

APPENDIX 1B

Mouse Carcinogenicity Study
Histopathology - Non-neoplastic Lesion Data

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APPEARS THIS WAY
ON ORIGINAL

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SUMMARY TABLES

PAGE H : 19/ 937

STUDY NO. : KM8929

TEST ARTICLE : B8610-023
 TEST SYSTEM : MOUSE, CARCINOGENICITY STUDY,
 SPONSOR : BYK GULDEN, HAMBURG

PATHOL. NO.: 92003
 DATE : 25-NOV-92
 PATHDATA SYSTEM V3.5d

NUMBER OF ANIMALS WITH NON-NEOPLASTIC LESIONS BY ORGAN/GROUP/SEX
 STATUS AT NECROPSY: KO, INCL. +

ORGAN/FINDING	SEX : DOSE GROUP: NO. ANIMALS:	K1	K2	N	M	H	MALE
PAROTID GLANDS	NO. EXAM.:	50	50	50	50	50	
- ROUND CELL INFILTR.		2	1				
SUBMANDIBULAR GLANDS	NO. EXAM.:	50	50	50	50	50	
- ROUND CELL INFILTR.		33	32	31	40	19	
CERVICAL LYMPH NODES	NO. EXAM.:	50	50	50	50	49	
- LYMPHOID HYPERPLASIA			2	2	3	1	
- HYPEREMIA		1	2			1	
- SINUSOIDAL DILATION						1	
LACRIMAL GLANDS	NO. EXAM.:	50	49	50	50	50	
- ROUND CELL INFILTR.		17	7	9	8	3	
PROSTATE	NO. EXAM.:	50	50	49	49	48	
- PROSTATITIS			1				
- ATROPHY		1		2			
SEMINAL VESICLES	NO. EXAM.:	50	50	50	50	50	
- DISTENSION			3	1		3	
- ATROPHY			1	3		1	
- VESICULITIS		1		1	3	2	
TESTES	NO. EXAM.:	50	50	50	50	50	
- TUBULAR ATROPHY		6	2	4	5	3	
- TUBULAR CALCIFICAT.				2			
EPIDIDYMIDES	NO. EXAM.:	50	50	50	50	50	
- ARTERITIS/PERIARTER.		1					
- ASPERMIA		2		3	3		
- SPERMATOCELE		1	1		1		

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TEST ARTICLE : B8610-023
TEST SYSTEM : MOUSE, CARCINOGENICITY STUDY,
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PATHOL. NO.: 92003
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PATHDATA SYSTEM V3.5d

NUMBER OF ANIMALS WITH NON-NEOPLASTIC LESIONS BY ORGAN/GROUP/SEX
STATUS AT NECROPSY: KO, INCL. +

ORGAN/FINDING	SEX :						MALE
	DOSE GROUP:	K1	K2	N	M	H	
	NO. ANIMALS:	50	50	50	50	50	
KIDNEYS	NO. EXAM.:	50	50	50	50	50	
- ROUND CELL INFILTR.		21	24	20	21	19	
- GLOMERULAR CYST(S)		9	7	3	5		
- CORTICAL CALCIFICAT.		15	17	15	15	9	
- TUBULAR ATROPHY		11	11	6	5	3	
- TUBULAR DILATION						5	
- GLOMERULONEPHROPATHY			3	1	1	2	
- TUBULAR PIGMENT					1		
- ARTERITIS/PERIARTER.					1		
- PROTEINACEOUS CASTS		1			1		
- OSSEOUS METAPLASIA						1	
- INFARCT		2		1			
- PELVIC DILATION			2				
.....							
URINARY BLADDER	NO. EXAM.:	50	50	50	50	50	
- ROUND CELL INFILTR.		3	10	8	4	3	
- DISTENSION		12	19	16	15	12	
- UROTHEL. HYPERPLASIA			1			1	
- CYSTITIS			1				
- COLLOID PLUG			1				
.....							
MESENT. LYMPH NODE	NO. EXAM.:	49	48	50	49	50	
- HYPEREMIA		32	34	31	33	21	
- LYMPHOID HYPERPLASIA		11	7	11	11	9	
- SINUSOIDAL DILATION		2	3				
- GRANULOCYTOSIS					1		
.....							
ADRENAL GLANDS	NO. EXAM.:	50	50	50	50	50	
- A-CELL PROLIFERATION		37	39	35	42	38	
- ALTERED CELL FOCUS		15	11	10	10	8	
- B-CELL PROLIFERATION		7	4	4	12	5	
- MEDULLARY HYPERPLAS.			1				
- NECROSIS			1				
- AMYLOIDOSIS		1					
.....							

