

Experiment Number: 97011-23

Test Type: 26-WEEK

Route: SKIN APPLICATION

Species/Strain: Mouse/FVB/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Transgenic model evaluation (Melphalan)

CAS Number: 148-82-3

Date Report Requested: 10/20/2014

Time Report Requested: 04:55:33

First Dose M/F: NA / NA

Lab: MBA

C Number: C97011C

Lock Date: 07/21/2000

Cage Range: All

Date Range: All

Reasons For Removal: All

Removal Date Range: All

Treatment Groups: All

Study Gender: Both

PWG Approval Date NONE

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FVB/N Mouse Male
VEHICLE CONTROL

DAY ON TEST															
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
8	8	8	8	8	8	8	4	8	8	8	8	8	8	8	8
6	6	6	6	6	6	6	4	6	6	6	6	6	6	6	6
ANIMAL ID	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	5	5	5	5	5	5	5	5	5	6	6	6	6	6	6
	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5

*TOTALS

Alimentary System

Intestine Large, Cecum	+	+	+	+	+	+	A	+	+	+	+	+	+	+	+	14
Intestine Large, Colon	+	+	+	+	+	+	A	+	+	+	+	+	+	+	+	14
Intestine Large, Rectum	+	+	+	+	+	+	A	+	+	+	+	+	+	+	+	14
Intestine Small, Duodenum	+	+	+	+	+	+	A	+	+	+	+	+	+	+	+	14
Intestine Small, Ileum	+	+	+	+	+	+	A	+	+	+	+	+	+	+	+	14
Intestine Small, Jejunum	+	+	+	+	+	+	A	+	+	+	+	+	+	+	+	14
Liver	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15
Hepatocyte, Necrosis, Focal								1								1 1.0
Infiltration Cellular, Polymorphnucr, Focal								1								1 1.0
Stomach, Forestomach	+	+	+	+	+	+	A	+	+	+	+	+	+	+	+	14
Stomach, Glandular	+	+	+	+	+	+	A	+	+	+	+	+	+	+	+	14

Cardiovascular System

Heart	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15
-------	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	----

Endocrine System

Adrenal Cortex	+	+	+	+	+	+	M	+	+	+	+	+	+	+	+	14
Atrophy	2	2	2	2	2	3		3	2	3	2	3	3	2	2	14 2.4
Hypertrophy, Focal	1											1	1			4 1.0
Zona Reticul, Vacuolization Cytoplasmic, Focal								1				1				2 1.0
Adrenal Medulla	+	+	+	+	+	+	M	+	+	+	+	+	+	+	+	14

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

1-4 ..Lesion qualified as:

X ..Lesion present

A ..Autolysis precludes evaluation

1) Minimal 3) Moderate

I ..Insufficient tissue

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VEHICLE CONTROL

DAY ON TEST															
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
ANIMAL ID	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	8	8	8	8	8	8	4	8	8	8	8	8	8	8	8
	6	6	6	6	6	6	4	6	6	6	6	6	6	6	6
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	5	5	5	5	5	5	5	5	5	6	6	6	6	6	6
	1	2	3	4	5	6	7	8	9	0	1	2	3	4	5
	*TOTALS														
Pituitary Gland	+	+	+	+	+	+	A	+	I	+	+	+	+	+	13
Thyroid Gland	+	+	+	+	+	+	A	+	+	+	+	+	+	+	14
General Body System															
NONE															
Genital System															
Epididymis	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15
Testes	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15
Germinal Epith, Degeneration							1					2		2	1.5
Hematopoietic System															
Bone Marrow	+	+	+	+	+	+	A	+	+	+	+	+	+	+	14
Lymph Node, Mandibular	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15
Lymph Node, Mediastinal	M	+	I	+	+	+	+	+	+	+	+	+	+	+	13
Lymph Node, Mesenteric	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15
Spleen	+	+	+	+	+	+	A	+	+	+	+	+	+	+	14
Hematopoietic Cell Proliferation	1	2	1	1	2	2	1	1	2	1	2	2	2	1	14
Pigmentation	1	1	1	1	1	1			1	1	1	1	1		10
Thymus	+	+	+	+	+	+	A	+	+	+	+	+	+	+	14
Integumentary System															
Mammary Gland	M	M	M	M	M	M	M	M	M	M	M	M	M	M	0
Skin	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15
Epidermis, Hyperplasia, Focal							1					1		1	1.0

