

Experiment Number: 20712 - 04

**P18: INCIDENCE RATES OF NON-NEOPLASTIC LESIONS BY ANATOMIC SITE (a) WITH
AVERAGE SEVERITY GRADES[b]**

Date Report Requested: 09/14/2020

Test Type: CHRONIC

Tris(Chloropropyl)phosphate

Time Report Requested: 10:18:26

Route: DOSED FEED

CAS Number: 13674-84-5

First Dose M/F: 10/25/11 / 10/24/11

Species/Strain: MICE/B6C3F1/N

Lab: BAT

Final 2_Mice

NTP Study Number: C20712
Lock Date: 02/26/2018
Cage Range: ALL
Date Range: ALL
Reasons For Removal: ALL
Removal Date Range: ALL
Treatment Groups: Include ALL
Study Gender: Both
TDMSE Version: 3.0.2.3_002
PWG Approval Date: NONE

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Species/Strain: MICE/B6C3F1/N

Lab: BAT

| B6C3F1/N MICE MALE | 0 ppm males | 1250 ppm males | 2500 ppm males | 5000 ppm males |
|--------------------|-------------|----------------|----------------|----------------|
|--------------------|-------------|----------------|----------------|----------------|

Disposition Summary

| | | | | |
|---|-----------|-----------|-----------|-----------|
| Animals Initially In Study | 50 | 50 | 50 | 50 |
| Early Deaths | | | | |
| Moribund Sacrifice | 1 | 1 | 1 | 3 |
| Natural Death | 10 | 5 | 7 | 3 |
| Survivors | | | | |
| Natural Death | 1 | | | 1 |
| Terminal Sacrifice | 38 | 44 | 42 | 43 |
| Animals Examined Microscopically | 50 | 50 | 50 | 50 |

ALIMENTARY SYSTEM

| | | | | |
|--------------------------------------|----------|---------|---------|---------|
| Esophagus | (50) | (50) | (50) | (50) |
| Gallbladder | (50) | (50) | (46) | (50) |
| Intestine Large, Cecum | (50) | (49) | (50) | (50) |
| Intestine Large, Colon | (50) | (50) | (50) | (50) |
| Intestine Large, Rectum | (50) | (50) | (50) | (50) |
| Intestine Small, Duodenum | (50) | (50) | (50) | (50) |
| Intestine Small, Ileum | (50) | (50) | (50) | (50) |
| Diverticulum | | | | 1 |
| Intestine Small, Jejunum | (50) | (50) | (50) | (50) |
| Liver | (50) | (50) | (50) | (50) |
| Basophilic Focus | 4 | 4 | 3 | 2 |
| Clear Cell Focus | 13 | 18 | 12 | 8 |
| Eosinophilic Focus | 13 | 16 | 9 | 15 |
| Fatty Change, Focal | 3 [1.3] | 4 [1.0] | 1 [2.0] | |
| Mixed Cell Focus | | 2 | 1 | 1 |
| Necrosis | 10 [1.5] | 8 [1.1] | 9 [1.1] | 8 [2.8] |
| Hepatocyte, Cytoplasmic Alteration | 1 [1.0] | | 2 [1.0] | |
| Hepatocyte, Vacuolation, Cytoplasmic | | | 1 [3.0] | |
| Mesentery | (1) | (1) | (1) | (2) |
| Fat, Necrosis | 1 [2.0] | | | |
| Pancreas | (50) | (50) | (50) | (50) |
| Salivary Glands | (50) | (50) | (50) | (50) |
| Stomach, Forestomach | (50) | (50) | (50) | (50) |

a - Number of animals examined microscopically at site and number of animals with lesion

b-Average severity grade(1-minimal;2-mild;3-moderate;4-marked)

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Species/Strain: MICE/B6C3F1/N