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TEST ARTICLE : B8610-023
TEST SYSTEM : MOUSE, CARCINOGENICITY STUDY,
SPONSOR : BYK GULDEN, HAMBURG
PATHOL. NO.: 92003
DATE : 25-NOV-92
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STATUS AT NECROPSY: KO, INCL. +

ORGAN/FINDING	SEX :						MALE
	DOSE GROUP:	K1	K2	N	M	H	
	NO. ANIMALS:	50	50	50	50	50	
PANCREAS	NO. EXAM.:	50	50	50	50	49	
- EDEMA				3	1	1	
- PANCREATITIS					1	1	
- CYSTIC DUCT				1			
- ISLET CELL HYPERPL.		2	2	1	2		
SPLEEN	NO. EXAM.:	50	50	50	50	50	
- LYMPHOID HYPERPLASIA		2			3	1	
- INCR. HEMATOPOIESIS		7	7	5	8	12	
- AMYLOIDOSIS		1					
- ATROPHY						2	
LIVER	NO. EXAM.:	50	50	50	50	50	
- KUPFFER CELL PROLIF.		6	9	10	9	26	
- SINGLE CELL NECROSIS		2	6	9	7	8	
- ROUND CELL INFILTR.		12	11	14	16	15	
- PATCHY NECROSIS		3	3	3	1	9	
- BASOPHILIC FOCUS		2	1	1	1	3	
- HEPATOC. HYPERPLASIA		11	8	10	13	13	
- FATTY CHANGE		1	1	3		8	
- INCREASED GLYCOGEN		3	8	1	1	1	
- HYPERTROPHY			3	3	10	6	
- PIGMENT ACCUMULAT.			1	1		1	
- NECROSIS, CENTRILOB.					1	5	
- GLYCOGEN FOCUS						1	
- EOSINOPHILIC FOCUS					1	1	
- LOBAR NECROSIS			1	1	1		
- HEMORRHAGE		1		1	1		
- SINUSOIDAL ECTASIA		3			1		
- HEMATOPOIESIS					1	1	
- INCREASED MITOSES		1	1		2	2	
- CONGESTION		4	4	3	2	2	
- INFLAMMATORY FOCI						1	
- AMYLOIDOSIS		1					
GALLBLADDER	NO. EXAM.:	50	50	50	50	50	
- DISTENSTION		1	1	2	3	5	

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TEST ARTICLE : B8610-023
TEST SYSTEM : MOUSE, CARCINOGENICITY STUDY,
SPONSOR : BYK GULDEN, HAMBURG

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NUMBER OF ANIMALS WITH NON-NEOPLASTIC LESIONS BY ORGAN/GROUP/SEX
STATUS AT NECROPSY: KO, INCL. +

ORGAN/FINDING	SEX :						MALE
	DOSE GROUP:	K1	K2	N	M	H	
	NO. ANIMALS:	50	50	50	50	50	
ESOPHAGUS	NO. EXAM.:	49	50	50	50	50	
- INFLAMMATION						1	
SKIN	NO. EXAM.:	50	50	49	49	50	
- EDEMA		2	1		1		
- INFLAMMATION			2			1	
- PENIS INFLAMMATION		2		1			
MAMMARY GLAND AREA	NO. EXAM.:	50	50	49	49	50	
- EDEMA		1			1		
- HEMORRHAGE				1			
SKELETAL MUSCLE	NO. EXAM.:	50	50	50	49	50	
- ATROPHY		4	2	6	3	5	
- HYALINE NECROSIS		1	1		3		
- ROUND CELL INFILTR.		1	2	1	1		
- SARCOsporIDIOSIS			1				
THYROID GLAND	NO. EXAM.:	50	49	50	50	50	
- CYSTIC FOLLICLES		8	9	9	8	8	
- AMYLOIDOSIS		1					
PARATHYROID GLANDS	NO. EXAM.:	48	45	45	45	42	
- CYSTS		2	2	1	1		
- AMYLOIDOSIS		1					
TIBIAL NERVE	NO. EXAM.:	48	48	50	50	43	
- DEGENER. NEUROPATHY		31	30	21	22	14	
THYMUS	NO. EXAM.:	47	46	50	47	45	
- INVOLUTION		26	24	33	31	27	
EYES	NO. EXAM.:	50	50	50	50	50	
- PETHISIS BULBI		1	1				
- KERATITIS		1					
- LENTICUL. DEGENERAT.						1	

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TEST ARTICLE : B8610-023
TEST SYSTEM : MOUSE, CARCINOGENICITY STUDY,
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PATHOL. NO.: 92003
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PATHDATA SYSTEM V3.5d

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STATUS AT NECROPSY: KO, INCL. +

