DAY

GESTATION DAYS:	0-7	7-14	0-14
ANIMAL NUMBER			
641265	21.0	25.3	23.2
641393	25.8	28.4	27.1
641402	20.3	20.3	20.3
641246	18.4	22.2	20.3
641320	22.6	25.7	24.1
641353	25.0	25.6	25.3
641401	23.1	24.3	23.7
641307	20.7	19.2	20.0
641368	24.9	26.0	25.5
641343	27.6	29.2	28.4
641254	24.9	25.8	25.4
641242 ^a	20.6	22.9	21.7
641284	23.9	24.8	24.4
641230	20.3	22.8	21.5
641273 ^a	24.7	25.5	25.1
641296	21.2	22.0	21.6
641248	27.4	31.0	29.2
641370	28.0	25.6	26.8
641243	23.6	27.2	25.4
641329	21.7	23.5	22.6

APPENDIX T
REPRODUCTIVE TOXICOLOGY EVALUATIONS
INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ MALE RATS

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ MALE RATS

GROUP: I-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641114	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641217	ALOPECIA LEFT LEG(S) SACRIFICED BY DESIGN TEST DAY 107	100	107
641204	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641158	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641100	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641060	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 108	100	108
641188	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 108	78	108
641106	SORE TAIL SACRIFICED BY DESIGN TEST DAY 108	100	108
641175	ALOPECIA RIGHT FRONT PAW(S) ALOPECIA RIGHT FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 108	85 107	108 108
641079	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641125	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641186	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641052	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 109	78	109
641140	ALOPECIA RIGHT FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 109	109	109
641135	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ MALE RATS

GROUP: I-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641174	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641107	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641072	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641053	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641168	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ MALE RATS

GROUP: III-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641087	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 107	78	107
641108	LUNG NOISE SACRIFICED BY DESIGN TEST DAY 107	85	85
641118	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641128	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641091	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641199	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641209	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641163	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641129	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641138	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641103	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641181	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641171	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641111	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 109	78	109
641062	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ MALE RATS

GROUP: III-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641156	ALOPECIA RIGHT FRONT LEG(S) ALOPECIA BOTH FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 110	78 85	78 110
641173	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641121	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641074	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641191	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ MALE RATS

GROUP: V-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641170	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641065	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641126	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641058	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641131	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641176	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641119	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641075	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641180	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641117	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641124	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641082	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 109	78	100
641151	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641195	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641122	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ MALE RATS

GROUP: V-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641184	ALOPECIA LEFT FRONT LEG(S) SORE LEFT FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 110	107 110	110 110
641142	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641120	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641179	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641164	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ MALE RATS

GROUP: VII-0 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641110	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641096	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641212	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641068	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641093	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 107		
641054	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641208	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641193	ALOPECIA BOTH FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 108	78	108
641207	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 108	78	108
641139	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 108		
641080	ALOPECIA LEFT FRONT PAW(S) ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 109	78 85	78 109
641078	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641177	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641112	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		
641210	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 109		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ MALE RATS

GROUP: VII-0 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641152	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641182	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641150	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641211	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 110		
641145	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 110	85	110

APPENDIX U

REPRODUCTIVE TOXICOLOGY EVALUATIONS

**INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS
DURING GESTATION**

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS
DURING GESTATION

EXPLANATORY NOTES

Notes

This appendix includes only signs from rats that delivered a litter.

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING GESTATION

GROUP: II-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641345	NO ABNORMALITIES DETECTED		
641236	NO ABNORMALITIES DETECTED		
641337	NO ABNORMALITIES DETECTED		
641328	NO ABNORMALITIES DETECTED		
641310	NO ABNORMALITIES DETECTED		
641324	NO ABNORMALITIES DETECTED		
641371	NO ABNORMALITIES DETECTED		
641332	ALOPECIA BOTH FRONT LEG(S)	0	21
641317	NO ABNORMALITIES DETECTED		
641255	NO ABNORMALITIES DETECTED		
641403	NO ABNORMALITIES DETECTED		
641388	NO ABNORMALITIES DETECTED		
641261	NO ABNORMALITIES DETECTED		
641306	NO ABNORMALITIES DETECTED		
641349	NO ABNORMALITIES DETECTED		
641346	ALOPECIA ABDOMEN	21	21
641308	NO ABNORMALITIES DETECTED		
641289	NO ABNORMALITIES DETECTED		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING GESTATION

GROUP: IV-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641378	ALOPECIA RIGHT SIDE(S)	7	21
641263	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S)	0 14	14 14
641297	NO ABNORMALITIES DETECTED		
641304	NO ABNORMALITIES DETECTED		
641351	NO ABNORMALITIES DETECTED		
641272	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S)	0 7	21 21
641258	NO ABNORMALITIES DETECTED		
641350	ALOPECIA LEFT FRONT LEG(S)	21	21
641376	NO ABNORMALITIES DETECTED		
641315	NO ABNORMALITIES DETECTED		
641286	NO ABNORMALITIES DETECTED		
641231	NO ABNORMALITIES DETECTED		
641338	NO ABNORMALITIES DETECTED		
641318	NO ABNORMALITIES DETECTED		
641373	ALOPECIA BOTH FRONT LEG(S) ALOPECIA BOTH HIND QUARTERS ALOPECIA BASE TAIL ALOPECIA ABDOMEN	0 0 0 21	21 21 21 21
641365	NO ABNORMALITIES DETECTED		
641348	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S)	7 14	21 21
641323	NO ABNORMALITIES DETECTED		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING GESTATION

GROUP: VI-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641271	NO ABNORMALITIES DETECTED		
641277	NO ABNORMALITIES DETECTED		
641233	ALOPECIA BOTH FRONT LEG(S) ALOPECIA BOTH FRONT PAW(S)	0 0	21 21
641238	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S)	7 14	21 21
641235	NO ABNORMALITIES DETECTED		
641392	NO ABNORMALITIES DETECTED		
641360	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S)	0 0	21 21
641239	NO ABNORMALITIES DETECTED		
641334	NO ABNORMALITIES DETECTED		
641347	NO ABNORMALITIES DETECTED		
641303	NO ABNORMALITIES DETECTED		
641339	NO ABNORMALITIES DETECTED		
641316	ALOPECIA BOTH FRONT PAW(S)	7	21

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING GESTATION

GROUP: VIII-0 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641265	NO ABNORMALITIES DETECTED		
641393	NO ABNORMALITIES DETECTED		
641402	NO ABNORMALITIES DETECTED		
641246	ALOPECIA BOTH FRONT LEG(S) ALOPECIA BOTH FRONT PAW(S)	7 14	21 21
641320	NO ABNORMALITIES DETECTED		
641353	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S)	0 21	21 21
641401	NO ABNORMALITIES DETECTED		
641307	NO ABNORMALITIES DETECTED		
641368	NO ABNORMALITIES DETECTED		
641343	NO ABNORMALITIES DETECTED		
641254	NO ABNORMALITIES DETECTED		
641284	NO ABNORMALITIES DETECTED		
641230	NO ABNORMALITIES DETECTED		
641296	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S)	7 21	21 21
641248	NO ABNORMALITIES DETECTED		
641370	NO ABNORMALITIES DETECTED		
641243	ALOPECIA BOTH FRONT PAW(S)	7	21
641329	ALOPECIA LEFT FRONT LEG(S)	14	21

APPENDIX V

REPRODUCTIVE TOXICOLOGY EVALUATIONS

**INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS
DURING LACTATION**

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING LACTATION

EXPLANATORY NOTES

Note

This appendix includes signs from lactating rats only.

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING LACTATION

GROUP: II-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641345	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 114		
641236	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 118		
641337	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 116		
641328	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 115		
641310	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 114		
641324	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 116		
641371	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 114		
641332	ALOPECIA BOTH FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 117	0	21
641317	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 115		
641255	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 118		
641403	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 116		
641388	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 117		
641261	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 117		
641306	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 114		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING LACTATION

GROUP: II-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641349	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 114		
641346	ALOPECIA ABDOMEN ALOPECIA LEFT FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 115	0 14	21 21
641308	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 118		
641289	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 116		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING LACTATION

GROUP: IV-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641378	ALOPECIA RIGHT SIDE(S) SACRIFICED BY DESIGN TEST DAY 114	0	14
641263	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 114	0 0	21 21
641297	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 115		
641304	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 115		
641351	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 114		
641272	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 115	0 0	21 21
641258	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 126		
641350	ALOPECIA LEFT FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 118	0	21
641376	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 118		
641315	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 120		
641286	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 118		
641231	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 117		
641338	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 115		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING LACTATION

GROUP: IV-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641318	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 117		
641373	ALOPECIA BOTH FRONT LEG(S) ALOPECIA BOTH HIND QUARTERS ALOPECIA BASE TAIL ALOPECIA ABDOMEN SACRIFICED BY DESIGN TEST DAY 118	0 0 0 0	21 21 21 21
641365	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 115		
641348	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 114	0 0	21 21
641323	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 113		

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING LACTATION

GROUP: VI-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641271	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 114		
641277	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 113		
641233	ALOPECIA BOTH FRONT LEG(S) ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 114	0 0	21 21
641238	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 114	0 0	21 21
641235	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 114		
641392	FOUND DEAD TEST DAY 107	13	13
641360	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 117	0 0	21 21
641239	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 116		
641334	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 124		
641347	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 118		
641303	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 116		
641339	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 115		
641316	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 114	0	21

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING LACTATION

GROUP: VIII-0 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641265	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 117		
641393	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 115		
641402	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 116		
641246	ALOPECIA BOTH FRONT LEG(S) ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 115	0 0	21 21
641320	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 118		
641353	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 114	0 0	21 21
641401	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 113		
641307	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 114		
641368	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 115		
641343	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 116		
641254	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 115		
641284	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 114		
641230	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 117	14	21

INDIVIDUAL CLINICAL OBSERVATIONS IN P₁ FEMALE RATS DURING LACTATION

GROUP: VIII-0 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
641296	ALOPECIA BOTH FRONT PAW(S) ALOPECIA BOTH FRONT LEG(S) SACRIFICED BY DESIGN TEST DAY 114	0 0	21 21
641248	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 118		
641370	NO ABNORMALITIES DETECTED SACRIFICED BY DESIGN TEST DAY 118		
641243	ALOPECIA BOTH FRONT PAW(S) SACRIFICED BY DESIGN TEST DAY 114	0	21
641329	ALOPECIA LEFT FRONT LEG(S) MASS 1 3.5 cm RIGHT SIDE(S) MASS 2 2.0 cm LEFT SIDE(S) SACRIFICED BY DESIGN TEST DAY 117	0 14 14	21 14 14

APPENDIX W

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL ANIMAL ESTROUS CYCLE PARAMETERS
DURING PREMATING AND COHABITATION IN P₁ FEMALE RATS

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL ANIMAL ESTROUS CYCLE PARAMETERS DURING PREMATING AND COHABITATION IN P₁ FEMALE RATS

EXPLANATORY NOTES

Footnotes:

- a* No evidence of copulation was found, but animal was pregnant. Data were excluded from statistical analysis since the animal was pregnant during part of the estrous cycle evaluation.
- b* No precoital interval could be calculated since no evidence of copulation was found.
- c* Mean cycle length was not calculable due to insufficient data.

Abbreviations:

--- = No Data

INDIVIDUAL ANIMAL ESTROUS CYCLE PARAMETERS
DURING PREMATING AND COHABITATION

GROUP II-0: 0 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF DAYS OF DATA COLLECTION	PERCENT OF DAYS IN ESTRUS	PERCENT OF DAYS IN DIESTRUS	PERCENT OF DAYS IN PROESTRUS	MEAN CYCLE LENGTH (days)	PRECOITAL INTERVAL (days)
641389	22	5%	82%	14%	4.50	1
641358	22	23%	64%	14%	4.00	1
641332	25	24%	64%	12%	4.00	4
641371	22	23%	77%	0%	4.00	1
641324	23	22%	65%	13%	5.00	2
641310	21	24%	71%	5%	4.00	1
641328	23	13%	74%	13%	5.33	2
641337	23	4%	74%	22%	4.67	3
641236	26	23%	58%	19%	4.75	5
641345	23	22%	61%	17%	4.50	2
641289	25	8%	76%	16%	6.67	4
641308	26	31%	65%	4%	4.20	5
641346	23	13%	74%	13%	4.00	2
641349 ^a	35	14%	80%	6%	4.50	--- ^b
641306	22	18%	59%	23%	5.00	1
641261	25	24%	72%	4%	4.00	4
641388	26	12%	88%	0%	10.50	5
641403	24	25%	50%	25%	4.00	3
641255	26	31%	62%	8%	4.75	5
641317 ^a	35	9%	74%	17%	4.00	--- ^b

INDIVIDUAL ANIMAL ESTROUS CYCLE PARAMETERS
DURING PREMATING AND COHABITATION

GROUP IV-0: 50 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF DAYS OF DATA COLLECTION	PERCENT OF DAYS IN ESTRUS	PERCENT OF DAYS IN DIESTRUS	PERCENT OF DAYS IN PROESTRUS	MEAN CYCLE LENGTH (days)	PRECOITAL INTERVAL (days)
641350	25	12%	76%	12%	5.00	5
641276	24	25%	67%	8%	4.75	4
641258	34	9%	76%	15%	5.40	13
641272	22	18%	68%	14%	5.00	2
641351	22	18%	68%	14%	4.00	2
641274	29	24%	69%	7%	6.67	8
641304	22	32%	50%	18%	4.00	1
641297	23	26%	74%	0%	4.00	2
641263	23	13%	87%	0%	---C	2
641378	21	10%	71%	19%	---C	1
641323	22	23%	77%	0%	4.00	1
641348	22	18%	68%	14%	5.00	1
641365	23	22%	61%	17%	4.00	2
641373	26	23%	65%	12%	4.75	5
641318	25	40%	48%	12%	5.25	4
641338	23	22%	61%	17%	5.00	2
641231	25	28%	52%	20%	4.20	4
641286	26	19%	65%	15%	5.00	5
641315	28	32%	64%	4%	4.20	7
641376	25	28%	64%	8%	5.67	5

INDIVIDUAL ANIMAL ESTROUS CYCLE PARAMETERS
DURING PREMATING AND COHABITATION

GROUP VI-0: 250 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF DAYS OF DATA COLLECTION	PERCENT OF DAYS IN ESTRUS	PERCENT OF DAYS IN DIESTRUS	PERCENT OF DAYS IN PROESTRUS	MEAN CYCLE LENGTH (days)	PRECOITAL INTERVAL (days)
641360	24	21%	54%	25%	4.00	4
641392	23	26%	61%	13%	4.00	2
641235	23	22%	52%	26%	4.00	2
641238	22	32%	50%	18%	4.50	1
641233	22	23%	68%	9%	4.00	1
641277	22	18%	73%	9%	4.67	1
641386	22	27%	68%	5%	4.00	1
641252	22	23%	64%	14%	4.00	1
641280	22	27%	55%	18%	4.00	1
641271	22	14%	68%	18%	5.00	1
641316	22	23%	68%	9%	4.00	1
641339	23	26%	70%	4%	4.00	2
641303	24	21%	54%	25%	4.00	3
641367	27	19%	70%	11%	4.60	6
641347	25	28%	48%	24%	4.00	4
641334	31	35%	61%	3%	5.00	10
641228	25	4%	68%	28%	7.00	4
641239	24	13%	75%	13%	6.33	3
641333	33	15%	79%	6%	4.00	---b
641381	23	26%	61%	13%	5.67	2

INDIVIDUAL ANIMAL ESTROUS CYCLE PARAMETERS
DURING PREMATING AND COHABITATION

GROUP VIII-0: 1000 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF DAYS OF DATA COLLECTION	PERCENT OF DAYS IN ESTRUS	PERCENT OF DAYS IN DIESTRUS	PERCENT OF DAYS IN PROESTRUS	MEAN CYCLE LENGTH (days)	PRECOITAL INTERVAL (days)
641343	24	13%	75%	13%	5.00	3
641368	23	26%	70%	4%	4.25	2
641307	22	23%	77%	0%	4.25	1
641401	22	27%	64%	9%	4.00	1
641353	22	23%	55%	23%	4.00	1
641320	26	12%	77%	12%	5.67	5
641246	22	27%	50%	23%	3.67	2
641402	22	23%	73%	5%	---	3
641393	23	22%	74%	4%	4.00	2
641265	25	24%	64%	12%	4.00	4
641329	25	24%	72%	4%	4.00	4
641243	22	41%	45%	14%	5.67	1
641370	26	15%	77%	8%	5.00	5
641248	26	27%	54%	19%	5.00	5
641296	21	5%	76%	19%	5.00	1
641273	22	36%	55%	9%	5.33	1
641230	24	29%	50%	21%	4.75	4
641284	22	36%	64%	0%	4.67	1
641242	27	33%	67%	0%	4.60	6
641254	23	22%	52%	26%	4.00	2

APPENDIX X

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL ANIMAL ESTROUS CYCLE STAGES
DURING PREMATING AND COHABITATION IN P₁ FEMALE RATS

REPRODUCTIVE TOXICOLOGY EVALUATIONS
INDIVIDUAL ANIMAL ESTROUS CYCLE STAGES
DURING PREMATING AND COHABITATION IN P₁ FEMALE RATS
EXPLANATORY NOTES

Notes:

Refer to footnotes located in Individual Animal Estrous Cycle Parameters During Premating and Cohabitation appendix for individual animal variation.

Test Days 50-70 = Premating
Test Days 71-85 = Cohabitation period

Abbreviations:

D = Diestrus
E = Estrus
P = Proestrus
X = No sample or stage could not be defined from sample.

H-24616: Subchronic Toxicity
90-Day Gavage Study in Rats with One-Generation Reproductive Evaluations

DuPont-4739

INDIVIDUAL ANIMAL ESTROUS CYCLE STAGES DURING PREMATING AND COHABITATION

GROUP II-0: 0 MG/KG/DAY

ANIMAL NUMBER	TEST DAYS																																			
	49	50	51	52	53	54	55	56	57	58	59	60	61	62	63	64	65	66	67	68	69	70	71	72	73	74	75	76	77	78	79	80	81	82	83	
641389	D	P	D	D	D	P	D	D	D	P	E	D	D	D	D	D	D	D	D	D	D	D	D													
641358	D	D	D	E	D	D	P	E	D	D	P	E	D	D	D	E	D	D	P	E	D	D														
641332	P	E	D	D	D	E	D	D	D	E	D	D	D	E	D	D	P	E	D	D	D	E	D	D	P											
641371	D	D	E	D	D	D	E	D	D	D	E	D	D	D	E	D	D	D	E	D	D	D														
641324	D	D	D	P	D	D	P	E	E	D	D	P	E	D	D	D	D	E	E	D	D	D	D													
641310	P	X	E	D	D	D	E	D	D	D	E	D	D	D	E	D	D	D	E	D	D	D	P													
641328	D	E	D	D	D	D	P	D	D	D	D	D	D	D	D	D	P	D	D	D	P	E	D	D												
641337	P	X	D	D	D	D	P	D	D	D	P	E	D	D	D	D	P	E	D	D	P	E	D	D	D	P										
641236	P	E	D	D	D	E	E	D	D	P	E	D	D	D	D	P	E	E	D	D	E	E	D													
641345	D	D	D	P	D	D	P	D	D	P	E	D	D	D	D	D	E	D	D	D	E	P	D	D	D											
641289	D	E	D	D	P	E	D	D	D	D	P	D	D	D	D	D	D	D	D	D	P	P	D	D	D	D										
641308	E	E	D	D	D	E	E	D	D	D	E	D	D	D	D	E	D	D	P	E	D	D	E	D	D	D	D									
641346	D	D	D	E	D	D	D	E	D	D	D	P	D	D	D	D	P	D	D	D	E	D	D	P												
641349	D	D	P	E	D	D	D	E	D	D	D	D	E	D	D	D	D	E	D	D	P	E	D	D	D	D	D	D	D	D	D	D	D	D	D	
641306	D	P	E	D	D	D	P	E	D	D	P	E	D	D	D	D	D	P	E	D	D	D	P													
641261	D	E	D	D	D	E	D	D	P	E	D	D	D	D	E	D	D	E	D	D	D	E	D	D	D	D										
641388	D	D	E	D	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	
641403	E	D	D	P	E	D	D	P	E	D	D	P	E	D	D	D	P	E	D	D	P	E	D	D	D	D	E									
641255	D	P	E	D	D	D	E	E	D	D	D	D	D	D	D	D	D	E	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D	
641317	D	D	D	D	D	D	P	P	D	D	P	E	D	D	D	P	E	D	D	P	E	D	D	P	D	D	D	D	D	D	D	D	D	D	D	D