Lab: BAT

| B6C3F1/N MICE MALE | 0 ppm males | 1250 ppm males | 2500 ppm males | 5000 ppm males |
|--|--|---|---|--|
| Cyst, Squamous Inflammation, Suppurative Ulcer Epithelium, Hyperplasia, Focal Stomach, Glandular | 1 [1.0] (50) | (50) | 1 [1.0] 1 [2.0] 1 [2.0] (50) | 1 (50) |
| CARDIOVASCULAR SYSTEM | | | | |
| Blood Vessel Heart Cardiomyopathy Polyarteritis Nodosa | (50) (50) 1 [1.0] 1 [2.0] | (49) (50) | (49) (50) | (49) (50) |
| ENDOCRINE SYSTEM | | | | |
| Adrenal Cortex Hyperplasia, Focal Hypertrophy, Focal Inflammation, Suppurative Necrosis Bilateral, Hypertrophy, Focal Adrenal Medulla Necrosis Islets, Pancreatic Parathyroid Gland Pituitary Gland Pars Distalis, Hyperplasia Pars Intermedia, Hyperplasia Thyroid Gland | (50) 1 [2.0] 14 [1.5] (50) (50) (42) (50) 1 [2.0] (50) | (50) 1 [2.0] 20 [1.2] 1 [4.0] 2 [2.0] (50) 1 [4.0] (50) (45) (46) 1 [1.0] (50) | (50) (50) 11 [1.3] 1 [2.0] (50) (50) (46) (48) (50) | (49) 3 [1.0] 6 [1.7] (49) (50) (46) (50) (50) |
| GENERAL BODY SYSTEM | | | | |
| None | | | | |

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b-Average severity grade(1-minimal;2-mild;3-moderate;4-marked)

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Lab: BAT

| B6C3F1/N MICE MALE | 0 ppm males | 1250 ppm males | 2500 ppm males | 5000 ppm males |
|------------------------------|-------------|----------------|----------------|----------------|
| GENITAL SYSTEM | | | | |
| Epididymis | (50) | (50) | (50) | (50) |
| Preputial Gland | (50) | (50) | (50) | (50) |
| Duct, Dilation | | | | 1 [3.0] |
| Prostate | (50) | (50) | (50) | (50) |
| Inflammation, Suppurative | 1 [1.0] | | 2 [3.0] | |
| Seminal Vesicle | (50) | (50) | (50) | (50) |
| Atrophy | | | 1 [4.0] | 1 [2.0] |
| Inflammation, Suppurative | | | 1 [3.0] | |
| Testis | (50) | (50) | (50) | (50) |
| Germinal Epithelium, Atrophy | 1 [1.0] | | 2 [3.0] | 1 [4.0] |

HEMATOPOIETIC SYSTEM

| | | | | |
|--------------------------------------|----------|----------|----------|----------|
| Bone Marrow | (50) | (50) | (50) | (50) |
| Hypocellularity | | | 1 [3.0] | |
| Lymph Node | (4) | (3) | (4) | (3) |
| Mediastinal, Hyperplasia, Lymphocyte | | 1 [3.0] | | |
| Lymph Node, Mandibular | (47) | (50) | (50) | (50) |
| Lymph Node, Mesenteric | (49) | (49) | (50) | (49) |
| Congestion | 1 [4.0] | | | 1 [4.0] |
| Extramedullary Hematopoiesis | 1 [2.0] | | 1 [3.0] | |
| Spleen | (50) | (50) | (50) | (50) |
| Erythrophagocytosis | | | 1 [3.0] | |
| Extramedullary Hematopoiesis | 23 [2.0] | 21 [2.0] | 18 [2.9] | 20 [2.4] |
| Hyperplasia, Lymphocyte | | | 1 [2.0] | 1 [3.0] |
| Thymus | (46) | (47) | (49) | (46) |
| Atrophy | 1 [3.0] | 1 [2.0] | 2 [3.5] | 1 [4.0] |
| Inflammation, Granulomatous | | | 1 [4.0] | |

INTEGUMENTARY SYSTEM

| | | | | |
|------|------|------|------|------|
| Skin | (50) | (50) | (50) | (50) |
|------|------|------|------|------|

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Species/Strain: MICE/B6C3F1/N

Lab: BAT

| B6C3F1/N MICE MALE | 0 ppm males | 1250 ppm males | 2500 ppm males | 5000 ppm males |
|--------------------|-------------|----------------|----------------|----------------|
|--------------------|-------------|----------------|----------------|----------------|

MUSCULOSKELETAL SYSTEM

| | | | | |
|-----------------------|------|------|------|---------|
| Bone | (50) | (50) | (50) | (49) |
| Femur, Increased Bone | | | | 1 [2.0] |
| Skeletal Muscle | (1) | (0) | (1) | (0) |

NERVOUS SYSTEM

| | | | | |
|--------------------------------|---------|------|------|------|
| Brain | (50) | (50) | (50) | (50) |
| Meninges, Polyarteritis Nodosa | 1 [2.0] | | | |

RESPIRATORY SYSTEM

| | | | | |
|-----------------------------------|---------|---------|---------|---------|
| Lung | (50) | (50) | (50) | (50) |
| Infiltration Cellular, Histiocyte | 1 [2.0] | 1 [1.0] | 1 [2.0] | |
| Inflammation, Granulomatous | | 1 [1.0] | | |
| Alveolus, Epithelium, Hyperplasia | 1 [1.0] | | 3 [2.0] | 1 [3.0] |
| Nose | (50) | (50) | (50) | (50) |
| Trachea | (50) | (50) | (49) | (50) |