ORGAN/FINDING	SEX :						MALE
	DOSE GROUP:	K1	K2	N	M	H	
	NO. ANIMALS:	50	50	50	50	50	
HARDERIAN GLANDS	NO. EXAM.:	49	50	49	50	50	
- DACRYOADENITIS		10	15	13	11	5	
BRAIN	NO. EXAM.:	50	50	50	50	50	
- CALCIFICATION		23	23	24	24	19	
- HEMORRHAGE		2					
- COMPRESSION		1					
PITUITARY GLAND	NO. EXAM.:	49	49	46	50	49	
- CYSTS		3	4	2	3		
- CONGESTION		1				1	
- HYPERPLASIA				1			
HEART	NO. EXAM.:	50	50	50	50	50	
- DISTENDED CHAMBERS		3	5	5	2	10	
- ARTERITIS/PERIARTER.				1			
- THROMBOENDOCARDITIS		2			1		
- CALCIFICATION		1					
LUNGS	NO. EXAM.:	50	50	50	50	50	
- EMPHYSEMA		3	3	1	2	1	
- CONGESTION		5	6	5	2	8	
- LIPOPROTEINOSIS		2	2	1		1	
- HEMORRHAGE, ALVEOLAR		1	3	3	2	1	
- ATELECTASIS			1		1	1	
- PERIVASCULAR CUFFING			3	2			
- INTERSTIT. PNEUMONIA				1	1		
- GRANULOCYTOSIS						2	
STOMACH	NO. EXAM.:	50	50	50	50	50	
- FUNDIC HYPERPLASIA				9	49	50	
- ULCER(S)		1		2	1	2	
- EROSION(S)		2	2	1			
- FOCAL HYPERKERATOSIS				2	1		
- GLANDULAR ECTASIA		7	8	4			
- GLANDULAR HYPERPLAS.		2	2	3			

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NUMBER OF ANIMALS WITH NON-NEOPLASTIC LESIONS BY ORGAN/GROUP/SEX
STATUS AT NECROPSY: KO, INCL. +

ORGAN/FINDING	SEX :	MALE				
	DOSE GROUP:	K1	K2	N	M	H
	NO. ANIMALS:	50	50	50	50	50
SMALL INTESTINE	NO. EXAM.:	50	50	50	50	50
- LYMPHOID HYPERPLASIA		1	3	3	2	1
LARGE INTESTINE	NO. EXAM.:	50	50	50	50	50
- NEMATODES IN LUMEN		1			3	1
BONE	NO. EXAM.:	50	50	50	50	50
- CALLUS		1		1	1	2
- ARTHROPATHY		2				
- STERNAL DEVIATION		2	3		3	3
- CYST		1				
NOSE	NO. EXAM.:	50	49	49	50	49
- RHINITIS					1	
ORAL CAVITY	NO. EXAM.:			2		1
- INFLAMMATION				2		1
LYMPH NODES	NO. EXAM.:	4	9	6	6	7
- LYMPHOID HYPERPLASIA		1		2	2	1
- HYPEREMIA					1	
BODY CAVITIES	NO. EXAM.:		1	2	1	2
- HEMORRHAGE			1			
BONE MARROW	NO. EXAM.:	50	50	50	50	42
- HYPERCELLULAR MARROW		2	4	9	3	4
- HYPOCELLULAR MARROW				1		
PREPUTIAL GLANDS	NO. EXAM.:			1		
- CYSTIC CHANGE				1		
- HYPERPLASIA				1		
ADIPOSE TISSUE	NO. EXAM.:	2	2	1	2	4
- FAT NECROSIS		1				2
- ANGIOECTASIS					1	

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STUDY NO. : KM8929

TEST ARTICLE : BB610-023
TEST SYSTEM : MOUSE, CARCINOGENICITY STUDY,
SPONSOR : BYK GULDEN, HAMBURG

PATHOL. NO.: 92003
DATE : 25-NOV-92
PATHDATA SYSTEM V3.5d

NUMBER OF ANIMALS WITH NON-NEOPLASTIC LESIONS BY ORGAN/GROUP/SEX
STATUS AT NECROPSY: KO, INCL. +

ORGAN/FINDING	SEX :	K1	K2	N	M	H	FEMALE
	DOSE GROUP: NO. ANIMALS:						
PAROTID GLANDS	NO. EXAM.:	50	50	50	50	50	
- ROUND CELL INFILTR.		6	3		1	3	
- ACINAR HYPERPLASIA		1					
- INTERSTITIAL EDEMA			1		1		
SUBMANDIBULAR GLANDS	NO. EXAM.:	50	50	50	50	50	
- ROUND CELL INFILTR.		28	38	38	31	33	
- SIALOADENITIS				1			
- INTERSTITIAL EDEMA			1		1		
CERVICAL LYMPH NODES	NO. EXAM.:	50	50	48	48	50	
- LYMPHOID HYPERPLASIA		2	5	8	2	3	
- HYPEREMIA		1		2	1	1	
LACRIMAL GLANDS	NO. EXAM.:	50	50	50	50	50	
- ROUND CELL INFILTR.		14	25	23	22	16	
- CYST(S)				1			
- DACRYOADENITIS		2		1			
- METAPL. HARDER. GLD.				1	2	1	
UTERUS	NO. EXAM.:	50	50	50	50	50	
- CYSTIC GLAND. HYPERPL		41	44	43	42	34	
- DISTENSION		3	4	6	9	10	
- HEMORRHAGE		1					
- PYOMETRA		1	1			1	
- ATROPHY					1		
OVARIES	NO. EXAM.:	50	50	50	50	50	
- CYST(S)		29	29	29	25	27	
- ABSCESS		1			1	1	
- ATROPHY			1				
- HEMORRHAGE			1		1		
- ANGIOMATOUS CHANGE						1	
VAGINA	NO. EXAM.:	50	50	50	50	48	
- INFLAMMATION		1	1				

