

TDMS No. 93020 - 03

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

Date Report Requested: 08/08/2005

Test Type: 90-DAY

DIISOPROPYLCARBODIIMIDE

Time Report Requested: 08:53:32

Route: SKIN APPLICATION

CAS Number: 693-13-0

First Dose M/F: 09/06/94 / 09/07/94

Species/Strain: RATS/F 344

Pathologist: BIERLEY, B. - STEDHAM, M. - SHANAHAN, P.

Lab: MBA

Final #2/Rats

C Number: C93020
Lock Date: 05/16/1995
Cage Range: ALL
Date Range: ALL
Reasons For Removal: ALL
Removal Date Range: ALL
Treatment Groups: Include ALL

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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| FISCHER 344 RATS MALE | 0 MG/KG | 10 MG/KG | 20 MG/KG | 40 MG/KG | 80 MG/KG | 160 MG/KG |
|--|---------|----------|----------|----------|----------|-----------|
| Disposition Summary | | | | | | |
| Animals Initially in Study | 10 | 10 | 10 | 10 | 10 | 10 |
| Early Deaths | | | | | | |
| Moribund Sacrifice | | | | | 10 | 3 |
| Natural Death | | | | | | 7 |
| Survivors | | | | | | |
| Terminal Sacrifice | 10 | 10 | 10 | 10 | | |
| Animals Examined Microscopically | 10 | 10 | 10 | 10 | 10 | 10 |
| ALIMENTARY SYSTEM | | | | | | |
| Liver | (10) | (0) | (0) | (10) | (10) | (10) |
| Centrilobular, Congestion | | | | | 4 [1.5] | 6 [2.3] |
| Centrilobular, Vacuolization Cytoplasmic | | | | | 1 [1.0] | 6 [1.5] |
| Periportal, Vacuolization Cytoplasmic | 1 [1.0] | | | | | |
| Pancreas | (10) | (0) | (0) | (10) | (10) | (9) |
| Acinus, Atrophy, Focal | 1 [1.0] | | | | | |
| Stomach, Glandular | (10) | (0) | (0) | (10) | (10) | (10) |
| Infiltration Cellular, Mixed Cell | | | | 1 [1.0] | | |
| Cyst | | | | 1 [2.0] | | |
| CARDIOVASCULAR SYSTEM | | | | | | |
| Heart | (10) | (10) | (10) | (10) | (10) | (10) |
| Cardiomyopathy | 4 [1.3] | 4 [1.0] | 6 [1.0] | 4 [1.0] | 5 [1.2] | 3 [2.7] |
| Artery, Inflammation, Chronic Active | | | | | 1 [1.0] | |
| ENDOCRINE SYSTEM | | | | | | |
| None | | | | | | |
| GENERAL BODY SYSTEM | | | | | | |
| None | | | | | | |

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| FISCHER 344 RATS MALE | 0 MG/KG | 10 MG/KG | 20 MG/KG | 40 MG/KG | 80 MG/KG | 160 MG/KG |
|--|-----------------|-----------------|-----------------|------------------|------------------|-----------------|
| GENITAL SYSTEM | | | | | | |
| Preputial Gland Inflammation, Chronic Active | (10) 2 [2.0] | (0) | (0) | (10) 2 [2.5] | (10) 1 [2.0] | (10) 4 [1.3] |
| HEMATOPOIETIC SYSTEM | | | | | | |
| Bone Marrow Hemorrhage | (10) | (0) | (0) | (10) | (10) | (10) 2 [2.5] |
| Lymph Node, Mandibular Hyperplasia, Lymphoid | (10) 1 [2.0] | (0) | (0) | (10) | (10) | (10) |
| Hyperplasia, Plasma Cell Hemorrhage | 1 [2.0] | | | 1 [1.0] | | |
| Lymph Node, Mesenteric Necrosis | (10) 1 [1.0] | (0) | (0) | (10) | (10) | (10) |
| Spleen Lymphoid Follicle, Depletion Cellular | (10) | (10) | (10) | (10) | (10) | (10) 1 [2.0] |
| Thymus Necrosis | (10) | (10) | (10) | (10) | (10) 2 [2.0] | (10) |
| Hemorrhage | | | | | 2 [2.5] | |
| Atrophy | | | | | 2 [2.5] | 3 [2.3] |
| INTEGUMENTARY SYSTEM | | | | | | |
| Skin Epidermis, Skin, Site Of Application, Hyperplasia | (10) | (10) 5 [1.0] | (10) 7 [1.0] | (10) 10 [1.6] | (10) 10 [2.0] | (10) 3 [2.3] |
| Epidermis, Skin, Site Of Application, Necrosis, Focal | | | | | | 9 [2.9] |
| Skin, Site Of Application, Inflammation, Chronic Active | | | | 1 [1.0] | 7 [1.7] | 10 [2.8] |
| MUSCULOSKELETAL SYSTEM | | | | | | |
| None | | | | | | |
| NERVOUS SYSTEM | | | | | | |