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+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

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Lab: MBA

FVB/N Mouse Male
VEHICLE CONTROL

DAY ON TEST															
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
8	8	8	8	8	8	8	4	8	8	8	8	8	8	8	8
6	6	6	6	6	6	4	6	6	6	6	6	6	6	6	6
ANIMAL ID	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
5	5	5	5	5	5	5	5	5	5	6	6	6	6	6	6
1	2	3	4	5	6	7	8	9	0	1	2	3	4	5	*TOTALS

Epidermis, Hyperplasia, Focal

1

1 1.0

Musculoskeletal System

NONE

Nervous System

NONE

Respiratory System

Lung

+

+

+

+

+

+

+

+

+

+

+

+

+

+

+

15

Perivascular, Infiltration Cellular,
Lymphocyte, Focal

1

1 1.0

Special Senses System

NONE

Urinary System

Kidney

+

+

+

+

+

+

A

+

+

+

+

+

+

+

+

14

Renal Tubule, Degeneration, Focal

1

1 1.0

Renal Tubule, Dilatation, Focal

1

4 1.0

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First Dose M/F: NA / NA

Lab: MBA

FVB/N Mouse Male 4.0 MG/KG	DAY ON TEST	0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0																														
		1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1																														
		8 8 8 8 8 8 4 8 8 4 8 7 8 8 8 3																														
		6 6 2 6 6 6 8 6 6 5 6 3 6 6 3																														
		0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0																														
ANIMAL ID																																
0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0																																
1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1																																
6 6 6 6 7 7 7 7 7 7 7 7 7 7 7 8																																
6 7 8 9 0 1 2 3 4 5 6 7 7 8 9 0																																
*TOTALS																																

Alimentary System

Intestine Large, Cecum	+	+	+	+	+	+	A	+	+	A	+	+	+	+	+	+	13
Intestine Large, Colon	+	+	+	+	+	+	A	+	+	A	+	+	+	+	+	+	13
Intestine Large, Rectum	+	+	+	+	+	+	A	+	+	A	+	+	+	+	+	+	13
Intestine Small, Duodenum	+	+	+	+	+	+	A	+	+	A	+	+	+	+	+	+	13
Epithelium, Necrosis	3	3		3	3		3	3		2		3	3		9	2.9	
Intestine Small, Ileum	+	+	+	+	+	+	A	+	+	A	+	+	+	+	A		12
Epithelium, Necrosis	3			2	3		2	3		2		3	2		8	2.5	
Intestine Small, Jejunum	+	+	+	+	+	+	A	+	+	A	+	+	+	+	A		12
Epithelium, Necrosis	3			2	3		2	3		2		3	2		8	2.5	
Liver	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+		15
Hepatocyte, Necrosis, Focal							1						1		2	1.0	
Pancreas															+		1
Stomach, Forestomach	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+		15
Stomach, Glandular	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+		15
Tooth	+	+			+		+	+				+		+			7
Inflammation, Chronic Active, Diffuse															4	1	4.0

Cardiovascular System

Heart	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15
-------	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	----

Endocrine System

Adrenal Cortex	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15
----------------	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	---	----

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Lab: MBA

**FVB/N Mouse Male
4.0 MG/KG**

DAY ON TEST															
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	8	8	8	8	8	8	4	8	8	4	8	7	8	8	3
	6	6	2	6	6	6	8	6	6	5	6	3	6	6	3
ANIMAL ID	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
	6	6	6	6	7	7	7	7	7	7	7	7	7	7	8
	6	7	8	9	0	1	2	3	4	5	6	7	8	9	0