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ORGAN/FINDING	SEX DOSE GROUP: NO. ANIMALS:	K1	K2	N	M	H	FEMALE
KIDNEYS	NO. EXAM.:	50	50	50	50	50	
- ROUND CELL INFILTR.		33	38		26	35	
- GLOMERULAR CYST(S)				1			
- CORTICAL CALCIFICAT.							2
- GLOMERULONEPHROPATHY		1		1	1		1
- ARTERITIS/PERIARTER.							1
- PROTEINACEOUS CASTS		1	1				
- AMYLOIDOSIS			2				
- HYAL. RESORP. BODIES					1		
- CELL CYLINDERS							1
- PELVIC DILATION			1		1		
- CONGESTION		1					
URINARY BLADDER	NO. EXAM.:	50	49	49	49	50	
- ROUND CELL INFILTR.		31	31	25	25	22	
- DISTENSION		16	10	11	11	11	
- EDEMA							1
MESENT. LYMPH NODE	NO. EXAM.:	48	50	48	47	47	
- HYPEREMIA		8	7	4	10	17	
- LYMPHOID HYPERPLASIA		10	6	5	7	5	
ADRENAL GLANDS	NO. EXAM.:	50	49	50	49	50	
- A-CELL PROLIFERATION		48	49	50	47	49	
- CEROID DEGENERATION		39	44	49	43	44	
- ALTERED CELL FOCUS		3	4	1			
- B-CELL PROLIFERATION			1	1			
- X-ZONE DEGENERATION		4	1			1	
- MEDULLARY HYPERPLAS.						1	
- CORTICAL CYST(S)			1	1			3

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ORGAN/FINDING	SEX : DOSE GROUP: NO. ANIMALS:	K1	K2	N	M	H	FEMALE
PANCREAS	NO. EXAM.:	50	49	50	50	50	
- EDEMA		1		1	1	1	
- PANCREATITIS		1					
- CYSTIC DUCT		1	3	2	2	3	
- EXOCRINE HYPERPLASIA							1
- ACINAR ATROPHY				2			
- ISLET CELL HYPERPL.		1					
.....							
SPLEEN	NO. EXAM.:	50	50	50	50	50	
- LYMPHOID HYPERPLASIA		10	12	12	11	10	
- INCR. HEMATOPOIESIS		21	10	5	13	20	
- CAPSULAR FIBROSIS		1				1	
.....							
LIVER	NO. EXAM.:	50	50	50	50	50	
- KUPFFER CELL PROLIF.		13	16	9	11	22	
- SINGLE CELL NECROSIS		6	8	13	10	21	
- ROUND CELL INFILTR.		26	26	24	24	24	
- PATCHY NECROSIS		2	1	2	4	8	
- BASOPHILIC FOCUS			2				
- HEPATOC. HYPERPLASIA		3	3	2	2	7	
- FATTY CHANGE		4	1	1	2	9	
- INCREASED GLYCOGEN		8	1	3		1	
- HYPERTROPHY				1	2	1	
- PIGMENT ACCUMULAT.			3	5	7	2	
- NECROSIS, CENTRILOB.			1			1	
- VACUOLATED FOCUS						1	
- GLYCOGEN FOCUS						1	
- EOSINOPHILIC FOCUS		1			1		
- LOBAR NECROSIS			1				
- HEMORRHAGE					2		
- CALCIFICATION			1				
- SINUSOIDAL ECTASIA		3	3	1	3	3	
- HEMATOPOIESIS		1	3	1	3	2	
- CAPSULAR FIBROSIS		1					
- INCREASED MITOSES		1					
- BILIARY CYST		2	1				
- CONGESTION		1	3	1	3	6	
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ORGAN/FINDING	SEX : DOSE GROUP: NO. ANIMALS:	K1	K2	N	M	H	FEMALE
GALLBLADDER	NO. EXAM.:	50	50	50	50	50	
- DISTENSION		2	4	3	5	1	
- ROUND CELL INFILTR.			1		2		
- EPITHELIAL HYPERPL.		1					
SKIN	NO. EXAM.:	49	50	48	50	49	
- EDEMA		1			1	1	
- ADNEXAL ATROPHY					1		
- INFLAMMATION			1	1	4		
- SQUAMOUS CYST(S)		1					
- TAIL NECROSIS					1		
MAMMARY GLAND AREA	NO. EXAM.:	49	50	48	50	49	
- LACTEAL CYSTS		11	11	16	14	14	
SKELETAL MUSCLE	NO. EXAM.:	50	49	50	50	50	
- ATROPHY		16	14	10	9	13	
- HYALINE NECROSIS		2	2	3			
- ROUND CELL INFILTR.		4	6	7	7	2	
- CALCIFICATION		1					
- ARTERITIS/PERIARTER.							1
THYROID GLAND	NO. EXAM.:	50	50	50	50	50	
- CYSTIC FOLLICLES		21	21	18	13	17	
- FOLLICULAR HYPERPL.		6	3	1	1	1	
PARATHYROID GLANDS	NO. EXAM.:	47	48	49	48	45	
- CYSTS			1		1		
TIBIAL NERVE	NO. EXAM.:	49	47	48	45	44	
- DEGENER. NEUROPATHY		30	36	38	30	30	
TONGUE	NO. EXAM.:	50	50	50	50	50	
- ARTERITIS/PERIARTER.							1