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| FISCHER 344 RATS MALE | 0 MG/KG | 10 MG/KG | 20 MG/KG | 40 MG/KG | 80 MG/KG | 160 MG/KG |
|---|---------|----------|----------|----------|----------|-----------|
| Brain | (10) | (10) | (10) | (10) | (10) | (10) |
| Edema, Focal | | | | | 5 [1.0] | 1 [1.0] |
| Hemorrhage | | | | | 1 [1.0] | 4 [2.3] |
| Congestion | | | | | | 1 [3.0] |
| Arteriole, Necrosis, Fibrinoid | | | | | | 1 [1.0] |
| Neuron, Necrosis | | | | | 8 [2.6] | |
| RESPIRATORY SYSTEM | | | | | | |
| Lung | (10) | (0) | (0) | (10) | (10) | (10) |
| Congestion | | | | | 5 [2.8] | 8 [2.8] |
| Hemorrhage | | | | | 1 [2.0] | |
| Alveolus, Infiltration Cellular, Histiocyte | 1 [1.0] | | | | 1 [1.0] | |
| Interstitial, Inflammation, Chronic Active | 1 [1.0] | | | | 1 [2.0] | |
| Perivascular, Infiltration Cellular, | 3 [1.7] | | | 6 [1.0] | 1 [2.0] | |
| Lymphocyte | | | | | | |
| Nose | (10) | (0) | (0) | (10) | (10) | (10) |
| Respiratory Epithelium, Inflammation, | 3 [1.3] | | | 1 [1.0] | 1 [2.0] | 5 [1.4] |
| Chronic Active | | | | | | |
| Respiratory Epithelium, Metaplasia, | | | | | | 3 [1.7] |
| Squamous | | | | | | |
| Squamous Epithelium, Inflammation, | | | | 1 [1.0] | | |
| Chronic | | | | | | |
| Squamous Epithelium, Inflammation, | | | | | | 1 [2.0] |
| Chronic Active | | | | | | |
| SPECIAL SENSES SYSTEM | | | | | | |
| Eye | (0) | (0) | (0) | (0) | (1) | (0) |
| Cornea, Bilateral, Degeneration, Focal | | | | | 1 [2.0] | |
| URINARY SYSTEM | | | | | | |
| Kidney | (10) | (0) | (0) | (10) | (10) | (10) |
| Nephropathy | 5 [1.0] | | | 2 [1.0] | | |

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Lab: MBA

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*** END OF MALE ***

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| Disposition Summary | | | | | | |
| Animals Initially in Study | 10 | 10 | 10 | 10 | 10 | 10 |
| Early Deaths | | | | | | |
| Moribund Sacrifice | | | | | 10 | 10 |
| Survivors | | | | | | |
| Accidently Killed | 1 | | | | | |
| Terminal Sacrifice | 9 | 10 | 10 | 10 | | |
| Animals Examined Microscopically | 10 | 10 | 10 | 10 | 10 | 10 |
| ALIMENTARY SYSTEM | | | | | | |
| Liver | (10) | (0) | (0) | (10) | (10) | (10) |
| Congestion | | | | | 1 [1.0] | |
| Centrilobular, Congestion | | | | | | 2 [1.0] |
| Mesentery | (0) | (0) | (0) | (1) | (0) | (0) |
| Fat, Necrosis | | | | 1 [3.0] | | |
| Pancreas | (10) | (0) | (0) | (10) | (10) | (10) |
| Acinus, Cytoplasmic Alteration, Focal | 1 [2.0] | | | | | |
| CARDIOVASCULAR SYSTEM | | | | | | |
| Heart | (10) | (10) | (10) | (10) | (10) | (10) |
| Cardiomyopathy | | | 1 [1.0] | 2 [1.0] | 2 [1.0] | |
| ENDOCRINE SYSTEM | | | | | | |
| Adrenal Cortex | (10) | (0) | (0) | (10) | (10) | (10) |
| Hypertrophy, Focal | 1 [2.0] | | | | | |
| GENERAL BODY SYSTEM | | | | | | |
| None | | | | | | |
| GENITAL SYSTEM | | | | | | |