INDIVIDUAL ANIMAL ESTROUS CYCLE STAGES DURING PREMATING AND COHABITATION

GROUP IV-0: 50 MG/KG/DAY

ANIMAL NUMBER	TEST DAYS																																	
	49	50	51	52	53	54	55	56	57	58	59	60	61	62	63	64	65	66	67	68	69	70	71	72	73	74	75	76	77	78	79	80	81	82
641350	D	E	D	D	D	E	D	D	D	E	D	D	D	D	P	D	D	D	D	D	D	P	D	D	D	D	P							
641276	E	E	D	D	D	X	E	D	D	D	E	D	D	D	D	P	E	D	D	D	P	D	D	D	D									
641258	D	D	P	E	D	D	P	E	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	P	D	D	D
641272	D	D	P	E	D	D	D	E	D	D	D	D	D	P	X	D	D	P	E	D	D	D	D	D										
641351	P	E	D	D	D	E	D	D	D	E	D	D	D	D	P	D	D	E	D	D	D	D	P											
641274	E	E	E	D	D	D	D	E	E	E	D	D	D	D	D	D	D	E	D	D	D	D	P	D	D	D	D	D	D					
641304	D	P	E	E	D	D	P	E	D	D	P	E	D	D	P	E	E	D	D	D	D	D												
641297	D	D	E	D	D	D	E	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	E											
641263	D	D	D	E	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D											
641378	D	E	E	D	D	D	P	X	D	D	D	D	D	P	D	D	D	D	D	D	D	D	P											
641323	D	D	E	D	D	D	E	D	D	D	E	D	D	D	D	E	D	D	D	D	D	D	D											
641348	D	P	E	D	D	D	P	E	D	D	D	P	E	D	D	D	D	E	D	D	D	D	P											
641365	D	D	P	E	D	D	P	E	D	D	D	P	E	D	D	D	D	D	D	E	D	D	P											
641373	D	D	E	D	D	D	P	E	D	D	D	E	D	D	D	D	P	E	D	D	D	D	D											
641318	E	D	D	D	E	E	D	D	P	E	E	E	E	E	E	D	D	E	D	D	D	D	P	D	D	D	E							
641338	D	D	E	E	D	D	D	P	E	D	D	D	D	P	E	D	D	P	E	D	D	D	D											
641231	E	D	D	P	E	D	D	P	E	D	D	D	E	D	D	P	E	D	D	D	P	E	D	D	D	P	D							
641286	P	E	D	D	D	D	E	D	D	D	P	E	D	D	D	P	E	D	D	D	D	P	D	D	D	D	D							
641315	D	D	E	D	D	D	E	E	D	D	D	E	E	D	D	D	E	E	D	D	D	D	D	E	D	D	D	D	E					
641376	D	D	P	E	X	D	D	D	D	E	D	D	D	D	D	E	D	D	D	P	E	D	D	D	D	D	D	E						

INDIVIDUAL ANIMAL ESTROUS CYCLE STAGES DURING PREMATING AND COHABITATION

GROUP VI-0: 250 MG/KG/DAY

ANIMAL NUMBER	TEST DAYS																																		
	49	50	51	52	53	54	55	56	57	58	59	60	61	62	63	64	65	66	67	68	69	70	71	72	73	74	75	76	77	78	79	80	81	82	83
641360	P	E	D	D	P	X	D	D	P	E	D	D	P	E	D	D	P	E	D	D	D	D	E	D	P										
641392	D	D	P	E	D	D	P	E	D	D		D	D	D	P	E	D	D	D	E	D	D	D												
641235	D	D	P	E	D	D	P	E	D	D	P	E	D	D	P	E	D	D	P	E	D	D	P												
641238	E	D	D	P	E	E	D	D	P	E	D	D	P	E	D	D	D	D	P	E	D	D	E												
641233	D	P	E	D	D	D	E	D	D	P	E	D	D	D	D	E	D	D	D	E	D	D	D												
641277	P	E	D	D	P	E	D	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D												
641386	D	E	D	D	D	E	D	D	P	E	D	D	D	D	E	D	D	D	D	D	D	E													
641252	D	E	D	D	P	E	D	D	P	E	D	D	D	E	D	D	D	E	D	D	D	P													
641280	D	P	E	D	D	P	E	D	D	E	E	D	D	D	P	E	D	D	P	E	D	D	D												
641271	D	P	E	D	D	D	P	D	D	D	D	P	D	D	D	D	E	E	D	D	D	P													
641316	D	P	E	D	D	D	E	D	D	D	E	D	D	D	P	E	D	D	D	E	D	D													
641339	D	D	D	E	D	D	D	E	D	D	P	E	D	D	D	E	D	D	D	D	D	D	E												
641303	E	D	D	D	P	D	D	P	E	D	D	P	E	D	D	P	E	D	D	P	E	D	D	P											
641367	P	D	D	D	D	D	E	D	D	D	E	D	D	D	E	D	D	D	P	E	D	D	D	P											
641347	P	E	D	D	P	E	D	D	P	E	D	D	P	E	D	D	E	E	D	D	D	P	D	D	E										
641334	E	E	D	D	D	E	E	D	D	D	E	D	D	D	D	E	E	D	D	D	E	D	D	D	D	D	E	D	D	D	D	D	D	D	
641228	D	D	D	P	P	D	D	D	D	D	P	P	P	D	D	D	D	D	P	E	D	D	D	D	P										
641239	E	D	D	D	P	D	D	D	D	D	D	D	D	P	P	D	D	D	D	E	D	D	D	E											
641333	D	X	X	P	D	D	D	E	D	D	P	E	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	
641381	D	D	D	E	D	D	P	E	E	E	D	D	D	D	D	P	E	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D

H-24616: Subchronic Toxicity
90-Day Gavage Study in Rats with One-Generation Reproductive Evaluations

DuPont-4739

INDIVIDUAL ANIMAL ESTROUS CYCLE STAGES DURING PREMATING AND COHABITATION

GROUP VIII-0: 1000 MG/KG/DAY

ANIMAL NUMBER	TEST DAYS																										
	49	50	51	52	53	54	55	56	57	58	59	60	61	62	63	64	65	66	67	68	69	70	71	72	73	74	75
641343	D	D	P	E	D	D	D	D	P	D	D	P	E	E	D	D	D	D	D	D	D	D	D	D			
641368	D	P	E	D	D	D	E	E	D	D	D	E	D	D	D	D	D	D	D	E	D	D	D				
641307	D	E	D	D	D	D	E	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D				
641401	D	E	D	D	D	D	E	D	D	E	D	D	P	E	D	D	D	D	D	D	D	D	D				
641353	D	P	E	D	D	D	E	D	D	P	E	D	D	D	D	D	D	D	D	D	D	D	D				
641320	P	E	D	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	
641246	D	D	P	E	X	D	D	P	E	E	D	D	D	D	D	D	D	D	D	P	E	D	D				
641402	D	E	D	D	D	E	X	D	D	D	P	X	D	D	D	D	D	D	D	D	E	D	D	D			
641393	D	D	D	E	D	D	D	E	D	D	P	E	D	D	D	D	D	D	D	D	D	D	D				
641265	E	D	D	D	D	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D		
641329	D	E	D	D	D	P	E	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D		
641243	E	E	D	D	D	P	E	D	D	D	E	E	E	E	D	D	P	E	E	D	D	D	D	D	D		
641370	D	D	P	E	D	D	D	D	P	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	
641248	E	D	D	P	E	E	E	D	D	P	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	
641296	D	P	D	D	D	D	X	P	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	
641273	E	D	D	D	D	D	E	D	D	D	E	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	
641230	P	E	D	D	D	D	E	X	D	P	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	
641284	D	D	E	D	D	D	E	E	D	D	D	E	E	D	D	D	D	D	D	D	D	D	D	D	D	D	
641242	E	E	D	D	D	E	E	D	D	D	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	D	
641254	D	D	P	E	D	D	P	E	D	D	P	E	D	D	D	D	D	D	D	D	D	D	D	D	D	D	

APPENDIX Y

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL ANIMAL SPERM MOTILITY DATA IN P₁ MALE RATS

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL ANIMAL SPERM MOTILITY DATA IN P₁ MALE RATS

EXPLANATORY NOTES

Footnotes:

a No sperm. Data excluded from calculations.

INDIVIDUAL ANIMAL SPERM MOTILITY DATA IN P₁ MALE RATS

GROUP I-0 0 MG/KG/DAY		GROUP III-0 50 MG/KG/DAY	
Animal Number	Percent Motile	Animal Number	Percent Motile
641114	90	641087	96
641217	97	641108	89
641204	93	641118	87
641158	95	641128	94
641100	96	641091	96
641060	86	641199	90
641188	91	641209	95
641106	92	641163	95
641175	94	641129	92
641079	95	641138	95

GROUP V-0 250 MG/KG/DAY		GROUP VII-0 1000 MG/KG/DAY	
Animal Number	Percent Motile	Animal Number	Percent Motile
641170	92	641110	95
641065	91	641096	91
641126	90	641212	93
641058	93	641068	92
641131	92	641093 ^a	---
641176	95	641054	75
641119	91	641208	97
641075	95	641193	92
641180	94	641207	85
641117	91	641139	91

APPENDIX Z

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL ANIMAL SPERM MORPHOLOGY DATA IN P₁ MALE RATS

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL ANIMAL SPERM MORPHOLOGY DATA IN P₁ MALE RATS

EXPLANATORY NOTES

Footnotes:

a No sperm. Data excluded from calculations.

INDIVIDUAL ANIMAL SPERM MORPHOLOGY DATA IN P₁ MALE RATS

GROUP I-0: 0 MG/KG/DAY

Animal Number	Total Normal	Tail Abnormal	Head Abnormal	Tail & Head Abnormal	Percent Normal
641114	199	0	1	0	99.5
641217	196	2	2	0	98.0
641204	200	0	0	0	100.0
641158	194	3	3	0	97.0
641100	198	0	2	0	99.0
641060	198	1	1	0	99.0
641188	196	2	2	0	98.0
641106	196	1	3	0	98.0
641175	199	1	0	0	99.5
641079	195	1	4	0	97.5

GROUP III-0: 50 MG/KG/DAY

Animal Number	Total Normal	Tail Abnormal	Head Abnormal	Tail & Head Abnormal	Percent Normal
641087	195	1	4	0	97.5
641108	199	0	1	0	99.5
641118	193	0	7	0	96.5
641128	194	4	2	0	97.0
641091	199	1	0	0	99.5
641199	194	0	6	0	97.0
641209	195	1	4	0	97.5
641163	196	1	3	0	98.0
641129	198	2	0	0	99.0
641138	198	1	1	0	99.0

INDIVIDUAL ANIMAL SPERM MORPHOLOGY DATA IN P₁ MALE RATS

GROUP V-0: 250 MG/KG/DAY

Animal Number	Total Normal	Tail Abnormal	Head Abnormal	Tail & Head Abnormal	Percent Normal
641170	199	1	0	0	99.5
641065	199	1	0	0	99.5
641126	196	4	0	0	98.0
641058	199	0	1	0	99.5
641131	197	2	1	0	98.5
641176	195	2	3	0	97.5
641119	190	2	8	0	95.0
641075	194	3	3	0	97.0
641180	194	2	4	0	97.0
641117	195	3	2	0	97.5

GROUP VII-0: 1000 MG/KG/DAY

Animal Number	Total Normal	Tail Abnormal	Head Abnormal	Tail & Head Abnormal	Percent Normal
641110	196	1	4	0	98.0
641096	196	1	3	0	98.0
641212	198	0	2	0	99.0
641068	198	1	1	0	99.0
641093 ^a	---	---	---	---	---
641054	195	1	4	0	97.5
641208	190	3	7	0	95.0
641193	198	1	1	0	99.0
641207	195	3	2	0	97.5
641139	192	2	6	0	96.0

APPENDIX AA

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL ANIMAL SPERM AND SPERMATID COUNTS IN P₁ MALE RATS

INDIVIDUAL ANIMAL SPERM AND SPERMATID COUNTS (Millions) IN P₁ MALE RATS

GROUP I-0: 0 MG/KG/DAY

Animal Number	Epididymal Sperm		Testicular Spermatids	
	Per cauda	Per g cauda	Per testis	Per g testis
641114	298.1	1226.2	140.9	93.8
641217	266.6	822.1	119.3	77.2
641204	302.8	1252.7	123.4	85.9
641158	347.2	1298.4	109.3	77.5
641100	355.4	1271.5	147.3	89.5
641060	301.6	1040.0	190.6	103.9
641188	298.1	1163.1	178.3	107.6
641106	319.2	812.3	190.6	102.9
641175	322.7	1195.4	157.8	101.2
641079	327.3	1016.2	144.4	98.5

GROUP III-0: 50 MG/KG/DAY

Animal Number	Epididymal Sperm		Testicular Spermatids	
	Per cauda	Per g cauda	Per testis	Per g testis
641087	288.8	1263.7	118.1	79.8
641108	370.6	1468.8	137.4	105.5
641118	306.3	1198.3	140.3	96.4
641128	308.6	1113.7	146.1	80.8
641091	334.4	1263.1	151.4	101.7
641199	313.3	1164.2	151.4	105.2
641209	370.6	1136.0	170.1	112.2
641163	336.7	1190.9	157.3	102.9
641129	281.7	1079.0	177.7	105.7
641138	333.2	950.0	175.4	102.2

INDIVIDUAL ANIMAL SPERM AND SPERMATID COUNTS (Millions) IN P₁ MALE RATS

GROUP V-0: 250 MG/KG/DAY

Animal Number	Epididymal Sperm		Testicular Spermatids	
	Per cauda	Per g cauda	Per testis	Per g testis
641170	370.6	1281.4	169.5	114.4
641065	316.8	1265.2	142.1	101.9
641126	286.4	1375.0	138.0	101.6
641058	286.4	1218.8	119.3	90.1
641131	320.3	1244.9	125.7	82.2
641176	242.0	1080.8	100.0	56.9
641119	353.1	1029.6	180.0	102.9
641075	322.7	1046.2	208.7	128.4
641180	376.4	1063.6	172.4	101.3
641117	339.0	1144.2	203.4	113.2

GROUP VII-0: 1000 MG/KG/DAY

Animal Number	Epididymal Sperm		Testicular Spermatids	
	Per cauda	Per g cauda	Per testis	Per g testis
641110	367.1	1275.9	173.6	99.4
641096	300.5	897.6	162.5	100.2
641212	312.1	1352.4	139.1	99.1
641068	264.2	1094.9	128.6	82.7
641093	272.4	986.2	107.6	91.2
641054	301.6	990.8	155.5	100.8
641208	343.7	1258.9	174.2	124.0
641193	316.8	1191.9	129.2	96.6
641207	351.9	1052.9	190.6	106.2
641139	382.3	1247.2	143.8	102.8

APPENDIX BB

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL MATING DATA AND GESTATION LENGTH: P₁ GENERATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL MATING DATA AND GESTATION LENGTH: P₁ GENERATION

EXPLANATORY NOTES

Note

Column labeled "Plug Observed"

- + indicates that a copulatory plug was observed
- indicates that no copulatory plug was observed

Column labeled "Litter Produced"

- + indicates that a litter was produced
- indicates that no litter was produced

Footnote

- a* Because a copulatory plug was not observed, the start of gestation is unknown and, therefore, gestation length cannot be determined.

Abbreviations

--- = No Data

INDIVIDUAL MATING DATA AND GESTATION LENGTH: P₁ GENERATION

GROUPS: II-0, I-0 DOSE: 0 MG/KG/DAY

FEMALE ANIMAL#	MALE #1 ANIMAL#	PLUG OBSERVED	MALE #2 ANIMAL#	PLUG OBSERVED	LITTER PRODUCED	GESTATION LENGTH (days)
641345	641114	+			+	21
641236	641217	+			+	22
641337	641204	+			+	22
641328	641158	+			+	22
641310	641100	+			+	22
641324	641060	+			+	23
641371	641188	+			+	22
641332	641106	+			+	22
641358	641175	+			-	--
641389	641079	+			-	--
641317	641125	-			+	--a
641255	641186	+			+	22
641403	641052	+			+	22
641388	641140	+			+	21
641261	641135	+			+	22
641306	641174	+			+	22
641349	641107	-			+	--a
641346	641072	+			+	22
641308	641053	+			+	22
641289	641168	+			+	21

INDIVIDUAL MATING DATA AND GESTATION LENGTH: P₁ GENERATION

GROUPS: IV-0, III-0 DOSE: 50 MG/KG/DAY

FEMALE ANIMAL#	MALE #1 ANIMAL#	PLUG OBSERVED	MALE #2 ANIMAL#	PLUG OBSERVED	LITTER PRODUCED	GESTATION LENGTH(days)
641378	641087	+			+	22
641263	641108	+			+	21
641297	641118	+			+	22
641304	641128	+			+	23
641274	641091	+			-	--
641351	641199	+			+	21
641272	641209	+			+	22
641258	641163	+			+	22
641276	641129	+			-	--
641350	641138	+			+	22
641376	641103	+			+	22
641315	641181	+			+	22
641286	641171	+			+	22
641231	641111	+			+	22
641338	641062	+			+	22
641318	641156	+			+	22
641373	641173	+			+	22
641365	641121	+			+	22
641348	641074	+			+	22
641323	641191	+			+	21

INDIVIDUAL MATING DATA AND GESTATION LENGTH: P₁ GENERATION

GROUPS: VI-0,V-0 DOSE: 250 MG/KG/DAY

FEMALE ANIMAL#	MALE #1 ANIMAL#	PLUG OBSERVED	MALE #2 ANIMAL#	PLUG OBSERVED	LITTER PRODUCED	GESTATION LENGTH(days)
641271	641170	+			+	22
641280	641065	+			-	--
641252	641126	+			-	--
641386	641058	+			-	--
641277	641131	+			+	21
641233	641176	+			+	22
641238	641119	+			+	22
641235	641075	+			+	21
641392	641180	+			+	22
641360	641117	+			+	22
641381	641124	+			-	--
641333	641082	-			-	--
641239	641151	+			+	22
641228	641195	+			-	--
641334	641122	+			+	23
641347	641184	+			+	23
641367	641142	+			-	--
641303	641120	+			+	22
641339	641179	+			+	22
641316	641164	+			+	22

INDIVIDUAL MATING DATA AND GESTATION LENGTH: P₁ GENERATION

GROUPS: VIII-0, VII-0 DOSE: 1000 MG/KG/DAY

FEMALE ANIMAL#	MALE #1 ANIMAL#	PLUG OBSERVED	MALE #2 ANIMAL#	PLUG OBSERVED	LITTER PRODUCED	GESTATION LENGTH (days)
641265	641110	+			+	22
641393	641096	+			+	22
641402	641212	+			+	22
641246	641068	+			+	22
641320	641093	+			+	22
641353	641054	+			+	22
641401	641208	+			+	21
641307	641193	+			+	22
641368	641207	+			+	22
641343	641139	+			+	22
641254	641080	+			+	22
641242	641078	+			-	--
641284	641177	+			+	22
641230	641112	+			+	22
641273	641210	+			-	--
641296	641152	+			+	22
641248	641182	+			+	22
641370	641150	+			+	22
641243	641211	+			+	22
641329	641145	+			+	22

APPENDIX CC

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL IMPLANTATION SITE DATA AND IMPLANTATION EFFICIENCY

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL IMPLANTATION SITE DATA AND IMPLANTATION EFFICIENCY

EXPLANATORY NOTES

Footnotes

- a* Number of pups born exceeded the number of implantation sites counted.
- b* No litter was delivered.
- c* No implantation site data recorded; Animal No. 641392 died on lactation day 13.