***TOTALS**

Atrophy	3	3	3	3	4	3	2	3	4	3	4	3	3	3	2	15	3.1
Hypertrophy, Focal	1	1	1	1	1				1	1	1	1	1	1	1	12	1.0
Zona Reticul, Vacuolization Cytoplasmic, Focal	1					1				1						3	1.0
Adrenal Medulla	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15	
Pituitary Gland	+	+	+	+	+	+		+	+		+	+	+	+	+	13	
Thyroid Gland	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15	
Cyst, Focal															1	1	1.0

General Body System

NONE

Genital System

Epididymis	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15	
Penis															+	1	
Testes	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15	
Germinal Epith, Degeneration	4	4	4	4	4	4	4	4	4	4	4	4	4	4	4	15	4.0

Hematopoietic System

Bone Marrow	+	+	+	+	+	+	A	+	+	+	+	+	+	+	+	14	
Lymph Node															+	1	
Lymph Node, Mandibular	+	+	+	+	+	+	+	+	+	+	+	+	+	+	+	15	
Lymph Node, Mediastinal	+	+	M	+	+	+	+	+		+	+	M	+	+	M	11	
Lymph Node, Mesenteric	+	+	+	+	+	+	A	+	+	+	+	+	+	+	M	13	

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

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Lab: MBA

FVB/N Mouse Male 4.0 MG/KG	DAY ON TEST	0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0															
		1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1															
		8 8 8 8 8 8 4 8 8 4 8 7 8 8 8 3															
		6 6 2 6 6 6 8 6 6 5 6 3 6 6 3															
		0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0															
ANIMAL ID		0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0															
1		1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1															
6		6 6 6 6 7 7 7 7 7 7 7 7 7 7 7 8															
6		7 8 9 0 1 2 3 4 5 6 7 7 8 9 0															
*TOTALS																	
Spleen		+ + + + + + + + + + + + + + + +															15
Hematopoietic Cell Proliferation		2 3		1 1	2		2 2		3		3 3	1					11 2.1
Lymph Follic, Depletion Cellular				3				2				4					3 3.0
Pigmentation		1 2		1 1 1				1		1 1		1 1					9 1.1
Thymus		+ + + + + + + + + + + + + + + M															14
Atrophy, Diffuse				4				4 4				4					4 4.0
Atrophy, Focal				2								1	2.0				
Integumentary System																	
Mammary Gland		M M M M M M M M M M M M M M M M															0
Skin		+ + + + + + + + + + + + + + + +															15
Subcut Tiss, Inflammation, Chronic Active, Focal												4		1	4.0		
Musculoskeletal System																	
NONE																	
Nervous System																	
NONE																	
Respiratory System																	
Lung		+ + + + + + + + + + + + + + + +															15
Alveolar Epith, Hyperplasia, Focal		1 4 1 1							2			5 1.8					
Alveolus, Inflammation, Chronic Active, Focal		1		1					1			3 1.0					
Inflammation, Chronic Active, Focal								1				1	1.0				

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Lab: MBA

**FVB/N Mouse Male
4.0 MG/KG**

DAY ON TEST															
	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
8	8	8	8	8	8	4	8	8	4	8	7	8	8	8	3
6	6	6	2	6	6	8	6	6	5	6	3	6	6	6	3
ANIMAL ID	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
1	1	1	1	1	1	1	1	1	1	1	1	1	1	1	1
6	6	6	6	6	7	7	7	7	7	7	7	7	7	7	8
6	7	8	9	0	1	2	3	4	5	6	7	8	9	0	

*TOTALS

Special Senses System

Eye

Retrobulbar, Inflammation, Chronic Active,
Focal

Urinary System

Kidney

+ + + + + + + + + + + + + + + + 15

Renal Tubule, Dilatation, Diffuse

2 2 2 2.0

Renal Tubule, Dilatation, Focal

2 2 2 2.0

END OF MALE DATA

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

1-4 ..Lesion qualified as:

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Lab: MBA

| FVB/N Mouse Female
VEHICLE CONTROL | DAY ON TEST | | | | | | | | | | | | | | | |
|---|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 |
| | | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 |
| | | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 |

