



Experimental Pathology Laboratories, Inc.

The Hamner Protocol No.: 10026
EPL Project No.: 304-466
FINAL REPORT – August 30, 2013

EPA FIBER PROJECT: SUBCHRONIC INHALATION EXPOSURE
OF RATS TO AMPHIBOLE ASBESTOS

(18 MONTHS POST-EXPOSURE)

THE HAMNER INSTITUTES FOR
HEALTH SCIENCES PROTOCOL NO. 10026

EPL PROJECT NO.: 304-466
EPL PATHDATA NO. 90188

FINAL PATHOLOGY REPORT

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BACKGROUND

The vermiculite mine near Libby, Montana was the world's leading source of vermiculite for 70 years until its closure in 1990. Vermiculite is used for insulation, as an absorbent material, and as a soil conditioner, and has applications in the construction, agricultural, horticultural and industrial markets. However, the Libby vermiculite ore coexists with a complex array of amphibole mineral types, primarily winchite, richterite, tremolite, and magnesioriebeckite with crystal forms (habits) ranging from asbestosiform to acicular/prismatic.

Occupational exposure to Libby vermiculite has been (and continues to be) associated with significant increases in asbestos, lung cancer, and pleural cancer compared to the rest of the U.S. population. For example, in addition to elevated rates of lung cancer and mesothelioma among Libby residents, 17.8% of 6,668 persons who lived or worked in the Libby area for at least 6 months before 1991 show (upon medical testing) pleural abnormalities (calcifications, thickenings, or plaques).

Furthermore, exposures to individuals outside of Libby have occurred, and are likely continuing; as asbestos-contaminated vermiculite ore from Libby was shipped to hundreds of locations around the nation for processing, and used as attic insulation in millions of homes throughout the United States. The health effects associated with former and current exposures from the asbestos contaminated vermiculite from the Libby mine continues to be a subject of intensive study and public health concern.



OBJECTIVE

The overall goal of this research is to improve the scientific basis for the risk assessment of asbestos-contaminated communities by conducting toxicology studies to help define key determinants of internal dose and provide critical insight on additional key health or pathologic endpoints. These types of toxicology studies can only be done in animals and to date, rodent inhalation studies have not been conducted with the amphibole asbestos that contaminates vermiculite from Libby, Montana (Libby amphibole or LA).

The specific aims of the study are to determine the biological potency of inhaled Libby amphibole (LA) fibers over the near-life span of the rat compared to the potency of inhaled amosite, a known fibrogenic amphibole asbestos fiber, and to develop fiber burden data to use in a dosimetry model of amphibole fiber deposition, clearance, and retention in the respiratory tract (head, trachea, lung lobes), pleura and lymph nodes.

INTRODUCTION

This Hamner Institutes for Health Sciences Study 10026 was conducted in male F344 rats. The in-life portion of this study consists of a 90-day inhalation exposure (65 exposures), 6 hours/day, 5 days per week, followed by necropsy time points at 1-Day, 1-Month, 3-Months and 18-Months after the last exposure day. The 1-Day, 1-Month, and 3-Months timepoints have already been reported (Willson and Wall, 2013).

Male F344 (CDF) rats were procured and housed a minimum of 10 days before commencing nose-only inhalation exposures. At the beginning of the inhalation exposures, the rats were about 10 weeks of age (about 200 grams). The rats were exposed in nose-only exposure tubes, to the LA2007 material, spanning a 10-fold concentration range, at low, medium, and high concentrations (1.0, 3.3, and 10.0 mg/m³), or a UICC amosite sample as a positive control fiber at 3.3 mg/m³, for 6 hr/day, 5 days/week, for 13 consecutive weeks (no exposure weekends). Rats exposed to filtered air served as the negative control.

Rats were necropsied at approximately 18 months after the end of the 13-week exposure (Table 1). Groups 1 – 5 were added by EPL for data collection purposes.



The critical endpoints examined in this study are outlined in Table 2. Rats reserved specifically for fiber burden analysis are shown in rows labeled "Fiber #". At each time point, 3 control rats and 6 amosite- or LA-exposed rats were necropsied and organs removed for the evaluation of clearance/biopersistence (via tissue fiber burden analysis). At the 1-day, 1-month, and 3-month time points, 8 other animals were necropsied, and bronchoalveolar lavage (BAL) fluid and blood were taken, and lungs, parietal pleura, and other organs were taken for histopathology. The BAL and lung histopathology were carried out in the same animal by tying off the left lung lobe and lavaging the right lung lobes, then removing the left lung lobe for histopathology. At the final (~18 month) necropsy, BAL was conducted on 8 of the surviving animals from the scheduled group of 50. The rest of the 50 rats assigned for final necropsy, including those with unscheduled early deaths, were reserved for histopathology only, and also for blood samples at final scheduled necropsy. To facilitate exposure and necropsy scheduling, animals were assigned to necropsy groups (Table 3). Table 4 details the animal numbers assigned to each dose group and exposure level for histopathology/BAL evaluations.

The study design is shown in Table 1 below:

Table 1. Study Design

Group Nos.	Test Material	Concentration (mg/m ³)	18-Months
1	Air Control	0.0	53
2	Amosite	3.3	56
3	LA	1.0	56
4	LA	3.3	56
5	LA	10.0	56

**Table 2. Critical Endpoints**

Group Numbers	Test Material	Exposure Level Conc. (mg/m ³)	End Point	No. of Rats
				18-Month
1	Air Control	0	Fiber #*	3
			BAL**	8
			Histopath	50
2	Amosite	3.3	Fiber #*	6
			BAL**	8
			Histopath	50
3	LA	1.0	Fiber #*	6
			BAL**	8
			Histopath	50
4	LA	3.3	Fiber #*	6
			BAL**	8
			Histopath	50
5	LA	10.0	Fiber #*	6
			BAL**	8
			Histopath	50

*Only fiber analysis is conducted on animals listed in "Fiber #" rows.

**BAL (Bronchoalveolar Lavage) is conducted on same set of animals as histopath; but only on 8 from each group of 50 at the final necropsy.

Table 3. Necropsy Group Assignment

Necropsy Group #	Description	Number of Animals
10	18-Months Post-exposure Period, Histopath /BAL (10 animals per dose group)	50
11	18-Months Post-exposure Period, Histopath /BAL (10 animals per dose group)	50
12	18-Months Post-exposure Period, Histopath /BAL (10 animals per dose group)	50
13	18-Months Post-exposure Period, Histopath /BAL (10 animals per dose group)	50
14	18-Months Post-exposure Period, Histopath /BAL (10 animals per dose group)	50
15	18-Months Post-exposure Period, Fiber Burden - Control, Amosite, and LA Low Dose Animals	15
16	18-Months Post-exposure Period, Fiber Burden - LA Mid and LA High Dose Animals	12

**Table 4. Dose Group, Exposure Level, End Point, Sentinel Animals and Animal Identification Number**

Dose Group	Exposure Level Conc. (mg/m ³)	End Point	Animal ID Numbers*	Sentinel Animals
			18-Month	
Air Control	0	Histopath / BAL	137-186	1-4
Amosite	3.3	Histopath / BAL	249-298	5-8
LA	1.0	Histopath / BAL	307, 319, 321, 349-395**, 397	9-12
LA	3.3	Histopath / BAL	419, 420, 423, 449-494***, 496	13-16
LA	10.0	Histopath / BAL	520, 549-597	17-20

* Per study protocol amendment #5, early mortality or moribundity caused reassignment of animals (within the same exposure group) for the 18-month time point with emphasis placed on maintaining 6 animals per group for fiber burden analysis. Animals 396, 398, 495, 497, 498, and 598 were reassigned to the fiber burden groups due to early deaths/moribundity of animals 319, 321, 419, 420, 423, and 520, who were reassigned to the histopathology/BAL groups. Animal #304 was reassigned to the 1-month necropsy fiber burden group, and animal #307 was reassigned to the 18-month histopathology/BAL group due to early moribundity of animal #364.

**Animal #364 was euthanized moribund during the 13-week inhalation exposure regimen.

***Animal #478 was found dead; too cannibalized and tissues too autolyzed for histopathology.



HISTOPATHOLOGY PROCEDURES

Necropsies were performed at The Hamner Institutes. Animals to be euthanized were deeply anesthetized with sodium pentobarbital (intraperitoneal injection, approximately 50 mg/kg) followed by additional injections, if necessary, but not exceeding the euthanasia dose of 200 mg/kg and exsanguinated by transection of the abdominal aorta.

For those animals reaching terminal sacrifice, 8 animals from each dose group had the right lungs lavaged for BAL cells, the remaining, (up to 42 animals), had left and right lung lobes collected for histopathology. For animals assigned to histopathology and BAL evaluation, the left lung was ligated and the right lung lobes were lavaged with a total of 15 ml phosphate-buffered saline (PBS) five times with approximately 3 ml each time. BAL cells were isolated and the total number of cells enumerated.

Once the right lung lobes were collected the ligature on the left lung lobe was removed. The trachea and left lung lobe were fixed *in situ* with 10% neutral buffered formalin (NBF) at approximately 30 cm of NBF pressure. The tissues of animals that had left and right lung lobes collected were fixed in the same manner. The nasal cavities were flushed with NBF. The head was removed, skinned, trimmed of excess tissue, and stored in NBF for approximately 3 days. The heads were then rinsed in running tap water, decalcified, and re-rinsed in water.

Cross sectional blocks of the nasal cavity were prepared (6 levels) and embedded in paraffin wax. The left lung lobe, trachea, sternum (including pleura), and relevant gross organ/tissue lesions were fixed with NBF for 48 hours, rinsed, and stored in 70% ethanol, embedded in paraffin wax, sectioned (approximately 5 micrometers), deparaffinized, and stained with H&E. An additional set of slides from the lungs, trachea, and sternum were stained with a collagen specific stain (Masson-Goldner's trichrome stain). Testes and epididymis were collected at necropsy, stored in fixative, and one section from the left testis and left epididymis evaluated histologically. A histological section was prepared from all tissues for which there was a gross observation made. Preparation of histological slides and microscopic examinations was performed by Experimental Pathology Laboratories, Inc. (EPL®) Durham, NC.

Hematoxylin and eosin stained microscopic slides of left lung, trachea, sternum, testis and epididymis and any additional gross lesions were evaluated via light



microscopy by an EPL pathologist and selected findings from the left lung are presented in the Results section. Trichrome stained microscopic slides of the left lungs, trachea, and sternum were evaluated by light microscopy by an EPL pathologist and the results incorporated into the histological assessment of these tissues.

During the light microscopic examination histopathologic diagnoses were recorded. Microscopic findings were graded using a subjective grading scale (1=minimal, 2=slight/mild, 3=moderate, 4=moderately severe, 5=severe). Mean severity grades for a finding are calculated by summing the severity grades in a group and dividing by the total number examined in the group. Some findings not suitable for grading were recorded as present (P). Gross findings at necropsy were correlated with histological findings when possible.

All lesions are summarized by treatment group and are presented in the Summary Tables. A tabulation of gross lesions observed at the time of necropsy or during gross trimming, along with corresponding microscopic findings, if any, is provided in the Macro/Micro Correlation Table. Animals with no visible lesions at necropsy are not included in this table. The Table of Individual Microscopic Findings (Animal Organ Finding Table; AOFT) are presented in Appendix B, along with the Summary and Macro/Micro Correlation tables.

RESULTS

Unless otherwise indicated, microscopic findings were diagnosed according to Pathology of the Fischer Rat (Boorman et al., 1990).

Cause of Death

Cause of Death (Appendix A) and Summary of Cause of Death Table 5: there was an increased incidence of mononuclear cell leukemia (MCL) in the LA 10 group ($n=7$; 14%) as compared to the control group ($n=3$; 6%). MCL was the main cause of death in the early death animals in all groups other than the LA 3.3 group. However, the historical control data for the incidence of MCL is 36% for male Fischer rats (NTP, 2011). There were a number of animals with a squamous cell carcinoma of the oral cavity as the primary cause of death in all groups other than the LA 10 group. The incidence of oral squamous cell carcinoma in control animals ($n=2$; 4%) was higher than that found in the National Toxicology Program (NTP) historical controls (0.4%) for male Fischer rats.



The incidences of oral mucosa squamous cell carcinoma in the fiber exposure groups were n=3 (6%), n=4 (8%), n=1 (2%), and n=0 (0%) for the amosite 3.3, LA 1.0, LA 3.3, and LA 10 mg/m³ groups, respectively. The other causes of death were responsible for only one or two animal deaths per exposure group (Table 5), an incidence of ≤ 4%. Except for MCL where the highest incidence in a fiber exposure group was below the historical control data, there did not appear to be a relationship between fiber exposure and cause of death.

**Table 5. Summary of Incidence of Cause of Death of Moribund Sacrifice or Early Death Animals (Group size n=50)**

	Control	Amosite 3.3 mg/m ³	Libby amphibole 1.0 mg/m ³	Libby amphibole 3.3 mg/m ³	Libby amphibole 10.0 mg/m ³
# of Early Deaths	10	10	19	9	12
Mononuclear cell leukemia	3	5	5	1	7
Oral mucosa squamous cell carcinoma	2	3	4	1	0
Pituitary adenoma	2	1	2	1	0
Peritoneal Mesothelioma	0	0	2	1	0
Nephropathy	0	0	1	0	2
Undetermined	0	1	1	1	0
Preputial gland carcinoma	0	0	0	2	0
Osteosarcoma	1	0	1	0	0
Salivary gland carcinoma	1	0	0	0	0
Subcutis fibroma	1	0	0	0	0
Mesentery schwannoma	0	0	1	0	0
Thyroid follicular cell carcinoma	0	0	1	1	0
Lung alveolar bronchiolar carcinoma	0	0	0	0	1
Mammary fibroadenoma	0	0	0	0	1
Meningioma	0	0	0	0	1
Adrenal cortex carcinoma	0	0	0	1	0
Subcutis Sarcoma	0	0	1	0	0



Gross and Microscopic correlations were performed and are detailed in the Correlation of Gross and Microscopic Findings Tables (Appendix B).

A variety of histopathologic changes was observed in both control and treated animals. Other than in the lung for the most part these changes were typical of the spontaneous type of microscopic pathology that may be observed for this age and strain of rat.

All eight animals diagnosed with mesothelioma had tumors in the epididymides and testes. In five animals it had spread to other organs in the abdomen [e.g. mesentery, body cavity, adipose tissue, spleen, lymph node and liver] (Table 6). Mesothelioma was not detected in the thoracic cavity of any rats. The distribution and morphology of mesotheliomas were characteristic of spontaneous neoplasms at these sites in the Fischer rat. These tumors are thought to originate in the tunica vaginalis.

Table 6. Sites of Mesothelioma Occurrence in Animals Diagnosed with Peritoneal Mesotheliomas

Animal number	Testes	Epididymides	Mesentery	Spleen	Lymph node	Body cavity	Adipose tissue	Liver
138	P	P	-	-	-	-	-	-
186	P	P	-	-	-	-	-	-
357	P	P	-	-	-	-	-	-
363	P	P	P	-	P	P	-	-
383	P	P	-	P	-	P	-	-
451	P	P	P	-	-	P	P	-
471	P	P	-	--	-	P	-	P
554	P	P	P	-	-	P	-	-

P = Mesothelioma diagnosed in this tissue/site

A summary of selected histological findings in the left lungs at 18-Months post-exposure is given in Table 7. Tables 8, 9 and 10 illustrate the histological findings seen in the trachea, left epididymis and left testes. Other histological findings are in the Summary Tables, Tables of Individual Microscopic Findings, and Macro/Micro Correlation Tables in Appendix B.

**Table 7. 18 month Post-exposure Summary of Incidence and Severity [] of Selected Findings in the Left Lung**

	Control	Amosite 3.3 mg/m ³	Libby amphibole 1.0 mg/m ³	Libby amphibole 3.3 mg/m ³	Libby amphibole 10.0 mg/m ³
Number of animals per group	50	50	50	49	50
Alveolus Inflammation	0	45 [1.0]	40 [0.9]	47 [1.1]	48 [1.1]
Interstitial Fibrosis	0	42 [0.8]	29 [0.6]	43 [0.9]	42 [0.9]
Pleural fibrosis	0	25 [0.5]	16 [0.3]	19[0.4]	22 [0.4]
Foreign Body*	0	47	40	48	49
Bronchiolization	0	5 [0.1]	5 [0.1]	0	0
Alveolar epithelium hyperplasia	1 [0.04]	9 [0.3]	6 [0.2]	6 [0.2]	6 [0.3]
Alveolar bronchiolar adenoma*	0	2	0	2	2
Alveolar bronchiolar carcinoma*	0	0	0	0	1
Mononuclear cell leukemia*	6	8	8	5	8

[] = Mean Severity Grade for the group is calculated by dividing the total of severity grades for a finding/lesion in a group by the number of that tissue examined in that group.

*Not graded

Lung

Air Control

The histopathological findings in the left lung of control rats were typical of the spontaneous type of microscopic pathology that may be observed for this age and strain of rat. Mononuclear cell leukemia (MCL) was evident in six animals. This was characterized by an infiltration of neoplastic mononuclear cells in the interstitium of the lung. One animal had mild alveolar epithelial hyperplasia. This finding is characterized by a proliferation of cuboidal type II alveolar cells without evidence of significant inflammation or macrophage infiltration as seen in more common spontaneous subpleural lesions.

Amosite 3.3 mg/m³

Minimal to mild alveolus inflammation was evident in the left lung in most animals from this exposure group. Alveolus inflammation was characterized by an infiltration of macrophages, lymphocytes and occasional neutrophils in alveoli and alveolar ducts resulting in a concomitant increase in the thickness of the alveolar and alveolar duct walls. A constituent of this process was occasional granulomas which are aggregates of macrophages with rare giant cells. This inflammation was associated with the presence of amosite fibers in alveolar macrophages. Minimal interstitial fibrosis was evident in most animals from this exposure group. Interstitial fibrosis was characterized by fibroblasts and collagen deposition in alveolar walls and within granulomas. The diagnosis of fibrosis was reinforced by positive staining with Masson's trichrome. Fibrosis was evident along alveolar walls frequently increasingly evident at branchpoints into alveolar sacs. Foreign body was diagnosed when amosite fibers were evident in sections. Amosite fibers were evident in most animals in the left lung from this exposure group. These were mostly contained in macrophages, some of which were completely engulfed, while others were partially engulfed; syncytial giant cells were occasionally evident. Amosite fibers were seen within alveolar macrophages and also within cells in the interstitium. Fibers were frequently seen in a subpleural location. Twenty-five animals (50%) had focal pleural fibrosis which in all but one animal was minimal. Pleural fibrosis was characterized by fibroblasts and collagen deposition in the pleura resulting in a focal area of thickening. The diagnosis of fibrosis was reinforced by positive staining with Masson's trichrome. In a few animals (five) minimal bronchiolization was diagnosed. Bronchiolization was characterized by the appearance of cells that resemble bronchiolar epithelial cells at locations distal to the terminal bronchioles and thus occurring in alveolar ducts and adjacent alveoli. The presence of cilia was a helpful diagnostic criterion.

Nine animals had minimal to moderate alveolar epithelium hyperplasia (see Air Control for lesion descriptions) and there were two animals with a finding of alveolar bronchiolar adenoma. Alveolar bronchiolar adenoma was characterized by a distinct mass that may cause compression of adjacent lung parenchyma. There is a proliferation of simple cuboidal or columnar cells distorting the alveolar architecture. The epithelium of this benign neoplasm is uniform with little pleomorphism or atypia. Eight

animals had MCL characterized by an infiltration of neoplastic mononuclear cells in the interstitium of the lung.

Libby amphibole 1.0 mg/m³

Minimal to mild alveolus inflammation, minimal interstitial fibrosis, and foreign body (Libby amphibole fiber) were evident in the left lung of most animals from this treatment group. In a few animals (five) minimal bronchiolization was diagnosed.

Six animals had alveolar epithelium hyperplasia (five minimal and one moderate), sixteen animals had minimal pleural fibrosis and there were no pulmonary neoplasms diagnosed. Eight animals had MCL. See Air Control and Amosite 3.3 mg/m³ for lesion descriptions.

Libby amphibole 3.3 mg/m³

Minimal to mild alveolus inflammation, minimal to mild interstitial fibrosis, and foreign body (Libby amphibole fiber) were evident in the left lung of most animals from this exposure group. Six animals had alveolar epithelium hyperplasia (three minimal, one mild, one moderate, and one moderately severe), nineteen animals had minimal pleural fibrosis and there were two alveolar bronchiolar adenomas present. Five animals had MCL. See Air Control and Amosite 3.3 mg/m³ for lesion descriptions.

Libby amphibole 10.0 mg/m³

Minimal to mild alveolus inflammation, minimal to mild interstitial fibrosis, and foreign body (Libby amphibole fiber) were evident in the left lung of most animals from this treatment group. Six animals had alveolar epithelium hyperplasia (one minimal, one mild, three moderate, and one moderately severe), twenty two animals had minimal pleural fibrosis and there were two animals with a finding of alveolar bronchiolar adenoma and one animal with an alveolar bronchiolar carcinoma. Alveolar bronchiolar carcinoma frequently present as masses which are often not well circumscribed and the normal alveolar architecture is obliterated. The epithelium may be multilayered or growing in solid clusters. The cells frequently exhibit anaplasia. Eight animals had MCL.

See Air Control and Amosite 3.3 mg/m³ for other lesion descriptions. The histological findings in the lung are slightly more severe in this exposure group compared to the amosite and the other LA exposure groups.

Table 8. 18 month Post Exposure Summary of Incidence and Severity of Selected Findings in the Trachea

	Control	Amosite 3.3 mg/m ³	Libby amphibole 1.0 mg/m ³	Libby amphibole 3.3 mg/m ³	Libby amphibole 10.0 mg/m ³
No. Examined	50	50	49	49	50
NAD	50	49	45	47	41
Squamous Metaplasia	-	1	-	-	-
Grade 1	-	1	-	-	-
Chronic Inflammation	-	-	4	2	9
Grade 1	-	-	3	1	9
Grade 2	-	-	1	1	-

NAD = Nothing abnormal discovered

Trachea

There were a few observations of minimal to mild chronic inflammation in each of the groups exposed to LA which indicates an exposure related effect. This was characterized by an infiltration of mononuclear cells in the submucosa of the trachea. One amosite animal had a focus of squamous metaplasia of the tracheal epithelium which was characterized by replacement of the respiratory epithelium by squamous epithelial cells.

Sternum

There were only two observations made for sternum which were blood vessel arteritis in one animal (LA 10.0) and MCL in one animal (Amosite 3.3). These lesions were considered unrelated to exposure and were characterized by an aggregation of perivascular inflammatory cells.

Left Epididymis and Left Testis

The findings in the left epididymis (Table 8) and left testis (Table 9) were typical for this strain of rat. In general, the incidence and severity of these findings in the fiber-exposed groups was similar to the air control.



In the epididymis, a few mesotheliomas were evident on the serosa. Hypospermia was routinely diagnosed which is characterized by a reduction in sperm in the epididymal tubules.

In the testis, interstitial cell adenomas were commonly diagnosed. A few animals had interstitial cell hyperplasia. Atrophy of the seminiferous tubules and edema were also regularly diagnosed. A few mesotheliomas were evident on the serosal surface of the testis.

Table 9. 18 month Post Exposure Summary of Incidence and Severity of Selected Findings in the Left Epididymis

	Control	Amosite 3.3 mg/m ³	Libby amphibole 1.0 mg/m ³	Libby amphibole 3.3 mg/m ³	Libby amphibole 10.0 mg/m ³
No. Examined	49	50	50	49	50
NAD	18	14	21	22	24
Mesothelioma	2	-	2	2	1
Hypospermia	28	36	27	25	26
Grade 1	1	-	-	-	-
Grade 2	3	-	6	4	-
Grade 3	8	14	10	11	8
Grade 4	15	22	11	10	18
Grade 5	1	-	-	-	-
Sperm Granuloma	1	-	-	-	-
Grade 3	1	-	-	-	-
Blood Vessels Arteritis	-	-	1	-	-
Grade 2	-	-	1	-	-

NAD = Nothing abnormal discovered

**Table 10. 18 month Post Exposure Summary of Incidence and Severity of Selected Findings in the Left Testis**

	Control	Amosite 3.3 mg/m ³	Libby amphibole 1.0 mg/m ³	Libby amphibole 3.3 mg/m ³	Libby amphibole 10.0 mg/m ³
No. Examined	49	50	49	49	50
NAD	2	1	3	1	1
Interstitial Cell Adenoma	31	38	34	33	37
Mesothelioma	2	-	3	2	-
Tubular Atrophy	25	16	15	12	14
Grade 1	10	5	2	7	10
Grade 2	9	4	11	4	2
Grade 3	2	2	1	-	-
Grade 4	3	5	1	1	1
Grade 5	1	-	-	-	1
Edema	13	6	4	8	8
Grade 1	2	4	1	4	5
Grade 2	10	2	2	4	3
Grade 3	1	-	1	-	-
Interstitial Cell Hyperplasia	-	5	1	1	1
Grade 1	-	3	-	1	-
Grade 2	-	2	-	-	-
Grade 3	-	-	1	-	1

NAD = Nothing abnormal discovered

CONCLUSIONS

Exposure of rats to amosite and various levels of LA for three months with an 18 month recovery period did not have an effect on survival of animals. After 18 months recovery period, post exposure histopathological changes persisted in the lungs of rats exposed to various exposure levels of LA and amosite. Fibers, though scant, could be identified in the majority of the animals from each exposure group. The foreign materials persisted and continued to be sequestered in macrophages and sometimes syncytial giant cells were evident. Pulmonary neoplasms, although only evident in treated groups, are within the current historical control incidence for male Fischer 344 rats which is 2.48% (range 0-8%) for alveolar bronchiolar adenomas or 1.2% (range 0-6%) for alveolar bronchiolar carcinomas (NTP, 2011). A few animals in each of the LA groups had minimal



to mild chronic inflammation in the trachea which may indicate a treatment associated effect.

As in the earlier time points previously reported, the most intense response was seen in the lungs of rats exposed to LA 10.0 mg/m³. In this group, there was a marginally increased incidence and severity of alveolus inflammation and the presence of foreign body in comparison to the other exposure groups. The incidence and severity of alveolus inflammation and interstitial fibrosis, and foreign body presence was similar in the amosite 3.3 mg/m³ group and the LA 3.3 mg/m³ group; the LA 1.0 mg/m³ group had a reduced incidence and severity of these histological findings.

The incidences of MCL were greater in the amosite, LA 1.0 and LA 10.0 groups than in the control group for animals in the moribund/early death category. The observed incidences of MCL were well within the historical control data range for this strain of rat. However, the incidences of squamous cell carcinomas in the air-control, amosite 3.3, and LA 1.0 mg/m³ groups were increased compared to historical control incidence. The significance of this finding is unclear. No pleural mesotheliomas were observed in any treatment groups, including the positive control group (Amosite 3.3 mg/m³). In animals dying before the end of the study, squamous cell carcinomas of the oral mucosa occurred at a higher incidence rate in the amosite and LA 1.0 groups than in the control group.

Post-Exposure Temporal Observations

Comparison of left lung histology to earlier time points in all groups:

The histological findings after 1 day and 3 months post-exposure were alveolus inflammation, interstitial fibrosis, the presence of foreign body, and bronchiolization and in the LA 10 mg/m³ group only, bronchiole epithelial hyperplasia. After 1 month post-exposure, the same findings were evident in all exposure groups as after 1 day post-exposure. After 3 months post-exposure, the same findings were evident as at the earlier sacrifices, other than the absence of bronchiole epithelial hyperplasia. Additionally, two rats from the LA 10 mg/m³ group had alveolar epithelial hyperplasia: one after one day post-exposure and one after three months post-exposure.



Experimental Pathology Laboratories, Inc.

The Hamner Protocol No.: 10026
EPL Project No.: 304-466
FINAL REPORT – August 30, 2013

The histological findings in the moribund sacrifice and terminal sacrifice animals can be summarized as follows: Alveolus inflammation, interstitial fibrosis, foreign body, and pleural fibrosis were features of lungs from all exposure groups. Bronchiolization was only present in a few rats. Alveolar epithelial hyperplasia was diagnosed in all exposure groups (also in one control). There were also alveolar bronchiolar adenomas diagnosed in Amosite 3.3 mg/m³, LA 3.3 mg/m³ and LA 10.0 mg/m³ groups and a single alveolar bronchiolar carcinoma diagnosed in the LA 10.0 mg/m³ exposure group.

A handwritten signature in black ink, appearing to read "GABRIELLE A. WILLSON".

GABRIELLE A. WILLSON, BVMS, Dip RC Path,
FRC Path, MRCVS
Pathologist

30 August 2013
Date

GAW/dc



Experimental Pathology Laboratories, Inc.

The Hamner Protocol No.: 10026
EPL Project No.: 304-466
FINAL REPORT – August 30, 2013

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<http://ntp.niehs.nih.gov/?objectid=92E705C7-F1F6-975E-72D23026B1645EB9>

Willson G, Wall H. March 6, 2013. Amended Final Report: EPA Fiber Project: Subchronic Inhalation Exposure of Rats to Amphibole Asbestos (1-Day Post-Exposure, 1-Month Post-Exposure, and 3-Months Post-Exposure).

QUALITY ASSURANCE FINAL CERTIFICATION

**QUALITY ASSURANCE FINAL CERTIFICATION**

Study Title: EPA Fiber Project: Subchronic Inhalation Exposure of Rats to Amphibole Asbestos - 18 Month timepoint

Client Study: Protocol No. 10026

EPL Principal Investigator: Dr. Gabrielle Willson

EPL Project Number: 304-466

EPL Pathologist: Dr. Gabrielle Willson

The following aspects of this study were inspected by the Quality Assurance Unit of Experimental Pathology Laboratories, Inc. Dates inspections were performed and findings reported to the EPL Principal Investigator and Management are indicated below.