SPECIAL SENSES SYSTEM

| | | | | |
|-----------------|---------|---------|---------|---------|
| Eye | (50) | (50) | (50) | (50) |
| Phthisis Bulbi | | | 1 [4.0] | |
| Harderian Gland | (50) | (50) | (50) | (50) |
| Hyperplasia | 2 [2.0] | 1 [1.0] | 3 [2.0] | 1 [1.0] |

URINARY SYSTEM

| | | | | |
|---------------------|---------|---------|---------|---------|
| Kidney | (49) | (50) | (50) | (50) |
| Infarct | | | 1 [1.0] | |
| Metaplasia, Osseous | 1 [2.0] | 1 [1.0] | 1 [1.0] | 1 [1.0] |

a - Number of animals examined microscopically at site and number of animals with lesion

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Lab: BAT

| B6C3F1/N MICE MALE | 0 ppm males | 1250 ppm males | 2500 ppm males | 5000 ppm males |
|--------------------------------------|-------------|----------------|----------------|----------------|
| Nephropathy, Chronic Progressive | 40 [1.1] | 45 [1.0] | 43 [1.2] | 45 [1.1] |
| Pigment | | | 1 [3.0] | |
| Capsule, Inflammation, Suppurative | | | 1 [2.0] | |
| Papilla, Cyst | | | 1 | |
| Renal Tubule, Cyst | 1 | | | |
| Renal Tubule, Cytoplasmic Alteration | | 28 [1.2] | 40 [1.2] | 48 [2.3] |
| Urinary Bladder | (49) | (48) | (50) | (50) |

*** END OF MALE ***

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Lab: BAT

| B6C3F1/N MICE FEMALE | 0 ppm females | 2500 ppm females | 5000ppm females | 10000ppm females |
|----------------------|---------------|------------------|-----------------|------------------|
|----------------------|---------------|------------------|-----------------|------------------|

Disposition Summary

| | | | | |
|---|-----------|-----------|-----------|-----------|
| Animals Initially In Study | 50 | 50 | 50 | 50 |
| Early Deaths | | | | |
| Moribund Sacrifice | 1 | 2 | 1 | 1 |
| Natural Death | 3 | 5 | 4 | 3 |
| Survivors | | | | |
| Terminal Sacrifice | 46 | 43 | 45 | 46 |
| Animals Examined Microscopically | 50 | 50 | 50 | 50 |

ALIMENTARY SYSTEM

| | | | | |
|--------------------------------------|----------|---------|---------|----------|
| Esophagus | (50) | (50) | (50) | (49) |
| Gallbladder | (48) | (48) | (50) | (50) |
| Intestine Large, Cecum | (50) | (50) | (50) | (50) |
| Intestine Large, Colon | (50) | (50) | (50) | (50) |
| Intestine Large, Rectum | (50) | (50) | (50) | (50) |
| Intestine Small, Duodenum | (50) | (50) | (50) | (50) |
| Intestine Small, Ileum | (50) | (50) | (50) | (50) |
| Intestine Small, Jejunum | (50) | (50) | (50) | (50) |
| Hyperplasia, Lymphocyte | | 1 [4.0] | | |
| Liver | (50) | (50) | (50) | (50) |
| Basophilic Focus | 2 | 1 | 5 | 2 |
| Clear Cell Focus | 3 | 2 | 3 | 8 |
| Eosinophilic Focus | 1 | 7 | 13 | 16 |
| Fatty Change, Focal | 1 [1.0] | | 1 [1.0] | |
| Inflammation, Chronic Active | | 1 [3.0] | | |
| Mixed Cell Focus | | | | 1 |
| Necrosis | 11 [1.2] | 8 [1.3] | 5 [1.4] | 3 [2.0] |
| Pigment | 1 [2.0] | | | |
| Bile Duct, Cyst | | 1 | | 1 |
| Bile Duct, Cyst, Multiple | | | 1 | |
| Hepatocyte, Cytoplasmic Alteration | | | 2 [1.0] | 48 [1.8] |
| Hepatocyte, Vacuolation, Cytoplasmic | 2 [3.5] | | | |
| Stellate Cell, Hyperplasia | | 1 [4.0] | | |
| Mesentery | (4) | (3) | (3) | (0) |