*TOTALS

Alimentary System

| | | | | | | | | | | | | | | | | |
|-----------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----|----|
| Intestine Large, Cecum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Intestine Large, Colon | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Intestine Large, Rectum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Intestine Small, Duodenum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Intestine Small, Ileum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Intestine Small, Jejunum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Liver | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Hepatocyte, Necrosis, Focal | 1 | 1 | 1 | 1 | 1 | 1 | 1 | | | | | 1 | | 8 | 1.0 | |
| Stomach, Forestomach | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Stomach, Glandular | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |

Cardiovascular System

| | | | | | | | | | | | | | | | | |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|
| Heart | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|

Endocrine System

| | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|----|-----|---|-----|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Subcapsular, Hyperplasia, Focal | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 2 | | | 10 | 1.1 | | |
| Zona Reticul, Vacuolization Cytoplasmic,
Diffuse | | | 2 | 2 | | | | | 4 | 3 | | | 4 | 2.8 | | |
| Zona Reticul, Vacuolization Cytoplasmic,
Focal | 2 | 2 | | | | 2 | 1 | 2 | | | 2 | 1 | 2 | | 8 | 1.8 |
| Adrenal Medulla | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Pituitary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

Experiment Number: 97011-23
Test Type: 26-WEEK
Route: SKIN APPLICATION
Species/Strain: Mouse/FVB/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL
Test Compound: Transgenic model evaluation (Melphalan)
CAS Number: 148-82-3

Date Report Requested: 10/20/2014
Time Report Requested: 04:55:34
First Dose M/F: NA / NA
Lab: MBA

| FVB/N Mouse Female
VEHICLE CONTROL | DAY ON TEST | | | | | | | | | | | | | | | | *TOTALS |
|---|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|----------------|
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | |
| | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | |
| | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | |
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | ANIMAL ID | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | |
| | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 | 9 | |
| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | | |
| Thyroid Gland | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| General Body System | | | | | | | | | | | | | | | | | |
| NONE | | | | | | | | | | | | | | | | | |
| Genital System | | | | | | | | | | | | | | | | | |
| Ovary | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Necrosis | | | | | | | | | | | | | | | | 1 | 4.0 |
| Rete Ovarii, Cyst | | | | | | | | | | | | | | | | 1 | 4.0 |
| Uterus | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Endometrium, Hyperplasia, Cystic | | 1 | 1 | 1 | 1 | 1 | 1 | | 1 | 3 | 1 | 2 | 1 | | | 10 | 1.3 |
| Hydrometra | | | | | | | | | 3 | | 4 | | | | | 3 | 3.0 |
| Hematopoietic System | | | | | | | | | | | | | | | | | |
| Bone Marrow | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Lymph Node, Mandibular | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Lymph Node, Mediastinal | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Lymph Node, Mesenteric | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Spleen | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Hematopoietic Cell Proliferation | | 2 | 2 | 2 | 2 | 2 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 15 | 1.8 |
| Pigmentation | | 2 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 15 | 1.1 |
| Thymus | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Integumentary System | | | | | | | | | | | | | | | | | |
| Mammary Gland | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

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Experiment Number: 97011-23

Test Type: 26-WEEK

Route: SKIN APPLICATION

Species/Strain: Mouse/FVB/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Transgenic model evaluation (Melphalan)

CAS Number: 148-82-3

Date Report Requested: 10/20/2014

Time Report Requested: 04:55:34

First Dose M/F: NA / NA

Lab: MBA

**FVB/N Mouse Female
VEHICLE CONTROL**

| DAY ON TEST | | | | | | | | | | | | | | | |
|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|
| | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 |
| 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 9 | 9 | 9 | 9 | 9 | 9 |
| 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | * |

***TOTALS**

| | | | | | | | | | | | | | | | |
|------------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|
| Mammary Gland | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Skin | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Epidermis, SOA, Hyperplasia, Focal | | | | | | | | | | | | | | | 1 1.0 |

Musculoskeletal System

| | | | | | | | | | | | | | | | |
|------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| NONE | | | | | | | | | | | | | | | |
|------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|