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TEST ARTICLE : B8610-023
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STATUS AT NECROPSY: KO, INCL. +

ORGAN/FINDING	SEX :						FEMALE
	DOSE GROUP:	K1	K2	N	M	H	
	NO. ANIMALS:	50	50	50	50	50	
THYMUS	NO. EXAM.:	48	49	49	46	47	
- INVOLUTION		15	18	22	19	20	
- HEMORRHAGE			1	2		1	
EYES	NO. EXAM.:	50	50	50	50	50	
- PHTHISIS BULBI			1				
- LENTICUL. DEGENERAT.					1		
HARDERIAN GLANDS	NO. EXAM.:	50	50	50	50	50	
- DACRYOADENITIS		29	26	20	17	18	
BRAIN	NO. EXAM.:	49	50	50	50	50	
- CALCIFICATION		13	20	16	17	10	
- HEMORRHAGE				1			
- COMPRESSION			1		2		
PITUITARY GLAND	NO. EXAM.:	49	50	49	50	50	
- CYSTS		2			1	1	
- CONGESTION			1			1	
- ANGIECTASIS		1	2	2	4		
- HYPERPLASIA		2	2	3	2	1	
HEART	NO. EXAM.:	50	50	50	50	50	
- DISTENDED CHAMBERS		4	3		4	10	
- ARTERITIS/PERIARTER.			1			2	
- FIBROSIS						1	
- EPICARDITIS						1	
- THROMBOENDOCARDITIS		3		1		2	
- CALCIFICATION		1					
- ROUND CELL INFILTR.						1	

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PATHOLOGY REPORT 42/92
SUMMARY TABLES

PAGE H : 30/ 937
STUDY NO. : KM8929

TEST ARTICLE : B8610-023
TEST SYSTEM : MOUSE, CARCINOGENICITY STUDY,
SPONSOR : BYE GULDEN, HAMBURG
PATHOL. NO.: 92003 ✓
DATE : 25-NOV-92
PATHDATA SYSTEM V3.5d

NUMBER OF ANIMALS WITH NON-NEOPLASTIC LESIONS BY ORGAN/GROUP/SEX
STATUS AT NECROPSY: KO, INCL. +

ORGAN/FINDING	SEX :	FEMALE				
	DOSE GROUP:	K1	K2	N	M	H
	NO. ANIMALS:	50	50	50	50	50
LUNGS	NO. EXAM.:	50	50	50	50	50
- GRANULOMA(S)		1		2		
- EMPHYSEMA		1	2	5	2	
- CONGESTION		8	4	1	5	11
- LIPOPROTEINOSIS					2	
- HEMORRHAGE, ALVEOLAR		6	4	3	4	4
- ATELECTASIS			1		2	1
- PERIVASCULAR CUFFING		4	3	1	2	1
- GRANULOCYTOSIS					1	
STOMACH	NO. EXAM.:	50	50	50	50	50
- FUNDIC HYPERPLASIA				24	50	50
- ULCER(S)		4	1			3
- ROUND CELL INFILTR.		2				
- EROSION(S)				2		
- FOCAL HYPERKERATOSIS		5		2		4
- GLANDULAR ECTASIA		2	6	5		
- INFLAMMATION		1				
- GLANDULAR HYPERPLAS.		4	3	5	1	
SMALL INTESTINE	NO. EXAM.:	50	50	50	50	50
- LYMPHOID HYPERPLASIA		1	1	4	4	2
- DIVERTICULUM					1	
LARGE INTESTINE	NO. EXAM.:	50	50	50	50	50
- ULCER(S)			1			
BONE	NO. EXAM.:	50	50	50	50	50
- CALLUS			1		1	
- ARTHROPATHY				1		
- STERNAL DEVIATION						1
- HEMORRHAGE					1	
- SCAR					1	
NOSE	NO. EXAM.:	49	50	50	50	50
- RHINITIS					3	

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PATHOLOGY REPORT 42/92
SUMMARY TABLES