Dates		
Area Inspected	Inspection	Reporting
EPL Project Sheets	Jan. 19, 2011; Feb. 3, 2011; May 17, 2011; Aug. 5, 2011; Aug. 16, 2011; May 29, 2012; Oct. 11, 2012; Oct. 24, 2012; Nov. 2, 2012; Dec. 12, 2012; Jan. 30, 2013	Jan. 19, 2011; Feb. 3, 2011; May 17, 2011; Aug. 5, 2011; Aug. 16, 2011; May 29, 2012; Oct. 11, 2012; Oct. 24, 2012; Nov. 2, 2012; Dec. 12, 2012; Jan. 30, 2013
Necropsy Records Review	Oct. 10, 2012; Nov. 6-7, 2012; Nov. 8, 2012; Nov. 12, 2012	Oct. 10, 2012; Nov. 7, 2012; Nov. 8, 2012; Nov. 12, 2012
Project Setup	May 26, 2011; Aug. 7, 2012; Aug. 20, 2011; Oct. 18, 2011; Oct. 22, 2012; Nov. 19, 2012; Nov. 28, 2012; Dec. 14, 2012; Jan. 16, 2013	May 26, 2011; Aug. 7, 2012; Aug. 20, 2011; Oct. 18, 2011; Oct. 22, 2012; Nov. 19, 2012; Nov. 28, 2012; Dec. 14, 2012; Jan. 16, 2013
Data Review	Oct. 12, 2012; Nov. 2, 2012; Nov. 19, 2012; Dec. 11, 2012; Jan. 11 & 14, 2013; Jan. 21-23, 2013; Feb. 1, 2013; Feb. 6, 2013	Oct. 12, 2012; Nov. 2, 2012; Nov. 19, 2012; Dec. 11, 2012; Jan. 14, 2013; Jan. 21-23, 2013; Feb. 1, 2013; Feb. 6, 2013
Draft Pathology Report	July 2, 3, 5, 8 & 9, 2013; July 17, 2013; July 29 &30, 2013; Aug. 16. 2013	July 9, 2013; July 17, 2013; July 30, 2013; Aug. 16. 2013
Final Pathology Report	August 30, 2013	August 30, 2013
Date of last quarterly facility inspection:	July 2013	

Jane T. Hollingsworth
EPL Quality Assurance Unit

Jane T. Hollingsworth, AB, RQAP-GLP

August 30, 2013
Date

APPENDIX A – CAUSE OF DEATH/MORIBUND SACRIFICE OR FOUND DEAD

Animal Number	Cause of Death/Moribund Sacrifice or Found Dead
Group 1 (Air Control)	
145	Mononuclear Cell Leukemia
152	Oral Mucosa Squamous Cell Carcinoma
154	Pituitary Gland, Pars Distalis Adenoma
157	Pituitary Gland, Pars Distalis Adenoma
169	Salivary Gland Carcinoma
170	Subcutis Fibroma
176	Mononuclear Cell Leukemia
177	Mononuclear Cell Leukemia
183	Oral Mucosa Squamous Cell Carcinoma
184	Spine Osteosarcoma
Group 2 (Amosite 3.3 mg/m³)	
252	Pituitary Gland, Pars Distalis Adenoma
256	Oral Mucosa Squamous Cell Carcinoma
265	Oral Mucosa Squamous Cell Carcinoma
271	Mononuclear Cell Leukemia
272	Mononuclear Cell Leukemia
273	Undetermined
288	Oral Mucosa Squamous Cell Carcinoma
292	Mononuclear Cell Leukemia
295	Mononuclear Cell Leukemia
297	Mononuclear Cell Leukemia
Group 3 (LA 1.0 mg/m³)	
319	Mesentery Schwannoma
321	Pituitary Gland, Pars Distalis Adenoma
349	Thyroid, Follicular Cell Carcinoma
354	Undetermined
355	Oral Mucosa Squamous Cell Carcinoma
357	Mesothelioma
359	Oral Mucosa Squamous Cell Carcinoma
367	Mononuclear Cell Leukemia
371	Mononuclear Cell Leukemia

Animal Number	Cause of Death/Moribund Sacrifice or Found Dead
Group 3 (LA 1.0 mg/m³) cont'd.	
373	Oral Mucosa Squamous Cell Carcinoma
374	Mononuclear Cell Leukemia
379	Oral Mucosa Squamous Cell Carcinoma
382	Subcutis Sarcoma
383	Mesothelioma
386	Mononuclear Cell Leukemia
387	Pituitary Gland, Pars Distalis Adenoma
390	Osteosarcoma Eye Socket
391	Nephropathy
397	Mononuclear Cell Leukemia
Group 4 (LA 3.3 mg/m³)	
419	Preputial Gland Carcinoma
420	Preputial Gland Carcinoma
451	Mononuclear Cell Leukemia
462	Pituitary Gland, Pars Distalis Adenoma
469	Adrenal Cortex Carcinoma
470	Thyroid Follicular Cell Carcinoma
471	Mesothelioma
473	Oral Mucosa Squamous Cell Carcinoma
478	Undetermined due to cannibalism and severe tissue autolysis
Group 5 (LA 10.0 mg/m³)	
520	Mononuclear Cell Leukemia
549	Alveolar Bronchiolar Carcinoma
552	Mononuclear Cell Leukemia
563	Mononuclear Cell Leukemia
567	Meningioma
568	Mononuclear Cell Leukemia
569	Mammary Fibroadenoma
574	Mononuclear Cell Leukemia
578	Mononuclear Cell Leukemia
582	Mononuclear Cell Leukemia
590	Nephropathy
595	Nephropathy

APPENDIX B – 18-MONTHS POST-EXPOSURE

SUMMARY TABLE

PATHOLOGY REPORT
SUMMARY TABLES

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

NUMBER OF ANIMALS WITH MICROSCOPIC FINDINGS BY ORGAN/GROUP/SEX Necropsy Status: TERMINAL SACRIFICE GROUP (K0), Incl. Deaths					
Sex	Males				
Dose Group No. Animals per Dose Group	1 50	2 50	3 50	4 49	5 50
LUNG, LEFT	No.Examined NAD	50 23	50 -	49 1	49 -
- Chronic Inflammation	Grade 1 Grade 2	13 12 1	- - -	- - -	3 3 -
- Mononuclear Cell Leukemia		6	8	8	5
- Alveolar Epithelium Hyperplasia	Grade 1 Grade 2 Grade 3 Grade 4	1 - 1 - - 1	9 6 2 1 - -	6 5 - 1 - -	6 3 1 1 3 1
- Intravascular Neutrophils	Grade 2 Grade 3 Grade 4	2 1 1 -	- - - -	- - - -	1 - - 1
- Infiltration Mixed Cell	Grade 1 Grade 2	2 1 1	- - -	1 - 1	- - -
- Pheochromocytoma		1	-	-	-
- Foreign Body		-	47	40	48
- Alveolus Inflammation	Grade 1 Grade 2	- - -	45 42 3	40 37 3	47 39 8
- Interstitial Fibrosis	Grade 1 Grade 2	- - -	42 42 -	29 29 -	43 41 2
- Bronchiolization	Grade 1	- -	5 5	5 5	- -
- Alveolar Bronchiolar Hyperplasia	Grade 2	- -	- -	- -	1 1
- Alveolar Bronchiolar Adenoma		-	2	-	2
- Congestion	Grade 2 Grade 3	2 1 1	1 - 1	- - -	1 - 1
- Osteosarcoma		1	-	1	-
- Infiltrate Mixed Cell Focal	Grade 1	-	-	-	1
- Pigmented Histiocytes	Grade 1	-	-	-	1
- Alveolar Bronchiolar Carcinoma		-	-	-	1

NAD = Nothing abnormal discovered

Group 1, AIR CONTROL, males: AMPHIBOLE ASBESTOS (0 mg/mg³)
Group 2, AMOSITE, males: AMPHIBOLE ASBESTOS (3.3 mg/mg³)
Group 3, LA, males: AMPHIBOLE ASBESTOS (1 mg/mg³)
Group 4, LA, males: AMPHIBOLE ASBESTOS (3.3 mg/mg³)
Group 5, LA, males: AMPHIBOLE ASBESTOS (10 mg/mg³)

PATHOLOGY REPORT
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NUMBER OF ANIMALS WITH MICROSCOPIC FINDINGS BY ORGAN/GROUP/SEX Necropsy Status: TERMINAL SACRIFICE GROUP (K0), Incl. Deaths					
Sex	Males				
Dose Group No. Animals per Dose Group	1 50	2 50	3 50	4 49	5 50
LUNG, LEFT	cont.d	50	50	49	50
- Pleura Fibrosis		-	25	16	19
Grade 1		-	24	16	19
Grade 2		-	1	-	-
TRACHEA	No.Examined NAD	50 49	50 45	49 47	50 41
- Squamous Metaplasia	Grade 1	- -	1 1	- -	- -
- Chronic Inflammation	Grade 1 Grade 2	- -	- -	4 3 1	2 1 1
EPIDIDYMIS, LEFT	No.Examined NAD	49 18	50 14	50 21	49 22
- Mesothelioma		2	-	2	2
- Hypospermia	Grade 1 Grade 2 Grade 3 Grade 4 Grade 5	28 1 3 8 15 1	36 - - 14 22 -	27 - 6 10 11 -	25 - 4 11 10 -
- Sperm Granuloma	Grade 3	1 1	- -	- -	- -
- Blood Vessels Arteritis	Grade 2	- -	- -	1 1	- -
TESTIS, LEFT	No.Examined NAD	49 2	50 1	49 3	49 1
- Interstitial Cell Adenoma		31	38	34	33
- Mesothelioma		2	-	3	2
- Tubular Atrophy	Grade 1 Grade 2 Grade 3 Grade 4 Grade 5	25 10 9 2 3 1	16 5 4 2 5 -	15 2 11 1 1 -	12 7 4 - 1 -
- Edema	Grade 1 Grade 2 Grade 3	13 2 10 1	6 4 2 -	4 1 2 1	8 4 4 -
- Interstitial Cell Hyperplasia	Grade 1 Grade 2 Grade 3	- - -	5 3 2 -	1 - -	1 - -

NAD = Nothing abnormal discovered

Group 1, AIR CONTROL, males: AMPHIBOLE ASBESTOS (0 mg/mg3)
Group 2, AMOSITE, males: AMPHIBOLE ASBESTOS (3.3 mg/mg3)
Group 3, LA, males: AMPHIBOLE ASBESTOS (1 mg/mg3)
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Sex	Males				
Dose Group No. Animals per Dose Group	1 50	2 50	3 50	4 49	5 50
STERNUM	No.Examined NAD	50 50	50 49	50 48	50 49
- Blood Vessel Arteritis	Grade 3	-	-	-	1
- Mononuclear Cell Leukemia		-	1	-	-
SKIN/SUBCUTIS	No.Examined	4	3	4	5
- Squamous Cell Carcinoma		2	-	-	1
- Fibroma		2	-	-	-
- Keratoacanthoma		-	1	1	-
- Fat Necrosis		-	1	-	-
- Epidermal Cyst		-	-	2	1
- Fibrosarcoma		-	1	-	-
- Sarcoma		-	-	1	1
- Basal Cell Adenoma		-	-	-	1
- Galactocoele Hemorrhage		-	-	-	1
- Fibroadenoma		-	-	-	1
PREPUTIAL GLANDS	No.Examined	3	2	3	3
- Adenoma		2	1	1	1
- Carcinoma		1	-	2	2
- Inflammation Chronic Active	Grade 2	-	-	-	1
- Cyst		-	-	-	1
- Inflammation Suppurative	Grade 3	-	1	-	-
SYSTEMIC NEOPLASMS	No.Examined	6	9	9	5
- Malignant Lymphoma		6	9	9	5
LIVER	No.Examined	11	19	16	15
- Bile Duct Hyperplasia	Grade 1	4	12	7	10
	Grade 2	-	1	1	-
	Grade 3	4	10	6	10
- Mixed Cell Focus		-	1	-	9
- Mononuclear Cell Leukemia		5	7	8	6
					7

NAD = Nothing abnormal discovered

Group 1, AIR CONTROL, males: AMPHIBOLE ASBESTOS (0 mg/mg3)
Group 2, AMOSITE, males: AMPHIBOLE ASBESTOS (3.3 mg/mg3)
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NUMBER OF ANIMALS WITH MICROSCOPIC FINDINGS BY ORGAN/GROUP/SEX Necropsy Status: TERMINAL SACRIFICE GROUP (K0), Incl. Deaths					
Sex	Males				
Dose Group No. Animals per Dose Group	1 50	2 50	3 50	4 49	5 50
LIVER cont.d	11	19	16	15	20
- Hepatocellular Adenoma	1	1	-	-	1
- Focal Necrosis	-	2	-	-	-
Grade 2	-	1	-	-	-
Grade 3	-	1	-	-	-
- Macrovesiculation	-	1	4	2	3
Grade 1	-	1	-	2	-
Grade 2	-	-	4	-	3
- Chronic Inflammation	-	1	-	-	-
Grade 1	-	1	-	-	-
- Basophilic Focus	-	-	-	-	1
- Hepatodiaphragmatic Nodule	-	1	-	1	3
- Eosinophilic Focus	-	2	-	1	1
- Granuloma	-	-	-	-	1
- Mesothelioma	-	-	-	1	-
- Congestion	1	-	-	-	-
Grade 3	1	-	-	-	-
SPLEEN No.Examined	11	22	15	15	22
- Mononuclear Cell Leukemia	6	9	7	6	9
- Pigment	2	-	-	-	-
Grade 2	2	-	-	-	-
- Extramedullary Hematopoesis	3	11	7	8	11
Grade 2	3	5	4	5	9
Grade 3	-	6	2	3	2
Grade 4	-	-	1	-	-
- Lymphoid Depletion	-	1	-	-	1
Grade 2	-	1	-	-	1
- Stromal Hyperplasia	-	-	-	1	-
Grade 3	-	-	-	1	-
- Congestion	-	1	-	-	1
Grade 2	-	-	-	-	1
Grade 3	-	1	-	-	-
- Atrophy	-	-	1	-	-
Grade 3	-	-	1	-	-
- Mesothelioma	-	-	1	-	-
LUNG No.Examined	2	2	4	2	9
- Mononuclear Cell Leukemia	2	2	3	1	4

Group 1, AIR CONTROL, males: AMPHIBOLE ASBESTOS (0 mg/mg3)
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Sex	Males				
Dose Group No. Animals per Dose Group	1 50	2 50	3 50	4 49	5 50
LUNG cont.d	2	2	4	2	9
- Interstitial Fibrosis	-	-	-	-	1
Grade 3	-	-	-	-	1
- Alveolar Epithelium Hyperplasia	-	-	-	-	2
Grade 3	-	-	-	-	2
- Alveolar Bronchiolar Adenoma	-	-	-	-	1
- Congestion	-	-	-	1	-
Grade 2	-	-	-	1	-
- Osteosarcoma	-	-	1	-	-
- Alveolar Bronchiolar Carcinoma	-	-	-	-	1
- Cystic Keratinizing Epithelioma	-	-	-	-	1
TESTES No.Examined	26	25	23	26	24
- Edema	4	1	1	2	-
Grade 2	2	1	1	1	-
Grade 3	2	-	-	1	-
- Tubular Atrophy	6	7	9	7	4
Grade 1	1	-	-	-	-
Grade 2	2	1	2	2	-
Grade 3	-	-	2	1	2
Grade 4	3	6	5	3	1
Grade 5	-	-	-	1	1
- Mesothelioma	1	-	1	-	1
- Interstitial Cell Adenoma	19	18	13	19	20
- Sperm Granuloma	-	-	1	-	-
- Interstitial Cell Hyperplasia	-	-	-	1	1
Grade 2	-	-	-	1	-
Grade 3	-	-	-	-	1
MAMMARY GLAND No.Examined	4	3	3	4	5
- Secretion	4	1	1	2	1
Grade 2	3	-	1	2	1
Grade 3	1	1	-	-	-
- Fibroadenoma	-	1	1	1	1
- Galactocole Hemorrhage	-	-	1	1	2
- Cyst	1	-	-	-	-
- Galactocoele	-	1	-	-	1
- Granuloma	-	-	-	-	1

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NUMBER OF ANIMALS WITH MICROSCOPIC FINDINGS BY ORGAN/GROUP/SEX Necropsy Status: TERMINAL SACRIFICE GROUP (K0), Incl. Deaths					
Sex	Males				
Dose Group No. Animals per Dose Group	1 50	2 50	3 50	4 49	5 50
MAMMARY GLAND cont.d	4	3	3	4	5
- Adenocarcinoma	-	-	1	-	-
KIDNEYS No.Examined	8	10	10	11	17
- Nephropathy	8	10	9	11	14
Grade 1	-	-	-	1	-
Grade 2	2	4	2	3	3
Grade 3	5	5	6	7	6
Grade 4	1	1	1	-	5
- Cyst	-	1	-	-	-
- Mononuclear Cell Leukemia	1	1	1	-	2
- Alveolar Bronchiolar Carcinoma	-	-	-	-	1
- Tubule Pigment	-	-	1	-	-
Grade 3	-	-	1	-	-
ADRENAL GLANDS No.Examined	2	-	-	3	2
- Phaeochromocytoma	1	-	-	-	-
- Cortex Vacuolization	1	-	-	-	1
Grade 2	1	-	-	-	1
- Cortex Focal Hyperplasia	-	-	-	1	1
Grade 2	-	-	-	1	1
- Cortex Adenoma	-	-	-	-	1
- Medulla Phaeochromocytoma	-	-	-	1	-
- Mononuclear Cell Leukemia	-	-	-	1	-
- Cortical Carcinoma	-	-	-	1	-
MESENTERY No.Examined	-	-	2	1	2
- Mesothelioma	-	-	1	1	1
- Blood Vessel Arteritis	-	-	-	-	1
Grade 3	-	-	-	-	1
- Schwannoma	-	-	1	-	-
LYMPH NODES No.Examined	3	4	5	5	8
- Plasma Cell Hyperplasia	2	4	2	5	2
Grade 2	1	3	1	4	1
Grade 3	1	1	1	1	1
- Pigment	-	-	1	-	2
Grade 2	-	-	1	-	1
Grade 3	-	-	-	-	1
- Mesothelioma	-	-	1	-	-

Group 1, AIR CONTROL, males: AMPHIBOLE ASBESTOS (0 mg/mg3)

Group 2, AMOSITE, males: AMPHIBOLE ASBESTOS (3.3 mg/mg3)

Group 3, LA, males: AMPHIBOLE ASBESTOS (1 mg/mg3)

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NUMBER OF ANIMALS WITH MICROSCOPIC FINDINGS BY ORGAN/GROUP/SEX Necropsy Status: TERMINAL SACRIFICE GROUP (K0), Incl. Deaths					
Sex	Males				
Dose Group No. Animals per Dose Group	1 50	2 50	3 50	4 49	5 50
LYMPH NODES cont.d	3	4	5	5	8
- Mononuclear Cell Leukemia	1	-	1	-	3
- Cyst	-	-	-	1	-
- Sinus Dilation Grade 2	-	-	-	-	1
BODY CAVITIES No.Examined	-	-	2	2	2
- Mesothelioma	-	-	2	2	1
- Alveolar Bronchiolar Carcinoma	-	-	-	-	1
EYES No.Examined	3	5	6	2	1
- Retinal Degeneration Grade 3	-	4	1	-	-
Grade 4	-	1	-	-	-
- Lens Cataract Grade 2	-	3	1	-	-
Grade 3	-	2	1	-	-
-	-	1	-	-	-
- Anterior Synechia	-	2	1	-	-
- Cornea Inflammation Grade 2	-	2	1	-	-
Grade 3	-	2	-	-	-
-	-	-	1	-	-
- Phthisis Bulbi	-	-	-	1	-
- Lens Mineralization Grade 3	1	-	-	-	-
Grade 4	1	-	-	-	-
- Panophthalmitis Grade 2	-	1	1	-	1
Grade 4	-	1	-	-	-
-	-	-	1	-	1
- Cornea Ulcer Grade 1	2	1	3	1	1
Grade 3	-	-	1	-	-
Grade 4	-	-	-	-	1
Grade 5	1	1	2	1	-
-	1	-	-	-	-
-	-	-	-	-	-
- Squamous Cell Carcinoma	1	-	2	-	-
- Socket Osteosarcoma	-	-	1	-	-
HARDERIAN GLANDS No.Examined	1	1	4	-	1
- Fibrosis Grade 2	-	1	-	-	-
Grade 4	-	1	-	-	-
- Inflammation Grade 1	-	-	1	-	1
Grade 4	-	-	1	-	-
-	-	-	-	-	1
- Squamous Cell Carcinoma	-	-	2	-	-

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 Group 2, AMOSITE, males: AMPHIBOLE ASBESTOS (3.3 mg/mg3)
 Group 3, LA, males: AMPHIBOLE ASBESTOS (1 mg/mg3)
 Group 4, LA, males: AMPHIBOLE ASBESTOS (3.3 mg/mg3)
 Group 5, LA, males: AMPHIBOLE ASBESTOS (10 mg/mg3)

PATHOLOGY REPORT
SUMMARY TABLES

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

NUMBER OF ANIMALS WITH MICROSCOPIC FINDINGS BY ORGAN/GROUP/SEX Necropsy Status: TERMINAL SACRIFICE GROUP (K0), Incl. Deaths					
Sex	Males				
Dose Group No. Animals per Dose Group	1 50	2 50	3 50	4 49	5 50
HARDERIAN GLANDS cont.d					
- Mononuclear Cell Leukemia	1 -	1 -	4 1	-	1 -
- Pigment	1 1	- -	- -	- -	- -
URINARY BLADDER No.Examined	1	-	-	-	-
- Distention	1 1	- -	- -	- -	- -
BONE No.Examined	1	-	-	-	1
- Osteosarcoma	1	-	-	-	-
- Alveolar Bronchiolar Carcinoma	-	-	-	-	1
THYROID GLAND No.Examined	1	1	1	1	-
- Follicular Cell Adenoma	-	1	-	-	-
- Follicular Cell Carcinoma	-	-	1	1	-
- C Cell Adenoma	1	-	-	-	-
GANGLION, GASS., RIGHT No.Examined	1	1	2	1	1
- Squamous Cell Carcinoma	1	1	2	1	-
- Meningioma Malignant	-	-	-	-	1
PITUITARY GLAND No.Examined	3	3	5	2	5
- Pars Distalis Adenoma	3	3	4	2	3
- Pars Distalis Hyperplasia Focal	-	-	1	-	1
Grade 3	-	-	1	-	1
- Pars Distalis Cyst	-	-	-	-	1
PALATE (ORAL MUCOSA) No.Examined NAD	3 -	3 -	5 1	1 -	1 -
- Squamous Cell Papilloma	1	-	-	-	-
- Squamous Cell Carcinoma	2	3	4	1	1
STOMACH No.Examined	-	-	1	-	-
- Ulceration	-	-	1	-	-
Grade 3	-	-	1	-	-
INTEST-LG, CECUM No.Examined	-	-	1	-	-
- Ulceration	-	-	1	-	-
Grade 4	-	-	1	-	-

NAD = Nothing abnormal discovered

Group 1, AIR CONTROL, males: AMPHIBOLE ASBESTOS (0 mg/mg3)

Group 2, AMOSITE, males: AMPHIBOLE ASBESTOS (3.3 mg/mg3)

Group 3, LA, males: AMPHIBOLE ASBESTOS (1 mg/mg3)

Group 4, LA, males: AMPHIBOLE ASBESTOS (3.3 mg/mg3)

Group 5, LA, males: AMPHIBOLE ASBESTOS (10 mg/mg3)

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SUMMARY TABLES

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

NUMBER OF ANIMALS WITH MICROSCOPIC FINDINGS BY ORGAN/GROUP/SEX Necropsy Status: TERMINAL SACRIFICE GROUP (K0), Incl. Deaths					
Sex	Males				
Dose Group No. Animals per Dose Group	1 50	2 50	3 50	4 49	5 50
INTEST-SM, ILEUM No.Examined	-	-	1	-	-
- Mononuclear Cell Leukemia	-	-	1	-	-
INTEST-LG, COLON No.Examined	-	-	1	-	1
- Adhesion	-	-	1	-	-
- Muscularis Focal Thickening	-	-	-	-	1
BRAIN No.Examined	-	-	1	-	-
- Mononuclear Cell Leukemia	-	-	1	-	-
HEART No.Examined	1	-	-	-	1
- Fibrosis Grade 2	1 1	-	-	-	-
- Alveolar Bronchiolar Carcinoma	-	-	-	-	1
DIAPHRAGM No.Examined	-	-	-	-	1
- Alveolar Bronchiolar Carcinoma	-	-	-	-	1
ADIPOSE TISSUE No.Examined	-	-	-	1	-
- Mesothelioma	-	-	-	1	-
PANCREAS No.Examined	-	-	1	-	-
- Mononuclear Cell Leukemia	-	-	1	-	-
GANGLION, GASS., LEFT No.Examined	-	-	2	-	-
- Squamous Cell Carcinoma	-	-	2	-	-
EPIDIDYMIDES No.Examined	-	-	1	-	1
- Mesothelioma	-	-	1	-	1
SALIVARY GLANDS No.Examined	1	-	-	1	-
- Sublingual Adenocarcinoma	1	-	-	1	-

Group 1, AIR CONTROL, males: AMPHIBOLE ASBESTOS (0 mg/mg3)
 Group 2, AMOSITE, males: AMPHIBOLE ASBESTOS (3.3 mg/mg3)
 Group 3, LA, males: AMPHIBOLE ASBESTOS (1 mg/mg3)
 Group 4, LA, males: AMPHIBOLE ASBESTOS (3.3 mg/mg3)
 Group 5, LA, males: AMPHIBOLE ASBESTOS (10 mg/mg3)

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 1, AIR CONTROL

ANIMAL NUMBER :

	137 MK0	138 MK0	139 MK0	140 MK0	141 MK0	142 MK0	143 MK0	144 MK0	145 MK0+	146 MK0
LUNG, LEFT	:	-	+	+	+	-	+	-	+	+
- Chronic Inflammation	.	.	2.	1.	.	.	1.	.	.	1.
- Mononuclear Cell Leukemia	.	.	.	M45.	M45.	.
- Alveolar Epithelium Hyperplasia	2.	.	.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	+	+	-	-	-	+	+	+	-
- Mesothelioma
- Hypospermia	.	4.	.	.	.	4.	4.	5.	2.	.
TESTIS, LEFT	:	+G	+G	+G	+	+	+G	+G	+	+G
- Interstitial Cell Adenoma	.	P.	P.	P.	.	.	P.	P.	.	.
- Mesothelioma	.	.	M26.
- Tubular Atrophy	.	.	.	1.	1.	2.	.	3.	5.	1.
- Edema	.	.	.	2.	2.	2.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	'	'	'	'	'	'	'	'
PREPUTIAL GLANDS	:	+G	'	'	+	'	'	'	'	'
- Adenoma	.	(P.	.	.	(P.
SYSTEMIC NEOPLASMS	:	'	'	'	+	'	'	'	'	'
- Malignant Lymphoma	N0.	.	.	.	N0.	.
LIVER	:	'	'	'	'	'	'	'	+G	'
- Mononuclear Cell Leukemia	M45.	.
SPLEEN	:	'	'	'	+G	'	'	'	+G	+G
- Mononuclear Cell Leukemia	M45.	.	.	.	M45.	.
- Pigment	2.
TESTES	:	+G	+G	'	'	+G	+G	'	'	'
- Edema	.	3.
- Tubular Atrophy	.	(2.
- Mesothelioma	.	.	M26.
- Interstitial Cell Adenoma	P.	P.	.	.	.
MAMMARY GLAND	:	'	'	'	'	'	'	'	+G	'
- Secretion	2.	.
KIDNEYS	:	'	'	'	'	'	'	'	+G	+G
- Nephropathy	3.	3.
PITUITARY GLAND	:	'	'	'	'	'	'	'	+G	'
- Pars Distalis Adenoma	P.	.

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 1, AIR CONTROL

ANIMAL NUMBER :

	147 MK0	148 MK0	149 MK0	150 MK0	151 MK0	152 MK0+ MK0	153 MK0	154 MK0+ MK0	155 MK0	156 MK0
LUNG, LEFT	:	-	+	-	+	-	+	-	+	+
- Chronic Inflammation	.	.	1.	.	1.	.	1.	.	.	.
- Mononuclear Cell Leukemia	M45.	.
- Intravascular Neutrophils	2.
- Congestion	3.	.	.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	-	-	+	+	+	-	+	-	-
- Hypospermia	.	.	.	4.	3.	4.	.	2.	.	.
TESTIS, LEFT	:	+	+	+G	+G	+G	-	+G	+	+G
- Interstitial Cell Adenoma	.	.	.	P.	P.	P.	.	P.	.	P.
- Tubular Atrophy	.	2.	1.	.	2.	4.	.	2.	1.	2.
- Edema	.	.	.	1.	.	2.	.	2.	2.	.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	'	'	'	'	'	'	'	+G
- Squamous Cell Carcinoma	P.
SYSTEMIC NEOPLASMS	:	'	'	'	'	'	+	'	'	'
- Malignant Lymphoma	NO.	.	NO.	.
LIVER	:	'	'	'	'	'	+G	'	+G	+G
- Bile Duct Hyperplasia	2.
- Mixed Cell Focus	P.
- Mononuclear Cell Leukemia	M45.	.
- Congestion	3.	.	.	.
SPLEEN	:	'	'	'	'	'	+G	'	'	+G
- Mononuclear Cell Leukemia	M45.	.
- Pigment	2.
TESTES	:	'	'	'	+G	+G	+G	'	+G	+G
- Edema	.	.	.	2.	2.	.
- Tubular Atrophy	.	.	.	(4.	.	.	.	1.	2.	.
- Interstitial Cell Adenoma	(P.	P.	P.	.	.	.
MAMMARY GLAND	:	'	'	'	'	'	'	'	'	'
- Secretion	2.	.
- Cyst	P.	.
KIDNEYS	:	'	'	'	'	'	'	'	'	'
- Nephropathy	+G	.
ADEXRAL GLANDS	:	'	'	'	'	'	'	'	'	'
- Cortex Vacuolization	+G	.
EYES	:	'	'	'	'	'	'	'	'	'
- Cornea Ulcer	+G	.	.	.
- Squamous Cell Carcinoma	(5.	.	.	.
HARDERIAN GLANDS	:	'	'	'	'	'	'	'	'	'
- Pigment	(2.	.	.	.
PITUITARY GLAND	:	'	'	'	'	'	'	'	'	'
- Pars Distalis Adenoma	+G	.	.

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
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PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 1, AIR CONTROL

ANIMAL NUMBER :

	147 MK0	148 MK0	149 MK0	150 MK0	151 MK0	152 MK0+	153 MK0	154 MK0+	155 MK0	156 MK0
--	------------	------------	------------	------------	------------	-------------	------------	-------------	------------	------------

PALATE (ORAL MUCOSA)	:	'	'	'	'	'	'	'	'	'
- Squamous Cell Carcinoma	:									M45.

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 1, AIR CONTROL

ANIMAL NUMBER :

	157 MK0+	158 MK0	159 MK0	160 MK0	161 MK0	162 MK0	163 MK0	164 MK0	165 MK0	166 MK0
LUNG, LEFT	:	+	-	+	-	-	-	+	+	+
- Chronic Inflammation	1.	1.	1.
- Infiltration Mixed Cell	.	.	.	1.
- Congestion	.	.	.	2.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	0	+	-	+	+	+	+	+	+
- Hypospermia	4.	3.	4.	4.	3.	4.
- Sperm Granuloma	.	.	.	3.
TESTIS, LEFT	:	0	+G	+	+G	+G	+G	+	+G	+G
- Interstitial Cell Adenoma	.	.	P.	.	P.	P.	P.	.	P.	P.
- Tubular Atrophy	.	.	.	3.	.	.	.	2.	.	.
- Edema	.	.	.	2.	.	.	.	2.	.	.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	'	'	'	'	'	'	'	+G
- Squamous Cell Carcinoma	P.
TESTES	:	'	'	'	+G	+G	+G	+G	'	+G
- Interstitial Cell Adenoma	P.	P.	P.	P.	.	P.
MAMMARY GLAND	:	+G	'	'	'	'	'	'	'	'
- Secretion	.	.	3.
PITUITARY GLAND	:	+G	'	'	'	'	'	'	'	'
- Pars Distalis Adenoma	.	.	P.
PALATE (ORAL MUCOSA)	:	'	'	'	'	'	'	'	'	+G
- Squamous Cell Papilloma	P.

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TEST ITEM	:	AMPHIBOLE ASBESTOS	PATHOL. NO.:	90188 GAW
TEST SYSTEM	:	RAT, , 18 MO, INHALATION	DATE :	29-AUG-13
SPONSOR	:	THE HAMNER	PathData®System V6.2d2	

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 1, AIR CONTROL

ANIMAL NUMBER :

	167 MK0	168 MK0	169 MK0+	170 MK0+	171 MK0	172 MK0	173 MK0	174 MK0	175 MK0	176 MK0+
LUNG, LEFT	:	-	-	-	+	-	-	-	-	+
- Mononuclear Cell Leukemia	M45.
- Intravascular Neutrophils	3.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	-	+	-	+	+	+	+	+	-
- Hypospermia	.	.	3.	.	3.	4.	4.	4.	3.	4.
TESTIS, LEFT	:	+G	+G	+G	+G	+G	+G	+G	+	-
- Interstitial Cell Adenoma	.	.	P.	P.	P.	P.	P.	P.	.	.
- Tubular Atrophy	.	.	2.	1.	4.	.
- Edema	.	.	2.	3.	.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	'	'	+G	'	'	+G	'	'
- Fibroma	P.	.	.	P.	.	.
PREPUTIAL GLANDS	:	'	'	'	'	'	'	'	'	+G
- Carcinoma	P.	.
SYSTEMIC NEOPLASMS	:	'	'	'	'	'	'	'	'	+
- Malignant Lymphoma	N0.	.
LIVER	:	'	'	+G	G	'	'	+G	'	+G
- Bile Duct Hyperplasia	.	.	.	2.
- Mononuclear Cell Leukemia	M45.	.
- Hepatocellular Adenoma	P.	.	.
SPLEEN	:	'	'	'	'	'	'	+G	'	+G
- Mononuclear Cell Leukemia	2.	2.	M45.
- Extramedullary Hematopoiesis	2.	2.	.
LUNG	:	'	'	'	G	'	'	'	'	+G
- Mononuclear Cell Leukemia	M45.
TESTES	:	'	+G	'	+G	+G	+G	+G	'	'
- Tubular Atrophy	(4.
- Interstitial Cell Adenoma	.	.	P.	.	P.	P.	P.	P.	.	.
KIDNEYS	:	'	'	+G	'	'	'	+G	'	+G
- Nephropathy	.	.	.	2.	.	.	.	3.	2.	.
- Mononuclear Cell Leukemia	M45.	.
LYMPH NODES	:	'	'	'	'	'	'	'	'	+G
- Plasma Cell Hyperplasia	(2.	.	.
BODY CAVITIES	:	'	'	'	G	'	'	'	'	'
EYES	:	'	'	'	+G	'	'	'	'	'
- Lens Mineralization	(3.
SALIVARY GLANDS	:	'	'	'	+	'	'	'	'	'
- Sublingual Adenocarcinoma	(P.