Abbreviations

--- = No data

INDIVIDUAL IMPLANTATION SITE DATA AND IMPLANTATION EFFICIENCY

GROUPS: II-0 DOSE: 0 MG/KG/DAY

Animal Number	Number of Implantation Sites	Implantation Efficiency
641345	13	100.00
641236	17	100.00
641337	9	88.89
641328	16	68.75
641310	17	94.12
641324	17	70.59
641371	17	---a
641332	16	93.75
641358 ^b	0	---
641389 ^b	0	---
641317	6	100.00
641255	12	---a
641403	17	100.00
641388	16	93.75
641261	11	100.00
641306	15	100.00
641349	19	94.74
641346	19	78.95
641308	15	93.33
641289	17	82.35

INDIVIDUAL IMPLANTATION SITE DATA AND IMPLANTATION EFFICIENCY

GROUPS: IV-0 DOSE: 50 MG/KG/DAY

Animal Number	Number of Implantation Sites	Implantation Efficiency
641378	18	94.44
641263	13	100.00
641297	15	86.67
641304	12	50.00
641274 ^b	0	---
641351	15	100.00
641272	14	92.86
641258	16	100.00
641276 ^b	0	---
641350	16	100.00
641376	9	77.78
641315	18	100.00
641286	13	--- ^a
641231	16	93.75
641338	16	81.25
641318	16	93.75
641373	17	94.12
641365	14	92.86
641348	15	100.00
641323	15	93.33

INDIVIDUAL IMPLANTATION SITE DATA AND IMPLANTATION EFFICIENCY

GROUPS: VI-0 DOSE: 250 MG/KG/DAY

Animal Number	Number of Implantation Sites	Implantation Efficiency
641271	15	100.00
641280 ^b	0	---
641252 ^b	0	---
641386 ^b	0	---
641277	13	100.00
641233	16	93.75
641238	17	76.47
641235	12	83.33
641392 ^c	---	---
641360	15	86.67
641381 ^b	0	---
641333 ^b	0	---
641239	18	88.89
641228 ^b	0	---
641334	18	88.89
641347	17	64.71
641367 ^b	0	---
641303	15	93.33
641339	17	82.35
641316	15	86.67

INDIVIDUAL IMPLANTATION SITE DATA AND IMPLANTATION EFFICIENCY

GROUPS: VIII-0 DOSE: 1000 MG/KG/DAY

Animal Number	Number of Implantation Sites	Implantation Efficiency
641265	16	75.00
641393	16	93.75
641402	15	93.33
641246	13	92.31
641320	17	88.24
641353	16	100.00
641401	15	100.00
641307	15	86.67
641368	17	100.00
641343	15	---a
641254	20	95.00
641242 ^b	0	---
641284	18	88.89
641230	14	100.00
641273 ^b	0	---
641296	17	88.24
641248	13	---a
641370	11	100.00
641243	15	100.00
641329	15	100.00

APPENDIX DD
REPRODUCTIVE TOXICOLOGY EVALUATIONS
INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

EXPLANATORY NOTES

Footnotes

- a* No litter delivered.
- b* Dam was found dead on lactation day 13 (test day 107). All pups were dead before day 7 of lactation.

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: II-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF PUPS					
		BORN	PRECULLING	POSTCULLING	DAY 7	DAY 14	DAY 21
		ALIVE	DAY 4	DAY 4			
641345	13	13	13	8	8	8	8
641236	17	17	16	8	8	8	8
641337	8	8	8	8	8	8	8
641328	11	11	10	8	8	8	8
641310	16	14	14	8	8	8	8
641324	12	12	12	8	8	8	8
641371	18	18	18	8	8	8	8
641332	15	15	15	8	8	8	8
641358 ^a	---	---	---	---	---	---	---
641389 ^a	---	---	---	---	---	---	---
641317	6	6	6	6	6	6	6
641255	16	16	16	8	8	8	8
641403	17	17	17	8	8	8	8
641388	15	15	14	8	8	8	8
641261	11	11	11	8	8	8	8
641306	15	13	13	8	8	8	8
641349	18	17	17	8	8	8	8
641346	15	15	15	8	8	8	8
641308	14	14	14	8	8	8	8
641289	14	14	13	8	8	8	8

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: II-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF MALE PUPS					
		BORN	PRECULLING	POSTCULLING			
		ALIVE	DAY 4	DAY 4	DAY 7	DAY 14	DAY 21
641345	6	6	6	4	4	4	4
641236	6	6	6	4	4	4	4
641337	5	5	5	5	5	5	5
641328	5	5	5	4	4	4	4
641310	9	9	9	4	4	4	4
641324	4	4	4	4	4	4	4
641371	8	8	8	4	4	4	4
641332	10	10	10	4	4	4	4
641358 ^a	---	---	---	---	---	---	---
641389 ^a	---	---	---	---	---	---	---
641317	3	3	3	3	3	3	3
641255	5	5	5	4	4	4	4
641403	9	9	9	4	4	4	4
641388	8	8	8	3	3	3	3
641261	3	3	3	3	3	3	3
641306	9	8	8	4	4	4	4
641349	8	8	8	4	4	4	4
641346	7	7	7	4	4	4	4
641308	8	8	8	4	4	4	4
641289	6	6	5	4	4	4	4

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: II-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF FEMALE PUPS					
		BORN ALIVE	PRECULLING DAY 4	POSTCULLING DAY 4	DAY 7	DAY 14	DAY 21
641345	7	7	7	4	4	4	4
641236	11	11	10	4	4	4	4
641337	3	3	3	3	3	3	3
641328	6	6	5	4	4	4	4
641310	7	5	5	4	4	4	4
641324	8	8	8	4	4	4	4
641371	10	10	10	4	4	4	4
641332	5	5	5	4	4	4	4
641358 ^a	---	---	---	---	---	---	---
641389 ^a	---	---	---	---	---	---	---
641317	3	3	3	3	3	3	3
641255	11	11	11	4	4	4	4
641403	8	8	8	4	4	4	4
641388	7	7	6	5	5	5	5
641261	8	8	8	5	5	5	5
641306	6	5	5	4	4	4	4
641349	10	9	9	4	4	4	4
641346	8	8	8	4	4	4	4
641308	6	6	6	4	4	4	4
641289	8	8	8	4	4	4	4

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: II-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	SEX RATIO (MALES)	NUMBER OF PUPS		LACTATION INDEX
		% BORN ALIVE	0-4 DAY VIABILITY	
641345	0.46	100.0	100.0	100.0
641236	0.35	100.0	94.1	100.0
641337	0.63	100.0	100.0	100.0
641328	0.45	100.0	90.9	100.0
641310	0.56	87.5	100.0	100.0
641324	0.33	100.0	100.0	100.0
641371	0.44	100.0	100.0	100.0
641332	0.67	100.0	100.0	100.0
641358 ^a	---	---	---	---
641389 ^a	---	---	---	---
641317	0.50	100.0	100.0	100.0
641255	0.31	100.0	100.0	100.0
641403	0.53	100.0	100.0	100.0
641388	0.53	100.0	93.3	100.0
641261	0.27	100.0	100.0	100.0
641306	0.60	86.7	100.0	100.0
641349	0.44	94.4	100.0	100.0
641346	0.47	100.0	100.0	100.0
641308	0.57	100.0	100.0	100.0
641289	0.43	100.0	92.9	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: II-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF MALE PUPS		LACTATION INDEX
	% BORN ALIVE	0-4 DAY VIABILITY	
641345	100.0	100.0	100.0
641236	100.0	100.0	100.0
641337	100.0	100.0	100.0
641328	100.0	100.0	100.0
641310	100.0	100.0	100.0
641324	100.0	100.0	100.0
641371	100.0	100.0	100.0
641332	100.0	100.0	100.0
641358 ^a	---	---	---
641389 ^a	---	---	---
641317	100.0	100.0	100.0
641255	100.0	100.0	100.0
641403	100.0	100.0	100.0
641388	100.0	100.0	100.0
641261	100.0	100.0	100.0
641306	88.9	100.0	100.0
641349	100.0	100.0	100.0
641346	100.0	100.0	100.0
641308	100.0	100.0	100.0
641289	100.0	83.3	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: II-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF FEMALE PUPS		LACTATION INDEX
	% BORN ALIVE	0-4 DAY VIABILITY	
641345	100.0	100.0	100.0
641236	100.0	90.9	100.0
641337	100.0	100.0	100.0
641328	100.0	83.3	100.0
641310	71.4	100.0	100.0
641324	100.0	100.0	100.0
641371	100.0	100.0	100.0
641332	100.0	100.0	100.0
641358 ^a	---	---	---
641389 ^a	---	---	---
641317	100.0	100.0	100.0
641255	100.0	100.0	100.0
641403	100.0	100.0	100.0
641388	100.0	85.7	100.0
641261	100.0	100.0	100.0
641306	83.3	100.0	100.0
641349	90.0	100.0	100.0
641346	100.0	100.0	100.0
641308	100.0	100.0	100.0
641289	100.0	100.0	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: IV-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF PUPS					
		BORN ALIVE	PRECULLING DAY 4	POSTCULLING DAY 4	DAY 7	DAY 14	DAY 21
641378	17	17	17	8	8	8	8
641263	13	13	13	8	8	8	8
641297	13	13	12	8	8	8	8
641304	6	6	6	6	6	6	6
641274 ^a	---	---	---	---	---	---	---
641351	15	14	14	8	8	8	8
641272	13	13	13	8	8	8	8
641258	16	16	16	8	8	8	8
641276 ^a	---	---	---	---	---	---	---
641350	16	16	15	8	8	8	8
641376	7	7	7	7	7	7	7
641315	18	18	18	8	8	8	8
641286	14	14	14	8	8	8	8
641231	15	15	14	8	8	8	8
641338	13	13	13	8	8	8	8
641318	15	15	15	8	8	8	8
641373	16	16	14	8	8	8	8
641365	13	13	13	8	8	8	8
641348	15	15	15	8	8	8	8
641323	14	14	14	8	8	8	8

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: IV-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF MALE PUPS					
		BORN ALIVE	PRECULLING DAY 4	POSTCULLING DAY 4	DAY 7	DAY 14	DAY 21
641378	9	9	9	4	4	4	4
641263	3	3	3	3	3	3	3
641297	7	7	6	4	4	4	4
641304	1	1	1	1	1	1	1
641274 ^a	---	---	---	---	---	---	---
641351	6	6	6	4	4	4	4
641272	5	5	5	4	4	4	4
641258	6	6	6	4	4	4	4
641276 ^a	---	---	---	---	---	---	---
641350	8	8	8	4	4	4	4
641376	4	4	4	4	4	4	4
641315	9	9	9	4	4	4	4
641286	8	8	8	4	4	4	4
641231	5	5	4	4	4	4	4
641338	4	4	4	4	4	4	4
641318	7	7	7	4	4	4	4
641373	9	9	8	4	4	4	4
641365	10	10	10	5	5	5	5
641348	8	8	8	4	4	4	4
641323	6	6	6	4	4	4	4

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: IV-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF FEMALE PUPS					
		BORN ALIVE	PRECULLING DAY 4	POSTCULLING DAY 4	DAY 7	DAY 14	DAY 21
641378	8	8	8	4	4	4	4
641263	10	10	10	5	5	5	5
641297	6	6	6	4	4	4	4
641304	5	5	5	5	5	5	5
641274 ^a	---	---	---	---	---	---	---
641351	9	8	8	4	4	4	4
641272	8	8	8	4	4	4	4
641258	10	10	10	4	4	4	4
641276 ^a	---	---	---	---	---	---	---
641350	8	8	7	4	4	4	4
641376	3	3	3	3	3	3	3
641315	9	9	9	4	4	4	4
641286	6	6	6	4	4	4	4
641231	10	10	10	4	4	4	4
641338	9	9	9	4	4	4	4
641318	8	8	8	4	4	4	4
641373	7	7	6	4	4	4	4
641365	3	3	3	3	3	3	3
641348	7	7	7	4	4	4	4
641323	8	8	8	4	4	4	4

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: IV-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF PUPS			LACTATION INDEX
	SEX RATIO (MALES)	% BORN ALIVE	0-4 DAY VIABILITY	
641378	0.53	100.0	100.0	100.0
641263	0.23	100.0	100.0	100.0
641297	0.54	100.0	92.3	100.0
641304	0.17	100.0	100.0	100.0
641274 ^a	---	---	---	---
641351	0.40	93.3	100.0	100.0
641272	0.38	100.0	100.0	100.0
641258	0.38	100.0	100.0	100.0
641276 ^a	---	---	---	---
641350	0.50	100.0	93.8	100.0
641376	0.57	100.0	100.0	100.0
641315	0.50	100.0	100.0	100.0
641286	0.57	100.0	100.0	100.0
641231	0.33	100.0	93.3	100.0
641338	0.31	100.0	100.0	100.0
641318	0.47	100.0	100.0	100.0
641373	0.56	100.0	87.5	100.0
641365	0.77	100.0	100.0	100.0
641348	0.53	100.0	100.0	100.0
641323	0.43	100.0	100.0	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: IV-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF MALE PUPS		LACTATION INDEX
	% BORN ALIVE	0-4 DAY VIABILITY	
641378	100.0	100.0	100.0
641263	100.0	100.0	100.0
641297	100.0	85.7	100.0
641304	100.0	100.0	100.0
641274 ^a	---	---	---
641351	100.0	100.0	100.0
641272	100.0	100.0	100.0
641258	100.0	100.0	100.0
641276 ^a	---	---	---
641350	100.0	100.0	100.0
641376	100.0	100.0	100.0
641315	100.0	100.0	100.0
641286	100.0	100.0	100.0
641231	100.0	80.0	100.0
641338	100.0	100.0	100.0
641318	100.0	100.0	100.0
641373	100.0	88.9	100.0
641365	100.0	100.0	100.0
641348	100.0	100.0	100.0
641323	100.0	100.0	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: IV-0 DOSE: 50 MG/KG/DAY

NUMBER OF FEMALE PUPS			
ANIMAL NUMBER	% BORN ALIVE	0-4 DAY VIABILITY	LACTATION INDEX
641378	100.0	100.0	100.0
641263	100.0	100.0	100.0
641297	100.0	100.0	100.0
641304	100.0	100.0	100.0
641274 ^a	---	---	---
641351	88.9	100.0	100.0
641272	100.0	100.0	100.0
641258	100.0	100.0	100.0
641276 ^a	---	---	---
641350	100.0	87.5	100.0
641376	100.0	100.0	100.0
641315	100.0	100.0	100.0
641286	100.0	100.0	100.0
641231	100.0	100.0	100.0
641338	100.0	100.0	100.0
641318	100.0	100.0	100.0
641373	100.0	85.7	100.0
641365	100.0	100.0	100.0
641348	100.0	100.0	100.0
641323	100.0	100.0	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VI-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF PUPS					
		BORN	PRECULLING	POSTCULLING	DAY 7	DAY 14	DAY 21
		ALIVE	DAY 4	DAY 4			
641271	15	14	13	8	8	8	8
641280 ^a	---	---	---	---	---	---	---
641252 ^a	---	---	---	---	---	---	---
641386 ^a	---	---	---	---	---	---	---
641277	13	13	13	8	8	8	8
641233	15	15	15	8	8	8	8
641238	13	12	12	8	8	7	7
641235	10	10	10	8	8	8	8
641392 ^b	13	13	12	8	0	---	---
641360	13	13	13	8	8	8	8
641381 ^a	---	---	---	---	---	---	---
641333 ^a	---	---	---	---	---	---	---
641239	16	16	16	8	8	8	8
641228 ^a	---	---	---	---	---	---	---
641334	16	16	16	8	8	8	8
641347	11	11	11	8	8	8	8
641367 ^a	---	---	---	---	---	---	---
641303	14	14	13	8	8	8	8
641339	14	14	14	8	8	8	8
641316	13	13	13	8	8	8	8

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VI-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF MALE PUPS						
	BORN	BORN	PRECULLING	POSTCULLING	DAY 7	DAY 14	DAY 21
		ALIVE	DAY 4	DAY 4			
641271	6	6	5	4	4	4	4
641280 ^a	---	---	---	---	---	---	---
641252 ^a	---	---	---	---	---	---	---
641386 ^a	---	---	---	---	---	---	---
641277	8	8	8	4	4	4	4
641233	9	9	9	4	4	4	4
641238	5	5	5	4	4	4	4
641235	4	4	4	4	4	4	4
641392 ^b	8	8	8	4	0	---	---
641360	4	4	4	4	4	4	4
641381 ^a	---	---	---	---	---	---	---
641333 ^a	---	---	---	---	---	---	---
641239	12	12	12	4	4	4	4
641228 ^a	---	---	---	---	---	---	---
641334	11	11	11	4	4	4	4
641347	7	7	7	4	4	4	4
641367 ^a	---	---	---	---	---	---	---
641303	8	8	8	4	4	4	4
641339	6	6	6	4	4	4	4
641316	6	6	6	4	4	4	4

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VI-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF FEMALE PUPS					
		BORN ALIVE	PRECULLING DAY 4	POSTCULLING DAY 4	DAY 7	DAY 14	DAY 21
641271	9	8	8	4	4	4	4
641280 ^a	---	---	---	---	---	---	---
641252 ^a	---	---	---	---	---	---	---
641386 ^a	---	---	---	---	---	---	---
641277	5	5	5	4	4	4	4
641233	6	6	6	4	4	4	4
641238	8	7	7	4	4	3	3
641235	6	6	6	4	4	4	4
641392 ^b	5	5	4	4	0	---	---
641360	9	9	9	4	4	4	4
641381 ^a	---	---	---	---	---	---	---
641333 ^a	---	---	---	---	---	---	---
641239	4	4	4	4	4	4	4
641228 ^a	---	---	---	---	---	---	---
641334	5	5	5	4	4	4	4
641347	4	4	4	4	4	4	4
641367 ^a	---	---	---	---	---	---	---
641303	6	6	5	4	4	4	4
641339	8	8	8	4	4	4	4
641316	7	7	7	4	4	4	4

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VI-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF PUPS			LACTATION INDEX
	SEX RATIO (MALES)	% BORN ALIVE	0-4 DAY VIABILITY	
641271	0.40	93.3	92.9	100.0
641280 ^a	---	---	---	---
641252 ^a	---	---	---	---
641386 ^a	---	---	---	---
641277	0.62	100.0	100.0	100.0
641233	0.60	100.0	100.0	100.0
641238	0.38	92.3	100.0	87.5
641235	0.40	100.0	100.0	100.0
641392 ^b	0.62	100.0	92.3	-
641360	0.31	100.0	100.0	100.0
641381 ^a	---	---	---	---
641333 ^a	---	---	---	---
641239	0.75	100.0	100.0	100.0
641228 ^a	---	---	---	---
641334	0.69	100.0	100.0	100.0
641347	0.64	100.0	100.0	100.0
641367 ^a	---	---	---	---
641303	0.57	100.0	92.9	100.0
641339	0.43	100.0	100.0	100.0
641316	0.46	100.0	100.0	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VI-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF MALE PUPS		LACTATION INDEX
	% BORN ALIVE	0-4 DAY VIABILITY	
641271	100.0	83.3	100.0
641280 ^a	---	---	---
641252 ^a	---	---	---
641386 ^a	---	---	---
641277	100.0	100.0	100.0
641233	100.0	100.0	100.0
641238	100.0	100.0	100.0
641235	100.0	100.0	100.0
641392 ^b	100.0	100.0	---
641360	100.0	100.0	100.0
641381 ^a	---	---	---
641333 ^a	---	---	---
641239	100.0	100.0	100.0
641228 ^a	---	---	---
641334	100.0	100.0	100.0
641347	100.0	100.0	100.0
641367 ^a	---	---	---
641303	100.0	100.0	100.0
641339	100.0	100.0	100.0
641316	100.0	100.0	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VI-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF FEMALE PUPS		LACTATION INDEX
	% BORN ALIVE	0-4 DAY VIABILITY	
641271	88.9	100.0	100.0
641280 ^a	---	---	---
641252 ^a	---	---	---
641386 ^a	---	---	---
641277	100.0	100.0	100.0
641233	100.0	100.0	100.0
641238	87.5	100.0	75.0
641235	100.0	100.0	100.0
641392 ^b	100.0	80.0	-
641360	100.0	100.0	100.0
641381 ^a	---	---	---
641333 ^a	---	---	---
641239	100.0	100.0	100.0
641228 ^a	---	---	---
641334	100.0	100.0	100.0
641347	100.0	100.0	100.0
641367 ^a	---	---	---
641303	100.0	83.3	100.0
641339	100.0	100.0	100.0
641316	100.0	100.0	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VIII-0 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF PUPS					
		BORN ALIVE	PRECULLING DAY 4	POSTCULLING DAY 4	DAY 7	DAY 14	DAY 21
641265	12	12	12	8	8	8	8
641393	15	14	14	8	8	8	8
641402	14	14	14	6	6	6	6
641246	12	12	12	8	8	8	8
641320	15	15	15	8	8	8	8
641353	16	15	15	8	8	8	8
641401	15	15	14	8	8	8	8
641307	13	13	13	8	8	8	8
641368	17	17	17	8	8	8	8
641343	18	18	18	8	8	8	8
641254	19	19	19	8	8	8	8
641242 ^a	---	---	---	---	---	---	---
641284	16	16	16	8	8	8	8
641230	14	14	14	8	8	8	8
641273 ^a	---	---	---	---	---	---	---
641296	15	13	13	8	8	8	8
641248	15	15	15	8	8	8	8
641370	11	11	11	8	8	8	8
641243	15	15	14	8	8	8	8
641329	15	15	15	8	8	8	8