PAGE H : 31/ 937
STUDY NO. : KM8929

TEST ARTICLE : B8610-023
TEST SYSTEM : MOUSE, CARCINOGENICITY STUDY,
SPONSOR : BYK GULDEN, HAMBURG

PATHOL. NO.: 92003 ✓
DATE : 25-NOV-92
PATHDATA SYSTEM V3.5d

NUMBER OF ANIMALS WITH NON-NEOPLASTIC LESIONS BY ORGAN/GROUP/SEX
STATUS AT NECROPSY: KO, INCL. +

ORGAN/FINDING	SEX :						FEMALE
	DOSE GROUP:	K1	K2	N	M	H	
	NO. ANIMALS:	50	50	50	50	50	
SPINAL CORD	NO. EXAM.:	50	50	50	50	50	
- ROUND CELL INFILTR.			1				
LYMPH NODES	NO. EXAM.:	10	6	9	16	10	
- LYMPHOID HYPERPLASIA			1	2	2	2	
- HYPEREMIA			1		1	2	
BONE MARROW	NO. EXAM.:	50	50	49	50	50	
- FIBRO-OSSEOUS LESION		20	26	24	21	18	
- HYPERCELLULAR MARROW		4	2	6	5	5	
- HYPOCELLULAR MARROW		1		1		2	
- GRANULOCYT. HYPERPL.			2			2	
ADIPOSE TISSUE	NO. EXAM.:	4	1	4	2	2	
- FAT NECROSIS		3	1	3	1	1	
- ANGIECTASIS						1	

APPEARS THIS WAY
ON ORIGINAL

APPENDIX 2A

**Sprague Dawley Rat Carcinogenicity Study
Histopathology - Tumor Data.**

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**APPEARS THIS WAY
ON ORIGINAL**

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PLACES Y8.204

Table 5

Summary of Primary Neoplasms

TUMOR	NUMERIC INCIDENCE OF NEOPLASTIC LESIONS											
	MALES						FEMALES					
	0	0.51	51	501	2001		0	0.51	51	501	2001	
GROUP	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day		mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	
BODY AS A WHOLE:												
THORACIC CAVITY:												
SQUAMOUS CELL CARCINOMA (M)		(3)	(3)	(6)	(11)			(1)	(3)	(6)		
ABDOMINAL CAVITY:												
SQUAMOUS CELL CARCINOMA (M)	(3)	(1)	(1)	(4)	(5)			(1)	(2)	(2)		
SCIRRHOUS (M)	1			1								
SCIRRHOUS CARCINOMA (M), metastasizing												
LEIOMYOSARCOMA (M), metastasizing												1
MALIGNANT LYMPHOMA, LYMPHOCTIC (M)									1			
HISTIOCYTIC SARCOMA (M)												
SARCOMA, NOS (M)				1	1							1
LIPOMA (B)												
INTODUMENTARY SYSTEM:-												
SKIN:												
SQUAMOUS CELL PAPILLOMA (B)	(70)	(64)	(42)	(66)	(70)		(70)	(55)	(50)	(46)	(70)	
SQUAMOUS CELL CARCINOMA (M)	3	4		7	4		2	1	1		2	
BASAL CELL TUMOR (B)									1			1

Figures in parentheses represent the number of animals from which this tissue was examined.

The absence of a numeral indicates that the lesion specified was not identified

(B) = benign, (M) = malignant

CG9024 | Study Completed

NDA 20,987
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PLACES VA.304

Table 5 (Cont'd)
Summary of Primary Neoplasms

TUMORS	NUMERIC INCIDENCE OF NEOPLASTIC LESIONS															
	GROUP	MALES						FEMALES								
		01 mg/kg /day	0.51 mg/kg /day	51 mg/kg /day	501 mg/kg /day	2001 mg/kg /day	(70)	(64)	(62)	(46)	(70)	01 mg/kg /day	0.51 mg/kg /day	51 mg/kg /day	501 mg/kg /day	2001 mg/kg /day
INTELGUMENTARY SYSTEM:-																
SKIN:																
BASAL CELL CARCINOMA (M)																
KERATOCARCINOMA/TRICHOEPITHELIOMA (B)																
SEBACEOUS ADENOMA (S)																
FIBROSA (B)																
FIBROSARCOMA (M)																
SCHWANNOMA (B)																
SCHWANNOMA (M)																
MYXOMA (B)																
HEMANGIOMA (B)																
LIPOMA (B)																
SARCOMA, NOS (M)																
LEIOMYOMA (B)																
HISTIOCYTIC SARCOMA (M)																
MYELOMONOCYTIC LEUKEMIA (M)																

Figures in parentheses represent the number of animals from which this tissue was examined.