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 1, AIR CONTROL

ANIMAL NUMBER :

	177 MK0+	178 MK0	179 MK0	180 MK0	181 MK0	182 MK0	183 MK0+	184 MK0	185 MK0	186 MK0
LUNG, LEFT	:	+	+	-	-	+	-	+	+	-
- Chronic Inflammation	.	.	1.
- Mononuclear Cell Leukemia	.	.	M45.	M26.
- Infiltration Mixed Cell	2.	.	.
- Pheochromocytoma	M44.
- Osteosarcoma	M59.	.	.	.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	+	+	+	-	+	-	-	+	+G
- Mesothelioma	NO.
- Hypospermia	.	4.	4.	2.	.	1.	.	.	3.	.
TESTIS, LEFT	:	+G	+G	+	+	+G	+G	+	+G	+G
- Interstitial Cell Adenoma	.	.	P.	P.	P.	P.	.	P.	P.	P.
- Mesothelioma	M26.
- Tubular Atrophy	.	4.	.	2.	.	1.	2.	1.	.	.
- Edema	.	.	.	2.	.	1.
STERNUM	:	-	-	-	-	-	-	-	-	-
SYSTEMIC NEOPLASMS	:	+	'	'	'	'	'	'	'	'
- Malignant Lymphoma	.	.	N0.
LIVER	:	+G	'	'	'	+G	'	+G	'	+G
- Bile Duct Hyperplasia	2.	2.
- Mononuclear Cell Leukemia	.	.	M45.	M26.
SPLEEN	:	+G	'	'	'	+G	'	'	'	+G
- Mononuclear Cell Leukemia	.	.	M45.	M26.
- Extramedullary Hematopoesis	2.
LUNG	:	+G	'	'	'	'	'	'	'	'
- Mononuclear Cell Leukemia	.	M45.
TESTES	:	'	+G	+G	'	+G	+G	'	'	+G
- Edema	.	.	.	3.
- Tubular Atrophy	.	.	.	4.
- Interstitial Cell Adenoma	.	.	P.	.	P.	P.	.	P.	P.	P.
MAMMARY GLAND	:	'	'	'	'	'	'	'	'	+G
- Secretion	2.
KIDNEYS	:	'	'	'	'	'	+G	'	'	'
- Nephropathy	3.	3.	3.	.	.	.
ARENAL GLANDS	:	'	'	'	'	'	+G	'	'	'
- Phaeochromocytoma	NO.
PARATHYROID GLAND	:	'	'	'	'	'	'	'	'	'
LYMPH NODES	:	+G	'	'	'	'	'	+G	'	'
- Plasma Cell Hyperplasia	3.	.	.
- Mononuclear Cell Leukemia	.	.	M45.
EYES	:	'	'	'	'	'	'	+G	'	'
- Cornea Ulcer	(4.	.	.	.

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SPONSOR	:	THE HAMNER	PathData®System V6.2d2	

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 1, AIR CONTROL

ANIMAL NUMBER :

	177 MK0+	178 MK0	179 MK0	180 MK0	181 MK0	182 MK0	183 MK0+	184 MK0+	185 MK0	186 MK0
URINARY BLADDER	:	:	:	:	:	:	:	+G	:	:
- Distention	3.		
PENIS	:	:	:	:	:	:	:	G	:	:
BONE	:	:	:	:	:	:	:	+G	:	:
- Osteosarcoma	No.		
THYROID GLAND	:	:	:	:	:	:	+			
- C Cell Adenoma	(P.			
GANGLION, GASS., RIGHT	:	:	:	:	:	:		+G		
- Squamous Cell Carcinoma		M10.		
PALATE (ORAL MUCOSA)	:	:	:	:	:	:		+G		
- Squamous Cell Carcinoma		No.		
HEART	:	:	:	:	:	:		+G		
- Fibrosis		2.		

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TEST SYSTEM	:	RAT, , 18 MO, INHALATION	DATE :	29-AUG-13
SPONSOR	:	THE HAMNER	PathData®System V6.2d2	

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 2, AMOSITE

ANIMAL NUMBER :

	249 MK0	250 MK0	251 MK0	252 MK0+	253 MK0	254 MK0	255 MK0	256 MK0+	257 MK0	258 MK0
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Foreign Body	.	P.	P.	P.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	.	1.	1.	1.	1.	1.	1.	1.	1.	1.
- Interstitial Fibrosis	.	1.	1.	1.	.	1.	1.	1.	1.	1.
- Bronchiolization	1.	.	.	1.	.
- Congestion	3.
- Pleura Fibrosis	.	1.	1.	.	1.	.	.	.	1.	1.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	+	+	+	-	+	-	-	+	-
- Hypospermia	.	3.	4.	4.	.	4.	.	.	3.	.
TESTIS, LEFT	:	+G	+G	+G	-	+G	+	+G	+G	+
- Interstitial Cell Adenoma	.	P.	.	P.	.	P.	P.	P.	P.	.
- Tubular Atrophy	.	.	3.	.	.	2.	.	2.	.	2.
- Edema	.	.	1.	.	.	1.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	+G	'	'	'	'	+G	G	'	'
- Keratoacanthoma	.	P.
- Fibrosarcoma	P.	.	.	.
LIVER	:	'	'	'	'	'	+G	+G	+G	'
- Bile Duct Hyperplasia	2.	.	2.	.
- Focal Necrosis	3.	.	.	.
SPLEEN	:	'	+G	'	'	'	+G	+G	'	+G
- Extramedullary Hematopoiesis	.	2.	.	2.	.	3.
- Lymphoid Depletion	2.	.
TESTES	:	'	+G	+G	'	+G	'	+G	'	+G
- Interstitial Cell Adenoma	.	(P.	P.	P.	P.	P.	P.	P.	P.	.
MAMMARY GLAND	:	'	'	'	'	'	'	'	'	'
- Fibroadenoma	P.	.	.
KIDNEYS	:	+G	'	'	'	'	+G	'	+G	'
- Nephropathy	.	2.	3.	.	4.	.
- Cyst	.	P.
LYMPH NODES	:	'	+G	'	'	'	+G	+G	'	'
- Plasma Cell Hyperplasia	.	2.	2.	(2.	.	.
EYES	:	+G	'	'	'	'	+G	'	'	'
- Retinal Degeneration	.	3.
- Lens Cataract	.	(2.
- Panophthalmitis	2.	.	.	.
- Cornea Ulcer	(4.	.	.	.
HARDERIAN GLANDS	:	'	'	'	'	'	'	+G	'	'
- Fibrosis	(2.	.	.
THYROID GLAND	:	'	'	'	'	+G	'	'	'	'
- Follicular Cell Adenoma	(P.
GANGLION,GASS.,RIGHT	:	'	'	'	'	'	'	+G	'	'
- Squamous Cell Carcinoma	P.	.	.

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 2, AMOSITE

ANIMAL NUMBER :

	249 MK0	250 MK0	251 MK0	252 MK0+	253 MK0	254 MK0	255 MK0	256 MK0+	257 MK0	258 MK0
PITUITARY GLAND	:	:	:	+	G	:	:	+	G	:
- Pars Distalis Adenoma	:	:	:		P.				P.	
PALATE (ORAL MUCOSA)	:	:	:	:	:	:	:	+	G	:
- Squamous Cell Carcinoma	:	:	:						NO.	

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TEST ITEM : AMPHIBOLE ASBESTOS	PATHOL. NO.: 90188 GAW
TEST SYSTEM : RAT, , 18 MO, INHALATION	DATE : 29-AUG-13
SPONSOR : THE HAMNER	PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 2, AMOSITE

ANIMAL NUMBER :

	259 MK0	260 MK0	261 MK0	262 MK0	263 MK0	264 MK0	265 MK0+	266 MK0	267 MK0	268 MK0
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Alveolar Epithelium Hyperplasia	.	2.	.	1.	.	1.	1.	.	.	1.
- Foreign Body	.	P.	P.	P.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	.	1.	1.	1.	1.	1.	1.	1.	1.	1.
- Interstitial Fibrosis	.	1.	1.	1.	1.	1.	1.	1.	1.	1.
- Pleura Fibrosis	.	1.	.	1.	1.	.	1.	.	1.	.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	-	+	-	+	-	+	+	+	+
- Hypospermia	.	.	3.	.	4.	.	3.	3.	4.	4.
TESTIS, LEFT	:	+	+G	+	+G	+	+G	+G	+G	+G
- Interstitial Cell Adenoma	.	.	P.	.	P.	P.	P.	P.	P.	P.
- Tubular Atrophy	.	1.	.	1.	.	.	3.	.	.	.
- Edema	.	2.	.	1.
- Interstitial Cell Hyperplasia	.	.	.	1.
STERNUM	:	-	-	-	-	-	-	-	-	-
SYSTEMIC NEOPLASMS	:	+	,	,	,	,	,	,	,	,
- Malignant Lymphoma
LIVER	:	+G	,	,	,	,	,	,	,	,
- Bile Duct Hyperplasia	2.	.	.
- Mononuclear Cell Leukemia	.	.	M45.
SPLEEN	:	+G	,	,	,	,	+G	+G	,	,
- Mononuclear Cell Leukemia	.	.	M45.
- Extramedullary Hematopoiesis	2.	.	.	.
- Congestion	3.	.	.
TESTES	:	,	+G	,	+G	,	+G	,	+G	+G
- Interstitial Cell Adenoma	.	.	P.	.	P.	P.	P.	P.	P.	P.
KIDNEYS	:	,	,	,	,	,	+G	,	,	,
- Nephropathy	3.	.	.	.
EYES	:	,	,	,	,	,	,	,	,	,
- Retinal Degeneration	4.	.
- Lens Cataract	(3.	.
- Anterior Synechia	P.	.
PALATE (ORAL MUCOSA)	:	,	,	,	,	,	,	,	,	,
- Squamous Cell Carcinoma	NO.	.	.	.
INTEST-SM, ILEUM	:	,	,	,	,	,	,	,	G	,
INTEST-SM, JEJUNUM	:	,	,	,	,	,	,	,	G	,
INTEST-SM, DUODENUM	:	,	,	,	,	,	,	,	G	,

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DATE : 29-AUG-13
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 2, AMOSITE

ANIMAL NUMBER :

	269 MK0	270 MK0	271 MK0+	272 MK0+	273 MK0+	274 MK0	275 MK0	276 MK0	277 MK0	278 MK0
LUNG, LEFT	:	+	+	+	+	+G	+	+	+	+
- Mononuclear Cell Leukemia	.	.	M45.	M45.	.	M45.
- Alveolar Epithelium Hyperplasia	3.
- Foreign Body	.	P.	P.	P.	P.	.	P.	P.	P.	P.
- Alveolus Inflammation	.	1.	1.	1.	1.	.	1.	1.	1.	1.
- Interstitial Fibrosis	.	1.	1.	.	.	.	1.	1.	1.	1.
- Bronchiolization	1.
- Alveolar Bronchiolar Adenoma	P.
- Pleura Fibrosis	.	.	.	1.	.	.	2.	1.	1.	1.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	+	+	-	+	+	+	+	+	+
- Hypospermia	.	4.	3.	.	4.	3.	3.	3.	3.	4.
TESTIS, LEFT	:	+G	+G	+	+G	+G	+G	+G	+G	+G
- Interstitial Cell Adenoma	.	P.	P.	.	P.	P.	P.	P.	P.	P.
- Tubular Atrophy	.	.	.	1.	4.	4.	.	4.	.	.
- Interstitial Cell Hyperplasia	.	.	.	1.
STERNUM	:	-	-	+	-	-	-	-	-	-
- Mononuclear Cell Leukemia	.	.	M45.
SYSTEMIC NEOPLASMS	:	'	'	+	+	'	+	'	'	'
- Malignant Lymphoma	.	.	NO.	NO.	NO.
LIVER	:	'	'	'	+G	+G	+G	+G	'	'
- Bile Duct Hyperplasia	3.	.	2.	.	.	.
- Mononuclear Cell Leukemia	.	.	M45.	.	M45.
- Hepatocellular Adenoma	.	.	.	P.
- Focal Necrosis	2.	.	.	.
- Macrovesiculation	1.	.	.	.
- Eosinophilic Focus	P.
SPLEEN	:	'	'	+G	+G	'	+G	+G	'	'
- Mononuclear Cell Leukemia	.	M45.	M45.	M45.	M45.
- Extramedullary Hematopoesis	2.	.	.	.
LUNG	:	'	'	'	+G	'	'	'	'	'
- Mononuclear Cell Leukemia	.	.	M45.
TESTES	:	+G	+G	'	'	'	'	'	+G	+G
- Tubular Atrophy	.	P.	P.	.	.	.	(P.	.	P.	.
- Interstitial Cell Adenoma
MAMMARY GLAND	:	'	'	'	'	'	'	'	+G	'
- Secretion	3.	.
KIDNEYS	:	'	'	'	'	+G	'	'	'	'
- Nephropathy	2.
BODY CAVITIES	:	'	'	'	'	G	'	'	'	'
PITUITARY GLAND	:	'	'	'	'	'	'	'	+G	'
- Pars Distalis Adenoma	P.	.

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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 2, AMOSITE

ANIMAL NUMBER :

	279 MK0	280 MK0	281 MK0	282 MK0	283 MK0	284 MK0	285 MK0	286 MK0	287 MK0	288 MK0+
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Mononuclear Cell Leukemia	M45.	.
- Alveolar Epithelium Hyperplasia	1.
- Foreign Body	.	P.								
- Alveolus Inflammation	.	1.	1.	1.	1.	1.	1.	1.	2.	1.
- Interstitial Fibrosis	.	1.	1.	1.	1.	1.	1.	1.	1.	1.
- Bronchiolization	1.
- Alveolar Bronchiolar Adenoma	.	.	.	P.
- Pleura Fibrosis	.	1.	.	1.	1.	1.	.	.	1.	.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	-	-	-	+	+	+	+	+	-
- Hypospermia	4.	3.	4.	4.	4.	.
TESTIS, LEFT	:	+	+G	+	+G	+G	+G	+G	+G	+
- Interstitial Cell Adenoma	.	.	P.	.	P.	P.	P.	P.	P.	.
- Tubular Atrophy	.	1.	1.
- Edema	.	1.	.	2.
- Interstitial Cell Hyperplasia	.	1.	.	2.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	'	+G	'	'	'	'	'	'
- Fat Necrosis	.	.	.	P.
SYSTEMIC NEOPLASMS	:	'	'	'	'	'	'	'	'	'
- Malignant Lymphoma	N0.	+
LIVER	:	'	'	+G	'	+G	'	+G	+G	+G
- Bile Duct Hyperplasia	.	.	.	1.	2.	.	2.	2.	2.	2.
- Mixed Cell Focus	P.
- Mononuclear Cell Leukemia	M45.	.	.	.
- Chronic Inflammation	.	.	.	1.
SPLEEN	:	'	'	+G	'	+G	'	+G	+G	+G
- Mononuclear Cell Leukemia	3.	3.	2.	.	3.	3.
- Extramedullary Hematopoiesis	.	.	.	3.	3.	2.	.	3.	3.	.
TESTES	:	+G	'	'	+G	+G	'	+G	+G	+G
- Edema	2.
- Tubular Atrophy	.	2.	.	.	(4.	.	.	(4.	.	.
- Interstitial Cell Adenoma	.	.	.	P.	.	.	P.	.	.	.
EYES	:	+G	'	'	+G	'	+G	'	'	'G
- Retinal Degeneration	.	4.	.	4.	.	4.	.	4.	.	.
- Lens Cataract	.	(2.
- Anterior Synechia	P.
- Cornea Inflammation	.	2.	.	2.	.	(2.	.	2.	.	.
PALATE (ORAL MUCOSA)	:	'	'	'	'	'	'	'	'	+
- Squamous Cell Carcinoma	N0.	.

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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 2, AMOSITE

ANIMAL NUMBER :

	289 MK0	290 MK0	291 MK0	292 MK0+	293 MK0	294 MK0	295 MK0+	296 MK0	297 MK0+	298 MK0
LUNG, LEFT	:	+	+	+	+	+	+G	+	+G	+
- Mononuclear Cell Leukemia	.	.	M45.	M45.	.	M45.	.	M45.	.	.
- Alveolar Epithelium Hyperplasia	1.	.	.	2.
- Foreign Body	.	P.	P.	.	P.	P.	.	P.	P.	P.
- Alveolus Inflammation	.	1.	1.	.	1.	1.	2.	.	1.	.
- Interstitial Fibrosis	.	1.	1.	.	1.	1.	.	1.	.	1.
- Bronchiolization	1.
- Pleura Fibrosis	1.	1.	.	1.	.	1.
TRACHEA	:	-	-	-	-	-	+G	-	-	-
- Squamous Metaplasia	1.	.	.	.
EPIDIDYMIS, LEFT	:	+	+	+	+	+	-	+	+	+
- Hypospermia	.	3.	4.	3.	4.	4.	4.	4.	4.	4.
TESTIS, LEFT	:	+G	+G	+	+G	+G	+G	+G	+G	+G
- Interstitial Cell Adenoma	.	P.	P.	.	P.	P.	P.	P.	P.	P.
- Tubular Atrophy	.	.	2.	4.	.	.	.	4.	.	.
- Interstitial Cell Hyperplasia	.	.	2.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	;	;	;	;	;	G	;	;	;
PREPUTIAL GLANDS	:	;	;	;	+G	+G	;	;	;	;
- Adenoma	P.	;	;	;	;	;
- Inflammation Suppurative	3.	;	;	;	;
SYSTEMIC NEOPLASMS	:	;	;	;	+	+	;	+	+	;
- Malignant Lymphoma	N0.	N0.	;	N0.	N0.	;
LIVER	:	;	;	;	+G	+G	+G	+G	;	;
- Bile Duct Hyperplasia	2.	2.	2.	.	;	;
- Mononuclear Cell Leukemia	.	.	M45.	M45.	.	.	M45.	.	;	;
- Hepatodiaphragmatic Nodule	P.	.	;	;
- Eosinophilic Focus	P.	.	.	;	;
SPLEEN	:	;	;	;	+G	+G	+G	+G	;	;
- Mononuclear Cell Leukemia	.	.	M45.	M45.	.	M45.	.	M45.	;	;
- Extramedullary Hematopoesis	3.	.	.	.	;	;
LUNG	:	;	;	;	;	;	+G	;	;	;
- Mononuclear Cell Leukemia	M45.	;	;	;
TESTES	:	;	;	;	+G	+G	+G	;	;	;
- Tubular Atrophy	(4.	(4.	.	;	;
- Interstitial Cell Adenoma	P.	.	.	.	;	;
MAMMARY GLAND	:	;	;	;	;	;	+	;	;	;
- Galactocoele	P.	;	;	;
KIDNEYS	:	;	;	;	+G	+G	+G	+G	;	;
- Nephropathy	2.	3.	3.	2.	;	;
- Mononuclear Cell Leukemia	M45.	.	;	;
LYMPH NODES	:	;	;	;	;	+G	;	;	;	;
- Plasma Cell Hyperplasia	3.	.	.	.	;	;

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PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 3, LA

ANIMAL NUMBER :

	307 MK0	319 MK0+	321 MK0+	349 MK0+	350 MK0	351 MK0	352 MK0	353 MK0	354 MK0+	355 MK0+
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Mononuclear Cell Leukemia	M45.	.	.
- Alveolar Epithelium Hyperplasia	.	.	1.
- Foreign Body	.	P.	P.	P.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	.	1.	1.	1.	1.	1.	1.	1.	1.	1.
- Interstitial Fibrosis	.	1.	.	1.	1.	1.	1.	.	.	1.
- Bronchiolization	.	.	.	1.	1.	1.
- Pleura Fibrosis	.	1.	1.	1.	.	.
TRACHEA	:	-	-	-	-	-	+	-	-	-
- Chronic Inflammation	1.	.	.	.
EPIDIDYMIS, LEFT	:	+	+	-	-	+	+	+	-	-
- Hypospermia	.	3.	4.	.	.	3.	3.	4.	.	.
TESTIS, LEFT	:	+	+G	+	+G	+G	+G	+	-	+G
- Interstitial Cell Adenoma	.	P.	P.	.	P.	P.	P.	.	.	.
- Tubular Atrophy	.	.	2.	2.	.	.	.	2.	.	1.
- Edema	2.	.	.
- Interstitial Cell Hyperplasia	3.
STERNUM	:	-	-	-	-	-	-	-	-	-
PREPUTIAL GLANDS	:		+G							
- Carcinoma	.		P.							
SYSTEMIC NEOPLASMS	:									+
- Malignant Lymphoma	.									N0.
LIVER	:	'	'	'	+G	'	'	+G	'	+G
- Bile Duct Hyperplasia	2.	.	.	2.	.	2.
- Mononuclear Cell Leukemia	M45.
SPLEEN	:	'	+G	'	'	'	'	+G	'	'
- Mononuclear Cell Leukemia	M45.
- Extramedullary Hematopoesis	.	.	.	4.
LUNG	:	'	G	'	'	'	'	'	'	'
TESTES	:	'	'	'	'	+G	+G	+G	+G	'
- Edema	2.	.	.
- Tubular Atrophy	4.	.	.
- Interstitial Cell Adenoma	P.	P.	.	(P.	.
KIDNEYS	:	'		+G						
- Nephropathy	.	.		3.						
MESENTERY	:	'	+	'	'	'	'	'	'	'
- Schwannoma	.	.	P.
BODY CAVITIES	:	'	G	'	'	'	'	'	'	'
EYES	:	'	'	'	'	'	'	'	'	+G
- Cornea Ulcer	(4.
- Squamous Cell Carcinoma	M10.
HARDERIAN GLANDS	:	'	'	'	+G	'	'	'	'	+G
- Inflammation	(1.	M10.
- Squamous Cell Carcinoma

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SPONSOR : THE HAMNER	PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 3, LA

ANIMAL NUMBER :

	307 MK0	319 MK0+	321 MK0+	349 MK0+	350 MK0	351 MK0	352 MK0	353 MK0	354 MK0+	355 MK0+
THYROID GLAND	:	:	:	+G	:	:	:	:	:	:
- Follicular Cell Carcinoma	(P.					
GANGLION, GASS., RIGHT	:	:	:	:	:	:	:	:	:	+G
- Squamous Cell Carcinoma	M10.
PITUITARY GLAND	:	:	:	+G	:	:	:	:	:	+G
- Pars Distalis Adenoma	P.					.
- Pars Distalis Hyperplasia Focal	3.
PALATE (ORAL MUCOSA)	:	:	:	-G	:	:	:	:	:	+G
- Squamous Cell Carcinoma	NO.
STOMACH	:	:	:	:	:	:	:	:	:	G
INTEST-LG, CECUM	:	:	:	:	:	:	:	:	:	G
INTEST-SM, ILEUM	:	:	:	:	:	:	:	:	:	G
INTEST-LG, COLON	:	:	:	+G	:	:	:	:	:	
- Adhesion	P.					

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TEST SYSTEM : RAT, , 18 MO, INHALATION	DATE : 29-AUG-13
SPONSOR : THE HAMNER	PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 3, LA

ANIMAL NUMBER :

	356 MK0	357 MK0+	358 MK0	359 MK0+	360 MK0	361 MK0	362 MK0	363 MK0	365 MK0	366 MK0
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Mononuclear Cell Leukemia	M45.	M45.
- Alveolar Epithelium Hyperplasia	.	.	1.	3.	.	.
- Foreign Body	.	P.	P.	P.	P.	P.	.	P.	P.	P.
- Alveolus Inflammation	.	1.	1.	1.	1.	1.	.	1.	1.	1.
- Interstitial Fibrosis	.	1.	1.	1.	1.	1.	.	1.	1.	1.
- Pleura Fibrosis	.	1.	.	1.	1.	1.
TRACHEA	:	+	-	-	-	-	-	-	-	-
- Chronic Inflammation	.	1.
EPIDIDYMIS, LEFT	:	+	+G	+	-	-	+	-	+G	+
- Mesothelioma	.	.	M45.
- Hypospermia	.	3.	4.	.	4.	.	4.	3.	.	.
TESTIS, LEFT	:	+G	+G	+G	+	+	+G	+G	+G	+
- Interstitial Cell Adenoma	.	P.	P.	P.	.	P.	P.	P.	P.	P.
- Mesothelioma	.	.	M45.	M26.	.	.
- Tubular Atrophy	1.
STERNUM	:	-	-	-	-	-	-	-	-	-
PREPUTIAL GLANDS	:	,	,	,	,	,	,	,	,	+G
- Carcinoma	(P.	.
SYSTEMIC NEOPLASMS	:	,	,	,	,	,	,	,	,	,
- Malignant Lymphoma	.	.	NO.	NO.	NO.	.
LIVER	:	,	,	,	,	,	+G	,	+G	+G
- Mononuclear Cell Leukemia	M26.	M45.	.
- Macrovesiculation	2.
SPLEEN	:	,	,	,	,	,	+G	,	+G	+G
- Mononuclear Cell Leukemia	M45.	.	.
- Extramedullary Hematopoiesis	2.	3.	.	.	.
TESTES	:	+G	,	+G	G	,	+G	+G	+G	,
- Mesothelioma	M26.	.	.
- Interstitial Cell Adenoma	.	P.	P.	.	.	P.	P.	.	.	P.
KIDNEYS	:	,	,	+G	,	,	,	,	,	,
- Nephropathy	.	.	2.	3.	.	.
MESENTERY	:	,	,	,	,	,	,	,	+G	,
- Mesothelioma	M26.	.	.
PARATHYROID GLAND	:	,	,	,	,	,	G	,	,	,
LYMPH NODES	:	,	,	,	,	,	+G	,	+G	,
- Pigment	2.	.	.	.
- Mesothelioma	M26.	.	.
BODY CAVITIES	:	,	,	,	,	,	,	,	+G	,
- Mesothelioma	M26.	.	.
EYES	:	,	,	,	,	,	+G	,	,	,
- Cornea Ulcer	(4.	.	.	.

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 3, LA

ANIMAL NUMBER :

	356 MK0	357 MK0+	358 MK0	359 MK0+	360 MK0	361 MK0	362 MK0	363 MK0	365 MK0	366 MK0
PALATE (ORAL MUCOSA)	:	:	:	+						
- Squamous Cell Carcinoma	NO.					
GANGLION, GASS., LEFT					+G					
- Squamous Cell Carcinoma	M10.					
EPIDIDYMIDES						+G				
- Mesothelioma	NO.				

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 3, LA

ANIMAL NUMBER :

	367 MK0+	368 MK0	369 MK0	370 MK0	371 MK0+	372 MK0	373 MK0+	374 MK0+	375 MK0	376 MK0
GENERAL OBSERVATIONS	:	G	'	'	'	'	'	'	'	'
LUNG, LEFT	:	+G	-	+	+	+G	+G	+	+	+
- Mononuclear Cell Leukemia	.	M45.	.	.	M45.	.	.	M45.	.	.
- Alveolar Epithelium Hyperplasia	.	.	.	1.	.	1.
- Foreign Body.	.	.	.	P.	.	P.	P.	.	P.	P.
- Alveolus Inflammation.	.	.	.	1.	1.	.	2.	1.	.	1.
- Interstitial Fibrosis.	.	.	.	1.	.	1.	.	.	1.	1.
- Pleura Fibrosis.	.	.	.	1.	.	1.
TRACHEA	:	-	-	-	-	-	-	-	-	+
- Chronic Inflammation	1.
EPIDIDYMIS, LEFT	:	+	+	-	-	-	+	-	-	+
- Hypospermia	.	4.	3.	.	.	.	3.	.	3.	4.
TESTIS, LEFT	:	+G	+G	+	+	0	+G	+G	-G	+G
- Interstitial Cell Adenoma	.	P.	P.	.	.	P.	P.	.	P.	P.
- Tubular Atrophy.	.	3.	.	2.	2.	.	.	2.	.	.
- Edema	2.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	G	'	'	'	'	'	'	'
SYSTEMIC NEOPLASMS	:	+	'	'	'	'	+	'	'	'
- Malignant Lymphoma	.	N0.					N0.			N0.
LIVER	:	+G	+G	'	'	'	+G	'	'	+G
- Bile Duct Hyperplasia.	.	.	2.
- Mixed Cell Focus	.	.	P.
- Mononuclear Cell Leukemia	.	M45.	.	.	M45.	.	M45.	.	M45.	.
- Macrovesiculation	.	.	2.
SPLEEN	:	+G	+G	'	'	'	+G	'	'	+G
- Mononuclear Cell Leukemia	.	M45.	.	M45.	.	M45.	.	M45.	.	M45.
- Extramedullary Hematopoiesis	.	.	2.
LUNG	:	+G	'	'	'	'	+G	'	'	'
- Mononuclear Cell Leukemia	.	M45.	.	M45.	.	M45.	.	M45.	.	M45.
TESTES	:	G	+G	+G	'	'	+G	'	'	+G
- Tubular Atrophy.	.	.	(4.	(4.	.	.
- Interstitial Cell Adenoma	.	.	.	(P.	.	P.	.	P.	.	.
MAMMARY GLAND	:	'	+G	'	'	'	'	'	'	'
- Fibroadenoma	.	.	P.
- Adenocarcinoma	.	.	P.
KIDNEYS	:	+G	'	'	'	'	'	'	'	'
- Nephropathy	.	2.
- Mononuclear Cell Leukemia	.	M45.
LYMPH NODES	:	'	'	'	'	'	G	+G	'	+G
- Plasma Cell Hyperplasia	3.	.	.	.
- Mononuclear Cell Leukemia	M45.	.	.	.
EYES	:	'	'	'	'	'	'	+G	'	'
- Squamous Cell Carcinoma	M10.	.	.	.

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TEST SYSTEM	:	RAT, , 18 MO, INHALATION	DATE :	29-AUG-13
SPONSOR	:	THE HAMNER	PathData®System V6.2d2	

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 3, LA

ANIMAL NUMBER :

	367 MK0+	368 MK0	369 MK0	370 MK0	371 MK0+	372 MK0	373 MK0+	374 MK0+	375 MK0	376 MK0
HARDERIAN GLANDS	:	:	:	:	+	G	:	+	G	:
- Squamous Cell Carcinoma	M10.	.
- Mononuclear Cell Leukemia	M45.
GANGLION, GASS., RIGHT	:	:	:	:	:	:	:	:	+	G
- Squamous Cell Carcinoma	M10.	.
PALATE (ORAL MUCOSA)	:	:	:	:	:	:	:	:	+	G
- Squamous Cell Carcinoma	NO.	.
STOMACH	:	:	:	:	G	:	:	:	:	:
INTEST-LG, CECUM	:	:	:	:	+	G	:	:	:	:
- Ulceration	4.
INTEST-SM, ILEUM	:	:	:	:	+	G	:	:	:	:
- Mononuclear Cell Leukemia	M45.
BRAIN	:	:	:	:	+	G	:	:	:	:
- Mononuclear Cell Leukemia	M45.
PANCREAS	:	:	:	:	+	G	:	:	:	:
- Mononuclear Cell Leukemia	M45.