Nervous System

| | | | | | | | | | | | | | | | |
|------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| NONE | | | | | | | | | | | | | | | |
|------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|

Respiratory System

| | | | | | | | | | | | | | | | |
|------------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|
| Lung | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Alveolar Epith, Hyperplasia, Focal | | | | | | | | | | | | | | | 2 1.0 |

Special Senses System

| | | | | | | | | | | | | | | | |
|-----------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----------|
| NONE | | | | | | | | | | | | | | | |
| Urinary System | | | | | | | | | | | | | | | |
| Kidney | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Renal Tubule, Dilatation, Diffuse | | | | | | | | | | | | | | | 2 1.0 |
| Renal Tubule, Dilatation, Focal | | | | | | | | | | | | | | | 1 1.0 |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

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- 3) Moderate
- 2) Mild
- 4) Marked

Experiment Number: 97011-23
Test Type: 26-WEEK
Route: SKIN APPLICATION
Species/Strain: Mouse/FVB/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL
Test Compound: Transgenic model evaluation (Melphalan)
CAS Number: 148-82-3

Date Report Requested: 10/20/2014
Time Report Requested: 04:55:34
First Dose M/F: NA / NA
Lab: MBA

| FVB/N Mouse Female
4.0 MG/KG | DAY ON TEST | 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
|---|-------------|---------------------------------|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|--|
| | | 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 8 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 5 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| | | 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| ANIMAL ID | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 0 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| 1 1 1 1 2 2 2 2 2 2 2 2 2 2 2 2 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| 9 9 9 9 0 0 0 0 0 0 0 0 0 0 0 0 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |
| 6 7 8 9 0 1 2 3 4 5 6 7 8 9 0 0 | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | | |

*TOTALS

Alimentary System

| | | | | | | | | | | | | | | | | | |
|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---------|
| Intestine Large, Cecum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Intestine Large, Colon | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Intestine Large, Rectum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Intestine Small, Duodenum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Epithelium, Necrosis | 3 | 3 | 3 | 2 | 3 | 2 | 3 | 2 | 3 | 3 | 3 | 3 | 3 | 2 | 3 | 2 | 15 2.7 |
| Intestine Small, Ileum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Epithelium, Necrosis | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 1 | 2 | 1 | 3 | 2 | 2 | 2 | 2 | 2 | 15 1.9 |
| Intestine Small, Jejunum | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Epithelium, Necrosis | 2 | 2 | 3 | 2 | 3 | 3 | 2 | 2 | 2 | 3 | 3 | 2 | 2 | 3 | 3 | 3 | 15 2.5 |
| Liver | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Hepatocyte, Necrosis, Focal | | | | | | | | | | | | | | | | | 1 1.0 |
| Hepatocyte, Vacuolization Cytoplasmic,
Focal | | | | | | | | | | | | | | | | | 2 1 2.0 |

Cardiovascular System

| | | | | | | | | | | | | | | | | | |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|
| Heart | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
|-------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|----|

Endocrine System

| | | | | | | | | | | | | | | | | | |
|--------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------|
| Adrenal Cortex | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 14 |
| Hypertrophy, Focal | | | | | | | | | | | | | | | | | 1 1.0 |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

X ..Lesion present

A ..Autolysis precludes evaluation

I ..Insufficient tissue

BLANK ..Not examined microscopically

1-4 ..Lesion qualified as:

1) Minimal 3) Moderate

2) Mild 4) Marked

Experiment Number: 97011-23

Test Type: 26-WEEK

Route: SKIN APPLICATION

Species/Strain: Mouse/FVB/N

P09: NON-NEOPLASTIC LESIONS BY INDIVIDUAL ANIMAL

Test Compound: Transgenic model evaluation (Melphalan)