The absence of a numeral indicates that the lesion specified was not identified
(B) = benign, (M) = malignant

CG9074 (Study Completed)

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PLACES V8.204

Table 5 (Cont'd)
Summary of Primary Neoplasms

TUMOR	NUMERIC INCIDENCE OF NEOPLASTIC LESIONS											
	MALES						FEMALES					
	0	0.5	5	50	200	200	0	0.5	5	50	200	200
GROUP	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day
INTEGUMENTARY SYSTEM:-												
SKIN:												
LEUKEMIA (M), lymphoid	(70)	(64)	(62)	(66)	(70)	(70)	(55)	(50)	(46)	(70)		
MAMMARY GLAND:				1								
FIBROADENOMA (B)	(43)	(36)	(36)	(32)	(38)	(68)	(63)	(65)	(63)	(67)		
CYSTADENOMA (B)	5	4	1	1	4	43	47	47	46	43		
ADENOMA (B), ductal		1				4	3	2	1	2		
ADENOCARCINOMA (M)	1	1				9	10	10	13	11		
CLITORAL GLAND:						(4)		(4)	(1)	(1)		
SQUAMOUS CELL PAPILLOMA (B)									1			
HEMATOPOIETIC AND LYMPHATIC SYSTEMS:-												
BONE MARROW:												
MYELOMONOCYTTIC LEUKEMIA (M)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)		
MALIGNANT LYMPHOMA, LYMPHOBLASTIC (M)	2						1	3	1	2		
HISTIOCYTIC SARCOMA (M)	1			2								

Figures in parentheses represent the number of animals from which this tissue was examined.
The absence of a numeral indicates that the lesion specified was not identified
(B) - benign, (M) - malignant

CS9034 (Study Completed)

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PLACES V8.204

Table 3 (Cont'd)
Summary of Primary Neoplasms

TUMOR	NUMERIC INCIDENCES OF NEOPLASTIC LESIONS											
	MALES						FEMALES					
	01	0.51	51	501	2001		01	0.51	51	501	2001	
	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day		mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	
HEMATOPOIETIC and LYMPHATIC SYSTEMS:-												
SPLEEN:	(70)	(60)	(61)	(59)	(70)		(69)	(51)	(50)	(44)	(70)	
HEMANGIOMA (B)					1						1	
HEMANGIOCARCOMA (M)				2								
HISTIOCYTIC SARCOMA (M)	1											
MYELOMONOCYTIC LEUKEMIA (M)	3							2			2	
LEUKEMIA (M), lymphoid				1					1			
THYMUS:	(55)	(54)	(47)	(35)	(57)		(50)	(42)	(44)	(29)	(60)	
THYMOMA (B)			1				1			1		
HISTIOCYTIC SARCOMA (M)												
MYELOMONOCYTIC LEUKEMIA (M)	1											
LEUKEMIA (M), lymphoid	(2)			(3)	(1)		(1)		(1)		(1)	
LYMPH NODE(S):				1								
HISTIOCYTIC SARCOMA (M)	1											
LEUKEMIA (M), lymphoid												
MYELOMONOCYTIC LEUKEMIA (M)	1										1	

figures in parentheses represent the number of animals from which this tissue was examined.

The absence of a numeral indicates that the lesion specified was not identified

(B) = benign, (M) = malignant

089024 (study completed)

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Table 5 (Cont'd)
Summary of Primary Neoplasms

TUMORS	GROUP	NUMERIC INCIDENCE OF NEOPLASTIC LESIONS												
		MALES					FEMALES							
		0	0.51	51	501	2001	0	0.51	51	501	2001			
		mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day
HEMATOPOIETIC and LYMPHATIC SYSTEMS:-														
LYMPH NODE - MANDIBULAR:		(69)	(56)	(60)	(55)	(69)	(69)	(46)	(47)	(41)	(70)			
MYELOMONOCYTIC LEUKEMIA (M)		2									1			
HISTIOCYTIC SARCOMA (M)		1				1				1				
LYMPH NODE - MESENTERIC:		(66)	(59)	(59)	(56)	(67)	(66)	(46)	(49)	(40)	(68)			
SCHWANNOMA (M)						1								
HEMANGIOMA (S)		1	1	2										
MYELOMONOCYTIC LEUKEMIA (M)		3									2			
HISTIOCYTIC SARCOMA (M)							1			1				
MUSCULOSKELETAL SYSTEM and SOFT TISSUES:-														
BONE:		(1)				(1)				(1)				
OSTEOSARCOMA (M), metastasizing		1				1				1				
SKELTAL MUSCLE:		(70)	(61)	(62)	(58)	(70)	(70)	(48)	(49)	(42)	(70)			
MALIGNANT LYMPHOMA, LYMPHOIDIC (M)														
HISTIOCYTIC SARCOMA (M)										1				
LEUKEMIA (M), Lymphoid					1									

Figures in parentheses represent the number of animals from which this tissue was examined.
The absence of a numeral indicates that the lesion specified was not identified
(B) - benign, (M) - malignant

089034 (Study Completed)