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TEST ITEM : AMPHIBOLE ASBESTOS
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PATHOL. NO.: 90188 GAW
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 3, LA

ANIMAL NUMBER :

	377 MK0	378 MK0	379 MK0+	380 MK0	381 MK0	382 MK0+	383 MK0+	384 MK0	385 MK0	386 MK0+
LUNG, LEFT	:	+	+	+	+	+	0	+	+	+
- Mononuclear Cell Leukemia	M45.
- Alveolar Epithelium Hyperplasia	.	1.
- Infiltration Mixed Cell	.	.	2.
- Foreign Body	.	P.	P.	.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	.	2.	1.	.	1.	1.	1.	1.	1.	1.
- Interstitial Fibrosis	.	1.	1.	.	1.	.	1.	1.	1.	.
- Bronchiolization	1.	.	.	.
- Pleura Fibrosis	1.	.	.	1.	1.	.
TRACHEA	:	-	-	-	-	-	0	-	-	-
EPIDIDYMIS, LEFT	:	+	+	+	+	-	+	+G	+G	+
- Mesothelioma	No.	.	.
- Hypospermia	.	2.	3.	4.	2.	.	4.	4.	.	4.
- Blood Vessels Arteritis	2.	.	.
TESTIS, LEFT	:	+G	+G	+G	+G	-	+	+G	+G	+G
- Interstitial Cell Adenoma	.	P.	P.	P.	P.	.	.	P.	P.	P.
- Mesothelioma	M26.
- Tubular Atrophy	.	.	.	2.	.	.	4.	.	.	2.
- Edema	3.	.	.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	;	;	;	;	;	;	+G	;	;
- Sarcoma	P.	.	.
PREPUTIAL GLANDS	:	;	;	;	;	;	;	;	+G	;
- Adenoma	(P.	.	.
SYSTEMIC NEOPLASMS	:	;	;	;	;	;	;	;	;	;
- Malignant Lymphoma	+
NO.										No.
LIVER	:	;	;	;	;	;	;	;	;	+G
- Mononuclear Cell Leukemia	M45.
SPLEEN	:	;	;	;	;	;	AG	+G	;	+G
- Mononuclear Cell Leukemia	M45.
- Extramedullary Hematopoesis	3.	.	.	.
- Mesothelioma	M26.	.	.
LUNG	:	;	;	;	;	;	G	;	;	+G
- Mononuclear Cell Leukemia	M45.
TESTES	:	+G	'	G	+G	'	+G	G	+G	'
- Tubular Atrophy	.	.	.	(2.	.	4.	.	(4.	.	2.
- Interstitial Cell Adenoma	.	P.
MAMMARY GLAND	:	;	;	;	;	;	;	;	;	+G
- Secretion	2.
KIDNEYS	:	;	;	;	;	;	;	;	;	+G
- Nephropathy	3.
- Tubule Pigment	3.	.	.	.
BODY CAVITIES	:	;	;	;	;	;	;	;	;	+G
- Mesothelioma	M26.	.	.

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SPONSOR : THE HAMNER	PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 3, LA

ANIMAL NUMBER :

	377 MK0	378 MK0	379 MK0+	380 MK0	381 MK0	382 MK0+	383 MK0+	384 MK0	385 MK0	386 MK0+
EYES	:	:	:	+G	:	+G	:	:	:	:
- Retinal Degeneration	:	.	.	.	4.					
- Lens Cataract	:	.	.	.	(2.					
- Anterior Synechia	:	.	.	.	P.					
- Cornea Inflammation	:	.	.	.	3.					
- Panophthalmitis	:	.	.	4.	.					
.....
PITUITARY GLAND	:	:	:	:	:	:	:	:	:	+G
- Pars Distalis Adenoma	:	P.				
.....
PALATE (ORAL MUCOSA)	:	:	:	+G	:	:	:	:	:	:
- Squamous Cell Carcinoma	:	.	.	NO.						
.....
GANGLION, GASS., LEFT	:	:	:	+G	:	:	:	:	:	:
- Squamous Cell Carcinoma	:	.	.	M10.						
.....

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
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DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 3, LA

ANIMAL NUMBER :

	387 MK0+	388 MK0	389 MK0	390 MK0+	391 MK0+	392 MK0	393 MK0	394 MK0	395 MK0	397 MK0+
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Mononuclear Cell Leukemia	M45.
- Foreign Body	.	P.	P.	P.	P.	P.	P.	P.	P.	.
- Alveolus Inflammation	.	1.	1.	1.	1.	2.	1.	1.	1.	.
- Interstitial Fibrosis	.	1.	1.	1.	.	.	1.	1.	.	.
- Bronchiolization	.	1.
- Osteosarcoma	M65.
- Pleura Fibrosis	.	.	1.	1.	.	.	.	1.	1.	.
TRACHEA	:	-	-	+	-	-	-	-	-	-
- Chronic Inflammation	.	.	.	2.
EPIDIDYMIS, LEFT	:	-	+	+	-	+	-	+	+	-
- Hypospermia	.	.	2.	2.	2.	.	3.	2.	.	.
TESTIS, LEFT	:	+G	+G	+G	+	+G	+G	+G	+G	+G
- Interstitial Cell Adenoma	.	.	P.	P.	.	P.	P.	P.	P.	P.
- Tubular Atrophy	.	.	.	2.	2.	2.
- Edema	.	1.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	+G	+G	'	'	'	'	'	+G
- Keratoacanthoma	.	.	.	P.
- Epidermal Cyst	.	.	P.	P.	.
SYSTEMIC NEOPLASMS	:	'	'	'	'	'	'	'	'	'
- Malignant Lymphoma	+
N0.	N0.
LIVER	:	'	'	'	+G	+G	'	+G	+G	'
- Bile Duct Hyperplasia	.	.	.	1.	.	2.	2.	.	.	.
- Mononuclear Cell Leukemia	M45.
- Macrovesiculation	2.	.	2.	.	.	.
SPLEEN	:	'	'	'	+G	'	+G	+G	'	+G
- Mononuclear Cell Leukemia	M45.
- Extramedullary Hematopoesis	2.	2.	.	.	.
- Atrophy	.	.	.	3.
LUNG	:	'	'	'	+G	'	'	'	'	'
- Osteosarcoma	.	.	.	M65.
TESTES	:	'	'	'	+G	'	0G	'	+G	'
- Tubular Atrophy	.	.	.	(3.	.	(3.	.	P.	P.	.
- Interstitial Cell Adenoma
- Sperm Granuloma	.	.	P.
MAMMARY GLAND	:	'	'	'	'	'	'	'	'	'
- Galactocole Hemorrhage	P.	.	.
KIDNEYS	:	+G	'	'	'	+G	'	+G	'	+G
- Nephropathy	.	3.	.	4.	.	3.	.	3.	.	.
ADRENAL GLANDS	:	'	'	'	'	G	'	'	'	'
LYMPH NODES	:	'	'	'	+G	'	'	'	'	'
- Plasma Cell Hyperplasia	.	.	.	2.

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TEST ITEM	:	AMPHIBOLE ASBESTOS	PATHOL. NO.:	90188 GAW
TEST SYSTEM	:	RAT, , 18 MO, INHALATION	DATE :	29-AUG-13
SPONSOR	:	THE HAMNER	PathData®System V6.2d2	

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 3, LA

ANIMAL NUMBER :

	387	388	389	390	391	392	393	394	395	397
	MK0+	MK0	MK0	MK0+	MK0+	MK0	MK0	MK0	MK0	MK0+
EYES	:	'	'	'	+G	'	'	'	'	'
- Cornea Ulcer.	1.					
- Socket Osteosarcoma	(NO.					
PITUITARY GLAND		+G	'	'	+G	'	'	'	'	'
- Pars Distalis Adenoma.	.	.	.	P.	P.					
STOMACH		'	'	'	G	+G	'	'	'	'
- Ulceration	3.				
INTEST-LG, CECUM		'	'	'	G	'	'	'	'	'
INTEST-SM, ILEUM		'	'	'	G	'	'	'	'	'
INTEST-SM, JEJUNUM		'	'	'	G	'	'	'	'	'
INTEST-LG, COLON		'	'	'	G	'	'	'	'	'
BRAIN		'	'	'	'	'	'	'	'	'

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
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PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 4, LA

ANIMAL NUMBER :

	419 MK0+	420 MK0+	423 MK0	449 MK0	450 MK0	451 MK0+	452 MK0	453 MK0	454 MK0	455 MK0
LUNG, LEFT	:	+	+	+	+	+	+	+G	+	+
- Mononuclear Cell Leukemia	.	.		M45.	.	M26.
- Foreign Body	.	.		P.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	.	.		1.	1.	1.	1.	1.	1.	2.
- Interstitial Fibrosis	.	.		.	1.	1.	.	1.	1.	1.
- Alveolar Bronchiolar Adenoma	P.	.	P.
- Infiltrate Mixed Cell Focal	.	.		1.
- Pleura Fibrosis	1.	.	1.	1.	.
TRACHEA	:	-	-	-	-	+	-	-	-	-
- Chronic Inflammation	1.
EPIDIDYMIS, LEFT	:	+	-	+	+	+	+G	-	-	+
- Mesothelioma	NO.	.	.	.
- Hypospermia	.	4.	.	3.	4.	4.	.	.	.	4.
TESTIS, LEFT	:	+G	+G	+G	+G	+	+G	+	+G	+G
- Interstitial Cell Adenoma	.	P.	P.	P.	P.	P.	P.	.	P.	P.
- Mesothelioma	M26.
- Tubular Atrophy	.	.	.	1.	.	.	.	2.	.	.
- Edema	2.	.	.
STERNUM	:	-	-	-	-	-	-	-	-	-
PREPUTIAL GLANDS	:	+G	+G	'	'	'	'	'	'	+G
- Adenoma	P.
- Carcinoma	.	P.	(P.
SYSTEMIC NEOPLASMS	:	'	'	+	'	'	'	'	'	'
- Malignant Lymphoma	NO.
LIVER	:	'	+G	+G	'	'	+G	+G	'	'
- Bile Duct Hyperplasia	2.	.
- Mononuclear Cell Leukemia	.	.	.	M45.	.	P.
- Macrovesiculation	1.	.	.
- Hepatodiaphragmatic Nodule	P.
- Eosinophilic Focus	.	.	.	P.
SPLEEN	:	'	+G	+G	'	'	+G	+G	'	'
- Mononuclear Cell Leukemia	.	.	.	M45.	.	M26.
- Extramedullary Hematopoiesis	.	.	.	3.	.	.	2.	.	.	.
LUNG	:	G	'	'	'	'	+G	'	'	'
- Mononuclear Cell Leukemia	M26.
TESTES	:	0G	0G	+G	+G	+G	0G	+G	'	+G
- Tubular Atrophy	.	.	.	(4.	.	.	P.	P.	.	(P.
- Interstitial Cell Adenoma	P.	.	.	.
MAMMARY GLAND	:	'	'	+G	'	'	'	'	'	'
- Fibroadenoma	.	.	.	P.
KIDNEYS	:	'	+G	'	'	'	+G	+G	'	'
- Nephropathy	.	3.	3.	3.	3.	2.	3.	.	.	.
ADRENAL GLANDS	:	'	'	'	'	'	+G	'	'	'
- Mononuclear Cell Leukemia	M26.

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TEST ITEM : AMPHIBOLE ASBESTOS
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 4, LA

ANIMAL NUMBER :

	419 MK0+	420 MK0+	423 MK0	449 MK0	450 MK0	451 MK0+	452 MK0	453 MK0	454 MK0	455 MK0
MESENTERY	:	:	:	:	:	+G	:	:	:	:
- Mesothelioma.	M26.
LYMPH NODES	:	:	:	+G	:			+G		
- Plasma Cell Hyperplasia	.	.	.	2.	.			2.	.	.
BODY CAVITIES	:	:	:			+G				
- Mesothelioma.	.	.	.			M26.
URINARY BLADDER	:	:	:			G				
ADIPOSE TISSUE	:	:	:			+G				
- Mesothelioma.	.	.	.			M26.
SALIVARY GLANDS	:	:	:			G				

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TEST SYSTEM : RAT, , 18 MO, INHALATION	DATE : 29-AUG-13
SPONSOR : THE HAMNER	PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 4, LA

ANIMAL NUMBER :

	456 MK0	457 MK0	458 MK0	459 MK0	460 MK0	461 MK0	462 MK0+	463 MK0	464 MK0	465 MK0
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Mononuclear Cell Leukemia	.	.	M45.	.	M45.
- Alveolar Epithelium Hyperplasia	.	2.
- Foreign Body	.	P.	P.	P.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	.	2.	2.	1.	2.	1.	1.	1.	1.	1.
- Interstitial Fibrosis	.	1.	1.	1.	1.	1.	1.	1.	1.	1.
- Pleura Fibrosis	.	1.	1.	.	1.	.	.	.	1.	.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	+	+	-	+	+	-	-	+	-
- Hypospermia	.	4.	4.	.	4.	3.	2.	.	3.	.
TESTIS, LEFT	:	+G P.	+G P.	+	+G P.	+G P.	+	+	+G P.	+
- Interstitial Cell Adenoma	1.	.	1.
- Tubular Atrophy	1.	.	2.
- Edema	1.	.	2.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	+G P.	+G P.	'	+G P.	'	'	'	'
- Epidermal Cyst
- Sarcoma
- Basal Cell Adenoma
SYSTEMIC NEOPLASMS	:	'	'	+	'	+	'	'	'	'
- Malignant Lymphoma	.	.	.	NO.	.	NO.
LIVER	:	'	'	+	'	+	'	'	+	'
- Bile Duct Hyperplasia	.	.	2.	2.	2.	2.	.	.	2.	.
- Mononuclear Cell Leukemia	.	.	M45.	.	M45.
- Macrovesiculation	.	.	.	1.
SPLEEN	:	+G M45.	'	+G M45.	+G M45.	'	'	'	+G 2.	'
- Mononuclear Cell Leukemia
- Extramedullary Hematopoesis	.	2.	.	2.	2.	.
TESTES	:	+G (5.	+G 2.	'	+G P.	+G P.	+G P.	0G P.	'	+G (3.
- Tubular Atrophy
- Interstitial Cell Adenoma
- Interstitial Cell Hyperplasia	.	.	.	(2.
MAMMARY GLAND	:	'	'	'	'	'	'	'	+G 2.	'
- Secretion
KIDNEYS	:	'	'	'	'	'	'	+G 3.	+G 3.	'
- Nephropathy	3.	3.	3.
ADEXRAL GLANDS	:	'	'	'	'	'	'	+G 2.	'	'
- Cortex Focal Hyperplasia	(P.	.	.
- Medulla Phaeochromocytoma
PITUITARY GLAND	:	'	'	'	'	'	'	'	+G P.	'
- Pars Distalis Adenoma

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 4, LA

ANIMAL NUMBER :

	466 MK0	467 MK0	468 MK0	469 MK0+	470 MK0+	471 MK0+	472 MK0	473 MK0+	474 MK0	475 MK0
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Mononuclear Cell Leukemia	.	.	.	M45.
- Foreign Body	.	.	.	P.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	.	.	.	1.	1.	1.	2.	1.	1.	1.
- Interstitial Fibrosis	.	.	.	1.	1.	1.	1.	1.	1.	2.
- Pigmented Histiocytes	1.	.	.	1.	2.
- Pleura Fibrosis	1.	.	.
TRACHEA	:	-	-	-	-	-	-	-	+	-
- Chronic Inflammation	2.	.
EPIDIDYMIS, LEFT	:	-	+	+	-	-	+	+	-	+
- Mesothelioma	M25.
- Hypospermia	.	.	.	3.	2.	.	.	2.	.	2.
TESTIS, LEFT	:	+	+G	+G	+	-	+G	+G	+G	+G
- Interstitial Cell Adenoma	.	.	P.	P.	.	P.	P.	.	P.	.
- Mesothelioma	NO.
- Tubular Atrophy	.	.	1.	.	2.
- Edema	.	.	2.	1.	.	.
- Interstitial Cell Hyperplasia	1.
STERNUM	:	-	-	-	-	-	-	0	-	-
SKIN/SUBCUTIS	:	+G	'	'	'	'	'	'	'	'
- Squamous Cell Carcinoma	.	.	P.
SYSTEMIC NEOPLASMS	:	+	'	'	'	'	'	'	'	+
- Malignant Lymphoma	.	.	NO.	NO.
LIVER	:	+G	'	'	'	'	+G	'	+G	'
- Bile Duct Hyperplasia	2.	2.	.
- Mononuclear Cell Leukemia	.	.	M45.	M45.	.
- Mesothelioma	M25.
SPLEEN	:	+G	'	'	'	'	'	+G	'	+G
- Mononuclear Cell Leukemia	.	M45.	M45.	.
- Extramedullary Hematopoiesis	3.	.	.
TESTES	:	'	+G	'	'	'	'	+G	0G	'
- Interstitial Cell Adenoma	.	.	P.	.	.	.	P.	.	.	.
MAMMARY GLAND	:	'	'	'	'	'	'	+G	'	'
- Secretion	2.	.	.
KIDNEYS	:	'	'	'	+G	'	+G	'	+G	'
- Nephropathy	1.	2.	2.	.	.	.
ADEXRAL GLANDS	:	'	'	'	'	+P.	'	'	'	'
- Cortical Carcinoma	(P.
LYMPH NODES	:	'	'	'	'	'	'	+G	'	'
- Plasma Cell Hyperplasia	3.	.	.
- Cyst	P.	.	.
BODY CAVITIES	:	'	'	'	'	'	'	+G	'	'
- Mesothelioma	M25.	.	.	.

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TEST SYSTEM	: RAT, , 18 MO, INHALATION	DATE :	29-AUG-13
SPONSOR	: THE HAMNER	PathData®System V6.2d2	

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 4, LA

ANIMAL NUMBER :

	466 MK0	467 MK0	468 MK0	469 MK0+	470 MK0+	471 MK0+	472 MK0	473 MK0+	474 MK0	475 MK0
EYES	:	:	:	:	:	:	:	:	+G	:
- Cornea Ulcer.	4.		
THYROID GLAND	:	:	:	:	:	:	:	+G		
- Follicular Cell Carcinoma	(NO.		
GANGLION, GASS, RIGHT	:	:	:	:	:	:	:	+G		
- Squamous Cell Carcinoma	M10.		
PITUITARY GLAND	:	:	:	:	:	:	:	+G		
- Pars Distalis Adenoma	P.		
PALATE (ORAL MUCOSA)	:	:	:	:	:	:	:	+G		
- Squamous Cell Carcinoma	NO.		

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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 4, LA

ANIMAL NUMBER :

	476 MK0	477 MK0	479 MK0	480 MK0	481 MK0	482 MK0	483 MK0	484 MK0	485 MK0	486 MK0
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Alveolar Epithelium Hyperplasia	.	.	1.	1.	1.	.
- Foreign Body.	.	P.								
- Alveolus Inflammation.	1.	2.	1.	1.	1.	1.	1.	1.	1.	1.
- Interstitial Fibrosis.	1.	2.	1.	1.	1.	1.	1.	1.	1.	1.
- Pleura Fibrosis.	1.	1.	1.	1.	1.	1.	1.	1.	1.	1.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	-	+	-	-	-	-	+	+	+
- Hypospermia	.	.	3.	.	.	.	4.	3.	3.	.
TESTIS, LEFT	:	+	+G	+	+G	+	+G	+G	+G	+
- Interstitial Cell Adenoma	.	P.	.	P.	P.	P.	P.	P.	P.	.
- Tubular Atrophy.	2.	.	1.	2.	1.
- Edema	.	.	2.	1.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	'	'	'	'	'	G	'	'
LIVER	:	'	'	'	'	'	'	+G	'	+G
- Bile Duct Hyperplasia.	2.	2.	2.	.
SPLEEN	:	'	'	'	'	'	'	+G	+G	'
- Extramedullary Hematopoiesis.	3.	.	.	.
- Stromal Hyperplasia	3.	.	.	.
TESTES	:	'	+G	'	+G	'	+G	+G	+G	+G
- Interstitial Cell Adenoma	.	P.	.	P.	P.	P.	P.	P.	P.	(P.
LYMPH NODES	:	'	'	'	+G	'	'	0G	'	'
- Plasma Cell Hyperplasia	.	.	.	2.
EYES	:	'	'	'	'	'	'	+G	'	'
- Phthisis Bulbi	(P.	.	.	.
SALIVARY GLANDS	:	'	'	'	'	'	'	+	'	'
- Sublingual Adenocarcinoma	(P.	.	.	.

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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 4, LA

ANIMAL NUMBER :

	487 MK0	488 MK0	489 MK0	490 MK0	491 MK0	492 MK0	493 MK0	494 MK0	496 MK0
LUNG, LEFT	:	+	+	+	+	+	+	+	+
- Alveolar Epithelium Hyperplasia	.	.	4.	.	3.
- Foreign Body	.	P.							
- Alveolus Inflammation	.	1.	1.	1.	1.	1.	2.	1.	1.
- Interstitial Fibrosis	.	1.	1.	1.	1.	1.	1.	1.	1.
- Pleura Fibrosis	.	1.	.	1.	.	1.	.	.	1.
TRACHEA	:	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	+	+	+	+	-	+	-	-
- Hypospermia	.	3.	4.	3.	3.	4.	.	3.	.
TESTIS, LEFT	:	+G	+G	+G	+G	+	+G	+	+
- Interstitial Cell Adenoma	.	P.	P.	P.	P.	.	P.	.	P.
- Tubular Atrophy	4.	.	1.	.
- Edema	1.	.
STERNUM	:	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	+G	'	'	'	'	'	'
- Galactocoele Hemorrhage	.	.	P.
LIVER	:	'	'	'	+G	'	'	'	'
- Bile Duct Hyperplasia	2.
SPLEEN	:	'	'	'	+G	'	'	'	'
- Extramedullary Hematopoiesis	2.
LUNG	:	'	'	'	+G	'	'	'	'
- Congestion	2.
TESTES	:	+G	+G	+G	+G	+G	'	+G	'
- Edema	.	.	3.	2.
- Tubular Atrophy	.	(4.	4.	(2.
- Interstitial Cell Adenoma	P.	P.	P.	P.	P.
MAMMARY GLAND	:	'	+	'	'	'	'	'	'
- Galactocole Hemorrhage	.	.	P.
LYMPH NODES	:	G	+G	'	'	'	'	'	'
- Plasma Cell Hyperplasia	.	.	2.

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DATE : 29-AUG-13
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 5, LA

ANIMAL NUMBER :

	520 MK0+	549 MK0+	550 MK0	551 MK0	552 MK0+	553 MK0	554 MK0	555 MK0	556 MK0	557 MK0
LUNG, LEFT	: +G	+	+	+	+	+	+	+	+	+
- Chronic Inflammation	1.	1.
- Mononuclear Cell Leukemia	M45.	.	.	M45.
- Alveolar Epithelium Hyperplasia	3.	.	.
- Foreign Body	P.	P.	P.	P.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	1.	1.	1.	1.	.	1.	1.	2.	1.	1.
- Interstitial Fibrosis	1.	1.	1.	.	.	1.	1.	1.	1.
- Alveolar Bronchiolar Adenoma	P.
- Congestion	3.
- Alveolar Bronchiolar Carcinoma	P.
- Pleura Fibrosis	1.	.	1.	1.	1.	1.	.	.	1.
TRACHEA	:	-	-	-	-	-	-	-	-	-
EPIDIDYMIS, LEFT	:	-	+	-	+	-	+	+	-	-
- Mesothelioma	M26.
- Hypospermia	4.	.	4.	.	4.	3.	.	.	4.
TESTIS, LEFT	:	+G	+G	+	+G	-G	+G	+G	+	+G
- Interstitial Cell Adenoma	P.	P.	.	P.	.	P.	P.	.	P.	P.
- Tubular Atrophy	1.	2.	1.	.	.	.	2.	.	.	.
- Edema	1.	.	.	.	1.	.	.	.
STERNUM	:	-	-	-	-	-	-	-	-	-
PREPUTIAL GLANDS	:	+G	'	'	'	+G	'	'	+G	'
- Adenoma	P.	(P.
- Carcinoma	2.	.	.
- Inflammation Chronic Active	P.	.
- Cyst
SYSTEMIC NEOPLASMS	:	+	,	,	,	+	,	,	,	,
- Malignant Lymphoma	N0.	N0.
LIVER	:	+G	+G	'	'	+G	'	+G	+G	+G
- Bile Duct Hyperplasia	2.	.	.	.	2.	.	2.	.
- Mononuclear Cell Leukemia	M45.	.	.	M45.
- Macrovesiculation	2.	.
- Basophilic Focus	P.	.	.	.
- Hepatodiaphragmatic Nodule	P.	.	.	.
- Eosinophilic Focus	P.	.	.	.
SPLEEN	:	+G	+G	'	'	+G	'	+G	'	+G
- Mononuclear Cell Leukemia	M45.	.	.	M45.
- Extramedullary Hematopoiesis	2.	.	3.	.	.	2.	.
- Lymphoid Depletion	2.
LUNG	:	+G	+G	'	'	+G	'	'	'	'
- Mononuclear Cell Leukemia	M45.	.	.	M45.
- Alveolar Bronchiolar Carcinoma	N0.
- Cystic Keratinizing Epithelioma	P.
TESTES	:	0	0G	'	'	+G	'	+G	'	+G
- Tubular Atrophy	(3.	3.	.	.	.
- Mesothelioma	M26.	.	.	.
- Interstitial Cell Adenoma	P.	.	.	P.	P.	.
- Interstitial Cell Hyperplasia	3.	.	.	.

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SPONSOR : THE HAMNER	PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 5, LA

ANIMAL NUMBER :

	520 MK0+	549 MK0+	550 MK0	551 MK0	552 MK0+	553 MK0	554 MK0	555 MK0	556 MK0	557 MK0
MAMMARY GLAND	:	:	:	:	+G	:	:	G	:	:
- Secretion.	2.
KIDNEYS		+G	+G		+G			+G		
- Nephropathy	.	2.	.		4.			3.		
- Alveolar Bronchiolar Carcinoma	.	.	M09.
ADRENAL GLANDS								+G		
- Cortex Vacuolization	2.		
- Cortex Focal Hyperplasia	2.		
MESENTERY								+G		
- Mesothelioma	M26.		
LYMPH NODES		:	+G				+G		G	
- Pigment	.	.	.					3.	.	.
- Sinus Dilation	.	.	2.	
BODY CAVITIES		G	+G					+G		
- Mesothelioma	.	.	.					M26.		
- Alveolar Bronchiolar Carcinoma	.	.	M09.					.		
BONE		+G								
- Alveolar Bronchiolar Carcinoma	.	M09.								
SEMINAL VESICLES			G							
PITUITARY GLAND		+G	+G		+G			+G		
- Pars Distalis Adenoma	.	.	.		P.			P.		
- Pars Distalis Hyperplasia Focal	.	3.
- Pars Distalis Cyst	.	.	P.	
INTEST-LG, COLON		+G								
- Muscularis Focal Thickening	.	P.								
HEART		+G								
- Alveolar Bronchiolar Carcinoma	.	M09.								
DIAPHRAGM		+G								
- Alveolar Bronchiolar Carcinoma	.	M09.								
EPIDIDYMIDES					+G					
- Mesothelioma	(NO.					

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 5, LA

ANIMAL NUMBER :

	558 MK0	559 MK0	560 MK0	561 MK0	562 MK0	563 MK0+	564 MK0	565 MK0	566 MK0	567 MK0+
GENERAL OBSERVATIONS	:	:	:	:	:	G	:	:	:	:
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Chronic Inflammation	:	1.
- Mononuclear Cell Leukemia	:	M45.
- Alveolar Epithelium Hyperplasia	:	1.	3.	.	4.
- Foreign Body	:	P.	P.	P.	P.	P.	P.	P.	P.	P.
- Alveolar Inflammation	:	1.	1.	1.	1.	1.	1.	1.	1.	1.
- Interstitial Fibrosis	:	1.	1.	1.	1.	1.	1.	1.	1.	1.
- Alveolar Bronchiolar Hyperplasia	:	2.	.
- Pleura Fibrosis	:	1.	.	1.	1.	.	1.	.	.	.
TRACHEA	:	+	-	-	-	-	-	-	-	-
- Chronic Inflammation	:	1.
EPIDIDYMIS, LEFT	:	+	+	+	-	+	-	+	-	+
- Hypospermia	:	3.	4.	4.	.	3.	.	3.	.	3.
TESTIS, LEFT	:	+G	+G	+G	+G	+G	+	+G	+	+G
- Interstitial Cell Adenoma	:	P.	P.	P.	P.	P.	P.	.	P.	P.
- Tubular Atrophy	:	1.	.	1.	.	.
- Edema	:	.	.	.	1.	.	.	2.	.	.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	+G	,	,	,	,	,	,	,	,
- Squamous Cell Carcinoma	:	P.
PREPUTIAL GLANDS	:	0G	,	,	,	,	,	,	,	,
SYSTEMIC NEOPLASMS	:	,	+	,	,	,	+	,	,	,
- Malignant Lymphoma	:	NO.	.	NO.	.	NO.	.	NO.	.	NO.
LIVER	:	,	,	,	,	,	+G	,	,	+G
- Mononuclear Cell Leukemia	:	M45.	.	.	.
- Hepatodiaphragmatic Nodule	:	P.
SPLEEN	:	+G	+G	,	,	,	+G	,	,	,
- Mononuclear Cell Leukemia	:	.	M45.	.	.	.	M45.	.	.	.
- Extramedullary Hematopoesis	:	2.
TESTES	:	+G	+G	+G	+G	+G	,	+G	,	,
- Tubular Atrophy	:	(4.	.	.	P.	P.	P.	P.	P.	.
- Interstitial Cell Adenoma	:	.	.	.	P.	P.	P.	P.	P.	.
MAMMARY GLAND	:	,	,	,	,	,	,	,	,	+G
- Galactocoele	:	P.
- Granuloma	:	P.
KIDNEYS	:	+G	,	,	,	,	+G	,	,	,
- Nephropathy	:	2.	2.	.	.	.
LYMPH NODES	:	+G	,	,	,	,	,	,	,	,
- Pigment	:	2.
EYES	:	,	,	,	,	,	,	,	,	+G
- Panophthalmitis	:	4.
- Cornea Ulcer	:	(3.

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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 5, LA

ANIMAL NUMBER :

	558 MK0	559 MK0	560 MK0	561 MK0	562 MK0	563 MK0+	564 MK0	565 MK0	566 MK0	567 MK0+
HARDERIAN GLANDS	:	:	:	:	:	:	:	:	:	+G
- Inflammation.	(4.
GANGLION, GASS., RIGHT	:	:	:	:	:	:	:	:	:	+G
- Meningioma Malignant	P.
PITUITARY GLAND	:	:	:	:	:	:	:	:	:	+G
- Pars Distalis Adenoma.	P.

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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 5, LA

ANIMAL NUMBER :

	568 MK0+	569 MK0+	570 MK0	571 MK0	572 MK0	573 MK0	574 MK0+	575 MK0	576 MK0	577 MK0
LUNG, LEFT	:	+	+	+	+	+G	+	+	+	+
- Mononuclear Cell Leukemia	.	.	.	M45.	.	.	.	M45.	.	.
- Alveolar Epithelium Hyperplasia	3.
- Foreign Body	.	.	.	P.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	.	.	.	1.	1.	1.	2.	1.	1.	2.
- Interstitial Fibrosis	.	.	.	1.	1.	1.	1.	1.	1.	1.
- Pleura Fibrosis	1.	.	1.	1.	1.
TRACHEA	:	-	-	-	+	-	-	-	-	+
- Chronic Inflammation	1.	1.
EPIDIDYMIS, LEFT	:	+	+	-	+	-	+	-	+	-
- Hypospermia	.	4.	4.	.	4.	.	4.	.	4.	.
TESTIS, LEFT	:	+G	+G	+	+G	+	+G	+	+G	+
- Interstitial Cell Adenoma	.	P.	P.	.	P.	.	P.	P.	P.	.
- Tubular Atrophy	.	4.	.	1.	.	.	1.	.	.	1.
- Edema	.	.	.	2.	.	1.	.	.	.	1.
STERNUM	:	-	-	-	-	-	-	-	+G	-
- Blood Vessel Arteritis	3.	.
SKIN/SUBCUTIS	:	'	+G	'	'	'	'	'	'	'
- Fibroadenoma	.	.	P.
SYSTEMIC NEOPLASMS	:	+	'	,	,	,	,	,	,	,
- Malignant Lymphoma	.	.	N0.	N0.	.	.
LIVER	:	+G	+G	'	'	+G	'	+G	+G	'
- Bile Duct Hyperplasia	.	.	2.	.	1.	.	.	2.	.	.
- Mixed Cell Focus	.	.	P.
- Mononuclear Cell Leukemia	.	.	M45.	.	.	.	M45.	.	.	.
- Macrovesiculation	2.
- Hepatodiaphragmatic Nodule	.	.	P.
- Granuloma	.	.	.	2.	.	2.
SPLEEN	:	+G	+G	'	'	+G	'	+G	+G	'
- Mononuclear Cell Leukemia	.	M45.	.	.	.	M45.
- Extramedullary Hematopoiesis	.	.	2.	2.	2.	.	.	3.	.	.
LUNG	:	+G	'	,	,	,	,	,	+G	'
- Mononuclear Cell Leukemia	.	M45.	.	.	.	M45.	.	.	M45.	.
TESTES	:	'	'	'	'	+G	'	+G	'	'
- Interstitial Cell Adenoma	P.	.	P.	P.	P.	.
MAMMARY GLAND	:	'	'	'	'	'	'	+G	'	'
- Galactocole Hemorrhage	P.	.	.	.
KIDNEYS	:	+G	'	'	'	'	'	+G	'	'
- Nephropathy	3.	.	.
- Mononuclear Cell Leukemia	.	.	M45.
ADRENAL GLANDS	:	'	'	'	'	'	'	+G	'	'
- Cortex Adenoma	(P.	.	.
MESENTERY	:	'	'	'	'	'	'	+G	'	'
- Blood Vessel Arteritis	3.	.	.