CAS Number: 148-82-3

Date Report Requested: 10/20/2014

Time Report Requested: 04:55:34

First Dose M/F: NA / NA

Lab: MBA

| FVB/N Mouse Female
4.0 MG/KG | DAY ON TEST | | | | | | | | | | | | | | | | |
|---|-------------|---------|---|---|---|---|---|---|---|---|---|---|---|---|----|--------|--|
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | |
| | | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | |
| | | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | |
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | |
| | | 1 | 1 | 1 | 1 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | 2 | |
| | | 9 | 9 | 9 | 9 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 1 | |
| | | 6 | 7 | 8 | 9 | 0 | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 0 | |
| | | | | | | | | | | | | | | | | | |
| | | *TOTALS | | | | | | | | | | | | | | | |
| Subcapsular, Hyperplasia, Focal | | | | | | 1 | | | 1 | | | | | | 3 | 1.0 | |
| Zona Reticul, Vacuolization Cytoplasmic,
Diffuse | | | 2 | | | | | | | 2 | | | | | 2 | 2.0 | |
| Zona Reticul, Vacuolization Cytoplasmic,
Focal | | | | | 2 | 2 | 1 | 1 | 1 | | 2 | 1 | 1 | 2 | 10 | 1.4 | |
| Adrenal Medulla | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Pituitary Gland | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Thyroid Gland | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| General Body System | | | | | | | | | | | | | | | | | |
| NONE | | | | | | | | | | | | | | | | | |
| Genital System | | | | | | | | | | | | | | | | | |
| Ovary | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Periovary Tiss, Hemorrhage, Focal | | | | | | | | | | | 1 | | | | 1 | 1.0 | |
| Uterus | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Endometrium, Hyperplasia, Cystic | | 1 | 2 | 2 | 2 | 1 | | 2 | 2 | 2 | 1 | 1 | 1 | | 11 | 1.5 | |
| Hematopoietic System | | | | | | | | | | | | | | | | | |
| Bone Marrow | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Lymph Node, Mandibular | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Lymph Node, Mediastinal | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Lymph Node, Mesenteric | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Spleen | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 | |
| Hematopoietic Cell Proliferation | | 2 | 2 | 2 | 2 | 2 | 2 | 3 | 3 | 3 | 2 | 3 | 3 | 3 | 3 | 15 2.5 | |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

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Experiment Number: 97011-23

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CAS Number: 148-82-3

Date Report Requested: 10/20/2014

Time Report Requested: 04:55:34

First Dose M/F: NA / NA

Lab: MBA

| FVB/N Mouse Female
4.0 MG/KG | DAY ON TEST | | | | | | | | | | | | | | | |
|---|-------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-----|
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| | ANIMAL ID | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| | | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 | 8 |
| | | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 | 5 |
| | | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 |
| Pigmentation | | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 | 1 |
| Thymus | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Atrophy, Focal | | | | | | | | 1 | | | | | | | 1 | 2 |
| Integumentary System | | | | | | | | | | | | | | | | 1.0 |
| Mammary Gland | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Skin | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Musculoskeletal System | | | | | | | | | | | | | | | | |
| NONE | | | | | | | | | | | | | | | | |
| Nervous System | | | | | | | | | | | | | | | | |
| NONE | | | | | | | | | | | | | | | | |
| Respiratory System | | | | | | | | | | | | | | | | |
| Lung | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Alveolar Epith, Hyperplasia, Focal | | 2 | | | | | | | 3 | 1 | 1 | 1 | 1 | 1 | 1 | 7 |
| Alveolus, Inflammation, Chronic Active, Focal | | | | | | | | | 1 | | 1 | | | | | 2 |
| Special Senses System | | | | | | | | | | | | | | | | |
| NONE | | | | | | | | | | | | | | | | |
| Urinary System | | | | | | | | | | | | | | | | |
| Kidney | | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 15 |
| Renal Tubule, Dilatation, Focal | | | | | | | | | 1 | | | | | | | 1 |
| | | | | | | | | | | | | | | | | 1.0 |

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

+ ..Tissue examined microscopically

M ..Missing tissue

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Experiment Number: 97011-23

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First Dose M/F: NA / NA

Lab: MBA

** END OF REPORT **

* ..Total animals with tissue examined microscopically; Total animals with lesion and mean severity grade

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