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Lab: BAT

| B6C3F1/N MICE FEMALE | 0 ppm females | 2500 ppm females | 5000ppm females | 10000ppm females |
|-------------------------|---------------|------------------|-----------------|------------------|
| Fat, Necrosis | 4 [2.8] | 3 [3.0] | 3 [3.7] | |
| Pancreas | (50) | (50) | (50) | (50) |
| Cyst | | 2 | 1 | 1 |
| Acinus, Atrophy | 1 [1.0] | | | |
| Acinus, Hyperplasia | | | | 1 [1.0] |
| Salivary Glands | (50) | (49) | (49) | (48) |
| Stomach, Forestomach | (50) | (50) | (50) | (50) |
| Epithelium, Hyperplasia | | | | 1 [1.0] |
| Stomach, Glandular | (50) | (50) | (50) | (50) |
| Epithelium, Hyperplasia | 1 [1.0] | | | |

CARDIOVASCULAR SYSTEM

| | | | | |
|---------------------------------------|---------|---------|---------|---------|
| Blood Vessel | (50) | (50) | (50) | (50) |
| Heart | (50) | (50) | (50) | (50) |
| Fibrosis | | | | 1 [1.0] |
| Infiltration Cellular, Mixed Cell | | | | 1 [1.0] |
| Polyarteritis Nodosa | 2 [1.5] | 1 [1.0] | | |
| Epicardium, Inflammation, Suppurative | | 1 [1.0] | | |
| Myocardium, Mineral | 1 [2.0] | | | |
| Myocardium, Necrosis | | | 1 [1.0] | |
| Valve, Degeneration | | 1 [3.0] | 1 [4.0] | |
| Valve, Mineral | | 1 [3.0] | | |

ENDOCRINE SYSTEM

| | | | | |
|--------------------------|---------|---------|---------|---------|
| Adrenal Cortex | (50) | (50) | (50) | (50) |
| Degeneration, Cystic | 1 [2.0] | | | |
| Hyperplasia, Focal | | 1 [1.0] | 1 [1.0] | |
| Hypertrophy, Focal | 1 [1.0] | | 1 [1.0] | 1 [1.0] |
| Vacuolation, Cytoplasmic | | 1 [1.0] | | |
| Adrenal Medulla | (50) | (50) | (50) | (49) |
| Hyperplasia, Focal | 1 [1.0] | | 1 [2.0] | |
| Islets, Pancreatic | (50) | (50) | (50) | (50) |
| Parathyroid Gland | (43) | (42) | (41) | (43) |

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Lab: BAT

| B6C3F1/N MICE FEMALE | 0 ppm females | 2500 ppm females | 5000ppm females | 10000ppm females |
|------------------------------|---------------|------------------|-----------------|------------------|
| Hyperplasia | | | 1 [1.0] | |
| Pituitary Gland | (50) | (50) | (50) | (49) |
| Pars Distalis, Hyperplasia | 17 [2.0] | 5 [2.0] | 4 [2.0] | 4 [2.8] |
| Pars Intermedia, Hyperplasia | 1 [2.0] | | | |
| Thyroid Gland | (49) | (50) | (50) | (48) |

GENERAL BODY SYSTEM

None

GENITAL SYSTEM

| | | | | |
|-----------------------------------|----------|----------|----------|----------|
| Clitoral Gland | (50) | (50) | (49) | (49) |
| Ovary | (49) | (50) | (50) | (49) |
| Hemorrhage | | | 1 [4.0] | |
| Follicle, Cyst | 8 | 9 | 6 | 11 |
| Rete Ovarii, Cyst | | 2 | | |
| Uterus | (50) | (50) | (50) | (50) |
| Adenomyosis | | | | 1 [1.0] |
| Dilation | 13 [3.3] | 18 [3.2] | 10 [3.4] | 8 [3.1] |
| Inflammation, Suppurative | 3 [2.0] | 1 [2.0] | | 1 [2.0] |
| Cervix, Inflammation, Suppurative | | 1 [2.0] | | 1 [2.0] |
| Cervix, Metaplasia, Mucous Cell | 1 [2.0] | | | |
| Endometrium, Hyperplasia, Cystic | 45 [1.8] | 41 [1.9] | 41 [1.9] | 40 [1.7] |
| Vagina | (0) | (0) | (1) | (0) |

HEMATOPOIETIC SYSTEM

| | | | | |
|-------------------------|---------|---------|------|---------|
| Bone Marrow | (50) | (50) | (50) | (50) |
| Fibrosis | | 2 [1.5] | | 1 [1.0] |
| Lymph Node | (4) | (5) | (5) | (3) |
| Hyperplasia, Lymphocyte | | 1 [3.0] | | |
| Lumbar, Hemorrhage | 1 [3.0] | | | |
| Lymph Node, Mandibular | (50) | (50) | (48) | (48) |