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 5, LA

ANIMAL NUMBER :

	568 MK0+	569 MK0+	570 MK0	571 MK0	572 MK0	573 MK0	574 MK0+	575 MK0	576 MK0	577 MK0
LYMPH NODES	: +G	'	'	'	'	'	+G	+G	'	'
- Plasma Cell Hyperplasia	: .						.	2.		
- Mononuclear Cell Leukemia	:M45.						M45.	.		
SEMINAL VESICLES	:	'	'	'	'	'	'	G	'	'

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TEST ITEM : AMPHIBOLE ASBESTOS
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PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
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TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 5, LA

ANIMAL NUMBER :

	578 MK0+	579 MK0	580 MK0	581 MK0	582 MK0+	583 MK0	584 MK0	585 MK0	586 MK0	587 MK0
LUNG, LEFT	:	+	+	+	+	+G	+	+	+	+G
- Mononuclear Cell Leukemia	.	.	M45.	.	M45.	M45.
- Foreign Body	.	P.	P.	P.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	.	2.	1.	1.	2.	1.	1.	2.	1.	1.
- Interstitial Fibrosis	.	2.	1.	.	1.	2.	1.	1.	1.	.
- Alveolar Bronchiolar Adenoma	P.
- Pleura Fibrosis	.	.	.	1.	1.	.	.	1.	1.	.
TRACHEA	:	-	-	-	-	-	-	+	+	-
- Chronic Inflammation	1.	1.	.
EPIDIDYMIS, LEFT	:	-	+	-	-	-	-	+	+	+
- Hypospermia	.	.	4.	.	.	.	4.	3.	4.	4.
TESTIS, LEFT	:	+	+G	+	+G	+	+G	+G	+G	+G
- Interstitial Cell Adenoma	.	.	P.	.	P.	P.	P.	P.	P.	P.
- Tubular Atrophy	.	1.	.	.	1.
- Interstitial Cell Hyperplasia	.	.	.	3.
STERNUM	:	-	-	-	-	-	-	-	-	-
SKIN/SUBCUTIS	:	'	'	'	'	'	'	+G	'	'
- Keratoacanthoma	P.	.	.
SYSTEMIC NEOPLASMS	:	+	'	'	'	+	'	'	'	+
- Malignant Lymphoma	.	.	NO.	.	NO.	NO.
LIVER	:	+G	'	'	'	'	'	+G	'	+G
- Bile Duct Hyperplasia	1.	2.	1.
- Mixed Cell Focus	P.	.	.
- Mononuclear Cell Leukemia	.	.	M45.	M45.	.
- Hepatocellular Adenoma	P.	.	.
SPLEEN	:	+G	'	'	+G	+G	'	+G	'	+G
- Mononuclear Cell Leukemia	.	M45.	.	M45.	M45.	.
- Extramedullary Hematopoesis	.	.	.	2.
- Congestion	2.	.	.	.
LUNG	:	'	'	'	'	'	'	+G	'	+G
- Interstitial Fibrosis	3.	.	.	.
- Alveolar Epithelium Hyperplasia	3.	.	.
TESTES	:	'	+G	+G	'	'	'	+G	'	+G
- Tubular Atrophy	(5.	.	.
- Interstitial Cell Adenoma	.	.	P.	(P.	.	.	P.	.	P.	.
MAMMARY GLAND	:	'	'	'	'	'	'	'	'	+G
- Galactocole Hemorrhage	P.
KIDNEYS	:	+G	'	'	+G	'	'	+G	'	+G
- Nephropathy	.	.	.	3.	.	3.	3.	3.	3.	.
- Mononuclear Cell Leukemia	.	.	M45.
LYMPH NODES	:	+G	'	'	'	'	'	+G	'	+G
- Mononuclear Cell Leukemia	.	M45.
PALATE (ORAL MUCOSA)	:	'	'	'	'	'	'	'	'	+G
- Squamous Cell Carcinoma	P.	.	.

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TEST ITEM : AMPHIBOLE ASBESTOS	PATHOL. NO.: 90188 GAW
TEST SYSTEM : RAT, , 18 MO, INHALATION	DATE : 29-AUG-13
SPONSOR : THE HAMNER	PathData®System V6.2d2

TABLE OF INDIVIDUAL MICROSCOPIC FINDINGS (AOFT)
DOSE GROUP : 5, LA

ANIMAL NUMBER :

	588 MK0	589 MK0	590 MK0+	591 MK0	592 MK0	593 MK0	594 MK0	595 MK0+	596 MK0	597 MK0
LUNG, LEFT	:	+	+	+	+	+	+	+	+	+
- Alveolar Epithelium Hyperplasia	2.
- Intravascular Neutrophils	.	.	.	4.
- Foreign Body	.	P.	P.	.	P.	P.	P.	P.	P.	P.
- Alveolus Inflammation	.	1.	2.	.	1.	1.	2.	1.	1.	1.
- Interstitial Fibrosis	.	1.	1.	.	1.	1.	2.	1.	1.	1.
- Pleura Fibrosis	1.	.	1.	.	1.	1.
TRACHEA	:	-	+	-	+	-	-	+	-	+
- Chronic Inflammation	.	.	1.	.	1.	.	.	1.	.	1.
EPIDIDYMIS, LEFT	:	+	-	-	-	+	-	+	-	+
- Hypospermia	.	3.	.	.	4.	.	4.	.	3.	4.
TESTIS, LEFT	:	+G	+	+	+G	+G	+G	+	+G	+G
- Interstitial Cell Adenoma	.	P.	.	P.	P.	P.	.	P.	P.	P.
- Tubular Atrophy	.	.	1.	.	.	.	5.	.	.	.
- Edema	.	.	2.
STERNUM	:	-	-	-	-	-	-	-	-	-
LIVER	:	'	'	'	'	'	'	+G	'	+G
- Bile Duct Hyperplasia	2.	2.	2.	2.
- Macrovesiculation	2.	.	.
SPLEEN	:	'	'	'	'	'	'	+G	'	+G
- Extramedullary Hematopoiesis	2.	2.	2.	2.
LUNG	:	'	'	'	'	'	'	+G	'	'
- Alveolar Epithelium Hyperplasia	.	.	3.
- Alveolar Bronchiolar Adenoma	P.	.	.	.
TESTES	:	+G	'	'	'	'	'	+G	'	+G
- Interstitial Cell Adenoma	.	P.	(P.	P.	P.	P.
MAMMARY GLAND	:	'	'	'	'	'	'	+G	'	'
- Fibroadenoma	P.	.	.	.
KIDNEYS	:	'	'	'	'	+G	'	+G	'	+G
- Nephropathy	4.	4.	4.	4.	4.	4.
LYMPH NODES	:	'	'	'	'	'	+G	'	'	'
- Plasma Cell Hyperplasia	3.

MACRO/MICRO CORRELATION TABLE

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 1, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 137

TESTIS (LEFT)

- 01: Pale mass, 2.5 x 1.5 1.5 mm.

- Interstitial Cell Adenoma.

PREPUTIAL GLANDS (INGUINAL GLANDS)

- 01: Right preputial gland, enlarged, 1.2 x 0.9 x 0.7 cm, green,soft.

- Adenoma, unilateral.

TESTES

- 01: Right, small.

- Tubular Atrophy, unilateral, grade 2.

ANIMAL NO: 138

TESTIS (LEFT)

- 01: Multiple pale nodules, 2-4mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, multiple pale nodules, 2mm-4mm.

- Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT).

- 02: Right, pale mass, 8 x 5 x 4 mm.

- Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT).

ANIMAL NO: 139

TESTIS (LEFT)

- 01: Pale area, 4 x 4 mm.

- Interstitial Cell Adenoma.

ANIMAL NO: 140

SKIN/SUBCUTIS

- 01: Skin anterior to scrotum, ulceration, 14 x 10 mm.

- SEE UNDER: PREPUTIAL GLANDS (INGUINAL GLANDS) .

PREPUTIAL GLANDS (INGUINAL GLANDS)

Finding 01 in SKIN/SUBCUTIS

- Adenoma, unilateral.

SPLEEN

- 01: Spleen enlarged, 60 x 15 x 10 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS) .

ANIMAL NO: 142

TESTIS (LEFT)

- 01: Pale mass, 30 x 15 x 12 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right testis, pale mass, 20 x 10 x 10 mm.

- Interstitial Cell Adenoma, bilateral.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 1, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 143

TESTIS (LEFT)

- 01: Pale mass, 33 x 20 x 18 mm.
- TESTES
- 01: Right, pale mass, 3 x 2 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 145

TESTIS (LEFT)

- 01: Flacid.

LIVER

- 01: Mottled and rough pitted surface on dorsal side.

SPLEEN

- 01: Enlarged- 55 x 15 x 8 mm.

- Tubular Atrophy, grade 1.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

MAMMARY GLAND

- 01: Lactating.

- Secretion, grade 2.

KIDNEYS

- 01: Dark brown with granular pitted surface.

- Nephropathy, bilateral, grade 3.

PITUITARY GLAND

- 01: Mass- 10 x 7 x 5 mm. Black and red.

- Pars Distalis Adenoma.

ANIMAL NO: 146

TESTIS (LEFT)

- 01: Pale area, 2 x 2mm.

- Tubular Atrophy, grade 1.

SPLEEN

- 01: Dark area 2 mm x 1 mm.

- Pigment, grade 2.

KIDNEYS

- 01: Pale and slightly pitted.

- Nephropathy, bilateral, grade 3.

ANIMAL NO: 149

TESTIS (LEFT)

- 01: Pale mass, 30 x 15 x 10 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, small, 17 x 8 x 8 mm.

- Tubular Atrophy, unilateral, grade 4.

- 02: Right, pale area, 2 x 2 mm.

- Tubular Atrophy, unilateral, grade 4.

ANIMAL NO: 150

TESTIS (LEFT)

- 01: Small, 15 x 9 x 4 mm.

- Tubular Atrophy, grade 2.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Mottled, surface irregular.

- Pigment, grade 2.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 1, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

TESTES

- 01: Right, pale mass, 25 x 15 x 9 mm.

- Interstitial Cell Adenoma, unilateral.

ANIMAL NO: 151

TESTIS (LEFT)

- 01: Small, 16 x 8 x 4 mm.

TESTES

- 01: Right, pale mass, 30 x 17 x 15 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 152

EYES

- 01: Protruding right eye, enlarged and crusted over.

- 02: Right eye socket, pale firm area that extends to hard palate of mouth. .5 cm on roof of mouth visible.

HARDERIAN GLANDS

- 01: Right, discolored, pale to reddish areas.

PALATE (ORAL MUCOSA)

Finding 02 in EYES

- Cornea Ulcer, unilateral, grade 5.

- Squamous Cell Carcinoma (site of primary neoplasm:SYSTEMIC NEOPLASMS). SEE UNDER: PALATE (ORAL MUCOSA).

- Pigment, unilateral, grade 2.

- Squamous Cell Carcinoma (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 153

TESTIS (LEFT)

- 01: Pale area, 10 x 9 mm.

TESTES

- 01: Right, pale area, 15 x 11 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 154

LIVER

- 01: Appeared to be 3 x normal size.

TESTES

- 01: Right, flaccid, smaller than left.

MAMMARY GLAND

- 01: Mass, 8 x 12 x 20 mm, white, contained white soft material, found in right inguinal area.

- 02: prominent, lactating.

KIDNEYS

- 01: Appeared to be 3 x normal size.

ADRENAL GLANDS

- 01: appeared to be 3 x normal size.

PITUITARY GLAND

- 01: Mass, 4 x 5 x 7 mm, red, lobulated.

- Congestion, grade 3.

- Tubular Atrophy, bilateral, grade 1.

- Cyst.

- Secretion, grade 2.

- Nephropathy, bilateral, grade 4.

- Cortex Vacuolization, Vacuolization, focal, bilateral, grade 2.

- Pars Distalis Adenoma.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 1, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 155

SKIN/SUBCUTIS

- 01: Left, inguinal, area mass 15 x 10 10 mm. - Squamous Cell Carcinoma.

LIVER

- 01: Surface irregular and pale. - Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 48 x 10 x 7 mm. - Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 156

TESTIS (LEFT)

- 01: Pale, mass 20 x 15 x 10 mm. - Interstitial Cell Adenoma.

TESTES

- 01: Right testis, small, flaccid 18 x 10 x 7 mm. - Tubular Atrophy, bilateral, grade 2.

ANIMAL NO: 157

MAMMARY GLAND

- 01: Prominent, lactating. - Secretion, grade 3.

PITUITARY GLAND

- 01: Mass, 10 x 8 x 5 m, red. - Pars Distalis Adenoma.

ANIMAL NO: 158

TESTIS (LEFT)

- 01: Pale area 16 x 11 mm. - Interstitial Cell Adenoma.

ANIMAL NO: 160

TESTIS (LEFT)

- 01: Pale area 5 x 4 mm. - Interstitial Cell Adenoma.

TESTES

- 01: Right, pale, mass 20 x 10 x 10 mm. - Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 161

TESTIS (LEFT)

- 01: Pale mass 20 x 10 x 7 mm. - Interstitial Cell Adenoma.

TESTES

- 01: Right, pale mass, 23 x 15 x 7 mm. - Interstitial Cell Adenoma, bilateral.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 1, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 162

TESTIS (LEFT)

- 01: Pale mass, 25 x 12 x 12 mm.
TESTES
- 01: Right, pale mass, 25 x 15 x 11 mm.

- Interstitial Cell Adenoma.
- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 163

TESTIS (LEFT)

- 01: Pale mass, 25 x 14 x 13 mm.
TESTES
- 01: Right, pale mass, 15 x 10 x 9 mm.

- Interstitial Cell Adenoma.
- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 165

TESTIS (LEFT)

- 01: Left testis, pale, mass 20 x 10 x 9 mm.
PALATE (ORAL MUCOSA)
- 01: Pale area, 8 x 6 mm.

- Interstitial Cell Adenoma.
- Squamous Cell Papilloma.

ANIMAL NO: 166

TESTIS (LEFT)

- 01: Left, testis, pale mass 20 x 13 x 12 mm. - Interstitial Cell Adenoma.
SKIN/SUBCUTIS
- 01: Right, ventral scrotum, pale, ulcerated mass, 20 x 18 x 7 mm.

- Squamous Cell Carcinoma.

TESTES

- 01: Right, pale area, 15 x 10 mm. - Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 167

TESTIS (LEFT)

- 01: Left testis, pale mass 15 x 15 x 7 mm. - Tubular Atrophy, grade 2.

ANIMAL NO: 168

TESTIS (LEFT)

- 01: Pale area, 15 x 10 mm. - Interstitial Cell Adenoma.
TESTES
- 01: Right, pale area, 5 x 5 mm. - Interstitial Cell Adenoma, bilateral.

TEST ITEM : AMPHIBOLE ASBESTOS	PATHOL. NO.: 90188 GAW
TEST SYSTEM : RAT, , 18 MO, INHALATION	DATE : 29-AUG-13
SPONSOR : THE HAMNER	PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 1, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 169

TESTIS (LEFT)

- 01: Yellow and white focal discoloration.

- Interstitial Cell Adenoma.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

- 02: Appeared smaller than normal.

- No corresponding finding.

KIDNEYS

- 01: Bilateral, focal area, tan, 2 mm, midway between poles.

- Nephropathy, bilateral, grade 2.

BODY CAVITIES

- 01: Oral cavity, lesion, 4 x 8 x 17 mm, tan, rough surface, located beneath the tongue, left attached to lower jaw.

- SEE UNDER: SALIVARY GLANDS.

EYES

- 01: Right, white and opaque.

- Lens Mineralization, unilateral, grade 3.

SALIVARY GLANDS

Finding 01 in BODY CAVITIES

- Sublingual Adenocarcinoma, unilateral.

ANIMAL NO: 170

TESTIS (LEFT)

- 01: Firm, yellow, white and black focal areas, bilateral.

- Interstitial Cell Adenoma.

SKIN/SUBCUTIS

- 01: Left axillary region, mass, 23 x 19 x 12 mm, firm, white, smooth (interior and exterior).

- Fibroma.

LIVER

- 01: Pale.

- No corresponding finding.

LUNG

- 01: Pale.

- No corresponding finding.

ANIMAL NO: 171

TESTIS (LEFT)

- 01: Pale mass, 17 x 15 x 12 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, small, flaccid, 15 x 9 x 6 mm.

- Tubular Atrophy, unilateral, grade 4.

ANIMAL NO: 172

TESTIS (LEFT)

- 01: Pale area 9 x 7 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Pale mass, 20 x 15 x 11 mm.

- Interstitial Cell Adenoma, bilateral.

TEST ITEM : AMPHIBOLE ASBESTOS	PATHOL. NO.: 90188 GAW
TEST SYSTEM : RAT, , 18 MO, INHALATION	DATE : 29-AUG-13
SPONSOR : THE HAMNER	PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY	DOSE GROUP 1, MALE
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<u>NECROPSY OBSERVATION</u>	<u>CORRESPONDING MICROSCOPIC FINDING</u>
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ANIMAL NO: 173

TESTIS (LEFT)	- Interstitial Cell Adenoma.
- 01: Pale area 4 x 5 mm.	
SKIN/SUBCUTIS	- Fibroma.
- 01: Right, lateral thoracic, pale subcutaneal mass, 20 x 15 x 7mm. Cut surface firm and white.	
SPLEEN	- Extramedullary Hematopoiesis, grade 2.
- 01: Slightly glandular surface.	
TESTES	- Interstitial Cell Adenoma, bilateral.
- 01: Right testis, pale mass 23 x 14 11 mm.	

ANIMAL NO: 174

TESTIS (LEFT)	- Interstitial Cell Adenoma.
- 01: Pale mass 22 x 18 x 14 mm.	
LIVER	- Hepatocellular Adenoma.
- 01: Pale mass, left median 10 x 9 x 7 mm.	
SPLEEN	- Extramedullary Hematopoiesis, grade 2.
- 01: Surface irregular.	
TESTES	- Interstitial Cell Adenoma, bilateral.
- 01: Right testis, pale area 2 x 2 mm.	
KIDNEYS	- Nephropathy, bilateral, grade 3.
- 01: Slightly mottled.	
LYMPH NODES	- Plasma Cell Hyperplasia, unilateral, grade 2.
- 01: Adjacent pancreas, enlarged 5 x 3 x 2 mm.	

ANIMAL NO: 175

PREPUTIAL GLANDS (INGUINAL GLANDS)	
- 01: Ventral scrotum, ulcerated mass, 20 x 12 x 10 mm.	- Carcinoma, bilateral.

ANIMAL NO: 176

LIVER	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- 01: Red, with 1-2 mm raised, pale areas.	
SPLEEN	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- 01: Enlarged, 85 x 25 x 18 mm.	
LUNG	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- 01: 1-3 red focal areas throughout.	
KIDNEYS	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- 01: Black, bilateral.	

TEST ITEM : AMPHIBOLE ASBESTOS	PATHOL. NO.: 90188 GAW
TEST SYSTEM : RAT, , 18 MO, INHALATION	DATE : 29-AUG-13
SPONSOR : THE HAMNER	PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY	DOSE GROUP 1, MALE
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NECROPSY OBSERVATION		CORRESPONDING MICROSCOPIC FINDING
ANIMAL NO:	177	
TESTIS (LEFT)		
- 01: Flaccid, with red discoloration. Small.	- Tubular Atrophy, grade 4.	
LIVER		
- 01: Enlarged. Weighed 30.2 g, pale and mottled.	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).	
SPLEEN		
- 01: Enlarged and pale. 6.5 cm x 1.8 cm x 1.3 cm.	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).	
LUNG		
- 01: Pale and mottled.	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).	
LYMPH NODES		
- 01: Mandibular Lymph Node- Enlarged and red.	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).	
ANIMAL NO:	178	
TESTIS (LEFT)		
- 01: Pale mass 25 x 15 x 14 mm.	- Interstitial Cell Adenoma.	
TESTES		
- 01: Right, pale mass, 12 x 10 x 8 mm.	- Interstitial Cell Adenoma, bilateral.	
ANIMAL NO:	179	
TESTES		
- 01: Right, small, 18 x 10 x 7 mm.	- Tubular Atrophy, bilateral, grade 4.	
ANIMAL NO:	181	
TESTIS (LEFT)		
- 01: Pale mass, 25 x 15 x 14 mm.	- Interstitial Cell Adenoma.	
LIVER		
- 01: Slightly pale.	- Bile Duct Hyperplasia, grade 2.	
SPLEEN		
- 01: Enlarged, 45 x 12 x 7 mm.	- Extramedullary Hematopoiesis, grade 2.	
TESTES		
- 01: Right, pale mass, 25 x 15 x 14 mm.	- Interstitial Cell Adenoma, bilateral.	
KIDNEYS		
- 01: Granular surface.	- Nephropathy, bilateral, grade 3.	
ADRENAL GLANDS		
- 01: Right, dark.	- Phaeochromocytoma, Malignant, bilateral (malignant neoplasm).	
- 02: Left, enlarged, irregular, 10 x 7 x 7 mm.	- Phaeochromocytoma, Malignant, bilateral (malignant neoplasm).	
PARATHYROID GLAND		
- 01: Red foci 1mm.	- SEE UNDER: THYROID GLAND (BOTH LOBES).	
THYROID GLAND (BOTH LOBES)		
Finding 01 in PARATHYROID GLAND	- C Cell Adenoma, unilateral.	

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 1, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 182

TESTIS (LEFT)

- 01: Pale area, 3 x 4 mm.

TESTES

- 01: Right, pale mass, 15 x 10 x 10 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 183

TESTIS (LEFT)

- 01: Yellow and white focal discoloration; contained fluid.

- Tubular Atrophy, grade 2.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

MAMMARY GLAND

- 01: Lactating.

- Secretion, grade 2.

KIDNEYS

- 01: Bilateral, granular, pitted surface.

- Nephropathy, bilateral, grade 3.

LYMPH NODES

- 01: Mandibular LN, enlarged, 10 x 4 x 3 mm.

- Plasma Cell Hyperplasia, bilateral, grade 3.

EYES

- 01: Right, protruded, red.

- Cornea Ulcer, unilateral, grade 4.

GASSERIAN GANGLION (RIGHT)

- 01: Enlarged, 13 x 4 x 3 mm, appears to be pressing on right side of pituitary gland.

- Squamous Cell Carcinoma (site of primary neoplasm:PALATE (ORAL MUCOSA)).

PALATE (ORAL MUCOSA)

- 01: Right, lesion, 10 x 6 x 2 mm, firm.

- Squamous Cell Carcinoma (malignant neoplasm).

ANIMAL NO: 184

URINARY BLADDER

- 01: Bladder full, urinary bladder distended, ~2.5 cm.

- Distention, grade 3.

PENIS

- 01: Crusty whitish precipitate around prepuce.

- No corresponding finding.

BONE

- 01: Mass located inside right rib cage, attached to spinal column.

- Osteosarcoma (malignant neoplasm).

ANIMAL NO: 185

TESTIS (LEFT)

- 01: Pale mass, 20 x 15 x 14 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale area, 4 x 3 mm.

- Interstitial Cell Adenoma, bilateral.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 1, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

HEART

- 01: Pale area, 3 x 4 mm.

- Fibrosis, grade 2.

ANIMAL NO: 186

EPIDIDYMIS (LEFT)

- 01: Multiple pale nodules, 1 - 2 mm.

- Mesothelioma (malignant neoplasm).

TESTIS (LEFT)

- 01: Multiple pale nodules, surface, 1-2mm.

- Mesothelioma (site of primary neoplasm:
EPIDIDYMIS, LEFT).

- 02: Pale mass 15 x 8 x 7 mm.

- Mesothelioma (site of primary neoplasm:
EPIDIDYMIS, LEFT).

LIVER

- 01: Surface granular.

- Mononuclear Cell Leukemia (site of primary
neoplasm:EPIDIDYMIS, LEFT).

SPLEEN

- 01: Enlarged, 60 x 15 x 7 mm.

- Mononuclear Cell Leukemia (site of primary
neoplasm:EPIDIDYMIS, LEFT).

TESTES

- 01: Right, pale mass 28 x 15 x 11 mm.

- Interstitial Cell Adenoma, bilateral.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 249

TESTIS (LEFT)

- 01: Pale mass, 2 x 1.2 x 1.0 mm.

- Interstitial Cell Adenoma.

SKIN/SUBCUTIS

- 01: Scrotum ulceration, 1.8 x 1.2 1.2 cm.

- Keratoacanthoma.

KIDNEYS

- 01: Right, pale cyst, 3 x 3 x 3 mm.

- Nephropathy, bilateral, grade 2.

EYES

- 01: Left, opacity, 3 x 2 mm.

- Lens Cataract, unilateral, grade 2.

ANIMAL NO: 250

TESTIS (LEFT)

- 01: Small.

- Tubular Atrophy, grade 3.

SPLEEN

- 01: Surface granular, enlarged 40 x 10 x 7 mm.

- Extramedullary Hematopoiesis, grade 2.

TESTES

- 01: Right, pale mass, 25 x 15 x 14mm.

- Interstitial Cell Adenoma, unilateral.

LYMPH NODES

- 01: Mediastinal, prominent.

- Plasma Cell Hyperplasia, bilateral, grade 2.

ANIMAL NO: 251

TESTIS (LEFT)

- 01: Left, pale mass, 30 x 15 x 15 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale mass, 10 x 10 x 8 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 252

THYROID GLAND (BOTH LOBES)

- 01: Left, enlarged, 5 x 4 x 3 mm.

- Follicular Cell Adenoma, unilateral.

PITUITARY GLAND

- 01: mass, 10 x 6 x 4 mm, red and tan.

- Pars Distalis Adenoma.

ANIMAL NO: 253

TESTIS (LEFT)

- 01: Pale area, 7 x 7 x 5 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale mass, 30 x 17 x 12 mm.

- Interstitial Cell Adenoma, bilateral.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 254

LIVER

- 01: Pale with lobes rounded.
- SPLEEN
- 01: Enlarged, slight 32 x 10 x 7 mm.

- Bile Duct Hyperplasia, grade 2.

- Extramedullary Hematopoiesis, grade 2.

ANIMAL NO: 255

TESTIS (LEFT)

- 01: Pale area, 3 x 3 mm.

- Interstitial Cell Adenoma.

SKIN/SUBCUTIS

- 01: Base of tail, ulcerated area, 18 x 15 x 8 mm.

- Fibrosarcoma.

LIVER

- 01: Left lobe nodule 3 x 3 mm.

- Focal Necrosis, grade 3.

SPLEEN

- 01: Enlarged 40 x 10 x 6 mm.

- Extramedullary Hematopoiesis, grade 3.

TESTES

- 01: Pale area, 4 x 4 mm.

- Interstitial Cell Adenoma, bilateral.

KIDNEYS

- 01: Granular and pale.

- Nephropathy, bilateral, grade 3.

LYMPH NODES

- 01: Mediastinal, prominent.

- Plasma Cell Hyperplasia, bilateral, grade 2.

ANIMAL NO: 256

TESTIS (LEFT)

- 01: Black and white discoloration, focal; (near rete end).

- Interstitial Cell Adenoma.

SKIN/SUBCUTIS

- 01: Mass, below left axillary, 47 x 38 x 18 mm, firm, lobulated, pale tan.

- SEE UNDER: MAMMARY GLAND.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

MAMMARY GLAND

Finding 01 in SKIN/SUBCUTIS

- Fibroadenoma.

LYMPH NODES

- 01: Mandibular, left, enlarged, 8 x 4 x 3 mm.

- Plasma Cell Hyperplasia, unilateral, grade 2.

- 02: Mandibular, right enlarged, 10 x 8 x 4 mm, cystic.

- Plasma Cell Hyperplasia, unilateral, grade 2.

EYES

- 01: Right, red, bulged out of socket.

- Cornea Ulcer, ulcer, unilateral, grade 4.

HARDERIAN GLANDS

- 01: Right, enlarged, 8 x 5 x 3 mm, pale areas.

- Fibrosis, unilateral, grade 2.

GASSERIAN GANGLION (RIGHT)

- 01: Right, enlarged, 14 x 4 x 2 mm.

- Squamous Cell Carcinoma.

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

PITUITARY GLAND

- 01: Focus, white, raised, 2 mm dia, right side.

- Pars Distalis Adenoma.

PALATE (ORAL MUCOSA)

- 01: distorted area, smooth, 15 x 5 x 1 mm.

- Squamous Cell Carcinoma (malignant neoplasm).

ANIMAL NO: 257

TESTIS (LEFT)

- 01: Pale mass, 25 x 13 x 10 mm.

- Interstitial Cell Adenoma.

SPLEEN

- 01: Enlarged, 48 x 13 x 7 mm.

- Lymphoid Depletion, grade 2.

TESTES

- 01: Pale mass, 25 x 16 x 11 mm.

- Interstitial Cell Adenoma, bilateral.

KIDNEYS

- 01: Pale, granular.

- Nephropathy, bilateral, grade 4.

ANIMAL NO: 259

LIVER

- 01: Left median lobe, pale, irregular surfaces.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 45 x 15 x 7 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 260

TESTIS (LEFT)

- 01: Pale mass, 28 x 12 x 10 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale area, 4 x 5 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 262

TESTIS (LEFT)

- 01: Pale mass, 25 x 15 x 12 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale area, 4 x 4 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 264

TESTIS (LEFT)

- 01: Pale mass, 25 x 12 x 10 mm.

- Interstitial Cell Adenoma.

SPLEEN

- 01: Enlarged, 50 x 12 x 7 mm.

- Extramedullary Hematopoiesis, grade 2.

TESTES

- 01: Right, small, 20 x 10 x 8 mm.

- Interstitial Cell Adenoma, bilateral.

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

KIDNEYS

- 01: Slightly pitted.

- Nephropathy, bilateral, grade 3.

ANIMAL NO: 265

TESTIS (LEFT)

- 01: Yellow, white and black focal discoloration, firm.

- Interstitial Cell Adenoma.

LIVER

- 01: Irregular, pale areas throughout.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Enlarged, 50 x 14 x 7 mm.

- Congestion, grade 3.

PALATE (ORAL MUCOSA)

- 01: Left side, has a raised firm area, 7 x 6 x 2 mm, extends up into the left eye socket.

- Squamous Cell Carcinoma (malignant neoplasm).

INTESTINE-SMALL, ILEUM

- 01: Yellowish in color.

- No corresponding finding.

INTESTINE-SMALL, JEJUNUM

- 01: Yellowish in color.

- No corresponding finding.

INTESTINE-SMALL, DUODENUM

- 01: Yellowish in color.

- No corresponding finding.

ANIMAL NO: 266

TESTIS (LEFT)

- 01: Pale mass, 25 x 15 x 11 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale area, 4 x 5 x mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 267

TESTIS (LEFT)

- 01: Pale mass, 34 x 15 x 12 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale area, 7 x 5 x 4 mm.

- Interstitial Cell Adenoma, bilateral.

EYES

- 01: Left, opacity, 3 x 4 mm.

- Lens Cataract, unilateral, grade 3.