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VIII-0 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF MALE PUPS					
		BORN ALIVE	PRECULLING DAY 4	POSTCULLING DAY 4	DAY 7	DAY 14	DAY 21
641265	10	10	10	6	6	6	6
641393	8	7	7	4	4	4	4
641402	2	2	2	2	2	2	2
641246	5	5	5	4	4	4	4
641320	9	9	9	4	4	4	4
641353	9	8	8	4	4	4	4
641401	6	6	5	4	4	4	4
641307	10	10	10	5	5	5	5
641368	9	9	9	4	4	4	4
641343	9	9	9	4	4	4	4
641254	7	7	7	4	4	4	4
641242 ^a	---	---	---	---	---	---	---
641284	6	6	6	4	4	4	4
641230	4	4	4	4	4	4	4
641273 ^a	---	---	---	---	---	---	---
641296	9	7	7	4	4	4	4
641248	5	5	5	4	4	4	4
641370	4	4	4	4	4	4	4
641243	6	6	6	4	4	4	4
641329	10	10	10	4	4	4	4

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VIII-0 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	BORN	NUMBER OF FEMALE PUPS					
		BORN ALIVE	PRECULLING DAY 4	POSTCULLING DAY 4	DAY 7	DAY 14	DAY 21
641265	2	2	2	2	2	2	2
641393	7	7	7	4	4	4	4
641402	12	12	12	4	4	4	4
641246	7	7	7	4	4	4	4
641320	6	6	6	4	4	4	4
641353	7	7	7	4	4	4	4
641401	9	9	9	4	4	4	4
641307	3	3	3	3	3	3	3
641368	8	8	8	4	4	4	4
641343	9	9	9	4	4	4	4
641254	12	12	12	4	4	4	4
641242 ^a	---	---	---	---	---	---	---
641284	10	10	10	4	4	4	4
641230	10	10	10	4	4	4	4
641273 ^a	---	---	---	---	---	---	---
641296	6	6	6	4	4	4	4
641248	10	10	10	4	4	4	4
641370	7	7	7	4	4	4	4
641243	9	9	8	4	4	4	4
641329	5	5	5	4	4	4	4

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VIII-0 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	SEX RATIO (MALES)	NUMBER OF PUPS		LACTATION INDEX
		% BORN ALIVE	0-4 DAY VIABILITY	
641265	0.83	100.0	100.0	100.0
641393	0.53	93.3	100.0	100.0
641402	0.14	100.0	100.0	100.0
641246	0.42	100.0	100.0	100.0
641320	0.60	100.0	100.0	100.0
641353	0.56	93.8	100.0	100.0
641401	0.40	100.0	93.3	100.0
641307	0.77	100.0	100.0	100.0
641368	0.53	100.0	100.0	100.0
641343	0.50	100.0	100.0	100.0
641254	0.37	100.0	100.0	100.0
641242 ^a	---	---	---	---
641284	0.38	100.0	100.0	100.0
641230	0.29	100.0	100.0	100.0
641273 ^a	---	---	---	---
641296	0.60	86.7	100.0	100.0
641248	0.33	100.0	100.0	100.0
641370	0.36	100.0	100.0	100.0
641243	0.40	100.0	93.3	100.0
641329	0.67	100.0	100.0	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VIII-0 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	NUMBER OF MALE PUPS		LACTATION INDEX
	% BORN ALIVE	0-4 DAY VIABILITY	
641265	100.0	100.0	100.0
641393	87.5	100.0	100.0
641402	100.0	100.0	100.0
641246	100.0	100.0	100.0
641320	100.0	100.0	100.0
641353	88.9	100.0	100.0
641401	100.0	83.3	100.0
641307	100.0	100.0	100.0
641368	100.0	100.0	100.0
641343	100.0	100.0	100.0
641254	100.0	100.0	100.0
641242 ^a	---	---	---
641284	100.0	100.0	100.0
641230	100.0	100.0	100.0
641273 ^a	---	---	---
641296	77.8	100.0	100.0
641248	100.0	100.0	100.0
641370	100.0	100.0	100.0
641243	100.0	100.0	100.0
641329	100.0	100.0	100.0

INDIVIDUAL PUP SURVIVAL: F₁ GENERATION

GROUP: VIII-0 DOSE: 1000 MG/KG/DAY

NUMBER OF FEMALE PUPS			
ANIMAL NUMBER	% BORN ALIVE	0-4 DAY VIABILITY	LACTATION INDEX
641265	100.0	100.0	100.0
641393	100.0	100.0	100.0
641402	100.0	100.0	100.0
641246	100.0	100.0	100.0
641320	100.0	100.0	100.0
641353	100.0	100.0	100.0
641401	100.0	100.0	100.0
641307	100.0	100.0	100.0
641368	100.0	100.0	100.0
641343	100.0	100.0	100.0
641254	100.0	100.0	100.0
641242 ^a	---	---	---
641284	100.0	100.0	100.0
641230	100.0	100.0	100.0
641273 ^a	---	---	---
641296	100.0	100.0	100.0
641248	100.0	100.0	100.0
641370	100.0	100.0	100.0
641243	100.0	88.9	100.0
641329	100.0	100.0	100.0

APPENDIX EE
REPRODUCTIVE TOXICOLOGY EVALUATIONS
INDIVIDUAL PUP WEIGHTS: F₁ GENERATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL PUP WEIGHTS: F₁ GENERATION

EXPLANATORY NOTES

Footnotes

- a* No litter delivered.
- b* No live pups on lactation day 7.
- c* Found dead test day 107.
- d* One pup was incorrectly designated as female. Data were excluded from calculations.

Individual Pup Weights (grams): F1 Generation
Lactation Day 0

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>										
		1	2	3	4	5	6	7	8	9	10	11
641345	♂	6.7	7.4	7.2	7.7	7.1	7.4					
	♀	6.1	6.4	6.9	6.2	6.4	7.2	6.8				
641236	♂	6.8	6.8	7.0	6.1	6.8	6.2					
	♀	6.1	5.9	6.0	6.3	6.1	6.3	5.9	6.1	6.2	6.8	5.4
641337	♂	8.0	7.3	8.1	7.8	6.8						
	♀	7.6	7.2	6.9								
641328	♂	7.6	9.0	7.7	7.5	8.2						
	♀	7.1	7.6	7.7	7.9	6.8	8.2					
641310	♂	6.7	7.1	6.8	7.3	6.5	6.7	5.9	7.0	6.5		
	♀	5.8	6.5	6.3	6.8	6.8						
641324	♂	7.3	6.8	7.2	7.6							
	♀	6.4	7.1	7.5	5.7	7.2	6.6	7.2	7.0			
641371	♂	7.3	6.2	7.4	6.9	6.5	6.4	6.9	6.4			
	♀	6.4	6.2	6.2	5.2	5.7	6.6	5.5	6.2	6.3	6.2	
641332	♂	6.8	6.9	7.6	7.2	6.8	7.6	7.4	6.6	6.9	7.2	
	♀	7.1	6.8	6.9	6.9	6.8						
641358 ^a	♂											
	♀											
641389 ^a	♂											
	♀											

Individual Pup Weights (grams): F1 Generation
Lactation Day 0

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>										
		1	2	3	4	5	6	7	8	9	10	11
641317	♂	7.1	7.9	7.6								
	♀	8.0	7.2	7.0								
641255	♂	6.8	6.6	7.0	6.5	6.2						
	♀	6.0	5.4	6.0	6.4	6.3	5.9	6.4	6.2	5.7	6.0	6.1
641403	♂	7.1	7.0	6.7	6.6	6.5	6.5	6.5	7.5	6.3		
	♀	6.3	6.6	6.9	6.4	6.6	5.9	6.4	6.0			
641388	♂	6.6	7.2	7.5	7.8	7.3	7.2	7.2	7.4			
	♀	6.7	7.1	6.5	6.8	6.6	6.6	6.6				
641261	♂	7.8	7.0	7.5								
	♀	5.6	6.9	6.8	7.7	6.8	6.7	6.5	6.3			
641306	♂	6.7	6.8	6.7	6.6	6.7	6.8	7.2	6.3			
	♀	6.1	6.1	6.1	6.7	6.0						
641349	♂	6.7	4.4	4.8	5.5	5.5	5.7	5.2	5.8			
	♀	5.0	5.8	5.4	5.3	5.8	5.0	4.2	6.0	5.2		
641346	♂	6.2	5.7	6.3	6.4	6.9	6.7	6.1				
	♀	5.5	6.0	5.6	6.1	5.8	6.5	5.9	6.4			
641308	♂	7.5	7.1	6.9	5.4	6.1	4.0	6.6	7.5			
	♀	6.2	6.6	7.5	6.6	6.3	6.3					
641289	♂	3.5	5.9	6.0	5.6	5.9	5.8					
	♀	5.7	5.2	5.4	5.0	5.3	5.2	5.6	5.8			

Individual Pup Weights (grams): F₁ Generation
Lactation Day 0

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	Pup Number									
		1	2	3	4	5	6	7	8	9	10
641378	♂	6.8	7.0	6.4	6.4	6.1	7.2	6.5	6.7	6.9	
	♀	6.3	6.3	6.3	6.5	7.4	5.9	6.5	6.2		
641263	♂	6.0	5.6	6.0							
	♀	5.2	5.6	5.6	5.6	5.4	5.7	5.5	5.6	5.8	5.8
641297	♂	7.5	7.6	7.9	7.8	8.1	7.8	7.1			
	♀	7.2	6.9	6.8	7.2	7.3	7.2				
641304	♂	7.9									
	♀	7.3	8.0	7.6	7.5	6.1					
641274 ^a	♂										
	♀										
641351	♂	6.0	5.7	6.1	6.1	5.7	5.9				
	♀	6.0	5.8	5.6	5.5	6.1	5.4	5.5	5.7		
641272	♂	7.3	7.3	7.1	6.6	6.7					
	♀	7.1	6.5	6.2	7.2	6.8	6.5	6.2	6.3		
641258	♂	7.1	7.5	7.1	7.3	6.9	7.6				
	♀	7.2	7.2	6.9	7.2	6.7	6.5	6.5	5.2	6.5	6.9
641276 ^a	♂										
	♀										
641350	♂	6.8	7.0	6.1	6.6	6.7	7.1	6.7	7.3		
	♀	6.8	6.5	6.4	6.4	6.3	6.5	5.7	6.7		

Individual Pup Weights (grams): F1 Generation
Lactation Day 0

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>									
		1	2	3	4	5	6	7	8	9	10
641376	♂	7.1	7.3	6.7	6.7						
	♀	6.6	5.8	5.8							
641315	♂	6.3	6.7	7.0	6.8	6.8	6.4	6.7	5.4	5.7	
	♀	5.8	6.1	6.4	5.9	6.3	6.4	5.4	6.1	6.2	
641286	♂	7.6	6.1	6.9	6.9	6.7	7.4	7.1			
	♀	6.7	6.1	6.4	6.6	6.3	6.4	6.5			
641231	♂	6.4	7.3	6.7	6.9	7.3					
	♀	6.1	6.9	6.4	6.1	5.9	6.9	6.3	6.2	6.2	5.0
641338	♂	6.5	6.8	6.5	7.0						
	♀	6.2	6.4	6.0	5.7	6.2	6.8	6.1	6.2	6.1	
641318	♂	6.3	6.7	6.0	6.0	6.1	6.5	6.6			
	♀	5.8	6.0	6.3	5.9	5.5	6.0	5.9	5.7		
641373	♂	7.2	6.9	7.0	7.1	6.7	7.3	6.7	6.7	7.0	
	♀	6.8	6.5	6.5	6.8	6.6	6.7	6.7			
641365	♂	7.6	7.6	8.0	7.3	7.7	8.0	7.6	7.4	7.8	8.1
	♀	7.5	7.6	7.0							
641348	♂	7.1	6.9	6.4	6.3	6.6	6.7	7.1	6.8		
	♀	6.3	6.5	6.5	6.5	6.3	6.6	6.1			
641323	♂	6.1	6.0	6.7	6.1	6.3	6.7				
	♀	5.9	6.1	6.3	6.2	5.8	6.0	5.7	5.7		

Individual Pup Weights (grams): F₁ Generation
Lactation Day 0

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	Pup Number								
		1	2	3	4	5	6	7	8	9
641271	♂	6.9	6.6	6.4	6.0	6.4	6.6			
	♀	6.6	6.0	6.6	6.2	5.9	5.8	6.8	5.8	
641280 ^a	♂									
	♀									
641252 ^a	♂									
	♀									
641386 ^a	♂									
	♀									
641277	♂	6.7	6.0	5.9	6.6	6.0	6.4	6.0	6.6	
	♀	5.8	6.0	6.2	6.1	6.0				
641233	♂	5.7	6.3	6.7	6.1	6.5	6.2	6.5	6.2	6.7
	♀	5.9	5.8	5.8	5.8	5.9	5.7			
641238	♂	7.5	6.9	7.3	7.1	7.1				
	♀	7.1	6.2	5.4	6.2	6.2	6.9	6.6		
641235	♂	6.2	6.2	5.5	6.1					
	♀	5.5	5.8	5.9	5.8	5.8	5.5			
641392 ^{b, c}	♂	6.7	6.7	6.4	6.4	6.5	5.7	6.0	5.8	
	♀	6.4	5.9	6.2	5.4	6.4				
641360	♂	7.4	6.9	7.5	6.3					
	♀	7.3	7.2	6.9	6.5	6.6	7.3	6.7	7.5	5.8

Individual Pup Weights (grams): F1 Generation
Lactation Day 0

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>											
		1	2	3	4	5	6	7	8	9	10	11	12
641381 ^a	♂												
	♀												
641333 ^a	♂												
	♀												
641239	♂	5.8	5.8	6.1	6.9	5.9	6.6	6.7	6.2	5.8	6.0	5.4	5.8
	♀	6.2	5.8	4.9	5.8								
641228 ^a	♂												
	♀												
641334	♂	6.2	7.2	6.7	7.3	6.9	6.5	7.3	6.4	7.4	6.3	6.6	
	♀	5.6	7.2	6.1	6.4	5.4							
641347	♂	8.3	8.6	7.7	8.4	8.7	7.3	8.1					
	♀	8.2	7.8	8.6	7.5								
641367 ^a	♂												
	♀												
641303	♂	7.7	6.9	7.2	6.8	7.1	7.1	6.8	7.1				
	♀	6.8	7.0	6.4	6.6	6.9	7.3						
641339	♂	7.2	7.1	6.9	7.0	7.3	6.8						
	♀	6.7	6.6	7.0	6.5	6.5	6.8	6.7	6.5				
641316	♂	7.0	6.4	6.7	6.6	6.7	6.9						
	♀	5.8	6.3	6.4	6.7	6.0	5.7	6.6					

Individual Pup Weights (grams): F1 Generation
Lactation Day 0

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>											
		1	2	3	4	5	6	7	8	9	10	11	12
641265	♂	7.0	7.3	7.5	7.1	7.5	7.0	7.4	7.4	6.0			
	♀	6.5	6.2	6.2									
641393	♂	6.5	7.0	6.8	5.9	6.2	5.7	7.0					
	♀	6.3	6.2	6.3	6.7	6.2	6.6	6.1					
641402	♂	5.9	5.8										
	♀	5.9	6.1	6.1	5.8	6.1	5.9	5.6	5.7	6.0	6.0	6.1	5.7
641246	♂	6.3	6.6	6.7	6.2	6.5							
	♀	6.1	5.8	6.2	6.0	6.3	5.7	6.1					
641320	♂	6.2	6.5	6.7	6.3	6.5	7.0	6.5	6.3	6.5			
	♀	6.4	5.7	6.5	6.2	6.0	6.0						
641353	♂	7.2	6.6	7.9	7.2	7.3	6.7	7.0	6.2				
	♀	6.8	5.5	7.3	6.2	5.9	5.6	4.2					
641401	♂	6.1	5.7	6.1	6.5	6.1	5.5						
	♀	6.5	6.2	5.8	6.0	6.0	5.9	5.9	6.0	5.6			
641307	♂	6.0	7.1	5.7	7.3	6.4	6.9	5.3	6.5	6.0	5.2		
	♀	6.2	5.4	5.9									
641368	♂	5.7	6.5	6.1	6.7	6.4	6.1	5.9	6.7	6.3			
	♀	6.0	5.9	6.3	6.3	5.9	5.7	5.9	6.0				
641343	♂	6.8	6.7	6.6	5.8	6.5	7.0	6.7	6.6	6.9			
	♀	6.9	6.9	6.5	6.3	6.5	6.8	6.0	5.2	6.6			

Individual Pup Weights (grams): F1 Generation
Lactation Day 0

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>											
		1	2	3	4	5	6	7	8	9	10	11	12
641254	♂	6.2	6.1	4.8	6.5	6.2	6.0	5.6					
	♀	5.8	5.7	5.9	5.3	5.5	5.8	5.7	6.4	5.6	5.5	6.4	5.3
641242 ^a	♂												
	♀												
641284	♂	7.8	7.5	7.1	6.8	8.1	7.2						
	♀	6.7	6.6	6.4	5.6	7.4	6.9	6.9	6.7	7.6	7.0		
641230	♂	6.0	6.4	6.6	6.4								
	♀	6.2	6.2	5.5	5.4	6.0	6.2	5.5	5.9	6.0	5.6		
641273 ^a	♂												
	♀												
641296	♂	5.5	6.0	6.5	6.5	6.2	6.2	6.3					
	♀	5.1	5.3	5.8	5.9	5.3	5.7						
641248	♂	7.1	8.4	7.6	7.4	7.6							
	♀	6.7	7.1	6.6	7.0	5.8	6.3	6.0	7.0	6.3	6.2		
641370	♂	6.8	7.2	7.2									
	♀	6.5	6.6	6.7	6.7	6.3	7.0	6.3	6.3				
641243	♂	6.2	6.8	7.3	5.9	6.3	6.6						
	♀	6.1	6.0	5.9	6.5	5.6	6.4	6.1	6.5	5.8			
641329	♂	5.8	6.7	6.2	6.7	6.9	6.3	6.4	6.2	5.9	6.2		
	♀	6.5	6.4	6.7	6.6	6.5							

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Preculling

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	Pup Number									
		1	2	3	4	5	6	7	8	9	10
641345	♂	12.1	10.8	11.9	11.4	11.4	11.6				
	♀	10.3	11.0	11.0	11.4	9.6	11.4	10.3			
641236	♂	10.4	10.9	11.1	11.7	11.3	10.1				
	♀	9.5	9.0	9.7	9.2	10.5	8.3	10.0	9.4	10.4	10.7
641337	♂	12.8	13.2	14.7	14.8	13.1					
	♀	13.5	14.2	14.2							
641328	♂	14.6	12.6	15.1	13.7	13.3					
	♀	12.9	13.7	13.5	13.9	13.1					
641310	♂	10.4	9.8	10.5	10.9	11.4	11.6	9.5	11.6	11.6	
	♀	11.2	10.3	10.7	11.2	11.2					
641324	♂	12.0	12.5	13.0	13.6						
	♀	13.6	12.0	9.5	13.3	13.1	12.7	11.8	14.2		
641371	♂	11.5	9.0	9.8	11.5	11.3	9.5	11.7	9.4		
	♀	9.6	9.8	10.1	9.2	8.1	10.5	10.3	10.2	9.8	7.9
641332	♂	10.7	9.9	9.4	10.5	9.5	11.1	10.8	9.3	10.2	11.6
	♀	10.0	9.9	10.0	9.1	9.3					
641358 ^a	♂										
	♀										
641389 ^a	♂										
	♀										

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Preculling

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>										
		1	2	3	4	5	6	7	8	9	10	11
641317	♂	15.0	14.6	12.7								
	♀	13.5	14.3	14.1								
641255	♂	9.9	10.4	10.4	10.7	10.7						
	♀	9.9	9.4	9.5	8.8	8.8	8.5	9.8	8.6	10.1	8.6	8.4
641403	♂	11.1	11.0	11.6	11.4	11.4	11.1	11.2	11.6	10.8		
	♀	9.7	11.4	11.4	11.0	10.3	12.0	11.2	10.8			
641388	♂	11.1	12.4	11.6	12.9	12.4	12.2	12.1	12.0			
	♀	11.3	11.4	11.4	11.2	11.0	11.5					
641261	♂	12.5	12.6	11.8								
	♀	10.0	11.0	11.9	11.2	11.8	10.6	12.6	11.0			
641306	♂	10.1	11.5	10.8	11.4	10.9	10.9	9.7	9.8			
	♀	9.5	11.0	9.6	9.4	10.3						
641349	♂	11.9	8.3	10.2	9.3	10.8	10.5	9.6	10.7			
	♀	9.1	11.0	8.6	9.2	10.4	9.9	9.6	10.5			
641346	♂	12.6	11.6	12.7	10.3	11.7	11.5	11.7				
	♀	11.9	10.5	10.6	11.0	11.5	11.3	11.1	11.8			
641308	♂	11.5	11.5	10.8	12.3	10.0	11.0	7.5	12.1			
	♀	11.2	10.8	12.2	12.6	10.7	10.5					
641289	♂	9.2	10.1	8.2	10.2	9.5						
	♀	9.2	8.8	8.1	7.5	8.1	9.6	9.1	8.6			