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|---|------------------|------------------|------------------|------------------|
| Hyperplasia, Lymphocyte Lymph Node, Mesenteric | 25 [1.4] (49) | 22 [1.8] (49) | 25 [1.8] (50) | 33 [1.9] (49) |
| Hemorrhage | 1 [4.0] | | | |
| Hyperplasia, Lymphocyte | 35 [2.1] | 26 [2.5] | 27 [2.3] | 35 [2.3] |
| Hyperplasia, Plasma Cell | | | | 1 [3.0] |
| Spleen | (49) | (49) | (50) | (50) |
| Extramedullary Hematopoiesis | 10 [1.7] | 11 [2.2] | 20 [2.1] | 13 [2.0] |
| Hyperplasia, Lymphocyte | 38 [2.8] | 36 [3.2] | 31 [2.8] | 39 [2.9] |
| White Pulp, Atrophy | | | | 1 [4.0] |
| Thymus | (48) | (48) | (47) | (48) |
| Atrophy | 3 [3.3] | 5 [3.0] | 3 [3.7] | |
| Hyperplasia, Lymphocyte | 18 [1.7] | 16 [1.6] | 17 [1.9] | 16 [1.8] |

INTEGUMENTARY SYSTEM

| | | | | |
|---------------|------|------|------|------|
| Mammary Gland | (50) | (49) | (50) | (50) |
| Skin | (50) | (50) | (50) | (50) |

MUSCULOSKELETAL SYSTEM

| | | | | |
|-----------------------------|------|---------|------|---------|
| Bone | (50) | (50) | (50) | (50) |
| Femur, Fibro-Osseous Lesion | | 2 [2.5] | | 1 [2.0] |
| Skeletal Muscle | (0) | (1) | (1) | (0) |

NERVOUS SYSTEM

| | | | | |
|--|---------|---------|---------|---------|
| Brain | (50) | (50) | (50) | (50) |
| Cyst Epithelial Inclusion | | | 1 | |
| Inflammation, Suppurative | | 1 [1.0] | | |
| Meninges, Infiltration Cellular, Mononuclear Cell | 2 [1.5] | | | 1 [2.0] |
| Neuron, Necrosis | 1 [4.0] | 1 [1.0] | 1 [2.0] | |
| Thalamus, Fibrosis | | 1 [2.0] | | |

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|----------------------|---------------|------------------|-----------------|------------------|
|----------------------|---------------|------------------|-----------------|------------------|

RESPIRATORY SYSTEM

| | | | | |
|-----------------------------------|---------|---------|------|---------|
| Lung | (50) | (50) | (50) | (50) |
| Infiltration Cellular, Histiocyte | 1 [3.0] | | | |
| Inflammation, Granulomatous | | 1 [3.0] | | |
| Pigment | | | | 1 [2.0] |
| Alveolus, Epithelium, Hyperplasia | 1 [1.0] | | | |
| Nose | (50) | (50) | (50) | (50) |
| Trachea | (50) | (50) | (50) | (49) |

SPECIAL SENSES SYSTEM

| | | | | |
|---|---------|------|---------|---------|
| Eye | (50) | (50) | (50) | (50) |
| Posterior Chamber, Inflammation, Chronic Active | | | 1 [3.0] | |
| Harderian Gland | (50) | (50) | (50) | (50) |
| Atrophy | | | 1 [2.0] | |
| Hyperplasia | 2 [2.0] | | 2 [1.5] | 1 [1.0] |

URINARY SYSTEM

| | | | | |
|------------------------------------|----------|----------|----------|----------|
| Kidney | (50) | (50) | (50) | (49) |
| Metaplasia, Osseous | 1 [1.0] | | | |
| Nephropathy, Chronic Progressive | 13 [1.0] | 13 [1.0] | 11 [1.2] | 13 [1.0] |
| Left, Atrophy | | | 1 [2.0] | |
| Papilla, Inflammation, Suppurative | | | 1 [2.0] | |
| Papilla, Necrosis | | | 1 [3.0] | |
| Renal Tubule, Necrosis | | | 1 [2.0] | |
| Urinary Bladder | (49) | (49) | (50) | (49) |
| Degeneration | 1 [4.0] | | | |

*** END OF REPORT ***

a - Number of animals examined microscopically at site and number of animals with lesion

b-Average severity grade(1-minimal;2-mild;3-moderate;4-marked)