ANIMAL NO: 268

TESTIS (LEFT)

- 01: Pale mass, 20 x 11 x 10 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Pale mass 25 x 14 x 10 mm.

- Interstitial Cell Adenoma, bilateral.

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 269

TESTIS (LEFT)

- 01: Pale mass, 20 x 12 x 10 mm.
- TESTES
- 01: Right, pale mass, 22 x 15 x 14 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 270

TESTIS (LEFT)

- 01: Pale mass, 15 x 15 x 10 mm.
- TESTES
- 01: Right, pale areas, few, 2-4mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 271

SPLEEN

- 01: Enlarged, 43 x 12 x 5 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 272

TESTIS (LEFT)

- 01: Fluid filled. White, yellow and black discoloration, focal.

- Interstitial Cell Adenoma.

LIVER

- 01: Mottled.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Hepatocellular Adenoma.

- 02: Left median lobe, raised area, 3 x 2 x 1 mm dia.
- 03: Caudate lobe, tan focal area, 2 mm dia.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 75 x 22 x 15 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

LUNG

- 01: Pale, 1-2 mm red foci throughout.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

KIDNEYS

- 01: Dark.

- Nephropathy, bilateral, grade 2.

BODY CAVITIES

- 01: Thoracic cavity, red fluid, small amount.
- 02: Abdominal cavity, red fluid, small amount.

- No corresponding finding.

- No corresponding finding.

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 273

LUNG, LEFT

- 01: Red.
- 02: Cranial lobe (ventral), focal area, tan, 3 mm dia.

- Alveolar Epithelium Hyperplasia, grade 3.
- Alveolar Bronchiolar Adenoma.

TESTIS (LEFT)

- 01: Black and white discoloration, focal; filled with fluid.

- Interstitial Cell Adenoma.

LIVER

- 01: Red.
- 02: Right posterior lobe, tan area, 3 mm dia.
- 03: Caudate lobes, 3 x 4 (? units), tan areas.

- No corresponding finding.
- Bile Duct Hyperplasia, grade 3.
- Eosinophilic Focus.

ANIMAL NO: 274

TESTIS (LEFT)

- 01: Pale area, 10 x 10 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Mottled edges, rounded.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged 75 x 22 x 20 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 275

TESTIS (LEFT)

- 01: 20 x 13 x 11 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Slightly pale.

- Focal Necrosis, grade 2.

SPLEEN

- 01: Slightly irregular.

- Extramedullary Hematopoiesis, grade 2.

ANIMAL NO: 276

TESTIS (LEFT)

- 01: Flaccid, small 14 x 6 x 3 mm.

- Tubular Atrophy, grade 4.

TESTES

- 01: Right, pale mass, 24 x 15 x 12 mm.

- Interstitial Cell Adenoma, unilateral.

MAMMARY GLAND

- 01: Prominent.

- Secretion, grade 3.

PITUITARY GLAND

- 01: Enlarged, 4 x 3 x 3 mm, red.

- Pars Distalis Adenoma.

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 277

TESTIS (LEFT)

- 01: Pale mass, 15 x 15 x 10 mm.
- TESTES
- 01: Small, flaccid 14 x 9 x 7 mm.

- Interstitial Cell Adenoma.

- Tubular Atrophy, unilateral, grade 4.

ANIMAL NO: 278

TESTIS (LEFT)

- 01: Pale mass 25 x 15 x 10 mm.
- TESTES
- 01: Pale mass, 20 x 12 x 10 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 279

TESTES

- 01: Small, 15 x 8 x 7 mm.
- EYES
- 01: Left, Opacity.

- Tubular Atrophy, bilateral, grade 2.

- Lens Cataract, unilateral, grade 2.

ANIMAL NO: 280

TESTIS (LEFT)

- 01: Pale area, 15 x 10mm.

- Interstitial Cell Adenoma.

ANIMAL NO: 281

SKIN/SUBCUTIS

- 01: Right, lateral mass, 10 x 9 x 5 cm. Cut surface firm, pale tissue.

- Fat Necrosis.

LIVER

- 01: Slightly pale.

- Bile Duct Hyperplasia, grade 1.

SPLEEN

- 01: Enlarged, 50 x 12 x 5 mm.

- Extramedullary Hematopoiesis, grade 3.

ANIMAL NO: 282

TESTIS (LEFT)

- 01: Pale mass, 25 x 15 x 12 mm.
- TESTES
- 01: Pale area, 9 x 7 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 283

TESTIS (LEFT)

- 01: Pale mass, 25 x 16 x 14 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Slightly enlarged, 42 x 10 x 7 mm.

- Extramedullary Hematopoiesis, grade 3.

TESTES

- 01: Small, 16 x 7 x 7 mm.

- Tubular Atrophy, unilateral, grade 4.

EYES

- 01: Opacity.

- Cornea Inflammation, unilateral, grade 2.

ANIMAL NO: 284

TESTIS (LEFT)

- 01: Pale mass, 25 x 14 x 15 mm.

- Interstitial Cell Adenoma.

ANIMAL NO: 285

TESTIS (LEFT)

- 01: Pale mass, 25 x 18 x 11 mm.

- Interstitial Cell Adenoma.

SPLEEN

- 01: Slightly granular.

- Extramedullary Hematopoiesis, grade 2.

TESTES

- 01: Pale mass, 22 x 17 x 12 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 286

TESTIS (LEFT)

- 01: Pale mass, 25 x 15 x 11 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Granular, pale.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 95 x 20 x 10 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTES

- 01: Small, 15 x 9 x 9 mm.

- Tubular Atrophy, unilateral, grade 4.

ANIMAL NO: 287

TESTIS (LEFT)

- 01: Pale mass, 25 x 14 x 13 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Slightly rounded edges.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Slightly enlarged, 45 x 11 x 7 mm.

- Extramedullary Hematopoiesis, grade 3.

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

TESTES

- 01: Small, 15 x 10 x 7 mm.

- Tubular Atrophy, unilateral, grade 4.

ANIMAL NO: 288

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Dark.

- Extramedullary Hematopoiesis, grade 3.

EYES

- 01: Left eye socket, tan lesion, 6 x 4 x 3 mm. Cranial portion of lesion (located next to Harderian gland) was soft. The caudal end (next to molars teeth) was hard.

- SEE UNDER: PALATE (ORAL MUCOSA).

PALATE (ORAL MUCOSA)

Finding 01 in EYES

- Squamous Cell Carcinoma (malignant neoplasm).

ANIMAL NO: 289

TESTIS (LEFT)

- 01: Pale mass, 10 x 9 x 8 mm.

- Interstitial Cell Adenoma.

ANIMAL NO: 290

TESTIS (LEFT)

- 01: Pale area, 10 x 10 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale mass, 20 x 15 x 11 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 291

PREPUTIAL GLANDS (INGUINAL GLANDS)

- 01: Scrotum mass, 20 x 10 x 8 mm.

- Adenoma, bilateral.

LIVER

- 01: Pale.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 77 x 20 x 11 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 292

TESTIS (LEFT)

- 01: Yellow and white discoloration, focal.

- Interstitial Cell Adenoma.

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

PREPUTIAL GLANDS (INGUINAL GLANDS)

- 01: Lesion in area of right preputial gland, 15 x 13 x 5 mm, a portion of lesion was hollow, red and green material inside.

- Inflammation Suppurative, bilateral, grade 3.

LIVER

- 01: Pale.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 60 x 16 x 6 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

KIDNEYS

- 01: Discolored, dark brown, mottled.

- Nephropathy, bilateral, grade 2.

ANIMAL NO: 293

TESTIS (LEFT)

- 01: Pale mass, 20 x 9 x 9 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Mottled, surface granular.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Enlarged, 58 x 11 x 10 mm.

- Extramedullary Hematopoiesis, grade 3.

TESTES

- 01: Right, small, 10 x 6 x 6 mm.

- Tubular Atrophy, unilateral, grade 4.

KIDNEYS

- 01: Slightly pitted.

- Nephropathy, bilateral, grade 3.

LYMPH NODES

- 01: Mandibular LN, prominent.

- Plasma Cell Hyperplasia, bilateral, grade 3.

ANIMAL NO: 294

TESTIS (LEFT)

- 01: Pale area, 3 x 3 mm.

- Interstitial Cell Adenoma.

- 02: Small, 20 x 9 x 9 mm.

- No corresponding finding.

SKIN/SUBCUTIS

- 01: Ventral abdomen, red nodule, two, 2 x 2 x 2 mm each.

- SEE UNDER: MAMMARY GLAND.

LIVER

- 01: Nodule, 7 x 6 x 5 mm.

- Bile Duct Hyperplasia, grade 2.

TESTES

- 01: Right, small, 17 x 9 x 7 mm.

- Tubular Atrophy, unilateral, grade 4.

MAMMARY GLAND

Finding 01 in SKIN/SUBCUTIS

- Galactocoele.

KIDNEYS

- 01: Pale, mottled.

- Nephropathy, bilateral, grade 3.

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 2, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 295

LUNG, LEFT

- 01: Pale.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTIS (LEFT)

- 01: Black, yellow and white, focal, discoloration.

- Interstitial Cell Adenoma.

LIVER

- 01: Pitted, granular surface.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 24 x 75 x 22 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

LUNG

- 01: Pale.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTES

- 01: Right, flaccid.

- No corresponding finding.

KIDNEYS

- 01: Black, bilateral.

- Nephropathy, bilateral, grade 2.

ANIMAL NO: 296

TESTIS (LEFT)

- 01: Pale mass, 27 x 15 x 14 mm.

- Interstitial Cell Adenoma.

ANIMAL NO: 297

LUNG, LEFT

- 01: Pale, irregular white areas throughout.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTIS (LEFT)

- 01: Yellow and white discoloration, focal, bilateral.

- Interstitial Cell Adenoma.

SPLEEN

- 01: Enlarged, 70 x 20 x 20 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

KIDNEYS

- 01: Black, granular.

- Nephropathy, bilateral, grade 3.

ANIMAL NO: 298

TESTIS (LEFT)

- 01: Pale mass, 20 x 15 x 14 mm.

- Interstitial Cell Adenoma.

TEST ITEM : AMPHIBOLE ASBESTOS	PATHOL. NO.: 90188 GAW
TEST SYSTEM : RAT, , 18 MO, INHALATION	DATE : 29-AUG-13
SPONSOR : THE HAMNER	PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 319

TESTIS (LEFT)

- 01: Flaccid, with yellow and white, focal discoloration.

- Interstitial Cell Adenoma.

PREPUTIAL GLANDS (INGUINAL GLANDS)

- 01: Mass, 27 x 13 x 10 mm, bilateral. Mass is filled with soft, tan, caseous material.

- Carcinoma, bilateral.

SPLEEN

- 01: Enlarged; 60 x 17 x 8 mm, red.

- Extramedullary Hematopoiesis, grade 4.

LUNG

- 01: Pale.

- No corresponding finding.

MESENTERY

Finding 01 in BODY CAVITIES

BODY CAVITIES

- 01: Mass in Abdominal Cavity, 100 x 67 x 38 mm. Firm to soft; cystic. Contained red fluid. Mass is tan, red, yellow.

- Schwannoma.

- SEE UNDER: MESENTERY.

ANIMAL NO: 321

KIDNEYS

- 01: Brown; Bilateral.

- Nephropathy, bilateral, grade 3.

PITUITARY GLAND

- 01: Mass; 12 x 7 x 5 mm, black.

- Pars Distalis Adenoma.

ANIMAL NO: 349

TESTIS (LEFT)

- 01: Flaccid.

- Tubular Atrophy, grade 2.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

HARDERIAN GLANDS

- 01: Right, dark area, 2 mm dia.

- Inflammation, focal, unilateral, grade 1.

THYROID GLAND (BOTH LOBES)

- 01: Right, mass, 7 x 4 x 3 mm, firm, red.

- Follicular Cell Carcinoma, unilateral.

PALATE (ORAL MUCOSA)

- 01: Dark red area, 5 mm dia, in center of hard palate.

- No corresponding finding.

- 02: Dark red area, 5 mm dia, on the tongue.

- No corresponding finding.

INTESTINE-LARGE, COLON

- 01: Hard area adhered to duodenum, 30 mm dia.

- Adhesion.

ANIMAL NO: 350

TESTIS (LEFT)

- 01: Pale mass, 20 x 17 x 10 mm.

- Interstitial Cell Adenoma.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

TESTES

- 01: Right, small.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 351

TESTIS (LEFT)

- 01: pale mass, 10 x 10 x 8 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale area, 3 x 4 x 3 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 352

TESTIS (LEFT)

- 01: Pale area, 22 x 12 x 7 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Edges rounded, slightly granular.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 60 x 12 x 10 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTES

- 01: Right, small and flaccid.

- Tubular Atrophy, bilateral, grade 4.

ANIMAL NO: 353

TESTES

- 01: Right, 12 x 10 x 7 mm, mass.

- Interstitial Cell Adenoma, unilateral.

ANIMAL NO: 355

TESTIS (LEFT)

- 01: Yellow discoloration area, 3 x 4 mm.

- Interstitial Cell Hyperplasia, grade 3.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

EYES

- 01: Right eye, crusted over with blood.

- Cornea Ulcer, unilateral, grade 4.

- 02: Right eye socket, some pale soft tissue remained.

- Squamous Cell Carcinoma (site of primary neoplasm:PALATE (ORAL MUCOSA)).

HARDERIAN GLANDS

- 01: Pale, smooth, firm, 9 x 5 x 6 mm with dark 2 mm near center.

- Squamous Cell Carcinoma (site of primary neoplasm:PALATE (ORAL MUCOSA)).

GASSERIAN GANGLION (RIGHT)

- 01: Enlarged, two times size of left.

- Squamous Cell Carcinoma (site of primary neoplasm:PALATE (ORAL MUCOSA)).

PITUITARY GLAND

- 01: Small, pale, >1 mm black focus in center.

- Pars Distalis Hyperplasia Focal, Hyperplasia, focal, grade 3.

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

PALATE (ORAL MUCOSA)

- 01: There is a dark discoloration on right hard palate in area of rear molar teeth. That area appears distorted.

- Squamous Cell Carcinoma (malignant neoplasm).

STOMACH

- 01: Contents black and bloody.
- INTESTINE-LARGE, CECUM
- 01: Black contents.
- INTESTINE-SMALL, ILEUM
- 01: Black contents.

- No corresponding finding.

- No corresponding finding.

- No corresponding finding.

ANIMAL NO: 356

TESTIS (LEFT)

- 01: Pale mass, 30 x 16 x 14 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale mass, 27 x 16 x 15 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 357

EPIDIDYMIS (LEFT)

- 01: Brown nodules, 1-4 mm.

- Mesothelioma (site of primary neoplasm: SYSTEMIC NEOPLASMS).

TESTIS (LEFT)

- 01: Brown nodules, 1-4 mm.

- Mesothelioma (site of primary neoplasm: SYSTEMIC NEOPLASMS).

KIDNEYS

- 01: Black, bilateral, 1-4 mm.

- Nephropathy, bilateral, grade 2.

ANIMAL NO: 358

TESTIS (LEFT)

- 01: Pale mass, 25 x 15 x 11 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Pale mass, 25 x 15 x 10 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 359

TESTES

- 01: Right, flaccid.

- No corresponding finding.

EYES

- 01: Left, bulging, red, crusted over.

- Cornea Ulcer, ulcer, unilateral, grade 4.

GASSERIAN GANGLION (LEFT)

- 01: Enlarged, 5 x 10 x 3 mm.

- Squamous Cell Carcinoma (site of primary neoplasm:PALATE (ORAL MUCOSA)).

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 361

TESTIS (LEFT)

- 01: Pale mass, 23 x 13 x 7 mm.
- LIVER
- 01: Pale foci, 1 mm dia, few.
- SPLEEN
- 01: Enlarged, 42 x 10 x 7 mm.
- TESTES
- 01: Right, pale area, 4 x 4 mm.
- PARATHYROID GLAND
- 01: Left parathyroid, prominent.
- LYMPH NODES
- 01: Mediastinal LN, dark.

- Interstitial Cell Adenoma.
- Macrovesiculation, grade 2.
- Extramedullary Hematopoiesis, grade 2.
- Interstitial Cell Adenoma, bilateral.
- No corresponding finding.
- Pigment, bilateral, grade 2.

ANIMAL NO: 362

TESTIS (LEFT)

- 01: Pale area 10 x 7 mm.
- TESTES
- 01: Right, pale area, 10 x 6 mm.

- Interstitial Cell Adenoma.
- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 363

EPIDIDYMIS (LEFT)

- 01: Pale nodules, 1-3 mm.
- TESTIS (LEFT)
- 01: Many pale nodules, 1-3mm.
- 02: Pale mass, 14 x 15 x 9 mm.
- LIVER
- 01: Pale.
- SPLEEN
- 01: Slightly mottled.
- TESTES
- 01: Right, fluid filled.
- 02: Right, many nodules, 1-3 mm.
- MESENTERY
- 01: Pale nodules, 1-2 mm.
- LYMPH NODES
- 01: Mesentery LN, dark red.
- BODY CAVITIES
- 01: Abdominal wall, multiple pale nodules, 1-3 mm.

- No corresponding finding.
- Mesothelioma (site of primary neoplasm: EPIDIDYMIDES).
- Interstitial Cell Adenoma.
- Mononuclear Cell Leukemia (site of primary neoplasm:EPIDIDYMIDES) .
- Extramedullary Hematopoiesis, grade 3.
- Mesothelioma (site of primary neoplasm: EPIDIDYMIDES) .

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

EPIDIDYMIDES

- 01: Right, pale nodules, 1-3mm.

CORRESPONDING MICROSCOPIC FINDING

- Mesothelioma, bilateral (malignant neoplasm).

ANIMAL NO: 365

TESTIS (LEFT)

- 01: Pale mass, 30 x 14 x 13 mm.

LIVER

- 01: Granular surface.

SPLEEN

- 01: Enlarged, 66 x 18 x 8 mm.

KIDNEYS

- 01: Slightly pitted.

- Interstitial Cell Adenoma.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Nephropathy, bilateral, grade 3.

ANIMAL NO: 366

PREPUTIAL GLANDS (INGUINAL GLANDS)

- 01: Left inguinal area, ulcerated mass, 17 x 14 x 7 mm.

- Carcinoma, unilateral.

TESTES

- 01: Right, pale mass, 14 x 15 x 8 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 367

GENERAL OBSERVATIONS

- 01: All fat, skin, organs have yellow tint.

- Nothing abnormal discovered.

LUNG, LEFT

- 01: Red, irregular pale areas throughout.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTIS (LEFT)

- 01: Flaccid.

- Tubular Atrophy, grade 3.

- 02: White and yellow discoloration, focal.

- Interstitial Cell Adenoma.

LIVER

- 01: Mottled.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- 02: Raised nodules throughout, 1-3 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: 95 x 26 x 20 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

LUNG

- 01: Red, irregular pale areas throughout.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTES

- 01: White and yellow discoloration, focal.

- No corresponding finding.

Flaccid.

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

KIDNEYS

- 01: Black.

CORRESPONDING MICROSCOPIC FINDING

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 368

TESTIS (LEFT)

- 01: Pale mass, 20 x 12 x 10 mm.

- Interstitial Cell Adenoma.

SKIN/SUBCUTIS

- 01: Left, inguinal area mass, 20 x 10 x 7 mm.

- SEE UNDER: MAMMARY GLAND.

LIVER

- 01: Granular, pale.

- Macrovesiculation, grade 2.

SPLEEN

- 01: Irregular surface.

- Extramedullary Hematopoiesis, grade 2.

TESTES

- 01: Right, small, 12 x 7 x 4 mm, flaccid.

- Tubular Atrophy, unilateral, grade 4.

MAMMARY GLAND

- 01: Mammary tissue prominent.
Finding 01 in SKIN/SUBCUTIS

- Fibroadenoma.

- Adenocarcinoma.

ANIMAL NO: 369

TESTES

- 01: Right, pale mass, 30 x 15 x 12 mm.

- Interstitial Cell Adenoma, unilateral.

ANIMAL NO: 370

LYMPH NODES

- 01: Mediastinal, prominent.

- No corresponding finding.

ANIMAL NO: 371

LUNG, LEFT

- 01: Red, abnormal consistency.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- 02: Appeared firm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

LIVER

- 01: Red, mottled.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 3 times.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

LUNG

- 01: Red.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- 02: Abnormal consistency, appeared firm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

LYMPH NODES

- 01: Mesenteric lymph node, enlarged, 2 times.
- 02: Thymic lymph node, enlarged, red.

HARDERIAN GLANDS

- 01: Dark.

STOMACH

- 01: Abnormal contents, dark, filled with air.

INTESTINE-LARGE, CECUM

- 01: Abnormal contents, dark.

INTESTINE-SMALL, ILEUM

- 01: Abnormal content, dark.

BRAIN

- 01: Dark foci, less than a pin-point in size, frontal cerebral cortex (ventral).
- 02: Dorsal side cerebral, dark area, 3 x 4 mm.

PANCREAS

- 01: Abnormal color, dark.
- 02: Very small.

CORRESPONDING MICROSCOPIC FINDING

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- No corresponding finding.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- No corresponding finding.

- Ulceration, grade 4.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 372

LUNG, LEFT

- 01: Red foci, 2mm.

TESTIS (LEFT)

- 01: Pale area, 6 x 4 mm.

TESTES

- 01: Right, pale area, 8 x 8 mm.

- Alveolar Epithelium Hyperplasia, grade 1.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 373

TESTIS (LEFT)

- 01: Flaccid.
- 02: White discoloration, focal.

LYMPH NODES

- 01: Mandibular, enlarged, 7 x 6 x 3 mm.

EYES

- 01: Right, bulging out of socket.
- 02: Socket has some paleness.

HARDERIAN GLANDS

- 01: Right, mass, 12 x 8 x 4 (units), pale, firm.

- Tubular Atrophy, grade 2.

- Interstitial Cell Adenoma.

- Plasma Cell Hyperplasia, bilateral, grade 3.

- Squamous Cell Carcinoma (site of primary neoplasm:PALATE (ORAL MUCOSA)).

- Squamous Cell Carcinoma (site of primary neoplasm:PALATE (ORAL MUCOSA)).

- Squamous Cell Carcinoma (site of primary neoplasm:PALATE (ORAL MUCOSA)).

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

GASSERIAN GANGLION (RIGHT)

- 01: Two times size of left.

PALATE (ORAL MUCOSA)

- 01: Right, concave lesion, 12 x 5 mm,
center is red, firm.

ANIMAL NO: 374

TESTIS (LEFT)

- 01: Yellow and black focal discoloration.

LIVER

- 01: Caudate lobe, mottled, two pale areas,
approximately 7 x 5 mm and 3 x 4 mm,
appear firm, extend down into the lobe.

SPLEEN

- 01: Enlarged, 75 x 25 x 14 mm.
- 02: Tear.

CORRESPONDING MICROSCOPIC FINDING

- Squamous Cell Carcinoma (site of primary neoplasm:PALATE (ORAL MUCOSA)).

- Squamous Cell Carcinoma (malignant neoplasm).

- No corresponding finding.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- No corresponding finding.

ANIMAL NO: 375

TESTIS (LEFT)

- 01: Pale mass, 30 x 15 x 14 mm.

TESTES

- 01: Small, 15 x 9 x 4 mm.

- Interstitial Cell Adenoma.

- Tubular Atrophy, unilateral, grade 4.

ANIMAL NO: 376

TESTIS (LEFT)

- 01: pale mass, 15 x 11 x 10 mm.

- Interstitial Cell Adenoma.

ANIMAL NO: 377

TESTIS (LEFT)

- 01: Pale mass, 20 x 15 x 12 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale area, 6 x 5 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 378

TESTIS (LEFT)

- 01: 25 x 14 x 12 mm.

- Interstitial Cell Adenoma.

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CORRELATION TABLE: NECROPSY - MICROSCOPY	DOSE GROUP 3, MALE
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<u>NECROPSY OBSERVATION</u>	<u>CORRESPONDING MICROSCOPIC FINDING</u>
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ANIMAL NO: 379

TESTIS (LEFT)

- 01: Yellow, black and white discoloration, focal.

TESTES

- 01: Right, flaccid.

EYES

- 01: Left, bulging, red, crusted over.

PALATE (ORAL MUCOSA)

- 01: Left, lesion, 10 x 7 x 2 mm.

GASSERIAN GANGLION (LEFT)

- 01: Enlarged, 12 x 4 x 3 mm.

- Interstitial Cell Adenoma.

- No corresponding finding.

- Panophthalmitis, bilateral, grade 4.

- Squamous Cell Carcinoma (malignant neoplasm).

- Squamous Cell Carcinoma (site of primary neoplasm:PALATE (ORAL MUCOSA)).

ANIMAL NO: 380

TESTIS (LEFT)

- 01: Pale mass, 30 x 15 x 14 mm.

TESTES

- 01: Small, 15 x 10 x 6 mm.

- Interstitial Cell Adenoma.

- Tubular Atrophy, unilateral, grade 2.

ANIMAL NO: 381

EYES

- 01: Left, opacity.

- Lens Cataract, unilateral, grade 2.

ANIMAL NO: 382

SKIN/SUBCUTIS

- 01: Mass, 30 x 20 x 10, soft, pale. Located upper Lt. abdominal area beneath skin.

- Sarcoma.

SPLEEN

- 01: Enlarged, 56 x 13 x 10 mm.

- Autolytic

TESTES

- 01: Shrunken to flatness.

- No corresponding finding.

- Tubular Atrophy, bilateral, grade 4.

ANIMAL NO: 383

EPIDIDYMIS (LEFT)

- 01: Brown nodules attached.

- Mesothelioma (malignant neoplasm).

TESTIS (LEFT)

- 01: Fluid filled.

- Edema, grade 3.

- 02: Yellow and white discoloration, focal.

- Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT).

SPLEEN

- 01: Enlarged.

- Extramedullary Hematopoesis, grade 3.

TEST ITEM : AMPHIBOLE ASBESTOS	PATHOL. NO.: 90188 GAW
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CORRELATION TABLE: NECROPSY - MICROSCOPY	DOSE GROUP 3, MALE
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NECROPSY OBSERVATION		CORRESPONDING MICROSCOPIC FINDING
LUNG	- 01: Pale.	- No corresponding finding.
TESTES	- 01: Fluid filled. - 02: white and yellow focal discolorations.	- No corresponding finding. - No corresponding finding.
KIDNEYS	- 01: Black.	- Tubule Pigment, bilateral, grade 3.
BODY CAVITIES	- 01: Abdominal cavity, fluid filled, clear, dark brown fluid. - 02: Abdominal cavity, muscle wall, brown raised nodules, 1-3 mm. - 03: Peritoneal cavity, fluid filled, clear, dark brown fluid. - 04: Peritoneal cavity, muscle wall, brown raised nodules, 1-3 mm.	- No corresponding finding. - Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT). - Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT). - Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT).
PITUITARY GLAND	- 01: Mass, 2 x 3 x 4 mm, black.	- Pars Distalis Adenoma.

ANIMAL NO: 384

EPIDIDYMIS (LEFT)	- 01: Fat, pale nodule, 5 x 5 mm.	- Blood Vessels Arteritis, grade 2.
TESTIS (LEFT)	- 01: Pale mass, 20 x 15 x 12 mm.	- Interstitial Cell Adenoma.
PREPUTIAL GLANDS (INGUINAL GLANDS)	- 01: Ventral scrotal, area mass, 15 x 11 x 6 mm.	- Adenoma, unilateral.
TESTES	- 01: Right, small, flaccid, 15 x 7 x 4 mm.	- Tubular Atrophy, unilateral, grade 4.

ANIMAL NO: 385

TESTIS (LEFT)	- 01: Pale mass, 20 x 15 x 11 mm.	- Interstitial Cell Adenoma.
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ANIMAL NO: 386

TESTIS (LEFT)	- 01: Small 15 x 10 x 5 mm.	- Tubular Atrophy, grade 2.
LIVER	- 01: Mottled, pale, rounded edges.	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
SPLEEN	- 01: Dark, slightly enlarged 50 x 20 x 10 mm.	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
LUNG	- 01: Right, congested.	- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
TESTES	- 01: Right, small, 15 x 10 x 5 mm.	- Tubular Atrophy, bilateral, grade 2.

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

MAMMARY GLAND

- 01: Left side, secretion.
- KIDNEYS
- 01: Dark, pitted.

- Secretion, grade 2.
- Nephropathy, bilateral, grade 3.

ANIMAL NO: 387

TESTIS (LEFT)

- 01: Flaccid, bilateral.

- Edema, grade 1.

KIDNEYS

- 01: Dark brown, pitted surface, bilateral.

- Nephropathy, bilateral, grade 3.

PITUITARY GLAND

- 01: Mass, 8 x 5 x 3 mm, black.

- Pars Distalis Adenoma.

ANIMAL NO: 388

TESTIS (LEFT)

- 01: Pale mass, 10 x 9 x 7 mm.

- Interstitial Cell Adenoma.

SKIN/SUBCUTIS

- 01: Right, ventral scrotum mass, 20 x 20 x 15 mm.

- Epidermal Cyst.

ANIMAL NO: 389

TESTIS (LEFT)

- 01: Pale, mass 15 x 13 x 9 mm.

- Interstitial Cell Adenoma.

SKIN/SUBCUTIS

- 01: Right forelimb, pale mass, 25 x 14 x 10 mm.

- Keratoacanthoma.

TESTES

- 01: Right, small, flaccid, 11 x 7 x 3 mm.

- Tubular Atrophy, unilateral, grade 3.

ANIMAL NO: 390

LIVER

- 01: Small, pale.

- Bile Duct Hyperplasia, grade 1.

LUNG

- 01: Right cranial lobe, has 2 mm white focus, dorsal side close to bifurcation.

- Osteosarcoma (site of primary neoplasm: EYES).

LYMPH NODES

- 01: Mandibular lymph node, right, enlarged.

- Plasma Cell Hyperplasia, bilateral, grade 2.

- 02: Mesenteric lymph node, enlarged.

- Plasma Cell Hyperplasia, bilateral, grade 2.

EYES

- 01: Right Eye, 10 x 5 mm mass, tan, extending down into hard palate.

- Socket Osteosarcoma, unilateral (malignant neoplasm).

STOMACH

- 01: Stomach contained bedding and dark fluid.

- No corresponding finding.

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

INTESTINE-LARGE, CECUM
- 01: Contained dark black contents.
INTESTINE-SMALL, ILEUM
- 01: Contained dark black contents.
INTESTINE-SMALL, JEJUNUM
- 01: Contained dark black contents.
INTESTINE-LARGE, COLON
- 01: Contained dark black contents.

CORRESPONDING MICROSCOPIC FINDING

- No corresponding finding.
- No corresponding finding.
- No corresponding finding.
- No corresponding finding.

ANIMAL NO: 391

TESTIS (LEFT)

- 01: Dehydrated and flaccid.

- Tubular Atrophy, grade 2.

LIVER

- 01: Pale.

- Macrovesiculation, grade 2.

SPLEEN

- 01: Small.

- Atrophy, grade 3.

TESTES

- 01: Dehydrated and flaccid.

- No corresponding finding.

KIDNEYS

- 01: Pale with granular pitted surface.

- Nephropathy, bilateral, grade 4.

ADRENAL GLANDS

- 01: Enlarged.

- Nothing abnormal discovered.

PITUITARY GLAND

- 01: Mass, black, 5 x 6 x 3 mm.

- Pars Distalis Adenoma.

- 02: Indention in brain from pituitary mass.

- Pars Distalis Adenoma.

STOMACH

- 01: Glandular, contained a small amount of black substance.

- No corresponding finding.

- 02: Non-glandular, surface raised, erosions.

- Ulceration, grade 3.

ANIMAL NO: 392

TESTIS (LEFT)

- 01: Pale area, 12 x 10 x 7 mm.

- Interstitial Cell Adenoma.

ANIMAL NO: 393

TESTIS (LEFT)

- 01: Pale mass, 20 x 13 x 12 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Enlarged, 40 x 10 x 7 mm.

- Extramedullary Hematopoiesis, grade 2.

TESTES

- 01: Right, small, 15 x 10 x 4 mm.

- Tubular Atrophy, unilateral, grade 3.

KIDNEYS

- 01: Slightly pitted.