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Preculling

GROUP: IV-0

DOSE: 50 MG/KG/DAY

Animal Number	Pup Sex	Pup Number									
		1	2	3	4	5	6	7	8	9	10
641378	♂	9.9	10.3	10.5	10.4	10.9	11.6	10.5	11.5	9.5	
	♀	10.5	10.5	9.9	10.1	10.9	9.3	8.0	10.5		
641263	♂	9.6	10.0	10.0							
	♀	9.6	10.0	9.5	9.1	9.3	9.8	9.2	9.9	9.0	9.7
641297	♂	11.7	12.6	13.1	12.5	12.4	13.0				
	♀	11.2	11.6	12.4	12.3	12.2	11.2				
641304	♂	16.6									
	♀	16.0	15.5	15.5	12.7	15.2					
641274 ^a	♂										
	♀										
641351	♂	9.4	9.8	9.0	8.8	10.2	9.2				
	♀	8.6	10.0	9.4	9.9	9.5	9.2	9.5	9.6		
641272	♂	11.5	12.6	11.4	12.2	12.4					
	♀	12.6	11.2	11.3	12.0	11.4	11.2	12.2	10.2		
641258	♂	11.0	10.8	11.5	11.6	9.8	11.6				
	♀	11.4	11.5	8.5	12.0	11.0	10.9	11.1	9.1	10.3	10.2
641276 ^a	♂										
	♀										
641350	♂	12.0	10.0	11.3	12.2	11.6	10.1	11.6	10.1		
	♀	9.0	10.6	11.1	10.6	10.9	9.7	10.7			

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Preculling

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>									
		1	2	3	4	5	6	7	8	9	10
641376	♂	12.6	12.3	11.8	13.0						
	♀	12.0	10.8	10.0							
641315	♂	10.4	10.6	10.4	10.0	9.4	8.7	10.7	9.9	7.3	
	♀	9.4	10.5	11.0	9.4	10.1	9.4	8.8	9.3	9.7	
641286	♂	10.3	11.2	11.4	11.3	10.0	11.1	11.4	10.1		
	♀	9.6	10.5	10.6	9.9	10.4	10.4				
641231	♂	12.2	11.7	12.0	12.6						
	♀	10.9	12.0	11.5	11.3	12.0	11.2	8.9	11.7	11.5	11.1
641338	♂	12.1	11.0	11.8	12.3						
	♀	11.2	10.3	11.5	9.5	12.0	10.7	10.9	11.1	10.3	
641318	♂	10.1	9.0	10.0	9.8	9.9	10.1	9.4			
	♀	8.8	9.0	9.1	9.4	11.0	9.2	9.7	9.3		
641373	♂	9.9	10.7	10.0	10.4	10.3	11.0	9.6	10.3		
	♀	8.1	10.0	9.4	9.5	10.8	9.9				
641365	♂	11.3	12.7	11.3	12.8	12.5	12.3	11.3	10.5	12.1	12.9
	♀	11.9	11.7	12.0							
641348	♂	10.6	11.5	12.2	12.2	12.4	12.5	11.0	12.2		
	♀	10.8	11.3	11.0	12.1	12.1	11.6	11.2			
641323	♂	9.5	11.0	9.7	10.2	9.9	10.7				
	♀	9.9	10.4	10.0	9.4	10.0	9.1	9.2	10.2		

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Preculling

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	Pup Number								
		1	2	3	4	5	6	7	8	9
641271	♂	10.4	10.9	10.4	10.3	10.7				
	♀	9.7	10.0	10.8	10.0	9.4	9.9	10.1	9.3	
641280 ^a	♂									
	♀									
641252 ^a	♂									
	♀									
641386 ^a	♂									
	♀									
641277	♂	11.6	11.8	12.1	10.4	10.1	11.2	11.1	10.4	
	♀	11.0	10.9	10.5	10.7	10.4				
641233	♂	10.9	10.0	9.9	10.9	9.5	10.1	10.8	10.2	11.1
	♀	10.2	9.6	10.3	9.7	9.5	9.4			
641238	♂	13.3	13.5	12.4	13.5	12.9				
	♀	12.3	13.0	10.8	12.0	11.9	12.4	12.2		
641235	♂	10.5	10.7	10.5	9.7					
	♀	10.1	9.6	9.9	9.3	9.5	9.6			
641392 ^{b, c}	♂	6.8	6.5	6.4	7.1	6.8	6.9	5.9	6.6	
	♀	6.3	7.2	6.4	6.3					
641360	♂	11.4	11.8	12.1	10.1					
	♀	11.4	11.2	9.7	11.4	10.9	10.9	11.9	11.8	11.4

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Preculling

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	Pup Number											
		1	2	3	4	5	6	7	8	9	10	11	12
641381 ^a	♂												
	♀												
641333 ^a	♂												
	♀												
641239	♂	9.6	9.6	10.8	9.2	9.1	9.4	10.5	9.0	8.9	10.3	9.3	9.1
	♀	9.0	9.5	7.5	10.1								
641228 ^a	♂												
	♀												
641334	♂	9.9	9.0	9.6	8.9	10.6	8.4	9.7	10.0	10.7	9.5	8.8	
	♀	9.0	8.5	10.2	7.5	8.8							
641347	♂	14.1	11.7	14.7	13.9	13.0	14.2	13.9					
	♀	14.1	13.9	13.5	11.8								
641367 ^a	♂												
	♀												
641303	♂	12.3	12.0	11.8	12.1	12.5	11.4	11.3	12.7				
	♀	12.0	11.8	11.7	10.8	12.1							
641339	♂	11.9	11.6	11.3	11.2	12.1	11.5						
	♀	10.8	10.3	11.3	11.9	10.3	11.1	11.2	11.3				
641316	♂	11.4	11.4	11.7	11.2	11.0	10.9						
	♀	11.0	11.4	10.5	11.8	10.0	10.4	10.6					

Individual Pup Weights (grams): F₁ Generation
 Lactation Day 4 Preculling

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>											
		1	2	3	4	5	6	7	8	9	10	11	12
641265	♂	13.6	12.6	13.3	12.9	13.4	12.7	13.5	13.6	11.4	11.6		
	♀	11.6	13.0										
641393	♂	9.3	9.4	11.3	11.3	10.3	9.3	10.5					
	♀	10.6	10.6	11.3	10.9	10.3	11.1	11.1					
641402	♂	8.7	9.1										
	♀	9.8	10.1	8.9	9.3	9.0	7.3	8.9	8.9	8.5	8.6	9.2	8.7
641246	♂	10.9	10.6	10.9	11.8	11.7							
	♀	10.5	10.1	10.9	10.5	10.6	11.5	10.3					
641320	♂	11.5	11.2	10.7	10.9	10.1	10.1	10.0	12.1	11.6			
	♀	10.0	10.3	9.3	10.0	10.5	10.6						
641353	♂	10.4	10.7	10.7	10.0	11.1	10.2	10.5	11.3				
	♀	11.3	8.6	9.0	9.0	10.4	7.1	9.5					
641401	♂	8.2	9.1	9.7	10.1	10.2							
	♀	9.9	9.8	9.0	8.9	9.1	10.2	10.0	9.5	9.7			
641307	♂	10.8	11.9	11.3	11.5	11.3	10.3	10.0	10.2	10.7	9.6		
	♀	10.4	11.0	9.2									
641368	♂	10.9	9.1	10.2	9.6	7.7	10.6	10.0	11.3	10.1			
	♀	10.3	10.1	9.0	9.2	8.3	10.3	10.0	9.3				
641343	♂	10.7	12.1	12.3	11.5	10.8	9.4	11.6	11.3	11.1			
	♀	9.4	11.4	11.7	10.8	11.3	11.9	11.8	11.0	10.4			

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Preculling

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	Pup Number											
		1	2	3	4	5	6	7	8	9	10	11	12
641254	♂	8.7	6.8	11.1	7.6	10.3	10.4	9.8					
	♀	8.8	10.3	10.8	8.6	9.9	9.4	9.2	8.4	9.3	7.3	8.4	9.5
641242 ^a	♂												
	♀												
641284	♂	10.6	10.7	10.5	12.2	10.6	10.1						
	♀	10.0	8.8	9.8	10.1	10.2	10.8	11.4	10.6	10.9	11.6		
641230	♂	10.5	10.5	10.1	10.5								
	♀	10.1	10.2	10.3	10.4	10.1	10.5	9.6	9.1	8.3	9.7		
641273 ^a	♂												
	♀												
641296	♂	9.5	9.8	10.5	10.4	9.6	9.5	10.6					
	♀	8.9	8.6	9.4	10.1	10.0	9.6						
641248	♂	11.8	12.4	12.1	11.4	12.0							
	♀	10.9	9.8	10.5	11.1	10.6	11.0	10.5	11.3	9.7	9.9		
641370	♂	12.5	12.2	12.8	12.7								
	♀	11.8	12.3	11.3	12.0	11.8	11.3	12.0					
641243	♂	10.1	10.0	9.5	9.9	9.3	9.6						
	♀	9.6	9.6	10.1	9.4	8.8	9.1	9.8	10.3				
641329	♂	10.3	10.0	10.2	9.8	9.7	10.6	10.2	10.7	9.6	10.4		
	♀	10.8	10.8	10.0	10.6	9.9							

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Postculling

GROUP: II-0

DOSE: 0 MG/KG/DAY

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641345	♂	10.8	11.9	11.4	11.6	
	♀	11.0	11.4	11.4	10.3	
641236	♂	10.9	11.1	11.7	10.1	
	♀	9.0	9.2	10.5	10.7	
641337	♂	12.8	13.2	14.7	14.8	13.1
	♀	13.5	14.2	14.2		
641328	♂	14.6	12.6	15.1	13.7	
	♀	12.9	13.5	13.9	13.1	
641310	♂	9.8	10.9	11.4	11.6	
	♀	11.2	10.3	11.2	11.2	
641324	♂	12.0	12.5	13.0	13.6	
	♀	12.0	12.7	11.8	14.2	
641371	♂	11.5	9.8	11.5	11.3	
	♀	8.1	10.5	10.3	9.8	
641332	♂	9.5	11.1	10.8	10.2	
	♀	10.0	9.9	10.0	9.1	
641358 ^a	♂					
	♀					
641389 ^a	♂					
	♀					

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Postculling

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641317	♂	15.0	14.6	12.7		
	♀	13.5	14.3	14.1		
641255	♂	9.9	10.4	10.4	10.7	
	♀	9.4	8.8	9.8	8.6	
641403	♂	11.1	11.0	11.4	11.1	
	♀	11.0	10.3	11.2	10.8	
641388	♂	12.4	11.6	12.4	12.1	
	♀	11.3	11.4	11.2	11.5	
641261	♂	12.5	12.6	11.8		
	♀	11.0	11.2	11.8	10.6	12.6
641306	♂	11.4	10.9	9.7	9.8	
	♀	11.0	9.6	9.4	10.3	
641349	♂	11.9	8.3	10.2	9.3	
	♀	9.1	11.0	9.2	10.5	
641346	♂	12.6	12.7	10.3	11.7	
	♀	10.6	11.5	11.3	11.8	
641308	♂	11.5	10.0	11.0	7.5	
	♀	10.8	12.6	10.7	10.5	
641289	♂	9.2	8.2	10.2	9.5	
	♀	8.8	7.5	9.6	9.1	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Postculling

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641378	♂	9.9	10.3	11.5	9.5	
	♀	10.5	10.1	10.9	8.0	
641263	♂	9.6	10.0	10.0		
	♀	10.0	9.3	9.2	9.9	9.0
641297	♂	12.6	13.1	12.5	12.4	
	♀	11.6	12.4	12.3	11.2	
641304	♂	16.6				
	♀	16.0	15.5	15.5	12.7	15.2
641274 ^a	♂					
	♀					
641351	♂	9.8	9.0	10.2	9.2	
	♀	8.6	9.9	9.5	9.5	
641272	♂	12.6	11.4	12.2	12.4	
	♀	12.6	11.3	12.0	11.4	
641258	♂	11.0	10.8	11.5	9.8	
	♀	8.5	12.0	11.1	9.1	
641276 ^a	♂					
	♀					
641350	♂	10.0	11.3	12.2	11.6	
	♀	10.6	11.1	10.9	9.7	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Postculling

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641376	♂	12.6	12.3	11.8	13.0	
	♀	12.0	10.8	10.0		
641315	♂	10.4	10.6	9.4	8.7	
	♀	11.0	10.1	9.4	9.7	
641286	♂	10.3	11.4	11.1	11.4	
	♀	10.5	10.6	9.9	10.4	
641231	♂	12.2	11.7	12.0	12.6	
	♀	11.3	11.2	8.9	11.1	
641338	♂	12.1	11.0	11.8	12.3	
	♀	11.5	9.5	12.0	11.1	
641318	♂	9.0	10.0	9.8	9.9	
	♀	8.8	9.0	9.1	9.7	
641373	♂	9.9	10.7	10.3	11.0	
	♀	10.0	9.5	10.8	9.9	
641365	♂	12.7	11.3	12.8	10.5	12.1
	♀	11.9	11.7	12.0		
641348	♂	10.6	12.2	12.4	11.0	
	♀	10.8	11.3	11.0	12.1	
641323	♂	11.0	9.7	9.9	10.7	
	♀	10.4	10.0	9.4	9.2	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Postculling

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	Pup Number			
		1	2	3	4
641271	♂	10.9	10.4	10.3	10.7
	♀	9.7	10.0	10.8	9.3
641280 ^a	♂				
	♀				
641252 ^a	♂				
	♀				
641386 ^a	♂				
	♀				
641277	♂	11.6	10.1	11.2	10.4
	♀	10.9	10.5	10.7	10.4
641233	♂	10.9	9.5	10.8	10.2
	♀	9.6	10.3	9.7	9.4
641238	♂	13.5	12.4	13.5	12.9
	♀	13.0	12.0	11.9	12.4
641235	♂	10.5	10.7	10.5	9.7
	♀	10.1	9.3	9.5	9.6
641392 ^{b, c}	♂	6.8	6.5	6.4	6.9
	♀	6.3	7.2	6.4	6.3
641360	♂	11.4	11.8	12.1	10.1
	♀	11.4	11.2	11.4	11.4

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Postculling

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	Pup Number			
		1	2	3	4
641381 ^a	♂				
	♀				
641333 ^a	♂				
	♀				
641239	♂	9.6	9.2	9.1	8.9
	♀	9.0	9.5	7.5	10.1
641228 ^a	♂				
	♀				
641334	♂	9.6	10.6	8.4	9.7
	♀	9.0	10.2	7.5	8.8
641347	♂	14.7	13.0	14.2	13.9
	♀	14.1	13.9	13.5	11.8
641367 ^a	♂				
	♀				
641303	♂	12.1	12.5	11.4	11.3
	♀	12.0	11.8	11.7	10.8
641339	♂	11.9	11.6	11.3	11.2
	♀	10.8	11.9	10.3	11.1
641316	♂	11.4	11.4	11.7	11.0
	♀	11.0	11.8	10.0	10.4

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Postculling

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	Pup Number					
		1	2	3	4	5	6
641265	♂	13.6	12.9	13.4	12.7	13.5	11.4
	♀	11.6	13.0				
641393	♂	11.3	11.3	10.3	10.5		
	♀	10.6	10.6	10.9	11.1		
641402	♂	8.7	9.1				
	♀	7.3	8.9	8.5	9.2		
641246	♂	10.9	10.6	10.9	11.8		
	♀	10.1	10.9	10.5	10.6		
641320	♂	11.2	10.7	10.9	12.1		
	♀	10.3	10.0	10.5	10.6		
641353	♂	10.7	10.0	10.2	11.3		
	♀	8.6	9.0	9.0	9.5		
641401	♂	8.2	9.1	9.7	10.1		
	♀	9.8	9.0	9.5	9.7		
641307	♂	11.9	11.5	11.3	10.2	9.6	
	♀	10.4	11.0	9.2			
641368	♂	9.1	10.2	9.6	11.3		
	♀	10.3	10.1	8.3	9.3		
641343	♂	10.7	11.5	10.8	11.6		
	♀	11.7	11.9	11.8	11.0		

Individual Pup Weights (grams): F₁ Generation
Lactation Day 4 Postculling

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	Pup Number			
		1	2	3	4
641254	♂	6.8	7.6	10.3	10.4
	♀	8.6	9.9	9.2	8.4
641242 ^a	♂				
	♀				
641284	♂	10.5	12.2	10.6	10.1
	♀	10.0	10.1	10.8	10.6
641230	♂	10.5	10.5	10.1	10.5
	♀	10.2	10.3	10.4	10.1
641273 ^a	♂				
	♀				
641296	♂	9.5	10.5	10.4	9.6
	♀	8.9	9.4	10.0	9.6
641248	♂	12.4	12.1	11.4	12.0
	♀	9.8	10.5	11.3	9.7
641370	♂	12.5	12.2	12.8	12.7
	♀	12.3	11.3	11.8	11.3
641243	♂	10.1	10.0	9.5	9.3
	♀	9.6	10.1	9.1	10.3
641329	♂	10.6	10.2	9.6	10.4
	♀	10.8	10.0	10.6	9.9

Individual Pup Weights (grams): F1 Generation
Lactation Day 7

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>				
		1	2	3	4	5
641345	♂	18.2	18.7	17.5	18.6	
	♀	18.2	16.6	18.9	17.4	
641236	♂	17.8	19.4	16.3	14.2	
	♀	15.3	18.1	14.5	17.1	
641337	♂	20.3	19.5	21.2	19.6	20.5
	♀	19.2	20.2	20.5		
641328	♂	21.7	22.0	18.8	20.0	
	♀	20.4	20.3	19.6	20.4	
641310	♂	18.6	15.2	16.2	17.8	
	♀	17.8	18.0	16.7	17.6	
641324	♂	22.3	21.5	20.4	21.0	
	♀	19.0	20.4	21.4	23.3	
641371	♂	16.3	20.1	19.8	17.9	
	♀	18.2	15.0	17.5	17.6	
641332	♂	16.5	16.4	17.7	17.1	
	♀	17.2	15.2	16.6	15.5	
641358 ^a	♂					
	♀					
641389 ^a	♂					
	♀					

Individual Pup Weights (grams): F₁ Generation
Lactation Day 7

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>				
		1	2	3	4	5
641317	♂	20.8	21.5	18.9		
	♀	20.1	20.9	20.6		
641255	♂	16.6	16.2	17.2	16.7	
	♀	14.6	13.7	15.4	16.6	
641403	♂	18.6	18.5	18.3	19.2	
	♀	18.8	17.1	18.3	18.3	
641388	♂	20.0	19.6	19.6		
	♀	18.9	18.0	18.2	18.8	18.9
641261	♂	19.0	19.6	19.0		
	♀	18.1	17.1	18.8	18.3	17.8
641306	♂	15.7	19.0	17.9	16.6	
	♀	17.9	14.6	15.9	17.1	
641349	♂	16.5	14.2	16.5	19.7	
	♀	16.5	16.7	16.1	18.0	
641346	♂	21.2	20.3	21.0	17.7	
	♀	18.2	20.4	20.2	19.8	
641308	♂	17.5	18.6	11.5	15.0	
	♀	14.7	16.0	17.1	17.3	
641289	♂	16.0	13.9	14.7	15.1	
	♀	15.0	15.1	14.3	12.4	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 7

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641378	♂	16.3	16.4	16.8	18.9	
	♀	17.0	17.1	13.8	13.6	
641263	♂	17.1	15.9	16.0	15.4	
	♀	16.2	15.7	16.3	16.6	
641297	♂	19.2	18.4	18.2	19.0	
	♀	17.8	18.1	19.0	18.7	
641304	♂	25.5				
	♀	24.8	23.1	24.7	20.5	23.5
641274 ^a	♂					
	♀					
641351	♂	15.9	15.0	15.6	16.8	
	♀	15.7	16.3	13.8	15.2	
641272	♂	17.3	19.4	19.5	20.2	
	♀	18.7	19.8	18.0	17.3	
641258	♂	19.2	18.4	16.9	19.2	
	♀	14.0	18.8	20.1	15.9	
641276 ^a	♂					
	♀					
641350	♂	19.4	16.8	18.6	19.3	
	♀	17.7	17.6	18.1	15.9	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 7