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|--|-----------------|-----------------|-----------------|----------------------------|------------------|------------------------------|
| Clitoral Gland Inflammation, Chronic Active | (8) | (0) | (0) | (6) 1 [1.0] | (10) 2 [1.0] | (10) 1 [1.0] |
| Ovary Cyst | (10) | (0) | (0) | (10) 1 [2.0] | (10) | (10) |
| Uterus Hydrometra | (10) | (1) | (5) 5 [3.4] | (10) 4 [2.8] | (10) | (10) 1 [2.0] |
| HEMATOPOIETIC SYSTEM | | | | | | |
| Bone Marrow Hyperplasia, Focal, Lymphoid Hyperplasia, Focal, Histiocytic Hemorrhage Atrophy | (10) | (0) | (0) | (10) 1 [1.0] 2 [1.5] | (10) 1 [1.0] | (10) 2 [2.0] |
| Lymph Node, Mandibular Hemorrhage Hyperplasia, Lymphoid | (10) | (0) | (0) | (9) | (9) 1 [2.0] | (10) 2 [2.0] |
| Lymph Node, Mesenteric Hyperplasia, Histiocytic Hyperplasia, Lymphoid | (10) 1 [2.0] | (0) | (0) | (10) 1 [2.0] 1 [3.0] | (10) | (10) |
| Thymus Atrophy | (10) | (10) | (10) | (10) | (10) 4 [2.3] | (10) 1 [3.0] |
| INTEGUMENTARY SYSTEM | | | | | | |
| Skin Epidermis, Skin, Site Of Application, Hyperplasia Epidermis, Skin, Site Of Application, Necrosis, Focal Skin, Site Of Application, Inflammation, Chronic Active | (10) 1 [1.0] | (10) 2 [1.0] | (10) 3 [1.0] | (10) 5 [1.0] | (10) 10 [1.4] | (10) 10 [1.6] 10 [3.7] |
| MUSCULOSKELETAL SYSTEM | | | | | | |
| None | | | | | | |
| NERVOUS SYSTEM | | | | | | |

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|--|---------|----------|----------|----------|----------|-----------|
| Brain | (10) | (10) | (10) | (10) | (10) | (10) |
| Edema, Focal | | | | | 5 [1.0] | 2 [1.0] |
| Hemorrhage | | | | | 5 [2.2] | 6 [2.0] |
| Arteriole, Necrosis, Fibrinoid | | | | | 1 [1.0] | |
| Neuron, Necrosis | | | | | 8 [2.4] | 2 [4.0] |
| RESPIRATORY SYSTEM | | | | | | |
| Lung | (10) | (0) | (0) | (10) | (10) | (10) |
| Congestion | | | | | 5 [1.8] | 8 [2.5] |
| Hemorrhage | | | | | 1 [2.0] | |
| Interstitial, Inflammation, Chronic Active | 1 [2.0] | | | 3 [1.7] | | |
| Perivascular, Infiltration Cellular, Lymphocyte | 4 [1.3] | | | 5 [2.4] | 4 [2.3] | |
| Nose | (10) | (0) | (0) | (10) | (10) | (10) |
| Olfactory Epithelium, Atrophy | 1 [1.0] | | | | | |
| Olfactory Epithelium, Inflammation, Chronic Active | | | | 1 [1.0] | | |
| Respiratory Epithelium, Atrophy | | | | | 1 [1.0] | |
| Respiratory Epithelium, Inflammation, Chronic Active | 1 [1.0] | | | | | 2 [1.0] |
| Squamous Epithelium, Inflammation, Chronic Active | | | | | 1 [1.0] | |
| SPECIAL SENSES SYSTEM | | | | | | |
| None | | | | | | |
| URINARY SYSTEM | | | | | | |
| Kidney | (10) | (0) | (0) | (10) | (10) | (10) |
| Congestion | | | | | 1 [2.0] | 1 [2.0] |
| Medulla, Mineralization | 5 [1.0] | | | 6 [1.0] | 10 [1.0] | 3 [1.0] |
| Urinary Bladder | (10) | (0) | (0) | (10) | (10) | (10) |
| Serosa, Mineralization | | | | 1 [2.0] | | |

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