- Nephropathy, bilateral, grade 3.

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SUMMARY TABLES

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 3, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 394

TESTIS (LEFT)

- 01: Pale mass, 15 x 12 x 9 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Slightly enlarged, 37 x 10 x 9 mm.

- Extramedullary Hematopoiesis, grade 2.

TESTES

- 01: Right, pale mass, 25 x 15 x 12 mm.

- Interstitial Cell Adenoma, bilateral.

MAMMARY GLAND

- 01: Thorax, subcutis, red, nodule, 10 x 5 x 4 mm.

- Galactocole Hemorrhage.

ANIMAL NO: 395

TESTIS (LEFT)

- 01: Pale mass, 9 x 9 x 7 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale mass, 20 x 14 x 10 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 397

TESTIS (LEFT)

- 01: Yellow and white discoloration, focal.

- Interstitial Cell Adenoma.

SKIN/SUBCUTIS

- 01: Mass, 4 x 6 x 10 (units), firm,
epidermis.

- Epidermal Cyst.

LIVER

- 01: Enlarged.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- 02: Granular, pitted surface. Margins of lobes are uneven.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 15 x 30 x 90 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

KIDNEYS

- 01: Black.

- Nephropathy, bilateral, grade 3.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 4, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 419

TESTIS (LEFT)

- 01: Yellow and white focal discoloration.
- PREPUTIAL GLANDS (INGUINAL GLANDS)
- 01: Mass is 30 x 30 x 15 mm. Round and firm with a black crusted surface. Inside cuts-tan with some green exudate inside.

- Interstitial Cell Adenoma.

LUNG

- 01: Pale.

- No corresponding finding.

TESTES

- 01: Dark area.

- No corresponding finding.

KIDNEYS

- 01: Granular, pitted surface.

- Nephropathy, bilateral, grade 3.

ANIMAL NO: 420

TESTIS (LEFT)

- 01: Soft with white and yellow focal discoloration.

- Tubular Atrophy, grade 1.

PREPUTIAL GLANDS (INGUINAL GLANDS)

- 01: Right, mass, 25 x 18 x 10 mm. Tan; raw and exposed. Mass is firm with hard area raised above the skin (white exudate inside when sliced open).

- Carcinoma, unilateral.

LIVER

- 01: Pale, friable, and appeared smaller than normal.

- Eosinophilic Focus.

SPLEEN

- 01: Slightly enlarged, 40 x 11 x 7 mm.

- Extramedullary Hematopoesis, grade 3.

TESTES

- 01: Right, Flaccid.

- No corresponding finding.

ANIMAL NO: 423

TESTIS (LEFT)

- 01: Pale mass, 20 x 11 x 10 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Mottled, granular surface.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 88 x 25 x 15 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTES

- 01: Right, small, 13 x 6 x 5 mm.

- Tubular Atrophy, unilateral, grade 4.

MAMMARY GLAND

- 01: Right, ascilla subcutaneous mass, 40 x 25 x 10 mm.

- Fibroadenoma.

KIDNEYS

- 01: Dark.

- Nephropathy, bilateral, grade 3.

TEST ITEM : AMPHIBOLE ASBESTOS	PATHOL. NO.: 90188 GAW
TEST SYSTEM : RAT, , 18 MO, INHALATION	DATE : 29-AUG-13
SPONSOR : THE HAMNER	PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY	DOSE GROUP 4, MALE
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NECROPSY OBSERVATION	CORRESPONDING MICROSCOPIC FINDING
LYMPH NODES	
- 01: Mediastinal LN, prominent.	- Plasma Cell Hyperplasia, bilateral, grade 2.
ANIMAL NO: 449	
TESTIS (LEFT)	
- 01: Pale mass, 2.0 x 1.4 x 1.3 cm.	- Interstitial Cell Adenoma.
TESTES	
- 01: Right, pale mass, 3.0 x 1.5 x 1.5 cm.	- Interstitial Cell Adenoma, bilateral.
ANIMAL NO: 450	
TESTES	
- 01: Right, enlarged, pale mass, 23 x 15 x 12 mm.	- Interstitial Cell Adenoma, bilateral.
ANIMAL NO: 451	
EPIDIDYMIS (LEFT)	
- 01: Nodules, tan, 1-2 mm.	- Mesothelioma (malignant neoplasm).
TESTIS (LEFT)	
- 01: Firm, yellow, white and black discoloration, focal.	- Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT).
- 02: Tunical vaginalis, nodules, tan, 1-2 mm.	- Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT).
LIVER	
- 01: Dark brown.	- Mononuclear Cell Leukemia.
- 02: Dark red, focal areas throughout, 1-2 mm.	- Mononuclear Cell Leukemia.
- 03: Raised areas, tan, 2-3 mm.	- Mononuclear Cell Leukemia.
- 04: Diaphragmatic hernia, two, 8 x 4 x 2 mm, and 5 x 3 x 2 mm.	- Hepatodiaphragmatic Nodule.
SPLEEN	
- 01: Enlarged, 87 x 25 x 15 mm.	- Mononuclear Cell Leukemia (site of primary neoplasm:EPIDIDYMIS, LEFT).
LUNG	
- 01: Red.	- Mononuclear Cell Leukemia (site of primary neoplasm:EPIDIDYMIS, LEFT).
- 02: Cranial lobe, focus, dark red, 2 mm.	- Mononuclear Cell Leukemia (site of primary neoplasm:EPIDIDYMIS, LEFT).
TESTES	
- 01: Firm, focal, yellow, white and black discolorations.	- No corresponding finding.
KIDNEYS	
- 01: Black.	- Nephropathy, bilateral, grade 2.
ADRENAL GLANDS	
- 01: Enlarged, red.	- Mononuclear Cell Leukemia (site of primary neoplasm:EPIDIDYMIS, LEFT).

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 4, MALE

NECROPSY OBSERVATION

MESENTERY

- 01: Fat, brown granules.

BODY CAVITIES

- 01: Thoracic cavity, red fluid filled.

URINARY BLADDER

- 01: Distended, with brown fluid.

ADIPOSE TISSUE

- 01: Scrotal fat, nodules, tan, 1-2 mm.

CORRESPONDING MICROSCOPIC FINDING

- Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT).

- Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT).

- No corresponding finding.

- Mesothelioma (site of primary neoplasm: EPIDIDYMIS, LEFT).

ANIMAL NO: 452

LUNG, LEFT

- 01: Pale area, 9 x 5mm.

LIVER

- 01: Pale, rounded edges to lobes.

SPLEEN

- 01: Enlarged, 44 x 12 x 9 mm.

TESTES

- 01: Right, pale area, 12 x 10 x 10 mm.

KIDNEYS

- 01: Surface pitted, pale.

LYMPH NODES

- 01: Mediastinal, prominent.

- 02: Mesentery, prominent.

SALIVARY GLANDS

- 01: Right, dark area, 3 x 3mm.

- Alveolar Bronchiolar Adenoma.

- Macrovesiculation, grade 1.

- Extramedullary Hematopoiesis, grade 2.

- Interstitial Cell Adenoma, bilateral.

- Nephropathy, bilateral, grade 3.

- No corresponding finding.

- Plasma Cell Hyperplasia, bilateral, grade 2.

- No corresponding finding.

ANIMAL NO: 453

TESTIS (LEFT)

- 01: Pale area, 12 x 9 x 7 mm.

- Edema, grade 2.

ANIMAL NO: 455

TESTIS (LEFT)

- 01: Pale mass, 25 x 14 x 12 mm.

PREPUTIAL GLANDS (INGUINAL GLANDS)

- 01: Right, inguinal area mass, 20 x 13 x 7 mm.

TESTES

- 01: Right, pale mass, 12 x 11 x 10 mm.

- Interstitial Cell Adenoma.

- Adenoma, bilateral.

- Interstitial Cell Adenoma, unilateral.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 4, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 456

TESTIS (LEFT)

- 01: Enlarged, pale mass, 28 x 15 x 12 mm. - Interstitial Cell Adenoma.
- SPLEEN
- 01: Enlarged, 45 x 10 x 7 mm. - Extramedullary Hematopoiesis, grade 2.
- TESTES
- 01: Right, small and flaccid, 14 x 7 x 3 mm. - Tubular Atrophy, unilateral, grade 5.

ANIMAL NO: 457

TESTIS (LEFT)

- 01: Mass, 25 x 15 x 10 mm. - Interstitial Cell Adenoma.
- SKIN/SUBCUTIS
- 01: Right, lateral abdomen mass 45 x 40 x 20 mm. - Sarcoma.
- TESTES
- 01: Right, dark area, 4 x 3 mm. - Interstitial Cell Hyperplasia, unilateral, grade 2.

ANIMAL NO: 458

SKIN/SUBCUTIS

- 01: Left inguinal area, mass, 25 x 25 x 10 mm, fluid filled. - Epidermal Cyst.
- LIVER
- 01: Granular surface, pale. - Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- SPLEEN
- 01: Enlarged, 83 x 22 x 8 mm. - Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 459

TESTIS (LEFT)

- 01: Pale mass, 11 x 10 x 7 mm. - Interstitial Cell Adenoma.
- LIVER
- 01: Granular, pale. - Bile Duct Hyperplasia, grade 2.
- SPLEEN
- 01: Enlarged, 46 x 11 x 6 mm. - Extramedullary Hematopoiesis, grade 2.
- TESTES
- 01: Right, mass, pale, 20 x 15 x 10 mm. - Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 460

TESTIS (LEFT)

- 01: Pale mass, 20 x 10 x 10 mm. - Interstitial Cell Adenoma.

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 4, MALE

NECROPSY OBSERVATION

SKIN/SUBCUTIS

- 01: Nodule, 7 x 3 x 2mm, neck, below ear, right.

LIVER

- 01: Slightly pale.

SPLEEN

- 01: Enlarged, 60 x 15 x 10 mm.

TESTES

- 01: Right, pale mass, 25 x 15 x 10 mm.

KIDNEYS

- 01: Pitted, slight.

ADRENAL GLANDS

- 01: Pale 1mm foci, few.

CORRESPONDING MICROSCOPIC FINDING

- Basal Cell Adenoma.

- Bile Duct Hyperplasia, grade 2.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Interstitial Cell Adenoma, bilateral.

- Nephropathy, bilateral, grade 3.

- Medulla Phaeochromocytoma,
Phaeochromocytoma, unilateral.

ANIMAL NO: 461

TESTIS (LEFT)

- 01: Pale mass, 25 x 15 x 12 mm.

TESTES

- 01: Right, pale mass, 15 x 8 x 7 mm.

KIDNEYS

- 01: Pitted.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

- Nephropathy, bilateral, grade 3.

ANIMAL NO: 462

TESTES

- 01: Right, white and yellow discolorations.

MAMMARY GLAND

- 01: Prominent, lactating.

KIDNEYS

- 01: Granular, pitted surface.

PITUITARY GLAND

- 01: Mass, 8 x 6 x 6 mm, black.

- No corresponding finding.

- Secretion, grade 2.

- Nephropathy, bilateral, grade 3.

- Pars Distalis Adenoma.

ANIMAL NO: 464

TESTIS (LEFT)

- 01: Pale mass, 27 x 15 x 11 mm.

LIVER

- 01: Pale edges, slightly rounded.

SPLEEN

- 01: Slightly irregular.

TESTES

- 01: Right, small, 17 x 10 x 6 mm.

KIDNEYS

- 01: Slightly pitted.

- Interstitial Cell Adenoma.

- Bile Duct Hyperplasia, grade 2.

- Extramedullary Hematopoiesis, grade 2.

- Tubular Atrophy, unilateral, grade 3.

- Nephropathy, bilateral, grade 3.

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TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 4, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 466

SKIN/SUBCUTIS

- 01: Right, inguinal area, pale mass, 15 x 11 x 9 mm.

- Squamous Cell Carcinoma.

LIVER

- 01: Mottled surface, irregular.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 68 x 17 x 10 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 467

TESTIS (LEFT)

- 01: Pale mass, 25 x 14 x 12 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale mass, 20 x 14 x 12 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 468

TESTIS (LEFT)

- 01: pale mass, 15 x 12 x 12 mm.

- Interstitial Cell Adenoma.

ANIMAL NO: 469

KIDNEYS

- 01: Dark brown.

- Nephropathy, bilateral, grade 1.

- 02: Left, mass, 30 x 25 x 18 mm.

- SEE UNDER: ADRENAL GLANDS.

ADRENAL GLANDS

Finding 02 in KIDNEYS

- Cortical Carcinoma, unilateral.

ANIMAL NO: 470

THYROID GLAND (BOTH LOBES)

- 01: Left, mass 10 x 7 x 5 mm. Firm, appears to be lobulated. There is also a (10 x 7 x 4 mm) similar mass adjacent to Thyroid mass.

- Follicular Cell Carcinoma, unilateral (malignant neoplasm).

ANIMAL NO: 471

TESTIS (LEFT)

- 01: Filled with fluid; firm yellow lesions.

- Interstitial Cell Adenoma.

LIVER

- 01: Pale and friable, some adherence of nodules.

- Mesothelioma (site of primary neoplasm: TESTIS, LEFT).

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 4, MALE

NECROPSY OBSERVATION

KIDNEYS

- 01: Black.

BODY CAVITIES

- 01: Animal's abdominal cavity was filled with brown fluid and all muscle wall, mesentery, fat, pancreas, scrotal sac, epididymides were covered with tan nodules (pin point to 3 mm in diameter), including abdominal side of diaphragm.

CORRESPONDING MICROSCOPIC FINDING

- Nephropathy, bilateral, grade 2.

- Mesothelioma (site of primary neoplasm: TESTIS, LEFT).

ANIMAL NO: 472

TESTIS (LEFT)

- 01: Two pale areas, 4 x 5 mm each.

- Interstitial Cell Adenoma.

TESTES

- 01: Pale mass, 30 x 17 x 14 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 473

TESTIS (LEFT)

- 01: White, linear discoloration, 4 x 2 mm; slightly flaccid.

- Edema, grade 1.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Slightly enlarged, 50 x 11 x 6 mm.

- Extramedullary Hematopoiesis, grade 3.

TESTES

- 01: Slightly flaccid.

- No corresponding finding.

MAMMARY GLAND

- 01: Lactating.

- Secretion, grade 2.

KIDNEYS

- 01: Both- dark, irregular pale areas throughout.

- Nephropathy, bilateral, grade 2.

LYMPH NODES

- 01: Mandibular Lymph Node, appears to be cystic.

- Cyst, bilateral.

- 02: Cervical Lymph Node, right, enlarged, 7 x 5 x 3 mm and red.

- Plasma Cell Hyperplasia, bilateral, grade 3.

EYES

- 01: Right, bulging out of socket and red.

- Cornea Ulcer, bilateral, grade 4.

GASSERIAN GANGLION (RIGHT)

- 01: Enlarged (twice the size of the left).

- Squamous Cell Carcinoma (site of primary neoplasm: PALATE (ORAL MUCOSA)).

PITUITARY GLAND

- 01: Mass, 2 x 2 x 4 (unit), dark raised area.

- Pars Distalis Adenoma.

PALATE (ORAL MUCOSA)

- 01: Right, lesion, 9 x 4 mm, concave, and reddish.

- Squamous Cell Carcinoma (malignant neoplasm).

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 4, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 474

TESTIS (LEFT)

- 01: Pale mass, 20 x 15 x 10 mm.

- Interstitial Cell Adenoma.

ANIMAL NO: 475

LIVER

- 01: Pale.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Slightly mottled, 50 x 14 x 7 mm
(enlarged).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 477

TESTIS (LEFT)

- 01: Pale mass, 28 x 14 x 13 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Small, 15 x 8 x 7 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 480

TESTIS (LEFT)

- 01: Pale mass, 20 x 15 x 12 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale area, 10 x 10 mm.

- Interstitial Cell Adenoma, bilateral.

LYMPH NODES

- 01: Mandibular, prominent.

- Plasma Cell Hyperplasia, bilateral, grade 2.

ANIMAL NO: 482

TESTIS (LEFT)

- 01: Pale area, 5 x 5 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale mass, 20 x 12 x 10 mm.

- Interstitial Cell Adenoma, bilateral.

EYES

- 01: Left, opacity.

- Phthisis Bulbi, unilateral.

ANIMAL NO: 483

TESTIS (LEFT)

- 01: Pale mass, 30 x 17 x 14 mm.

- Interstitial Cell Adenoma.

SKIN/SUBCUTIS

- 01: Right side of head, skin, ulcerated mass, 18 x 15 x 11 mm.

- SEE UNDER: SALIVARY GLANDS.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 4, MALE

NECROPSY OBSERVATION

SPLEEN

- 01: Ventral, pale mass, 6 x 5 x 4 mm.

TESTES

- 01: Right, pale area 2 x 2 mm.

LYMPH NODES

- 01: Right, mandibular, enlarged, 5 x 5 mm.

SALIVARY GLANDS

Finding 01 in SKIN/SUBCUTIS

CORRESPONDING MICROSCOPIC FINDING

- Stromal Hyperplasia, grade 3.

- Interstitial Cell Adenoma, bilateral.

- No corresponding finding.

- Sublingual Adenocarcinoma, unilateral.

ANIMAL NO: 484

LIVER

- 01: Pale, granular surface.

SPLEEN

- 01: Enlarged, 45 x 9 x 6 mm.

TESTES

- 01: Right, pale mass, 20 x 10 x 7 mm.

- Bile Duct Hyperplasia, grade 2.

- Extramedullary Hematopoiesis, grade 3.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 485

TESTIS (LEFT)

- 01: pale mass, 25 x 14 x 15 mm.

TESTES

- 01: Right, pale area, 14 x 10 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 486

LIVER

- 01: Left, lobe, pale area, 5 x 4 mm.

TESTES

- 01: Right, pale area, 7 x 5 mm.

- Bile Duct Hyperplasia, grade 2.

- Interstitial Cell Adenoma, unilateral.

ANIMAL NO: 487

TESTIS (LEFT)

- 01: Pale mass, 18 x 10 x 7 mm.

TESTES

- 01: Right, small, 14 x 7 x 3 mm.

LYMPH NODES

- 01: Mediastinal LN, prominent.

- Interstitial Cell Adenoma.

- Tubular Atrophy, unilateral, grade 4.

- No corresponding finding.

ANIMAL NO: 488

TESTIS (LEFT)

- 01: Pale, 25 x 12 x 11 mm.

SKIN/SUBCUTIS

- 01: Right, subcutaneous tissue, red nodule, 7 x 4 x 3 mm.

- Interstitial Cell Adenoma.

- SEE UNDER: MAMMARY GLAND.

TEST ITEM : AMPHIBOLE ASBESTOS
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 4, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

TESTES

- 01: Right, amall, 16 x 9 x 7 mm.

MAMMARY GLAND

Finding 01 in SKIN/SUBCUTIS

LYMPH NODES

- 01: Mandibular LN, prominent.

- Tubular Atrophy, focal, bilateral, grade 4.

- Galactocole Hemorrhage.

- Plasma Cell Hyperplasia, bilateral, grade 2.

ANIMAL NO: 489

TESTIS (LEFT)

- 01: Pale mass, 25 x 15 x 12 mm.

LUNG

- 01: Congested.

TESTES

- 01: Right, small, 17 x 9 x 7 mm.

- Interstitial Cell Adenoma.

- Congestion, grade 2.

- Tubular Atrophy, unilateral, grade 2.

ANIMAL NO: 490

TESTIS (LEFT)

- 01: Pale mass, 25 x 15 x 10 mm.

LIVER

- 01: Slightly pale.

SPLEEN

- 01: Slightly enlarged, 43 x 10 x 6 mm.

TESTES

- 01: Right, pale mass, 20 x 15 x 9 mm.

- Interstitial Cell Adenoma.

- Bile Duct Hyperplasia, grade 2.

- Extramedullary Hematopoesis, grade 2.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 491

TESTIS (LEFT)

- 01: Pale mass, 28 x 15 x 14 mm.

TESTES

- 01: Right, pale area, 12 x 10 x 10 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 493

TESTIS (LEFT)

- 01: Pale mass, 25 x 15 x 15 mm.

TESTES

- 01: Right, pale mass, 22 x 14 x 12 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 520

LUNG, LEFT

- 01: Several (1-2)mm, red foci throughout.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTIS (LEFT)

- 01: Yellow, white, and black discoloration.

- Interstitial Cell Adenoma.

- 02: Some fluid.

- Interstitial Cell Adenoma.

PREPUTIAL GLANDS (INGUINAL GLANDS)

- 01: Left, mass, 30 x 20 x 10 mm, firm, tan. Crusted over (skin side); part of mass appeared to be necrotic.

- Adenoma, bilateral.

LIVER

- 01: Mottled.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 75 x 20 x 13 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

LUNG

- 01: Caudal lobe (dorsal cranial area) has (3mm dia) tan area.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

KIDNEYS

- 01: Dark.

- Nephropathy, bilateral, grade 2.

BODY CAVITIES

- 01: Thoracic cavity contained some red fluid; More fluid.

- No corresponding finding.

- 02: Adbominal cavity contained some red fluid.

- No corresponding finding.

PITUITARY GLAND

- 01: Black area, 1 x 2 x 1 mm.

- Pars Distalis Hyperplasia Focal, Hyperplasia, focal, grade 3.

ANIMAL NO: 549

TESTIS (LEFT)

- 01: White and yellow discoloration, focal.

- Interstitial Cell Adenoma.

LIVER

- 01: Irregular pale areas throughout.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Small.

- Lymphoid Depletion, grade 2.

LUNG

- 01: Nodules, 1-3 mm dia.

- Alveolar Bronchiolar Carcinoma (malignant neoplasm).

- 02: Mass (smooth flat side) nodules, 1-3 mm dia.

- Cystic Keratinizing Epithelioma.

- 03: Lung pluck, adhesion to spine.

- No corresponding finding.

TESTES

- 01: Small, flaccid.

- No corresponding finding.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

KIDNEYS

- 01: Granular surfaces.
- 02: Left, mass, 3 mm dia., raised, white.
- 03: Right, renal pelvis, dilated.
- 04: Right, mass, 15 x 13 x 5 mm, firm, white.

LYMPH NODES

- 01: Mesenteric LN, enlarged and red.

BODY CAVITIES

- 01: Thoracic cavity, blood inside, large clot below heart.
- 02: Thoracic cavity, lined with nodules, 1-3 mm dia.

BONE

- 01: Rib, nodules 1-3 mm dia.
- 02: Sternum, 1-3 mm dia.

SEMINAL VESICLES

- 01: Small, with irregular shaped edges.

PITUITARY GLAND

- 01: 2 x 3 mm red area.

INTESTINE-LARGE, COLON

- 01: Nodule, tan, 3 x 4 x 2 (units).

HEART

- 01: Nodules, 1-3 mm dia.

SKELETAL MUSCLE, DIAPHRAGM

- 01: Nodules, 1-3 mm dia.

CORRESPONDING MICROSCOPIC FINDING

- Alveolar Bronchiolar Carcinoma (site of primary neoplasm:LUNG).

- Sinus Dilation, bilateral, grade 2.

- No corresponding finding.

- Alveolar Bronchiolar Carcinoma (site of primary neoplasm:LUNG).

- Alveolar Bronchiolar Carcinoma (site of primary neoplasm:LUNG).
- Alveolar Bronchiolar Carcinoma (site of primary neoplasm:LUNG).

- No corresponding finding.

- Pars Distalis Cyst, Cyst.

- Muscularis Focal Thickening.

- Alveolar Bronchiolar Carcinoma (site of primary neoplasm:LUNG).

- Alveolar Bronchiolar Carcinoma (site of primary neoplasm:LUNG).

ANIMAL NO: 551

TESTIS (LEFT)

- 01: Pale mass, 28 x 15 x 13 mm.

SPLEEN

- 01: Enlarged, 50 x 15 x 7 mm.

TESTES

- 01: Right, pale mass, 23 x 18 x 15 mm.

- Interstitial Cell Adenoma.

- Extramedullary Hematopoiesis, grade 2.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 552

TESTIS (LEFT)

- 01: Flaccid.

PREPUTIAL GLANDS (INGUINAL GLANDS)

- 01: Right, mass, 18 x 14 x 6 mm, firm surface, soft inside, green exudate.

- No corresponding finding.

- Carcinoma, unilateral.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

LIVER

- 01: Mottled.

SPLEEN

- 01: Enlarged, 55 x 17 x 12 mm.

LUNG

- 01: Red, with black focal areas throughout.

MAMMARY GLAND

- 01: Lactating.

KIDNEYS

- 01: Granular, pitted surface.

PITUITARY GLAND

- 01: Mass, 6 x 5 x 3 mm, black.

ANIMAL NO: 553

TESTIS (LEFT)

- 01: Pale mass, 20 x 15 x 10 mm.

TESTES

- 01: Right, small.

ANIMAL NO: 554

TESTIS (LEFT)

- 01: Small, flaccid.

LIVER

- 01: Pale.

SPLEEN

- 01: Enlarged, 45 x 10 x 7 mm.

TESTES

- 01: Multiple pale nodules, 2-3mm.

- 02: Pale area 20 x 15 x 11 mm.

MESENTERY

- 01: Multiple pale nodules, 2-3mm, throughout.

LYMPH NODES

- 01: Thymic region, dark nodules, 2-4mm.

BODY CAVITIES

- 01: Abdominal wall, pale nodules, 2-3mm.

EPIDIDYMIDES

- 01: Right, multiple pale nodules, 2-3mm.

CORRESPONDING MICROSCOPIC FINDING

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Secretion, grade 2.

- Nephropathy, bilateral, grade 4.

- Pars Distalis Adenoma.

- Interstitial Cell Adenoma.

- Tubular Atrophy, unilateral, grade 3.

- Interstitial Cell Adenoma.

- Bile Duct Hyperplasia, grade 2.

- Extramedullary Hematopoesis, grade 3.

- Mesothelioma (site of primary neoplasm: EPIDIDYMIDES).

- Mesothelioma (site of primary neoplasm: EPIDIDYMIDES).

- Mesothelioma (site of primary neoplasm: EPIDIDYMIDES).

- Pigment, bilateral, grade 3.

- Mesothelioma (site of primary neoplasm: EPIDIDYMIDES).

- Mesothelioma, unilateral (malignant neoplasm).

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 555

PREPUTIAL GLANDS (INGUINAL GLANDS)

- 01: Left, green area 5 x 4 mm.
- Finding 01 in MAMMARY GLAND
- LIVER
- 01: Nodule adjacent diaphragm 17 x 10 x 5 mm.
- MAMMARY GLAND
- 01: Lactating.
- PITUITARY GLAND
- 01: Mass, dark, 8 x 5 x 4 mm.

- Inflammation Chronic Active, bilateral, grade 2.
- Cyst, bilateral.
- Hepatodiaphragmatic Nodule.
- SEE UNDER: PREPUTIAL GLANDS (INGUINAL GLANDS).
- Pars Distalis Adenoma.

ANIMAL NO: 556

TESTIS (LEFT)

- 01: Pale area, 9 x 8 x 6 mm.
- LIVER
- 01: Pale and enlarged.
- TESTES
- 01: Right, pale area, 6 x 5 x 4 mm.
- KIDNEYS
- 01: Pale and enlarged.
- ADRENAL GLANDS
- 01: Enlarged, red foci, 4 x 4 x 3 mm.
- LYMPH NODES
- 01: Mandibular, enlarged, 3 x 2 x 3 mm.
- 02: Thymic area, pale, 3 x 4 mm.

- Interstitial Cell Adenoma.
- Bile Duct Hyperplasia, grade 2.
- Interstitial Cell Adenoma, bilateral.
- Nephropathy, bilateral, grade 3.
- Cortex Focal Hyperplasia, Focal, Hypertrophy, bilateral, grade 2.
- No corresponding finding.
- No corresponding finding.

ANIMAL NO: 557

TESTIS (LEFT)

- 01: Pale mass, 25 x 14 x 11 mm.
- SPLEEN
- 01: Enlarged, 45 x 12 x 7 mm.
- TESTES
- 01: Right, pale mass, 30 x 15 x 12 mm.

- Interstitial Cell Adenoma.
- Extramedullary Hematopoiesis, grade 2.
- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 558

TESTIS (LEFT)

- 01: Pale mass, 23 x 12 x 10 mm.
- SKIN/SUBCUTIS
- 01: Ventral, ulceration 20 x 15 mm.
- PREPUTIAL GLANDS (INGUINAL GLANDS)
- 01: Mass, 30 x 30 x 15 mm.
- SPLEEN
- 01: Enlarged, 45 x 14 x 10 mm.

- Interstitial Cell Adenoma.
- Squamous Cell Carcinoma.
- No corresponding finding.
- Extramedullary Hematopoiesis, grade 2.

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TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
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PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

TESTES

- 01: Small, flaccid, 15 x 7 x 3 mm.

KIDNEYS

- 01: Granular.

LYMPH NODES

- 01: Mediastinal, dark.

CORRESPONDING MICROSCOPIC FINDING

- Tubular Atrophy, unilateral, grade 4.

- Nephropathy, bilateral, grade 2.

- Pigment, bilateral, grade 2.

ANIMAL NO: 559

TESTIS (LEFT)

- 01: Pale area, 11 x 6 mm.

SPLEEN

- 01: Enlarged, 55 x 11 x 7 mm.

TESTES

- 01: Right, pale mass, 12 x 12 x 6 mm.

- Interstitial Cell Adenoma.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 560

TESTIS (LEFT)

- 01: Small 15 x 12 x 4 mm.

- 02: Pale area, 2 x 3 mm.

TESTES

- 01: Right, 18 x 14 x 10 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 561

TESTIS (LEFT)

- 01: Pale area, 10 x 8 mm.

TESTES

- 01: Right, pale mass, 10 x 10 x 7 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 562

TESTIS (LEFT)

- 01: Pale mass, 25 x 14 x 12 mm.

TESTES

- 01: Right, pale mass, 28 x 18 x 14 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 563

GENERAL OBSERVATIONS

- 01: Whole body (skin, fat, and internal organs) have a yellow tint.

LIVER

- 01: Abnormal color (orange), granular-pitted surface.

- No corresponding finding.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

SPLEEN

- 01: Enlarged, 95 x 28 x 15 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

KIDNEYS

- 01: Black.

- Nephropathy, bilateral, grade 2.

PITUITARY GLAND

- 01: Focal area, red, approximately 3 mm dia.

- Pars Distalis Adenoma.

ANIMAL NO: 564

TESTIS (LEFT)

- 01: Pale mass, 25 x 11 x 11 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale mass, 20 x 12 x 10 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 566

TESTIS (LEFT)

- 01: Pale area, 10 x 9 mm.

- Interstitial Cell Adenoma.

MAMMARY GLAND

- 01: Subcutis neck, pale nodule, 2 x 2 x 1 mm.

- Galactocoele.

- 02: Subcutis abdomen, red nodule, 2 x 1 x 1 mm.

- Granuloma.

ANIMAL NO: 567

TESTIS (LEFT)

- 01: Black discoloration, focal.

- Interstitial Cell Adenoma.

LIVER

- 01: Diaphragmatic hernia, 15 x 10 x 6 mm.

- Hepatodiaphragmatic Nodule.

EYES

- 01: Right, bulging, red, crusted over.

- Cornea Ulcer, ulcer, unilateral, grade 3.

HARDERIAN GLANDS

- 01: Right, enlarged, 10 x 10 x 13 mm, red.

- Inflammation, focal, unilateral, grade 4.

GASSERIAN GANGLION (RIGHT)

- 01: Enlarged, 12 x 5 x 3 mm.

- Meningioma Malignant.

ANIMAL NO: 568

TESTIS (LEFT)

- 01: Discolored, yellow, firm areas.

- Interstitial Cell Adenoma.

LIVER

- 01: Liver is mottled, has granular and pitted surface, abnormal color (pale).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: 105 x 30 x 16 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

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TEST SYSTEM : RAT, , 18 MO, INHALATION
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DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

LUNG

- 01: Caudal lung lobe has two, 2 mm tan foci.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

KIDNEYS

- 01: Right kidney has a tumor present. 3 mm in diameter, both kidneys black.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

LYMPH NODES

- 01: Mandibular lymph nodes enlarged.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- 02: Mesenteric lymph nodes enlarged.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

- 03: Renal lymph nodes enlarged. (attached to mesentery).