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641376	♂	18.8	17.1	18.6	16.0	
	♀	13.2	18.3	16.8		
641315	♂	16.9	15.1	15.8	17.5	
	♀	16.7	17.7	16.7	15.5	
641286	♂	18.4	17.1	17.3	16.1	
	♀	16.5	16.4	16.4	17.5	
641231	♂	18.5	19.9	18.8	20.0	
	♀	14.6	18.0	18.5	17.5	
641338	♂	19.6	19.4	19.7	18.2	
	♀	19.8	18.8	16.1	18.8	
641318	♂	16.6	16.3	16.1	15.1	
	♀	12.9	13.7	16.1	14.6	
641373	♂	16.0	15.8	14.8	16.0	
	♀	15.7	15.2	14.8	15.3	
641365	♂	18.1	18.3	18.5	21.5	18.6
	♀	18.7	17.7	18.7		
641348	♂	22.1	19.1	19.2	20.7	
	♀	20.3	18.8	19.3	18.4	
641323	♂	16.8	18.1	17.1	16.6	
	♀	15.7	17.5	15.6	16.7	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 7

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	Pup Number			
		1	2	3	4
641271	♂	17.1	16.0	17.0	17.0
	♀	14.9	15.7	16.1	17.0
641280 ^a	♂				
	♀				
641252 ^a	♂				
	♀				
641386 ^a	♂				
	♀				
641277	♂	18.6	19.5	17.6	17.0
	♀	17.8	17.3	17.8	18.1
641233	♂	16.0	17.3	16.9	16.4
	♀	16.0	17.0	14.9	14.6
641238	♂	20.7	21.3	20.7	19.3
	♀	18.9	20.1	20.5	18.9
641235	♂	15.0	16.0	17.1	16.2
	♀	15.0	14.7	14.7	16.4
641392 ^{b, c}	♂				
	♀				
641360	♂	18.1	19.1	19.2	17.1
	♀	18.2	18.3	18.1	18.0

Individual Pup Weights (grams): F1 Generation
Lactation Day 7

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>			
		1	2	3	4
641381 ^a	♂				
	♀				
641333 ^a	♂				
	♀				
641239	♂	16.4	16.5	15.9	15.2
	♀	16.6	16.4	17.6	13.1
641228 ^a	♂				
	♀				
641334	♂	16.2	13.2	14.9	14.9
	♀	13.5	14.6	12.5	14.8
641347	♂	21.3	20.0	21.2	20.1
	♀	18.6	20.6	20.8	21.4
641367 ^a	♂				
	♀				
641303	♂	18.5	19.0	19.6	18.1
	♀	18.3	18.7	19.8	17.7
641339	♂	17.8	18.1	17.6	18.5
	♀	16.2	17.5	18.2	15.9
641316	♂	18.9	19.2	18.8	18.5
	♀	17.8	19.4	18.7	16.6

Individual Pup Weights (grams): F1 Generation
Lactation Day 7

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>					
		1	2	3	4	5	6
641265	♂	20.7	20.0	20.9	21.3	17.9	20.1
	♀	20.3	18.4				
641393	♂	18.3	17.2	17.9	19.4		
	♀	18.0	18.3	17.2	18.3		
641402	♂	15.2	16.3				
	♀	15.5	16.2	15.3	13.6		
641246	♂	18.1	18.1	19.5	17.9		
	♀	16.5	18.0	17.8	17.5		
641320	♂	19.6	16.5	17.3	17.5		
	♀	17.1	17.1	15.5	17.7		
641353	♂	17.8	16.2	17.8	16.9		
	♀	14.8	14.3	14.8	15.5		
641401	♂	15.3	14.2	14.4	13.0		
	♀	13.6	15.1	15.5	14.5		
641307	♂	19.2	18.8	19.0	15.8	17.0	
	♀	17.3	14.8	17.3			
641368	♂	15.8	18.1	18.0	16.3		
	♀	14.1	17.8	17.3	16.0		
641343	♂	18.5	17.9	19.6	19.8		
	♀	20.1	21.5	17.5	18.8		

Individual Pup Weights (grams): F₁ Generation
Lactation Day 7

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	Pup Number			
		1	2	3	4
641254	♂	13.4	18.3	12.7	18.6
	♀	16.5	15.8	14.7	12.6
641242 ^a	♂				
	♀				
641284	♂	16.8	20.0	17.1	16.9
	♀	18.2	17.3	15.7	17.1
641230	♂	17.0	16.4	17.0	16.9
	♀	16.3	16.8	16.5	16.8
641273 ^a	♂				
	♀				
641296	♂	16.1	16.4	17.9	17.2
	♀	15.8	15.3	15.4	16.6
641248	♂	18.9	19.1	18.2	19.0
	♀	15.9	18.6	16.9	15.9
641370	♂	20.5	20.1	20.2	20.3
	♀	18.3	20.1	18.8	19.1
641243	♂	16.5	16.9	15.5	16.4
	♀	15.0	16.2	16.9	17.7
641329	♂	17.8	16.4	17.3	16.7
	♀	17.1	17.1	16.9	16.3

Individual Pup Weights (grams): F1 Generation
Lactation Day 14

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641345	♂	38.4	37.4	36.3	39.3	
	♀	38.2	38.0	35.2	35.7	
641236	♂	37.9	38.7	37.8	34.0	
	♀	32.3	35.2	36.1	37.6	
641337	♂	38.7	40.0	39.5	37.7	35.9
	♀	37.5	36.7	38.6		
641328	♂	43.7	39.4	41.9	39.1	
	♀	40.3	40.1	40.8	39.8	
641310	♂	38.2	35.9	36.8	32.2	
	♀	36.3	35.5	36.6	37.1	
641324	♂	40.7	43.4	41.8	39.7	
	♀	42.4	42.8	40.1	38.9	
641371	♂	40.3	38.7	41.0	35.4	
	♀	36.5	33.0	36.9	38.0	
641332	♂	38.5	37.4	34.9	35.9	
	♀	36.1	35.9	34.0	35.5	
641358 ^a	♂					
	♀					
641389 ^a	♂					
	♀					

Individual Pup Weights (grams): F₁ Generation
Lactation Day 14

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641317	♂	37.3	36.3	33.6		
	♀	35.7	37.3	36.9		
641255	♂	34.5	35.0	32.9	36.5	
	♀	30.7	30.1	34.0	32.1	
641403	♂	41.7	41.9	41.5	41.8	
	♀	41.4	41.3	39.3	40.3	
641388	♂	39.9	40.2	40.0		
	♀	37.7	38.5	38.2	38.2	37.5
641261	♂	41.4	43.1	41.6		
	♀	38.8	38.8	38.6	41.1	40.7
641306	♂	38.7	37.4	36.2	35.7	
	♀	32.8	38.5	35.1	35.8	
641349	♂	39.5	43.0	33.8	39.1	
	♀	37.7	35.9	40.0	38.8	
641346	♂	36.3	40.6	41.4	41.6	
	♀	37.1	38.4	38.3	39.0	
641308	♂	28.8	34.7	39.2	36.9	
	♀	35.7	35.2	37.4	38.3	
641289	♂	33.4	35.7	32.5	32.9	
	♀	32.5	33.5	35.9	32.9	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 14

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641378	♂	36.9	40.8	37.3	38.1	
	♀	36.6	34.4	37.0	37.1	
641263	♂	35.5	36.8	36.4		
	♀	35.4	35.3	33.9	34.4	35.3
641297	♂	38.1	37.8	37.5	38.2	
	♀	36.4	35.6	37.6	37.3	
641304	♂	49.2				
	♀	46.1	47.3	47.5	45.9	42.3
641274 ^a	♂					
	♀					
641351	♂	35.9	35.0	33.6	37.3	
	♀	35.5	35.0	35.9	32.8	
641272	♂	39.2	40.9	42.1	40.8	
	♀	37.2	38.6	40.7	37.8	
641258	♂	40.1	39.1	36.9	40.3	
	♀	40.7	40.1	35.5	31.9	
641276 ^a	♂					
	♀					
641350	♂	39.2	40.9	39.6	36.3	
	♀	36.8	37.1	36.2	36.9	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 14

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641376	♂	36.0	33.8	32.8	35.3	
	♀	34.5	33.4	29.2		
641315	♂	35.5	38.7	40.9	36.9	
	♀	35.4	38.5	37.6	40.1	
641286	♂	38.1	36.0	39.7	34.7	
	♀	36.0	35.6	35.8	35.8	
641231	♂	37.7	38.3	38.2	38.2	
	♀	37.0	32.3	36.9	35.2	
641338	♂	38.9	39.9	38.2	39.1	
	♀	34.3	40.0	37.8	38.0	
641318	♂	33.6	34.7	33.9	32.1	
	♀	34.8	30.8	30.7	34.1	
641373	♂	35.4	36.2	35.1	34.8	
	♀	33.9	33.7	34.1	34.0	
641365	♂	38.8	37.6	38.5	41.8	38.0
	♀	37.1	37.0	38.9		
641348	♂	38.6	42.3	42.4	39.5	
	♀	39.2	37.6	37.9	39.9	
641323	♂	36.2	36.1	36.9	37.8	
	♀	34.9	35.6	33.7	36.0	

Individual Pup Weights (grams): F1 Generation
Lactation Day 14

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	Pup Number			
		1	2	3	4
641271	♂	35.6	32.3	34.5	34.3
	♀	32.7	34.5	34.3	29.6
641280 ^a	♂				
	♀				
641252 ^a	♂				
	♀				
641386 ^a	♂				
	♀				
641277	♂	40.0	38.0	38.3	40.1
	♀	36.9	38.8	38.2	38.5
641233	♂	36.3	37.6	35.2	36.2
	♀	33.4	34.2	34.7	35.7
641238	♂	42.3	40.8	38.2	41.2
	♀	40.1	39.9	39.2	
641235	♂	32.7	35.4	37.2	35.8
	♀	35.1	30.3	34.0	32.7
641392 ^{b, c}	♂				
	♀				
641360	♂	40.0	36.8	40.1	38.5
	♀	38.2	38.4	38.8	38.0

Individual Pup Weights (grams): F1 Generation
Lactation Day 14

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	Pup Number			
		1	2	3	4
641381 ^a	♂				
	♀				
641333 ^a	♂				
	♀				
641239	♂	38.6	34.5	36.6	34.7
	♀	37.1	39.5	32.9	26.3
641228 ^a	♂				
	♀				
641334	♂	30.5	34.1	34.4	34.0
	♀	30.7	33.7	30.2	34.2
641347	♂	42.3	39.6	42.7	41.5
	♀	41.5	36.7	42.1	41.3
641367 ^a	♂				
	♀				
641303	♂	38.5	38.5	39.8	40.6
	♀	42.2	38.2	40.7	37.8
641339	♂	39.1	39.3	39.0	38.5
	♀	39.1	38.1	36.1	35.1
641316	♂	37.7	39.2	40.1	39.3
	♀	39.0	39.5	35.7	37.5

Individual Pup Weights (grams): F1 Generation
Lactation Day 14

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>					
		1	2	3	4	5	6
641265	♂	44.3	39.9	39.6	43.4	43.3	41.8
	♀	38.2	43.1				
641393	♂	37.5	34.4	35.2	34.0		
	♀	35.6	33.4	32.3	34.9		
641402	♂	33.0	34.3				
	♀	33.0	34.5	30.8	33.9		
641246	♂	34.9	35.3	35.3	36.5		
	♀	34.8	34.9	32.4	33.6		
641320	♂	39.0	37.8	36.6	37.0		
	♀	36.4	35.7	35.4	36.6		
641353	♂	38.9	38.3	38.9	37.2		
	♀	35.9	34.8	35.7	35.4		
641401	♂	35.4	33.4	32.2	36.5		
	♀	35.6	32.8	36.1	35.0		
641307	♂	38.3	39.1	35.5	33.8	38.6	
	♀	36.2	35.9	32.3			
641368	♂	36.1	31.0	34.3	34.9		
	♀	32.1	32.0	34.9	32.9		
641343	♂	40.2	39.3	40.3	39.8		
	♀	39.7	39.4	39.9	37.3		

Individual Pup Weights (grams): F1 Generation
Lactation Day 14

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>			
		1	2	3	4
641254	♂	31.6	31.7	39.7	40.8
	♀	36.8	36.6	31.9	34.3
641242 ^a	♂				
	♀				
641284	♂	38.0	37.6	38.2	42.5
	♀	37.4	35.6	36.4	37.5
641230	♂	34.9	33.6	34.5	34.0
	♀	34.1	33.3	33.6	33.1
641273 ^a	♂				
	♀				
641296	♂	36.9	35.2	34.7	34.3
	♀	34.4	33.0	36.1	33.7
641248	♂	40.6	39.2	40.7	42.3
	♀	36.4	35.3	39.3	35.5
641370	♂	43.1	42.0	40.4	41.9
	♀	39.9	41.6	39.2	39.9
641243	♂	35.3	35.9	33.8	36.4
	♀	36.7	34.1	32.9	34.8
641329	♂	36.6	34.4	33.9	35.5
	♀	33.8	35.3	37.1	33.6

Individual Pup Weights (grams): F₁ Generation
Lactation Day 21

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641345	♂	65.4	59.1	60.1	65.6	
	♀	61.5	59.9	57.6	62.1	
641236	♂	58.1	64.8	65.8	59.3	
	♀	58.3	60.4	55.9	51.2	
641337	♂	59.4	59.8	63.9	56.8	60.8
	♀	62.8	58.7	59.0		
641328	♂	65.0	69.7	68.7	60.8	
	♀	63.6	63.9	59.8	63.4	
641310	♂	56.4	59.2	52.6	57.4	
	♀	54.3	55.9	65.8	60.0	
641324	♂	69.5	70.4	67.9	65.9	
	♀	67.9	65.5	61.3	66.1	
641371	♂	64.5	66.6	63.1	53.0	
	♀	58.4	58.4	57.4	60.9	
641332	♂	54.8	57.9	56.5	54.9	
	♀	57.9	57.2	53.2	56.6	
641358 ^a	♂					
	♀					
641389 ^a	♂					
	♀					

Individual Pup Weights (grams): F1 Generation
Lactation Day 21

Group: II-0

Dose: 0 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>				
		1	2	3	4	5
641317	♂	64.3	59.1	63.7		
	♀	62.2	62.0	60.1		
641255	♂	53.6	58.7	50.5	52.5	
	♀	51.0	48.2	50.0	50.7	
641403	♂	67.0	66.9	70.0	66.7	
	♀	64.3	67.4	63.0	63.7	
641388	♂	66.0	68.3	66.2		
	♀	61.1	59.8	60.8	61.2	61.4
641261	♂	68.6	68.4	64.7		
	♀	62.5	56.9	58.6	57.0	63.9
641306	♂	57.4	61.5	57.9	56.8	
	♀	53.7	62.7	55.4	58.1	
641349	♂	61.0	59.2	52.6	61.8	
	♀	60.1	55.6	59.0	60.9	
641346	♂	67.7	59.0	65.0	66.9	
	♀	58.8	59.9	59.4	59.7	
641308	♂	62.8	57.5	60.5	60.9	
	♀	46.1	61.6	57.1	54.5	
641289	♂	51.7	54.8	53.4	55.7	
	♀	54.6	50.7	49.5	51.9	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 21

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	Pup Number				
		1	2	3	4	5
641378	♂	60.6	58.0	59.8	58.5	
	♀	59.5	56.0	57.5	52.7	
641263	♂	57.2	56.2	56.1		
	♀	54.2	55.2	54.6	54.2	51.0
641297	♂	62.2	62.3	61.5	59.9	
	♀	60.0	63.9	62.4	59.6	
641304	♂	82.3				
	♀	72.2	72.4	76.3	73.9	65.3
641274 ^a	♂					
	♀					
641351	♂	56.1	52.5	51.4	50.2	
	♀	53.7	54.7	52.9	49.4	
641272	♂	61.2	60.1	59.6	60.3	
	♀	53.8	62.4	55.3	52.1	
641258	♂	60.2	64.2	65.3	63.0	
	♀	53.1	67.0	57.0	66.3	
641276 ^a	♂					
	♀					
641350	♂	56.7	64.4	61.5	61.0	
	♀	57.1	58.1	57.7	52.5	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 21

Group: IV-0

Dose: 50 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>				
		1	2	3	4	5
641376	♂	54.4	56.7	54.7	55.0	
	♀	57.5	46.1	49.4		
641315	♂	68.7	63.0	69.0	62.5	
	♀	63.2	61.6	67.8	65.6	
641286	♂	66.1	62.3	64.5	57.6	
	♀	59.1	61.5	63.3	57.9	
641231	♂	59.0	63.2	61.5	62.3	
	♀	54.3	51.8	57.1	56.8	
641338	♂	64.0	63.9	62.4	58.3	
	♀	59.8	57.0	60.0	57.7	
641318	♂	51.9	57.2	56.9	59.0	
	♀	55.7	55.6	51.5	51.4	
641373	♂	61.7	60.7	62.5	56.9	
	♀	58.7	57.1	58.0	56.3	
641365	♂	63.8	59.6	62.5	61.7	59.4
	♀	58.8	60.2	55.7		
641348	♂	70.3	72.9	63.7	60.1	
	♀	65.9	63.7	58.9	63.0	
641323	♂	57.5	57.5	53.3	54.3	
	♀	55.1	52.1	54.4	55.4	

Individual Pup Weights (grams): F₁ Generation
Lactation Day 21

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	Pup Number			
		1	2	3	4
641271	♂	57.8	52.5	56.0	57.4
	♀	58.3	46.9	53.2	52.7
641280 ^a	♂				
	♀				
641252 ^a	♂				
	♀				
641386 ^a	♂				
	♀				
641277	♂	56.3	59.4	59.6	61.2
	♀	58.6	58.3	56.2	58.4
641233	♂	56.0	60.3	56.3	58.0
	♀	51.2	54.4	52.6	56.0
641238	♂	61.1	61.7	70.1	68.6
	♀	61.8	63.4	61.0	
641235	♂	59.2	54.2	57.4	53.4
	♀	48.0	55.0	51.6	56.0
641392 ^{b, c}	♂				
	♀				
641360	♂	65.9	63.2	66.6	62.6
	♀	61.5	62.6	63.1	64.8

Individual Pup Weights (grams): F₁ Generation
Lactation Day 21

Group: VI-0

Dose: 250 mg/kg/day

Animal Number	Pup Sex	Pup Number			
		1	2	3	4
641381 ^a	♂				
	♀				
641333 ^a	♂				
	♀				
641239	♂	57.8	59.7	55.6	64.0
	♀	58.1	60.3	52.7	41.5
641228 ^a	♂				
	♀				
641334	♂	61.4	58.3	54.6	59.9
	♀	57.3	51.1	55.5	58.9
641347	♂	72.0	62.3	69.3	68.7
	♀	67.3	61.7	69.0	64.1
641367 ^a	♂				
	♀				
641303	♂	62.5	62.2	63.2	63.3
	♀	58.2	61.2	64.7	56.4
641339	♂	63.7	62.0	60.5	60.2
	♀	61.4	59.3	57.3	61.9
641316	♂	60.8	63.1	63.3	63.5
	♀	59.2	56.7	60.5	60.6

Individual Pup Weights (grams): F1 Generation
Lactation Day 21

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	<u>Pup Number</u>					
		1	2	3	4	5	6
641265	♂	71.3	67.5	68.5	64.6	64.0	62.2
	♀	65.4	60.2				
641393	♂	56.4	54.8	56.1	59.1		
	♀	52.4	56.9	53.7	54.8		
641402	♂	56.6	57.3				
	♀	52.0	52.2	56.7	50.0		
641246	♂	56.3	57.6	56.7	53.8		
	♀	56.0	52.1	54.2	52.6		
641320	♂	62.5	58.8	59.7	59.9		
	♀	58.2	57.6	56.0	54.6		
641353	♂	61.6	61.5	61.6	57.7		
	♀	56.5	53.4	54.1	55.3		
641401	♂	56.0	55.2	53.7	50.1		
	♀	54.6	48.8	54.8	52.8		
641307	♂	55.6	62.3	63.6	54.8	62.5	
	♀	51.8	55.9	56.2			
641368	♂	54.9	55.6	58.3	53.0		
	♀	56.8	58.3	53.2	53.8		
641343	♂	62.9	65.3	65.7	66.3		
	♀	64.9	64.4	60.6	64.2		

Individual Pup Weights (grams): F1 Generation
Lactation Day 21

Group: VIII-0

Dose: 1000 mg/kg/day

Animal Number	Pup Sex	Pup Number			
		1	2	3	4
641254	♂	65.0	53.6	63.8	53.0
	♀	51.9	57.7	56.3	55.6
641242 ^a	♂				
	♀				
641284	♂	64.7	64.4	62.4	67.0
	♀	59.9	62.7	60.4	58.7
641230	♂	55.7	53.6	59.1	55.7
	♀	52.3	54.0	54.4	52.7
641273 ^a	♂				
	♀				
641296	♂	52.8	56.0	56.2	53.0
	♀	50.8	50.0	55.0	51.2
641248	♂	64.7	64.8	61.9	63.4
	♀	56.2	58.5	58.3	61.2
641370	♂	62.0	69.0	64.4	62.2
	♀	59.0	60.9	64.0	56.9
641243	♂	61.3	60.7	57.3	62.5
	♀	54.7	55.1	61.0	51.0
641329	♂	54.3	56.8	54.2	55.0
	♀	53.4	56.0	50.4	52.3

APPENDIX FF

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL LITTER CLINICAL OBSERVATIONS: F₁ GENERATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL LITTER CLINICAL OBSERVATIONS: F₁ GENERATION

EXPLANATORY NOTES

Footnotes

- a* All pups were dead prior to lactation day 7.

INDIVIDUAL LITTER CLINICAL OBSERVATIONS: F₁ GENERATION

GROUP: II-0 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	LACTATION DAY	SIGN INCIDENCE	OBSERVATIONS
641345			NO ABNORMALITIES DETECTED
641236			NO ABNORMALITIES DETECTED
641337			NO ABNORMALITIES DETECTED
641328			NO ABNORMALITIES DETECTED
641310			NO ABNORMALITIES DETECTED
641324			NO ABNORMALITIES DETECTED
641371			NO ABNORMALITIES DETECTED
641332			NO ABNORMALITIES DETECTED
641317			NO ABNORMALITIES DETECTED
641255			NO ABNORMALITIES DETECTED
641403			NO ABNORMALITIES DETECTED
641388			NO ABNORMALITIES DETECTED
641261			NO ABNORMALITIES DETECTED
641306			NO ABNORMALITIES DETECTED
641349			NO ABNORMALITIES DETECTED
641346			NO ABNORMALITIES DETECTED
641308			NO ABNORMALITIES DETECTED
641289			NO ABNORMALITIES DETECTED

INDIVIDUAL LITTER CLINICAL OBSERVATIONS: F₁ GENERATION

GROUP: IV-0 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	LACTATION DAY	SIGN INCIDENCE	OBSERVATIONS
641378			NO ABNORMALITIES DETECTED
641263			NO ABNORMALITIES DETECTED
641297			NO ABNORMALITIES DETECTED
641304			NO ABNORMALITIES DETECTED
641351			NO ABNORMALITIES DETECTED
641272			NO ABNORMALITIES DETECTED
641258			NO ABNORMALITIES DETECTED
641350			NO ABNORMALITIES DETECTED
641376			NO ABNORMALITIES DETECTED
641315			NO ABNORMALITIES DETECTED
641286			NO ABNORMALITIES DETECTED
641231			NO ABNORMALITIES DETECTED
641338			NO ABNORMALITIES DETECTED
641318			NO ABNORMALITIES DETECTED
641373	DAY 0	1 OF 16	SWOLLEN HEAD
LITTER SUMMARY:		1	SWOLLEN HEAD
641365			NO ABNORMALITIES DETECTED
641348			NO ABNORMALITIES DETECTED
641323			NO ABNORMALITIES DETECTED

INDIVIDUAL LITTER CLINICAL OBSERVATIONS: F₁ GENERATION

GROUP: VI-0 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	LACTATION DAY	SIGN INCIDENCE	OBSERVATIONS
641271			NO ABNORMALITIES DETECTED
641277			NO ABNORMALITIES DETECTED
641233			NO ABNORMALITIES DETECTED
641238			NO ABNORMALITIES DETECTED
641235			NO ABNORMALITIES DETECTED
641392 ^a			NO ABNORMALITIES DETECTED
641360			NO ABNORMALITIES DETECTED
641239			NO ABNORMALITIES DETECTED
641334			NO ABNORMALITIES DETECTED
641347			NO ABNORMALITIES DETECTED
641303			NO ABNORMALITIES DETECTED
641339			NO ABNORMALITIES DETECTED
641316			NO ABNORMALITIES DETECTED

INDIVIDUAL LITTER CLINICAL OBSERVATIONS: F₁ GENERATION

GROUP: VIII-0

DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	LACTATION DAY	SIGN INCIDENCE	OBSERVATIONS
641265			NO ABNORMALITIES DETECTED
641393			NO ABNORMALITIES DETECTED
641402			NO ABNORMALITIES DETECTED
641246			NO ABNORMALITIES DETECTED
641320			NO ABNORMALITIES DETECTED
641353			NO ABNORMALITIES DETECTED
641401			NO ABNORMALITIES DETECTED
641307			NO ABNORMALITIES DETECTED
641368			NO ABNORMALITIES DETECTED
641343			NO ABNORMALITIES DETECTED
641254	DAY 21	1 OF 8	CORNEAL OPACITY LEFT EYE(S)
	LITTER SUMMARY:	1	CORNEAL OPACITY
641284			NO ABNORMALITIES DETECTED
641230			NO ABNORMALITIES DETECTED
641296			NO ABNORMALITIES DETECTED
641248			NO ABNORMALITIES DETECTED
641370			NO ABNORMALITIES DETECTED
641243			NO ABNORMALITIES DETECTED
641329			NO ABNORMALITIES DETECTED

APPENDIX GG

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHTS: F₁ GENERATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHTS: F₁ GENERATION

EXPLANATORY NOTES

Notes

Day of weaning = Test Day 1.

INDIVIDUAL BODY WEIGHTS (grams) OF F₁ MALE RATS

GROUP: I-1

DOSE: 0 MG/KG/DAY

DAYS ON TEST:	1	8	15	22	29	36	43
ANIMAL NUMBER							
644438	65.6	106.7	172.5	227.0	299.1	365.0	404.8
644439	59.3	101.1	163.5	216.1	280.0	332.9	390.2
644440	60.8	107.5	167.0	233.9	282.1	343.1	387.4
644441	60.8	112.8	184.0	264.6	336.7	413.9	477.2
644442	57.4	97.4	159.8	207.5	271.3	326.4	373.4
644443	65.9	109.5	167.0	205.8	272.1	335.2	388.5
644444	53.0	98.4	157.7	219.3	281.2	347.4	398.2
644445	54.9	96.7	165.5	234.0	295.6	362.0	412.6
644446	63.7	104.5	180.3	268.3	338.3	407.6	483.4
644447	52.5	95.5	153.8	215.4	281.9	341.4	389.7
644448	66.7	108.9	174.1	230.5	326.2	395.3	453.8
644449	66.2	104.9	167.2	223.8	291.3	351.9	415.4
644450	64.7	110.3	174.0	237.3	297.5	354.4	398.5
644451	56.8	99.2	156.7	211.7	267.7	330.7	378.2
644452	61.8	102.7	169.4	211.1	294.5	367.3	428.5
644453	66.9	110.9	183.4	254.0	308.3	363.2	401.8
644454	60.9	114.4	178.1	240.8	306.3	372.2	423.0
644455	55.7	95.0	159.0	228.9	285.5	335.6	376.2

INDIVIDUAL BODY WEIGHTS (grams) OF F₁ MALE RATS

GROUP: III-1

DOSE: 50 MG/KG/DAY

DAYS ON TEST:	1	8	15	22	29	36	43
ANIMAL NUMBER							
644508	58.5	108.9	176.8	226.7	322.9	399.2	468.4
644509	56.1	100.6	167.3	230.1	295.1	362.8	429.1
644510	59.9	113.8	175.8	259.1	328.9	392.1	459.2
644511	82.3	136.3	212.1	281.5	353.5	417.7	481.4
644512	50.2	87.8	148.3	205.2	277.0	345.1	403.8
644513	60.3	107.7	172.8	239.3	299.1	360.0	413.7
644514	63.0	103.8	165.3	230.1	290.6	345.4	398.1
644515	61.0	112.2	181.3	238.9	288.5	341.5	390.5
644516	55.0	96.4	156.2	200.0	262.0	319.4	368.7
644517	62.5	102.0	167.2	232.5	293.7	363.3	421.3
644518	57.6	102.7	167.9	233.6	293.7	366.6	421.4
644519	62.3	107.4	185.0	259.5	322.0	361.0	422.1
644520	58.3	101.3	161.8	222.8	285.2	345.5	409.3
644521	59.0	102.6	170.8	235.7	299.7	361.7	408.8
644522	56.9	103.8	163.7	220.4	271.8	317.6	361.2
644523	59.4	104.7	172.5	244.8	316.2	383.4	434.3
644524	60.1	97.2	161.9	223.5	290.6	370.6	431.8
644525	54.3	105.7	175.8	207.1	305.9	367.3	428.6

INDIVIDUAL BODY WEIGHTS (grams) OF F₁ MALE RATS

GROUP: V-1

DOSE: 250 MG/KG/DAY

DAYS ON TEST:	1	8	15	22	29	36	43
ANIMAL NUMBER							
644577	61.2	103.2	162.6	224.6	282.9	330.1	382.3
644578	57.4	99.4	168.7	223.9	300.1	374.5	425.1
644579	58.0	97.2	154.3	198.7	268.7	322.5	373.6
644580	68.6	114.1	184.5	236.7	329.9	414.6	465.8
644581	53.4	87.6	143.2	192.6	232.1	276.9	314.5
644582	62.6	103.8	173.3	231.6	283.3	330.8	376.5
644583	64.0	108.3	172.7	243.1	313.1	377.5	431.7
644584	59.9	104.0	168.9	232.1	293.2	356.2	379.4
644585	68.7	110.3	172.4	239.1	299.5	358.6	413.9
644586	63.3	111.7	180.7	250.5	310.0	365.2	405.6
644587	60.2	107.5	170.9	225.3	307.0	372.0	428.3
644588	63.5	98.9	161.1	222.8	281.3	322.6	358.5

INDIVIDUAL BODY WEIGHTS (grams) OF F₁ MALE RATS

GROUP: VII-1

DOSE: 1000 MG/KG/DAY

DAYS ON TEST:	1	8	15	22	29	36	43
ANIMAL NUMBER							
644625	62.2	116.2	185.8	261.0	325.6	401.7	457.2
644626	59.1	104.2	170.4	229.1	300.3	355.0	406.2
644627	57.3	96.6	155.2	224.8	287.9	345.9	395.2
644628	53.8	91.6	149.1	207.6	265.6	316.9	366.6
644629	59.9	110.7	176.4	240.7	304.6	364.1	413.2
644630	57.7	97.6	151.0	210.4	268.4	320.4	365.8
644631	50.1	80.9	135.0	193.1	244.0	304.0	344.3
644632	62.5	103.3	160.4	223.9	282.1	336.4	389.9
644633	53.0	94.4	154.5	206.3	295.2	361.7	423.3
644634	66.3	112.9	174.5	241.6	308.1	366.9	414.0
644635	53.0	86.5	150.9	218.9	287.3	351.7	403.9
644636	67.0	109.9	179.7	239.6	319.0	398.1	458.3
644637	55.7	98.6	157.2	195.0	262.0	330.2	382.7
644638	53.0	93.1	153.0	212.2	266.6	329.9	379.9
644639	63.4	110.4	179.2	245.9	306.8	367.4	424.3
644640	62.2	113.2	183.7	251.3	312.8	366.7	413.0
644641	62.5	103.8	168.6	230.9	289.7	346.4	402.4
644642	55.0	90.1	146.5	204.2	248.1	297.0	344.9

INDIVIDUAL BODY WEIGHTS (grams) OF F₁ FEMALE RATS

GROUP: II-1

DOSE: 0 MG/KG/DAY

DAYS ON TEST:	1	8	15	22	29	36	43
ANIMAL NUMBER							
644698	62.1	102.1	152.0	181.8	202.6	221.5	238.0
644699	51.2	93.4	136.9	161.3	189.9	209.9	231.1
644700	59.0	97.0	134.6	184.5	212.7	233.9	252.9
644701	63.4	102.5	149.5	198.6	217.8	249.9	278.4
644702	60.0	98.0	149.0	182.0	202.4	223.5	242.6
644703	66.1	97.5	140.0	178.8	206.8	230.5	260.4
644704	60.9	105.2	144.5	191.2	221.0	253.5	271.4
644705	56.6	95.7	148.9	190.3	219.3	246.5	256.9
644706	60.1	101.5	142.3	178.3	196.9	220.4	240.4
644707	50.7	96.8	145.1	185.1	205.9	233.7	254.9
644708	63.7	98.4	146.3	191.6	227.3	245.2	270.7
644709	61.4	101.8	150.2	184.0	207.0	228.8	245.0
644710	63.9	102.3	154.7	187.8	205.0	229.1	245.0
644711	58.1	97.0	146.1	179.5	208.0	225.5	231.3
644712	60.9	91.8	138.5	174.6	202.7	217.8	252.9
644713	59.7	107.1	156.2	194.4	216.8	232.4	245.6
644714	54.5	92.7	137.0	169.6	194.2	212.3	230.6
644715	51.9	88.7	128.5	165.8	181.8	212.0	213.1

INDIVIDUAL BODY WEIGHTS (grams) OF F₁ FEMALE RATS

GROUP: IV-1

DOSE: 50 MG/KG/DAY

DAYS ON TEST:	1	8	15	22	29	36	43
ANIMAL NUMBER							
644770	52.7	93.3	140.8	173.7	206.8	225.1	249.7
644771	51.0	86.3	129.2	163.4	184.9	197.3	225.9
644772	59.6	91.8	136.8	170.6	201.1	223.8	251.0
644773	65.3	103.9	158.3	204.3	231.7	256.1	285.1
644774	49.4	88.4	141.0	177.3	206.8	233.8	249.0
644775	52.1	85.9	127.5	165.3	190.2	205.5	231.3
644776	66.3	100.1	145.8	175.3	203.9	223.1	241.4
644777	52.5	96.0	146.3	176.8	201.1	218.1	235.2
644778	49.4	85.4	124.0	157.2	184.3	207.5	222.1
644779	65.6	101.3	151.6	165.4	174.7	204.8	216.1
644780	57.9	101.5	148.2	186.8	214.1	246.9	269.1
644781	56.8	101.1	155.0	193.4	217.2	238.6	257.1
644782	57.7	92.5	134.0	164.4	174.6	190.9	206.5
644783	51.4	89.3	143.0	186.8	211.7	240.9	272.6
644784	56.3	102.7	149.1	174.4	213.4	228.1	255.4
644785	55.7	93.5	140.5	177.2	200.0	222.1	235.3
644786	63.0	101.7	162.4	189.9	231.8	270.1	302.8
644787	55.4	99.4	154.3	198.4	225.8	259.2	273.1

INDIVIDUAL BODY WEIGHTS (grams) OF F₁ FEMALE RATS

GROUP: VI-1

DOSE: 250 MG/KG/DAY

DAYS ON TEST:	1	8	15	22	29	36	43
ANIMAL NUMBER							
644842	52.7	92.5	145.7	183.9	186.3	206.2	211.4
644843	58.4	99.2	151.8	197.9	228.1	254.2	283.7
644844	56.0	91.4	139.6	175.7	200.5	228.8	251.3
644845	61.0	96.1	140.1	177.5	201.3	220.2	234.9
644846	50.6	88.5	137.5	170.8	193.3	215.2	238.5
644847	64.8	103.6	158.3	204.3	227.4	257.4	285.8
644848	41.5	73.2	115.1	147.4	174.6	196.2	216.3
644849	58.9	96.8	139.5	169.1	208.3	224.5	243.7
644850	64.1	113.0	166.5	206.5	235.9	275.6	312.7
644851	56.4	100.4	146.4	180.6	202.0	218.0	233.5
644852	61.9	103.5	157.5	204.9	228.7	252.8	287.7
644853	60.6	93.6	139.0	170.8	186.4	198.4	215.3

INDIVIDUAL BODY WEIGHTS (grams) OF F₁ FEMALE RATS

GROUP: VIII-1

DOSE: 1000 MG/KG/DAY

DAYS ON TEST:	1	8	15	22	29	36	43
ANIMAL NUMBER							
644889	60.2	100.5	151.1	186.3	207.6	239.9	258.0
644890	54.8	95.1	141.8	175.0	185.1	198.4	203.9
644891	50.0	88.0	129.3	177.7	206.2	227.7	247.9
644892	52.6	86.3	130.6	168.3	189.3	216.1	228.0
644893	54.6	93.5	142.2	175.2	197.8	216.7	228.6
644894	55.3	87.7	122.6	154.1	172.7	188.0	210.0
644895	52.8	87.3	134.2	171.3	195.7	239.7	249.7
644896	56.2	86.5	139.2	179.3	204.2	226.5	248.5
644897	53.8	89.4	135.1	176.9	202.0	229.8	245.0
644898	64.2	104.5	154.2	183.5	214.1	242.6	257.8
644899	55.6	90.2	136.6	172.8	201.7	223.4	241.6
644900	58.7	97.9	153.0	187.9	214.4	243.7	265.0
644901	52.7	96.9	150.8	187.5	213.2	251.9	259.8
644902	51.2	87.4	134.9	161.2	182.6	204.3	229.4
644903	61.2	97.8	138.0	174.8	198.4	222.1	242.5
644904	56.9	101.7	147.5	178.9	197.5	220.5	238.1
644905	51.0	85.5	134.3	172.8	205.3	231.5	248.5
644906	52.3	89.4	136.5	161.6	174.2	211.4	218.9

APPENDIX HH

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHT GAINS: F₁ GENERATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL BODY WEIGHT GAINS: F₁ GENERATION

EXPLANATORY NOTES

Notes

Day of weaning = Test Day 1.

INDIVIDUAL BODY WEIGHT GAINS (grams) OF F₁ MALE RATS

GROUP: I-1

DOSE: 0 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644438	41.1	65.8	54.5	72.1	65.9	39.8
644439	41.8	62.4	52.6	63.9	52.9	57.3
644440	46.7	59.5	66.9	48.2	61.0	44.3
644441	52.0	71.2	80.6	72.1	77.2	63.3
644442	40.0	62.4	47.7	63.8	55.1	47.0
644443	43.6	57.5	38.8	66.3	63.1	53.3
644444	45.4	59.3	61.6	61.9	66.2	50.8
644445	41.8	68.8	68.5	61.6	66.4	50.6
644446	40.8	75.8	88.0	70.0	69.3	75.8
644447	43.0	58.3	61.6	66.5	59.5	48.3
644448	42.2	65.2	56.4	95.7	69.1	58.5
644449	38.7	62.3	56.6	67.5	60.6	63.5
644450	45.6	63.7	63.3	60.2	56.9	44.1
644451	42.4	57.5	55.0	56.0	63.0	47.5
644452	40.9	66.7	41.7	83.4	72.8	61.2
644453	44.0	72.5	70.6	54.3	54.9	38.6
644454	53.5	63.7	62.7	65.5	65.9	50.8
644455	39.3	64.0	69.9	56.6	50.1	40.6

INDIVIDUAL BODY WEIGHT GAINS (grams) OF F₁ MALE RATS

GROUP: III-1

DOSE: 50 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644508	50.4	67.9	49.9	96.2	76.3	69.2
644509	44.5	66.7	62.8	65.0	67.7	66.3
644510	53.9	62.0	83.3	69.8	63.2	67.1
644511	54.0	75.8	69.4	72.0	64.2	63.7
644512	37.6	60.5	56.9	71.8	68.1	58.7
644513	47.4	65.1	66.5	59.8	60.9	53.7
644514	40.8	61.5	64.8	60.5	54.8	52.7
644515	51.2	69.1	57.6	49.6	53.0	49.0
644516	41.4	59.8	43.8	62.0	57.4	49.3
644517	39.5	65.2	65.3	61.2	69.6	58.0
644518	45.1	65.2	65.7	60.1	72.9	54.8
644519	45.1	77.6	74.5	62.5	39.0	61.1
644520	43.0	60.5	61.0	62.4	60.3	63.8
644521	43.6	68.2	64.9	64.0	62.0	47.1
644522	46.9	59.9	56.7	51.4	45.8	43.6
644523	45.3	67.8	72.3	71.4	67.2	50.9
644524	37.1	64.7	61.6	67.1	80.0	61.2
644525	51.4	70.1	31.3	98.8	61.4	61.3

INDIVIDUAL BODY WEIGHT GAINS (grams) OF F₁ MALE RATS

GROUP: V-1

DOSE: 250 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644577	42.0	59.4	62.0	58.3	47.2	52.2
644578	42.0	69.3	55.2	76.2	74.4	50.6
644579	39.2	57.1	44.4	70.0	53.8	51.1
644580	45.5	70.4	52.2	93.2	84.7	51.2
644581	34.2	55.6	49.4	39.5	44.8	37.6
644582	41.2	69.5	58.3	51.7	47.5	45.7
644583	44.3	64.4	70.4	70.0	64.4	54.2
644584	44.1	64.9	63.2	61.1	63.0	23.2
644585	41.6	62.1	66.7	60.4	59.1	55.3
644586	48.4	69.0	69.8	59.5	55.2	40.4
644587	47.3	63.4	54.4	81.7	65.0	56.3
644588	35.4	62.2	61.7	58.5	41.3	35.9