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 569

TESTIS (LEFT)

- 01: Yellow, black, and white discoloration; bilateral.

- Interstitial Cell Adenoma.

- 02: Flaccid.

- No corresponding finding.

SKIN/SUBCUTIS

- 01: Left axillary region, mass, 85 x 80 x 32 mm, firm, lobulated, white and tan, necrotic area in center of mass.

- Fibroadenoma.

LIVER

- 01: Diaphragmatic hernia, 12 x 15 x 5 mm.

- Hepatodiaphragmatic Nodule.

SPLEEN

- 01: Enlarged, 50 x 15 x 7 mm.

- Extramedullary Hematopoiesis, grade 2.

ANIMAL NO: 571

TESTIS (LEFT)

- 01: pale area, 6 x 4 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale mass, 22 x 12 x 12 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 572

LUNG, LEFT

- 01: Pale area, 2 x 2 mm.

- Alveolar Epithelium Hyperplasia, grade 3.

LIVER

- 01: Slightly pale.

- Bile Duct Hyperplasia, grade 1.

SPLEEN

- 01: Slightly mottled.

- Extramedullary Hematopoiesis, grade 2.

ANIMAL NO: 573

TESTIS (LEFT)

- 01: Pale area, 10 x 10 mm.

- Interstitial Cell Adenoma.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

TESTES

- 01: Right, pale area, 7 x 5 mm.

- Interstitial Cell Adenoma, bilateral.

MAMMARY GLAND

- 01: Thorax subcutis, red nodules, 2 x 3 x 4 mm.

- Galactocole Hemorrhage.

ANIMAL NO: 574

LIVER

- 01: Pale, mottled + friable.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 9 x 2.7 x 1.5 cm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

LUNG

- 01: Mottled with pale areas.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

LYMPH NODES

- 01: Mesenteric lymph node, enlarged.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SEMINAL VESICLES

- 01: Small.

- No corresponding finding.

ANIMAL NO: 575

TESTIS (LEFT)

- 01: Pale area, 15 x 10 x 9 mm.

- Interstitial Cell Adenoma.

STERNUM

- 01: Red nodule, adjacent, 4 x 3 mm.

- Blood Vessel Arteritis, grade 3.

- 02: Red nodule, adjacent, 2 x 2 mm.

- Blood Vessel Arteritis, grade 3.

LIVER

- 01: Pale.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Mottled.

- Extramedullary Hematopoesis, grade 3.

TESTES

- 01: Right, pale areas (2) 2 x 4 mm.

- Interstitial Cell Adenoma, bilateral.

KIDNEYS

- 01: Mottled.

- Nephropathy, bilateral, grade 3.

ADRENAL GLANDS

- 01: Dark spots, 1 mm.

- Cortex Adenoma, Adenoma, unilateral.

MESENTERY

- 01: Multiple red, firm, nodules, 2-5mm diameter.

- Blood Vessel Arteritis, grade 3.

LYMPH NODES

- 01: Mediastinal, prominent.

- Plasma Cell Hyperplasia, bilateral, grade 2.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 576

TESTES

- 01: Right, pale mass, 25 x 15 x 11 mm. - Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 578

LIVER

- 01: Mottled, has granular appearance with pitted surface and abnormal color.

SPLEEN

- 01: Very enlarged, 90 x 13 x 14 mm.

KIDNEYS

- 01: Both kidneys are black.

LYMPH NODES

- 01: Renal lymph nodes, enlarged.
- 02: Mesenteric lymph nodes, enlarged.
- 03: Mandibular lymph nodes, enlarged.
- 04: Thymic lymph nodes, enlarged.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).
- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 579

TESTIS (LEFT)

- 01: Small, pale areas, 2-4mm.
TESTES

- 01: Pale mass, 22 x 14 x 10 mm.

- Interstitial Cell Adenoma.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 580

TESTES

- 01: Right, pale area, 9 x 7 mm.

- Interstitial Cell Adenoma, unilateral.

ANIMAL NO: 581

TESTIS (LEFT)

- 01: Pale area, 15 x 11 mm.

SPLEEN

- 01: Surface irregular.

KIDNEYS

- 01: Slightly mottled.

- Interstitial Cell Adenoma.

- Extramedullary Hematopoesis, grade 2.

- Nephropathy, bilateral, grade 3.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 582

SPLEEN

- 01: Dark.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

ANIMAL NO: 583

LUNG, LEFT

- 01: Pale area, 4 x 3 mm.

- Alveolar Bronchiolar Adenoma.

LUNG

- 01: Right, pale area, 6 x 5 mm.

- Interstitial Fibrosis, grade 3.

ANIMAL NO: 584

TESTIS (LEFT)

- 01: Pale mass, 30 x 15 x 12 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Slightly enlarged.

- Mixed Cell Focus.

SPLEEN

- 01: Slightly enlarged 40 x 11 x 7 mm.

- Congestion, grade 2.

TESTES

- 01: Pale mass, 25 x 15 x 11 mm.

- Interstitial Cell Adenoma, bilateral.

KIDNEYS

- 01: Slightly mottled.

- Nephropathy, bilateral, grade 3.

ANIMAL NO: 585

TESTIS (LEFT)

- 01: Pale mass, 30 x 15 x 14 mm.

- Interstitial Cell Adenoma.

SKIN/SUBCUTIS

- 01: Ventral chest, skin nodule, 7 x 5 x 4 mm.

- Keratoacanthoma.

TESTES

- 01: Small, flaccid, 18 x 9 x 7 mm.

- Tubular Atrophy, unilateral, grade 5.

ANIMAL NO: 586

TESTIS (LEFT)

- 01: Pale mass, 30 x 15 x 15 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Right, anterior lobe, pale mass, 15 x 15 x 9 mm.

- Hepatocellular Adenoma.

LUNG

- 01: Right, caudal lobe, pale area, 5 x 4 mm.

- Alveolar Epithelium Hyperplasia, grade 3.

KIDNEYS

- 01: Granular, pitted and pale.

- Nephropathy, bilateral, grade 3.

PALATE (ORAL MUCOSA)

- 01: Pale, raised area 5 x 7 mm.

- Squamous Cell Carcinoma.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 587

LUNG

- 01: Pale area, 3 x 7 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTIS (LEFT)

- 01: Pale area, 10 x 11 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Slightly pale.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

SPLEEN

- 01: Enlarged, 60 x 15 x 9 mm.

- Mononuclear Cell Leukemia (site of primary neoplasm:SYSTEMIC NEOPLASMS).

TESTES

- 01: Right, pale area, 4 x 11 mm.

- Interstitial Cell Adenoma, bilateral.

MAMMARY GLAND

- 01: Subcutaneous fat, red nodules 5 x 4 x 2 mm.

- Galactocole Hemorrhage.

- 02: Subcutaneous fat, red nodules, 3 x 3 x 2 mm.

- Galactocole Hemorrhage.

KIDNEYS

- 01: Surface, granular.

- Nephropathy, bilateral, grade 3.

ANIMAL NO: 588

TESTIS (LEFT)

- 01: Pale mass, 20 x 15 x 15 mm.

- Interstitial Cell Adenoma.

TESTES

- 01: Right, pale area, 15 x 11 mm.

- Interstitial Cell Adenoma, bilateral.

ANIMAL NO: 589

LUNG

- 01: Right , caudal lobe, pale area, 4 x 5 mm.

- Alveolar Epithelium Hyperplasia, grade 3.

ANIMAL NO: 591

TESTIS (LEFT)

- 01: Pale area, 2 x 3 mm.

- Interstitial Cell Adenoma.

KIDNEYS

- 01: Slightly granular.

- Nephropathy, bilateral, grade 4.

ANIMAL NO: 592

TESTIS (LEFT)

- 01: Pale mass, 12 x 10 x 7 mm.

- Interstitial Cell Adenoma.

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
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CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

LYMPH NODES

- 01: Mandibular LN, prominent.

- Plasma Cell Hyperplasia, bilateral, grade 3.

ANIMAL NO: 593

TESTIS (LEFT)

- 01: Pale area, 5 x 4 mm.

- Interstitial Cell Adenoma.

LUNG

- 01: Right caudal lobe, pale area, 4 x 4 mm.

- Alveolar Bronchiolar Adenoma.

MAMMARY GLAND

- 01: Right, lateral thorax, pale, soft mass,
45 x 47 x 22 mm. Cut surface reveals
milky liquid and soft pale tissue.

- Fibroadenoma, single.

KIDNEYS

- 01: Pale and pitted surface.

- Nephropathy, bilateral, grade 4.

ANIMAL NO: 594

TESTIS (LEFT)

- 01: Small, 15 x 8 x 7 mm.

- Tubular Atrophy, grade 5.

LIVER

- 01: Edges rounded, pale.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Enlarged, 40 x 10 x 7 mm.

- Extramedullary Hematopoiesis, grade 2.

TESTES

- 01: Right, pale mass, 20 x 20 x 18 mm.

- Interstitial Cell Adenoma, unilateral.

ANIMAL NO: 595

KIDNEYS

- 01: Brown, granular surface.

- Nephropathy, bilateral, grade 4.

ANIMAL NO: 596

TESTIS (LEFT)

- 01: Pale mass, 22 x 14 x 11 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Irregular surface.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Enlarged, 45 x 10 x 7 mm.

- Extramedullary Hematopoiesis, grade 2.

TESTES

- 01: Right, small, 15 x 10 x 7 mm.

- Interstitial Cell Adenoma, bilateral.

KIDNEYS

- 01: Pitted, slight.

- Nephropathy, bilateral, grade 4.

PATHOLOGY REPORT
SUMMARY TABLES

PAGE : 57/ 57
10026 EPL304-466

TEST ITEM : AMPHIBOLE ASBESTOS
TEST SYSTEM : RAT, , 18 MO, INHALATION
SPONSOR : THE HAMNER

PATHOL. NO.: 90188 GAW
DATE : 29-AUG-13
PathData@System V6.2d2

CORRELATION TABLE: NECROPSY - MICROSCOPY

DOSE GROUP 5, MALE

NECROPSY OBSERVATION

CORRESPONDING MICROSCOPIC FINDING

ANIMAL NO: 597

TESTIS (LEFT)

- 01: Left testis, pale mass 30 x 16 x 15 mm.

- Interstitial Cell Adenoma.

LIVER

- 01: Pale, edges rounded.

- Bile Duct Hyperplasia, grade 2.

SPLEEN

- 01: Slightly enlarged, 40 x 8 x 8 mm.

- Extramedullary Hematopoiesis, grade 2.

TESTES

- 01: Right, small, 20 x 10 x 9 mm.

- Interstitial Cell Adenoma, bilateral.

EXPLANATION OF CODES AND SYMBOLS

EXPLANATION OF CODES AND SYMBOLS

CODES AND SYMBOLS USED AT TABLE LEVEL:

AOFT = Animal Organ Finding Table

CODES AND SYMBOLS USED AT ANIMAL LEVEL:

M = Male Animal
K0 = Terminal Sacrifice Group
K1 . . . K9 = Interim Sacrifice Group 1 . . . 9
R1 . . . R9 = Recovery / Post-Treatment Group 1 . . . 9
+ = Intercurrent Death / Sacrificed Moribund

CODES AND SYMBOLS USED AT ORGAN LEVEL:

A = Organ autolytic, evaluation not possible
G = Gross finding evaluated histologically
0 = Tissue not present for histologic examination
' = Histologic examination not required
+ = Organ examined, findings present
- = Organ examined, no pathologic findings noted (AOFT only)
(= Only one of paired organs examined/present
! = No corresponding microscopic finding required
NAD = No abnormalities detected

CODES AND SYMBOLS USED AT FINDING LEVEL:

GRADE 1 = Minimal / very few / very small
GRADE 2 = Slight / mild / few / small
GRADE 3 = Moderate / moderate number / moderate size
GRADE 4 = Marked / many / large / moderately severe
GRADE 5 = Massive / extensive number / extensive size / severe
P = Finding present, severity not scored
B0 = Benign neoplasm
N0 = Malignant neoplasm
M = Metastasis
(= Finding unilateral in paired organs

CORRELATION TABLE: NECROPSY-MICROSCOPY:

Evaluation not required = No corresponding microscopic finding required

ATTACHMENT

**EPA FIBER PROJECT: SUBCHRONIC INHALATION EXPOSURE
OF RATS TO AMPHIBOLE ASBESTOS:**

**FINAL BRONCHOALVEOLAR LAVAGE CYTOLOGY REPORT
18 MONTHS POST-EXPOSURE**



Experimental Pathology Laboratories, Inc.

The Hamner Protocol No.: 10026
EPL Project No.: 304-466
FINAL REPORT – August 29, 2013

EPA FIBER PROJECT: SUBCHRONIC INHALATION EXPOSURE
OF RATS TO AMPHIBOLE ASBESTOS:
BRONCHOALVEOLAR LAVAGE CYTOLOGY

THE HAMNER INSTITUTES FOR
HEALTH SCIENCES PROTOCOL NUMBER: 10026

EPL PROJECT NO.: 304-466

FINAL BRONCHOALVEOLAR LAVAGE CYTOLOGY REPORT
(18 MONTHS POST-EXPOSURE)

Submitted to:

The Hamner Institutes for Health Sciences
6 Davis Drive
P.O. Box 12137
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Submitted by:

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August 29, 2013



EPA FIBER PROJECT: SUBCHRONIC INHALATION EXPOSURE
OF RATS TO AMPHIBOLE ASBESTOS:
BRONCHOALVEOLAR LAVAGE CYTOLOGY

THE HAMNER INSTITUTES FOR
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FINAL BRONCHOALVEOLAR LAVAGE CYTOLOGY REPORT
(18 MONTHS POST-EXPOSURE)

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EPA FIBER PROJECT: SUBCHRONIC INHALATION EXPOSURE
OF RATS TO AMPHIBOLE ASBESTOS:
BRONCHOALVEOLAR LAVAGE CYTOLOGY

THE HAMNER INSTITUTES FOR
HEALTH SCIENCES PROTOCOL NUMBER: 10026

EPL PROJECT NO.: 304-466

FINAL BRONCHOALVEOLAR LAVAGE CYTOLOGY REPORT
(18 MONTHS POST-EXPOSURE)

INTRODUCTION

This subchronic study was conducted to determine the biological potency of inhaled Libby amphibole (LA) fibers over the near-life span of the rat compared to the potency of inhaled amosite, a known fibrogenic amphibole asbestos fiber, and to develop fiber burden data to use in a dosimetry model of amphibole fiber deposition, clearance, and retention in the respiratory tract (head, trachea, lung lobes) and GI tract. As part of this study, Experimental Pathology Laboratories, Inc. (EPL[®]) was requested to perform the quantitative bronchoalveolar lavage (BAL) cytological evaluation of prepared cytological slides provided by The Hamner Institutes for Health Sciences.

The original experimental design pertinent to the BAL cytology analysis is summarized in Table 1. The results for the BAL cytology analysis for samples collected at 1 day, 1 month, and 3 months post-exposure have been separately reported (Willson and Wall, 2013). At the final (~18 months post-exposure) necropsy, BAL was conducted on 8 of the surviving animals from the scheduled group of 50 per dose. The present report only includes samples collected at 18 months post-exposure.

Table 1. Experimental Design

Group Number ¹	Test Material	Concentration (mg/m ³)	No. of rats to be necropsied at each time point ²				Total in Each Exp. Group	
			Post-Exposure Period					
			1 day	1 month	3 months	18 months		
1	Air Control	0.0	11	11	11	53	86	
2	Amosite	3.3	14	14	14	56	98	
3	LA	1.0	14	14	14	56	98	
4	LA	3.3	14	14	14	56	98	
5	LA	10.0	14	14	14	56	98	
Total in Each Post-Exp Period			67	67	67	277		
Total Animals in Study			478					

¹To facilitate the BAL cytology analysis, the exposure level (treatment) groups were numbered as follows: 1 – Air control, 2 – amosite 3.3 mg/m³, 3 – LA 1.0, 4 – LA 3.3 mg/m³, and 5 – LA 10 mg/m³.

²At each scheduled necropsy conducted at 1 day, 1 month, and 3 months post-exposure, the same 8 animals designated for histopathology were also designated to have the right lung lavaged for BAL cytology.

BRONCHOALVEOLAR LAVAGE ISOLATION AND CYTOLOGY PROCEDURES

Rat lungs were lavaged by means of a cannula inserted into the trachea. Three mL of room temperature phosphate buffered saline were introduced and drawn out of the lungs twice (total of 6 mL) and stored on ice (L1 sample). Lungs were then lavaged three more times (total of 9 mL) and stored on ice (L2 sample). The L1 and L2 samples were centrifuged at 200 x g for 10 minutes. Cell pellets from L1 and L2 samples were combined in 1 mL of F12 medium (Gibco F12 GlutaMax) and counted using a Coulter Counter Model ZM (Beckman Coulter, Brea, CA). Zap-oglobin was used to lyse red cells in the counting vial. Duplicate counts were recorded to obtain a mean count per sample.

Slides were prepared by adjusting the cell concentrations to 50,000/225 µL and centrifuging the 225 µL with 25 µL fetal bovine serum in a Shandon Cytospin (Thermo Fisher Scientific, Waltham, MA) for three minutes at 500 RPM. Slides were air dried followed by fixation and staining with Siemens Diff-Quik Stain Set (Fisher Scientific, Kalamazoo, MI) and provided to EPL for determination of the cell differential percentages. Three hundred (300) leukocytic cells per slide were enumerated and differentiated (Gao et al., 2006; Palmans et al., 2000) via a manual tagging method using digital photographic images of cells in photographic fields (40X total magnification). The fields for photography



were selected using a random pattern that prevented duplicate photography of the same fields.

The method of enumeration was performed via the application of manual tagging procedures available in the image analysis software, ImagePro Plus, v.5.0.2.9 (Media Cybernetics, Inc., Silver Spring, Maryland). Briefly, the cells enumerated included: macrophages, lymphocytes, neutrophils, and eosinophils. For each cell type a unique color code was assigned as the class color identifier. Using the appropriate class color each type of cell in each photograph was counted. As each cell was counted it was tagged with a unique number that had the color code for its class. The evaluation of photographic fields and counting was continued until a count of 300 cells was accumulated. Samples with less than 300 leukocytic cells were considered inadequate for evaluation. The absolute differential cell count for each photographic field was manually recorded.

Using an EXCEL spreadsheet, the absolute counts for each cell type observed in all photographic fields counted were totaled for each animal. The differential cell count data for animal 376 were recorded, but percentages were not calculated and the animal was excluded from the statistical analyses due to an inadequate sample.

For those animals for which 300 cells were counted, the total count of each cell type for each animal was divided by 300 and the result multiplied by 100 to obtain the cell differential percentage for each cell type enumerated. The cell counts recorded for individual photomicrographic images were included in the filed study raw data. The total cell counts for each cell type for each animal, and the calculated cell differential percentages for each animal were tabulated for inclusion in this report and for application to calculate the absolute (total) differential BAL cell counts used for the statistical analysis. The tabulated data for the differential counts of cells in BAL cytology preps were audited by the EPL Quality Assurance unit and the audited tabulated data were provided electronically to the Study Director for statistical analysis of the absolute (total) BAL differential cell counts that was completed by The Hamner Institutes for Health Sciences. The results of the statistical analysis were returned to EPL. Tabular and graphic information that document the statistical analysis was filed as raw data.

RESULTS AND DISCUSSION

The results for the differential cell count of up to 300 cells in BAL cytology samples from each animal are presented in Appendix A. The total cell counts that were calculated using percentages in Appendix A are presented in Appendix B. Means and standard deviations of total cell counts per major cell types (lymphocytes, neutrophils, and macrophages) are shown in Table 2. In general, amosite or LA fiber-exposed groups had lower mean lymphocytes and higher mean neutrophils compared to the air control group means, but there were no statistically significant differences in the total number of cells for each type of cell included in the differential count at the 18 month's post-exposure time point. The absence of differences in BAL cell types in animals that were dosed with Libby amphibole or Amosite when compared to animals in the Air Control at 18 months post-exposure suggests complete resolution of inflammatory cell responses that might have been associated with the earlier sampling time points (1 day, 1 month, and 3 months post-exposure).

Table 2. Summary of Total Cell Counts for Lymphocytes, Neutrophils, and Macrophages

Exposure Group and Concentration	Lymphocytes	Neutrophils	Macrophages ($\times 10^2$)
Air Control	11991 \pm 16607*	14379 \pm 13644	14939 \pm 3555
Amosite 3.3 mg/m ³	10784 \pm 15129	30647 \pm 14935	15510 \pm 3438
LA 1.0 mg/m ³	6931 \pm 7526**	15528 \pm 6803**	13958 \pm 2570**
LA 3.3 mg/m ³	1568 \pm 3024	43716 \pm 73747	14531 \pm 3112
LA 10 mg/m ³	3116 \pm 4123	41460 \pm 40470	13705 \pm 1943

*Mean \pm standard deviation, n=8

**n=7

CONCLUSIONS

The BAL cell differential cytology and total cell counts for individual cell types disclosed that Libby amphibole (LA) exposure of rats to 10.0, 3.3. or 1.0 mg/m³ for 90 days resulted in no changes in the proportion of cell types in BAL samples when compared to rats that were Air Control or exposed to Amosite 3.3 mg/m³ for 90 days when evaluated at the 18



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months post-exposure time point. In contrast to the 18 months post-exposure time point, neutrophils were increased in rats exposed to LA 10.0 mg/m³ or to Amosite 3.3 mg/m³ at 1 day, 1 month and 3 months post-exposure; neutrophils were increased in the LA 3.3 mg/m³ group at the 1 day and 3 months post-exposure; and macrophages were decreased in the LA 10.0 and LA 3.3 mg/m³ groups at 3 months post-exposure when these groups were compared to the Air Control group. The absence of differences in the cell types in BAL samples at the 18 months post-exposure suggests complete resolution of inflammatory cell responses that were associated with earlier sampling time points (1 day, 1 month, and 3 months post-exposure).

A handwritten signature in black ink that reads "Henry G. Wall".

HENRY G. WALL, D.V.M., Ph.D.
Diplomate, ACVP
Veterinary Pathologist

29 Aug 2013
DATE

HGW/dc



REFERENCES

Gao H, Hoesel LM, Guo R-F, Rancilio NJ, Sarma JV, Ward PA. 2006. Adenoviral-mediated overexpression of SOCS3 enhances IgG immune complex-induced acute lung injury. *J Immunol* 177:612-620.

Palmans E, Kips JC, Pauwels RA. 2000. Prolonged allergen exposure induces structural airway changes in sensitized rats. *Am J Respir Crit Care Med* 161:627-635.

G. Willson, H. Wall. March 6, 2013. Amended Final Report: EPA Fiber Project: Subchronic Inhalation Exposure of Rats to Amphibole Asbestos (1-Day Post-Exposure, 1-Month Post-Exposure, and 3-Months Post-Exposure).



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APPENDIX A

INDIVIDUAL ANIMAL BAL CELLS DIFFERENTIAL COUNTS AND PERCENTAGES – 18 MONTHS POST-EXPOSURE



Appendix A: Individual Animal BAL Cells Differential Counts And Percentages –18 Months Post-Exposure

Necropsy Interval (Months post-exposure)	Exposure Level Concentration (mg/m3)	GP*	ID	EOS	EOP	LYM	LYP	NEU	NEP	MAC	MAP	BAS	BAP	Total Cells Counted
18	Air Control	1	156	0	0.00	9	3.00	4	1.33	287	95.67	0	0.00	300
18	Air Control	1	158	0	0.00	0	0.00	3	1.00	297	99.00	0	0.00	300
18	Air Control	1	159	0	0.00	5	1.67	6	2.00	289	96.33	0	0.00	300
18	Air Control	1	160	0	0.00	2	0.67	3	1.00	295	98.33	0	0.00	300
18	Air Control	1	161	0	0.00	0	0.00	3	1.00	297	99.00	0	0.00	300
18	Air Control	1	162	0	0.00	2	0.67	0	0.00	298	99.33	0	0.00	300
18	Air Control	1	163	0	0.00	0	0.00	2	0.67	298	99.33	0	0.00	300
18	Air Control	1	164	0	0.00	0	0.00	0	0.00	300	100.00	0	0.00	300
18	Amosite 3.3	2	268	0	0.00	4	1.33	7	2.33	289	96.33	0	0.00	300
18	Amosite 3.3	2	269	0	0.00	0	0.00	4	1.33	296	98.67	0	0.00	300
18	Amosite 3.3	2	270	0	0.00	2	0.67	6	2.00	292	97.33	0	0.00	300
18	Amosite 3.3	2	274	0	0.00	8	2.67	7	2.33	284	94.67	1	0.33	300
18	Amosite 3.3	2	275	0	0.00	2	0.67	1	0.33	297	99.00	0	0.00	300
18	Amosite 3.3	2	276	0	0.00	1	0.33	8	2.67	291	97.00	0	0.00	300
18	Amosite 3.3	2	277	0	0.00	0	0.00	5	1.67	295	98.33	0	0.00	300
18	Amosite 3.3	2	278	0	0.00	0	0.00	8	2.67	292	97.33	0	0.00	300
18	LA 1.0	3	370	0	0.00	5	1.67	4	1.33	291	97.00	0	0.00	300
18	LA 1.0	3	372	0	0.00	0	0.00	4	1.33	295	98.33	1	0.33	300
18	LA 1.0	3	375	0	0.00	1	0.33	1	0.33	298	99.33	0	0.00	300
18	LA 1.0	3	376	0		14		10		134		0		158**
18	LA 1.0	3	377	0	0.00	1	0.33	5	1.67	294	98.00	0	0.00	300
18	LA 1.0	3	378	0	0.00	0	0.00	2	0.67	298	99.33	0	0.00	300

*Abbreviations: GP – group, ID – animal identification, EOS – eosinophil count, EOP – eosinophil percentage, LYM – lymphocyte count, LYP – lymphocyte percentage, NEU – neutrophil count, NEP – neutrophil percentage, MAC – macrophage count, MAP – macrophage percentage, BAS – basophil count, BAP – basophil percentage

**Inadequate sample for evaluation.



Appendix A: Individual Animal BAL Cells Differential Counts And Percentages –18 Months Post-Exposure (Continuation)

Necropsy Interval (Months post-exposure)	Exposure Level Concentration (mg/m3)	GP	ID	EOS	EOP	LYM	LYP	NEU	NEP	MAC	MAP	BAS	BAP	Total Cells Counted
18	LA 1.0	3	380	0	0.00	2	0.67	2	0.67	296	98.67	0	0.00	300
18	LA 1.0	3	395	0	0.00	2	0.67	6	2.00	292	97.33	0	0.00	300
18	LA 3.3	4	467	0	0.00	0	0.00	6	2.00	294	98.00	0	0.00	300
18	LA 3.3	4	468	0	0.00	1	0.33	0	0.00	299	99.67	0	0.00	300
18	LA 3.3	4	472	0	0.00	0	0.00	6	2.00	294	98.00	0	0.00	300
18	LA 3.3	4	474	0	0.00	0	0.00	47	15.67	253	84.33	0	0.00	300
18	LA 3.3	4	475	0	0.00	2	0.67	4	1.33	294	98.00	0	0.00	300
18	LA 3.3	4	476	0	0.00	0	0.00	5	1.67	295	98.33	0	0.00	300
18	LA 3.3	4	477	0	0.00	0	0.00	3	1.00	297	99.00	0	0.00	300
18	LA 3.3	4	496	0	0.00	0	0.00	2	0.67	298	99.33	0	0.00	300
18	LA 10.0	5	571	0	0.00	0	0.00	16	5.33	284	94.67	0	0.00	300
18	LA 10.0	5	572	0	0.00	1	0.33	5	1.67	294	98.00	0	0.00	300
18	LA 10.0	5	573	0	0.00	1	0.33	2	0.67	297	99.00	0	0.00	300
18	LA 10.0	5	575	0	0.00	0	0.00	5	1.67	295	98.33	0	0.00	300
18	LA 10.0	5	576	0	0.00	0	0.00	2	0.67	298	99.33	0	0.00	300
18	LA 10.0	5	577	0	0.00	2	0.67	21	7.00	277	92.33	0	0.00	300
18	LA 10.0	5	579	0	0.00	0	0.00	8	2.67	292	97.33	0	0.00	300
18	LA 10.0	5	580	0	0.00	1	0.33	5	1.67	294	98.00	0	0.00	300

*Abbreviations: GP – group, ID – animal identification, EOS – eosinophil count, EOP – eosinophil percentage, LYM – lymphocyte count, LYP – lymphocyte percentage, NEU – neutrophil count, NEP – neutrophil percentage, MAC – macrophage count, MAP – macrophage percentage, BAS – basophil count, BAP – basophil percentage

**Inadequate sample for evaluation.



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APPENDIX B

18 MONTHS POST-EXPOSURE TOTAL CELL COUNTS BY CELL TYPES



Appendix B: 18 Months Post-Exposure Total Cell Counts by Cell Types

Necropsy Interval (Months post-exposure)	Exposure Level Concentration (mg/m3)	ID	Eosinophils	Lymphocytes	Neutrophils	Macrophages	Basophils	Total Cells Counted
18	Air Control	156		39,627	17567.97	1263705.03		1,320,900
18	Air Control	158		0	15954	1579446		1,595,400
18	Air Control	159		36,020	43138	2077741.77		2,156,900
18	Air Control	160		12,078	18027	1772594.91		1,802,700
18	Air Control	161		0	9434	933966		943,400
18	Air Control	162		8,202	0	1215997.86		1,224,200
18	Air Control	163		0	10908.94	1617291.06		1,628,200
18	Air Control	164		0	0	1490100		1,490,100
18	Amosite 3.3	268		16,786	29406.93	1215780.93		1,261,974
18	Amosite 3.3	269		0	19521.74	1448278.26		1,467,800
18	Amosite 3.3	270		7,263	21680	1055057.2		1,084,000
18	Amosite 3.3	274		45,230	39470.2	1603709.8	5590.2	1,694,000
18	Amosite 3.3	275		10,479	5161.2	1548360		1,564,000
18	Amosite 3.3	276		6,518	52732.5	1915750		1,975,000
18	Amosite 3.3	277		0	35958.44	2117241.56		2,153,200
18	Amosite 3.3	278		0	41246.16	1503553.84		1,544,800
18	LA 1.0	370		21,376	17024	1241600		1,280,000
18	LA 1.0	372		0	20431.46	1510545.46	5069.46	1,536,046
18	LA 1.0	375		4,144	4143.81	1247286.81		1,255,574
18	LA 1.0	376						1931800*
18	LA 1.0	377		3,896	19716.02	1156988		1,180,600
18	LA 1.0	378		0	12263.01	1818036.99		1,830,300
18	LA 1.0	380		11,028	11027.53	1624009.53		1,646,065
18	LA 1.0	395		8,071	24092	1172437.18		1,204,600

*Inadequate sample for evaluation.



Appendix B: 18 Months Post-Exposure Total Cell Counts by Cell Types (Continuation)

Necropsy Interval (Months post-exposure)	Exposure Level Concentration (mg/m3)	ID	Eosinophils	Lymphocytes	Neutrophils	Macrophages	Basophils	Total Cells Counted
18	LA 3.3	467		0	24962	1223138		1,248,100
18	LA 3.3	468		4,694	0	1417706.08		1,422,400
18	LA 3.3	472		0	26202	1283898		1,310,100
18	LA 3.3	474		0	224942.85	1210557.15		1,435,500
18	LA 3.3	475		7,852	15587.6	1148560		1,172,000
18	LA 3.3	476		0	25509.25	1501990.75		1,527,500
18	LA 3.3	477		0	19943	1974357		1,994,300
18	LA 3.3	496		0	12578.58	1864821.42		1,877,400
18	LA 10.0	571		0	82828.2	1471171.8		1,554,000
18	LA 10.0	572		4,500	22773.79	1336426		1,363,700
18	LA 10.0	573		3,540	7186.42	1061874		1,072,600
18	LA 10.0	575		0	23949.47	1410150.53		1,434,100
18	LA 10.0	576		0	7508.02	1113091.98		1,120,600
18	LA 10.0	577		11,704	122283	1612912.77		1,746,900
18	LA 10.0	579		0	38891.22	1417708.78		1,456,600
18	LA 10.0	580		5,188	26255.74	1540756		1,572,200

*Inadequate sample for evaluation.