INDIVIDUAL BODY WEIGHT GAINS (grams) OF F₁ MALE RATS

GROUP: VII-1

DOSE: 1000 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644625	54.0	69.6	75.2	64.6	76.1	55.5
644626	45.1	66.2	58.7	71.2	54.7	51.2
644627	39.3	58.6	69.6	63.1	58.0	49.3
644628	37.8	57.5	58.5	58.0	51.3	49.7
644629	50.8	65.7	64.3	63.9	59.5	49.1
644630	39.9	53.4	59.4	58.0	52.0	45.4
644631	30.8	54.1	58.1	50.9	60.0	40.3
644632	40.8	57.1	63.5	58.2	54.3	53.5
644633	41.4	60.1	51.8	88.9	66.5	61.6
644634	46.6	61.6	67.1	66.5	58.8	47.1
644635	33.5	64.4	68.0	68.4	64.4	52.2
644636	42.9	69.8	59.9	79.4	79.1	60.2
644637	42.9	58.6	37.8	67.0	68.2	52.5
644638	40.1	59.9	59.2	54.4	63.3	50.0
644639	47.0	68.8	66.7	60.9	60.6	56.9
644640	51.0	70.5	67.6	61.5	53.9	46.3
644641	41.3	64.8	62.3	58.8	56.7	56.0
644642	35.1	56.4	57.7	43.9	48.9	47.9

INDIVIDUAL BODY WEIGHT GAINS (grams) OF F₁ FEMALE RATS

GROUP: II-1

DOSE: 0 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644698	40.0	49.9	29.8	20.8	18.9	16.5
644699	42.2	43.5	24.4	28.6	20.0	21.2
644700	38.0	37.6	49.9	28.2	21.2	19.0
644701	39.1	47.0	49.1	19.2	32.1	28.5
644702	38.0	51.0	33.0	20.4	21.1	19.1
644703	31.4	42.5	38.8	28.0	23.7	29.9
644704	44.3	39.3	46.7	29.8	32.5	17.9
644705	39.1	53.2	41.4	29.0	27.2	10.4
644706	41.4	40.8	36.0	18.6	23.5	20.0
644707	46.1	48.3	40.0	20.8	27.8	21.2
644708	34.7	47.9	45.3	35.7	17.9	25.5
644709	40.4	48.4	33.8	23.0	21.8	16.2
644710	38.4	52.4	33.1	17.2	24.1	15.9
644711	38.9	49.1	33.4	28.5	17.5	5.8
644712	30.9	46.7	36.1	28.1	15.1	35.1
644713	47.4	49.1	38.2	22.4	15.6	13.2
644714	38.2	44.3	32.6	24.6	18.1	18.3
644715	36.8	39.8	37.3	16.0	30.2	1.1

INDIVIDUAL BODY WEIGHT GAINS (grams) OF F₁ FEMALE RATS

GROUP: IV-1

DOSE: 50 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644770	40.6	47.5	32.9	33.1	18.3	24.6
644771	35.3	42.9	34.2	21.5	12.4	28.6
644772	32.2	45.0	33.8	30.5	22.7	27.2
644773	38.6	54.4	46.0	27.4	24.4	29.0
644774	39.0	52.6	36.3	29.5	27.0	15.2
644775	33.8	41.6	37.8	24.9	15.3	25.8
644776	33.8	45.7	29.5	28.6	19.2	18.3
644777	43.5	50.3	30.5	24.3	17.0	17.1
644778	36.0	38.6	33.2	27.1	23.2	14.6
644779	35.7	50.3	13.8	9.3	30.1	11.3
644780	43.6	46.7	38.6	27.3	32.8	22.2
644781	44.3	53.9	38.4	23.8	21.4	18.5
644782	34.8	41.5	30.4	10.2	16.3	15.6
644783	37.9	53.7	43.8	24.9	29.2	31.7
644784	46.4	46.4	25.3	39.0	14.7	27.3
644785	37.8	47.0	36.7	22.8	22.1	13.2
644786	38.7	60.7	27.5	41.9	38.3	32.7
644787	44.0	54.9	44.1	27.4	33.4	13.9

INDIVIDUAL BODY WEIGHT GAINS (grams) OF F₁ FEMALE RATS

GROUP: VI-1

DOSE: 250 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644842	39.8	53.2	38.2	2.4	19.9	5.2
644843	40.8	52.6	46.1	30.2	26.1	29.5
644844	35.4	48.2	36.1	24.8	28.3	22.5
644845	35.1	44.0	37.4	23.8	18.9	14.7
644846	37.9	49.0	33.3	22.5	21.9	23.3
644847	38.8	54.7	46.0	23.1	30.0	28.4
644848	31.7	41.9	32.3	27.2	21.6	20.1
644849	37.9	42.7	29.6	39.2	16.2	19.2
644850	48.9	53.5	40.0	29.4	39.7	37.1
644851	44.0	46.0	34.2	21.4	16.0	15.5
644852	41.6	54.0	47.4	23.8	24.1	34.9
644853	33.0	45.4	31.8	15.6	12.0	16.9

INDIVIDUAL BODY WEIGHT GAINS (grams) OF F₁ FEMALE RATS

GROUP: VIII-1

DOSE: 1000 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644889	40.3	50.6	35.2	21.3	32.3	18.1
644890	40.3	46.7	33.2	10.1	13.3	5.5
644891	38.0	41.3	48.4	28.5	21.5	20.2
644892	33.7	44.3	37.7	21.0	26.8	11.9
644893	38.9	48.7	33.0	22.6	18.9	11.9
644894	32.4	34.9	31.5	18.6	15.3	22.0
644895	34.5	46.9	37.1	24.4	44.0	10.0
644896	30.3	52.7	40.1	24.9	22.3	22.0
644897	35.6	45.7	41.8	25.1	27.8	15.2
644898	40.3	49.7	29.3	30.6	28.5	15.2
644899	34.6	46.4	36.2	28.9	21.7	18.2
644900	39.2	55.1	34.9	26.5	29.3	21.3
644901	44.2	53.9	36.7	25.7	38.7	7.9
644902	36.2	47.5	26.3	21.4	21.7	25.1
644903	36.6	40.2	36.8	23.6	23.7	20.4
644904	44.8	45.8	31.4	18.6	23.0	17.6
644905	34.5	48.8	38.5	32.5	26.2	17.0
644906	37.1	47.1	25.1	12.6	37.2	7.5

APPENDIX II

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL FOOD CONSUMPTION: F₁ GENERATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL FOOD CONSUMPTION: F₁ GENERATION

EXPLANATORY NOTES

Notes

Day of weaning = Test Day 1.

INDIVIDUAL FOOD CONSUMPTION (grams/day) BY F₁ MALE RATS

GROUP: I-1

DOSE: 0 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644438	13.8	22.0	25.8	31.5	31.3	31.6
644439	14.2	21.9	25.6	28.3	30.3	31.1
644440	14.6	21.6	25.7	27.6	30.5	31.0
644441	15.7	22.2	27.9	32.1	35.1	37.1
644442	12.9	20.4	24.3	27.2	32.0	30.1
644443	14.5	22.4	19.1	28.0	29.3	29.4
644444	13.1	19.9	25.3	29.9	30.7	30.7
644445	13.2	20.5	25.4	29.2	32.3	35.0
644446	13.4	22.6	28.2	32.5	33.0	39.6
644447	13.0	19.1	22.9	27.6	29.3	28.7
644448	13.6	21.1	25.8	34.1	36.0	36.7
644449	14.4	19.4	23.8	28.5	32.0	35.1
644450	14.7	21.8	25.4	30.8	30.2	30.4
644451	12.7	20.6	23.8	29.0	31.4	32.8
644452	13.4	20.0	22.3	29.9	32.3	34.8
644453	15.0	24.6	27.2	30.1	32.7	32.0
644454	14.8	22.1	27.4	32.9	33.9	32.6
644455	12.7	21.0	25.6	28.1	30.8	29.6

INDIVIDUAL FOOD CONSUMPTION (grams/day) BY F₁ MALE RATS

GROUP: III-1

DOSE: 50 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644508	16.0	24.7	27.5	36.3	37.7	40.6
644509	12.9	21.7	26.0	31.2	34.0	37.3
644510	16.1	23.1	29.4	33.8	35.0	39.6
644511	16.7	25.7	28.2	32.5	34.5	35.9
644512	11.9	19.8	23.3	29.5	30.6	31.8
644513	15.2	22.1	26.1	30.0	32.1	33.8
644514	14.0	20.7	27.8	29.4	31.2	31.2
644515	15.7	22.6	26.9	27.9	29.0	28.5
644516	12.1	18.9	18.4	26.6	29.0	29.7
644517	13.0	20.0	25.8	28.3	30.3	33.1
644518	13.2	21.2	26.3	29.9	31.7	31.6
644519	13.2	22.3	28.1	30.8	25.9	34.3
644520	13.7	20.9	24.5	28.8	31.6	34.2
644521	12.8	20.0	25.4	30.6	32.2	34.2
644522	13.5	20.8	24.5	27.3	28.1	28.5
644523	14.8	23.1	27.1	31.9	34.0	33.8
644524	12.5	20.5	25.8	30.2	35.7	37.7
644525	15.1	23.9	26.5	37.2	34.4	35.2

INDIVIDUAL FOOD CONSUMPTION (grams/day) BY F₁ MALE RATS

GROUP: V-1

DOSE: 250 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644577	12.0	19.1	22.3	25.8	26.8	28.4
644578	12.7	22.8	25.8	33.0	34.5	34.8
644579	12.6	19.0	22.5	28.7	29.0	29.8
644580	15.8	24.0	27.3	35.1	39.0	36.8
644581	12.6	19.7	22.7	26.9	29.6	29.8
644582	13.5	18.1	23.9	27.1	27.0	28.3
644583	14.8	22.8	27.0	31.3	34.7	35.0
644584	17.8	18.9	24.7	27.9	30.5	28.3
644585	13.6	22.8	26.5	28.8	31.8	31.7
644586	15.2	23.6	28.3	30.6	31.6	31.3
644587	14.0	22.6	25.8	32.4	33.6	34.9
644588	15.4	21.7	26.9	29.5	29.7	28.3

INDIVIDUAL FOOD CONSUMPTION (grams/day) BY F₁ MALE RATS

GROUP: VII-1

DOSE: 1000 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644625	14.8	22.3	29.3	32.6	35.6	36.8
644626	13.5	21.8	26.5	32.5	31.9	34.3
644627	13.9	22.7	26.4	30.1	31.7	32.5
644628	11.7	20.1	21.7	25.4	27.4	28.6
644629	14.4	22.5	27.4	31.1	34.1	31.3
644630	13.1	18.7	23.5	28.3	27.9	29.2
644631	10.9	16.7	25.4	23.6	27.9	28.7
644632	13.3	21.1	24.7	28.0	29.9	16.9
644633	12.6	20.8	24.8	32.5	34.0	37.4
644634	16.0	21.3	26.2	29.2	31.9	31.1
644635	11.4	20.5	25.7	32.2	34.1	35.5
644636	13.9	21.5	27.7	32.6	34.3	35.8
644637	13.1	18.5	19.9	30.6	34.2	36.9
644638	12.1	18.3	21.5	24.8	27.0	27.9
644639	15.1	22.9	27.7	31.7	33.9	33.4
644640	14.2	23.3	27.7	32.6	34.0	33.2
644641	13.3	19.8	23.8	26.3	27.9	29.2
644642	12.9	18.2	23.4	24.6	27.3	29.2

INDIVIDUAL FOOD CONSUMPTION (grams/day) BY F₁ FEMALE RATS

GROUP: II-1

DOSE: 0 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644698	13.8	19.5	20.0	20.6	19.1	19.4
644699	14.2	18.1	17.8	21.6	20.8	21.4
644700	14.0	18.9	18.5	22.5	22.2	25.4
644701	13.0	19.6	21.8	22.2	23.8	24.9
644702	18.1	18.0	20.7	21.4	19.8	20.6
644703	14.3	18.2	19.5	21.5	22.5	23.5
644704	9.1	20.3	23.5	23.8	25.4	26.5
644705	13.5	19.8	23.6	24.1	24.2	23.6
644706	17.5	17.7	18.5	19.1	19.7	18.9
644707	13.5	19.3	21.3	22.0	23.3	22.6
644708	12.9	18.7	21.1	23.9	21.9	23.2
644709	14.0	18.8	19.9	20.8	20.1	19.3
644710	12.9	18.2	19.0	19.7	20.6	20.2
644711	12.1	19.2	21.5	22.8	21.4	23.7
644712	10.3	16.8	20.0	20.7	18.1	21.8
644713	14.4	20.7	21.2	22.2	21.1	19.9
644714	10.7	17.4	18.3	19.5	19.6	19.1
644715	12.4	17.3	19.4	19.9	20.5	20.9

INDIVIDUAL FOOD CONSUMPTION (grams/day) BY F₁ FEMALE RATS

GROUP: IV-1

DOSE: 50 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644770	12.7	19.3	22.9	24.1	23.0	24.6
644771	12.5	17.3	19.3	19.2	18.4	19.6
644772	12.2	18.3	18.4	20.4	21.8	21.3
644773	12.7	21.5	23.3	23.5	24.5	26.2
644774	11.6	19.9	22.5	21.8	23.2	22.6
644775	10.6	16.4	18.8	19.7	19.6	21.5
644776	12.8	16.9	18.3	20.4	20.7	22.0
644777	11.7	19.2	20.1	19.1	19.1	20.8
644778	13.7	15.2	16.0	18.2	18.6	18.1
644779	13.1	19.5	18.0	17.2	19.8	18.8
644780	13.7	18.9	20.0	21.8	23.2	23.6
644781	13.4	19.3	3.6	20.6	23.7	24.2
644782	11.9	17.5	19.3	18.1	19.2	18.9
644783	11.5	17.5	20.5	20.4	22.4	24.2
644784	14.1	19.9	20.9	22.2	21.5	23.3
644785	13.0	19.0	20.4	21.6	21.0	20.7
644786	13.6	20.9	22.8	23.6	25.3	27.9
644787	13.3	20.3	22.4	22.6	24.0	24.4

INDIVIDUAL FOOD CONSUMPTION (grams/day) BY F₁ FEMALE RATS

GROUP: VI-1

DOSE: 250 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644842	12.3	19.5	21.8	18.9	18.8	17.3
644843	12.2	18.0	21.3	22.0	21.2	24.0
644844	13.2	18.6	20.3	21.1	20.6	22.2
644845	12.8	17.9	20.7	21.8	20.7	21.7
644846	12.0	17.0	19.5	19.6	19.7	20.6
644847	13.1	19.0	22.1	22.3	23.9	25.3
644848	12.7	18.5	21.2	21.8	23.9	25.5
644849	13.4	16.8	19.0	20.6	20.4	22.8
644850	15.1	21.3	22.7	24.4	29.6	28.9
644851	14.0	20.5	20.8	20.3	21.4	21.9
644852	13.7	20.2	23.1	24.4	23.4	23.9
644853	11.8	17.7	19.1	18.2	17.3	17.5

INDIVIDUAL FOOD CONSUMPTION (grams/day) BY F₁ FEMALE RATS

GROUP: VIII-1

DOSE: 1000 MG/KG/DAY

DAYS ON TEST:	1-8	8-15	15-22	22-29	29-36	36-43
ANIMAL NUMBER						
644889	12.5	18.0	19.5	21.0	22.9	23.1
644890	13.5	18.2	18.8	18.8	18.2	16.3
644891	13.0	17.9	20.7	22.5	23.2	22.2
644892	12.1	16.6	18.1	18.3	19.8	18.8
644893	12.2	17.9	18.9	19.6	19.2	18.4
644894	12.4	16.2	17.3	18.2	17.8	18.3
644895	11.8	17.0	19.5	21.1	23.1	20.3
644896	11.6	19.5	23.9	19.5	24.9	24.3
644897	13.1	17.5	20.2	22.4	23.0	22.8
644898	14.0	20.1	19.9	20.9	20.9	21.8
644899	11.7	17.7	19.8	21.9	21.6	21.3
644900	14.0	20.1	23.2	23.4	24.4	24.1
644901	14.1	19.4	21.6	21.7	25.2	23.8
644902	11.8	17.6	18.6	19.9	19.5	20.2
644903	13.2	17.0	20.7	9.7	21.2	21.2
644904	13.5	18.2	19.4	19.3	20.6	19.7
644905	12.0	17.9	20.8	24.0	23.7	24.1
644906	11.9	17.4	18.1	17.8	19.5	18.9

APPENDIX JJ

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL CLINICAL OBSERVATIONS: F₁ GENERATION

REPRODUCTIVE TOXICOLOGY EVALUATIONS

INDIVIDUAL CLINICAL OBSERVATIONS: F₁ GENERATION

EXPLANATORY NOTES

Notes

Day of weaning = Test Day 1.

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ MALE RATS

GROUP: I-1 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644438	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	30	30
644439	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644440	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	29	29
644441	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	30	30
644442	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	25	25
644443	HEAD CAUGHT IN CAGE AS CLOSING DOOR ^a PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	15 29	15 29
644444	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	33	33
644445	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	31	31
644446	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644447	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644448	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	30	30
644449	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644450	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	30	30

^a This clinical observation is not reported in the Summary Data Tables.

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ MALE RATS

GROUP: I-1 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644451	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644452	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644453	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644454	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	31	31
644455	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ MALE RATS

GROUP: III-1 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644508	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644509	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644510	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644511	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	38	38
644512	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	26	26
644513	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644514	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644515	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	26	26
644516	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644517	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	29	29
644518	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	29	29
644519	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644520	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	30	30
644521	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ MALE RATS

GROUP: III-1 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644522	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	30	30
644523	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644524	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	32	32
644525	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	30	30

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ MALE RATS

GROUP: V-1 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644577	PREPUTIAL SEPARATION INCOMPLETE SACRIFICED BY DESIGN TEST DAY 43	43	43
644578	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644579	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644580	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644581	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	39	39
644582	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644583	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644584	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644585	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	29	29
644586	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644587	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644588	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	29	29

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ MALE RATS

GROUP: VII-1 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644625	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644626	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	30	30
644627	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	29	29
644628	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	31	31
644629	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	29	29
644630	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644631	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	32	32
644632	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	29	29
644633	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	30	30
644634	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	30	30
644635	CORNEAL OPACITY LEFT EYE(S) PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	1 28	43 28
644636	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644637	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	33	33

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ MALE RATS

GROUP: VII-1 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644638	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644639	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	26	26
644640	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	27	27
644641	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28
644642	PREPUTIAL SEPARATION COMPLETE SACRIFICED BY DESIGN TEST DAY 43	28	28

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ FEMALE RATS

GROUP: II-1 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644698	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	20	20
644699	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	10	10
644700	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	14	14
644701	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644702	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	10	10
644703	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	10	10
644704	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644705	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644706	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644707	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644708	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644709	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	20	20
644710	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	12	12
644711	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	10	10

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ FEMALE RATS

GROUP: II-1 DOSE: 0 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644712	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644713	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644714	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	15	15
644715	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	9	9

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ FEMALE RATS

GROUP: IV-1 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644770	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644771	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	10	10
644772	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	17	17
644773	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	15	15
644774	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	20	20
644775	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644776	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644777	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644778	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	15	15
644779	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644780	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644781	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	19	19
644782	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	15	15
644783	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	18	18

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ FEMALE RATS

GROUP: IV-1 DOSE: 50 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644784	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	21	21
644785	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	18	18
644786	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	10	10
644787	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	10	10

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ FEMALE RATS

GROUP: VI-1 DOSE: 250 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644842	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644843	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644844	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644845	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644846	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	17	17
644847	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644848	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	21	21
644849	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	14	14
644850	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	21	21
644851	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	10	10
644852	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	17	17
644853	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ FEMALE RATS

GROUP: VIII-1 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644889	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644890	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	12	12
644891	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	16	16
644892	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644893	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644894	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	20	20
644895	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	12	12
644896	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	13	13
644897	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	9	9
644898	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	12	12
644899	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	15	15
644900	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644901	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	15	15
644902	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	10	10

INDIVIDUAL CLINICAL OBSERVATIONS IN F₁ FEMALE RATS

GROUP: VIII-1 DOSE: 1000 MG/KG/DAY

ANIMAL NUMBER	OBSERVATION	FIRST DAY OBSERVED	LAST DAY OBSERVED
644903	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	16	16
644904	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644905	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11
644906	VAGINAL PATENCY COMPLETE SACRIFICED BY DESIGN TEST DAY 43	11	11