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PLACES V9.304

Table 5 (Cont'd)
Summary of Primary Neoplasms

TUMORS	NUMERIC INCIDENCE OF NEOPLASTIC LESIONS											
	MALES						FEMALES					
	01 mg/kg /day	0.51 mg/kg /day	51 mg/kg /day	501 mg/kg /day	2001 mg/kg /day		01 mg/kg /day	0.51 mg/kg /day	51 mg/kg /day	501 mg/kg /day	2001 mg/kg /day	
RESPIRATORY SYSTEM:-												
TRACHEA:	(70)	(61)	(61)	(58)	(70)		(70)	(48)	(49)	(42)	(70)	
HISTIOCYTIC SARCOMA (M)	(70)	(61)	(61)	(58)	(70)		(70)	(48)	(48)	(42)	(70)	
LUNG:												
BRONCHIOALVEOLAR ADENOMA (B)	2							1		1	2	
MYELOMONOCYTIC LEUKEMIA (M)	1			3			2	1		2		
HISTIOCYTIC SARCOMA (M)				1						1		
LEUKEMIA (M), lymphoid												
CARDIOVASCULAR SYSTEM:-												
HEART:	(70)	(61)	(61)	(58)	(70)		(70)	(48)	(49)	(42)	(70)	
SCHWANNOMA (B), subendocardial											1	
MYELOMONOCYTIC LEUKEMIA (M)											1	
HISTIOCYTIC SARCOMA (M)							1				1	
LEUKEMIA (M), lymphoid				1							1	

Figures in parentheses represent the number of animals from which this tissue was examined.

The absence of a numeral indicates that the lesion specified was not identified
(B) - benign, (M) - malignant

OS9024 (Study Completed)

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PLACES V8.204

Table 5 (Cont'd)
Summary of Primary Neoplasms

TUMORS	NUMERIC INCIDENCE OF NEOPLASTIC LESIONS											
	MALES						FEMALES					
	01	0.51	51	501	2001		01	0.51	51	501	2001	
GROUP	ng/kg /day	ng/kg /day	ng/kg /day	ng/kg /day	ng/kg /day		ng/kg /day	ng/kg /day	ng/kg /day	ng/kg /day	ng/kg /day	
CARDIOVASCULAR SYSTEM:-												
AORTA:	(70)	(61)	(58)	(55)	(69)		(70)	(48)	(49)	(40)	(69)	
HISTIOCYTIC SARCOMA (M)										1		
DIGESTIVE SYSTEM:-												
TONGUE:					(1)			(1)	(1)			
SQUAMOUS CELL PAPILLOMA (B)					1							
SALIVARY GLAND - PAROTIDULAR:	(70)	(40)	(41)	(56)	(70)		(70)	(46)	(48)	(42)	(70)	
MYELOMONOCYTIC LEUKEMIA (M)	1											
HISTIOCYTIC SARCOMA (M)										1		
SALIVARY GLAND - SUBLINGUAL:	(70)	(59)	(60)	(55)	(70)		(70)	(45)	(48)	(41)	(68)	
HISTIOCYTIC SARCOMA (M)								1				
LIVER:	(70)	(70)	(70)	(70)	(70)		(70)	(70)	(70)	(70)	(70)	
HEPATOCELLULAR ADENOMA (B)		2	1	7	9							
HEPATOCELLULAR CARCINOMA (M)	1	2	2	6	6							
NEUROENDOCRINE CELL TUMOR (M), site of origin not identified			1	1	1							
CHOLANGIOMA (B)					1							

Figures in parentheses represent the number of animals from which this tissue was examined.

The absence of a numeral indicates that the lesion specified was not identified
(B) = benign, (M) = malignant

089024 (Study Completed)

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PLACES V8.204

Table 5 (Cont'd)
Summary of Primary Neoplasms

TUMOR	NUMERIC INCIDENCE OF NEOPLASTIC LESIONS															
	MALES						FEMALES									
	GROUP	01	0.51	51	501	2001	01	0.51	51	501	2001	01	0.51	51	501	2001
	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day
DIGESTIVE SYSTEM:-																
LIVER,	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)	(70)
SARCOMA, NOS (M)				4	1											
HISTIOCYTIC SARCOMA (M)	1															
MYELOMONOCYTIC LEUKEMIA (M)	2															
LEUKEMIA (M), Lymphoid				1												
PANCREAS,	(59)	(61)	(62)	(58)	(70)	(70)	(70)	(50)	(51)	(62)	(70)	(70)	(70)	(70)	(70)	(70)
ISLET CELL ADENOMA (B)	6	3	1	4												
ISLET CELL CARCINOMA (M)	2		1	2	1											
ACINAR CELL ADENOMA (B)	4	2	1	2	1											
ACINAR CELL CARCINOMA (M)																
HISTIOCYTIC SARCOMA (M)	1			1												
MYELOMONOCYTIC LEUKEMIA (M)	2															
MALIGNANT LYMPHOMA, LYMPHOBLASTIC (M)																
LEUKEMIA (M), Lymphoid				1												

Figures in parentheses represent the number of animals from which this tissue was examined.

The absence of a numeral indicates that the lesion specified was not identified
(M) = benign, (M) = malignant

089024 (Study Completed)

PLACES V8.204

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PLACES V8.204

Table 5 (Cont'd)
Summary of Primary Neoplasms

TUMORS	NOMERIC INCIDENCE OF NEOPLASTIC LESIONS												
	GROUP	MALES						FEMALES					
		0 mg/kg /day	0.51 mg/kg /day	51 mg/kg /day	501 mg/kg /day	2001 mg/kg /day	(n)	0 mg/kg /day	0.51 mg/kg /day	51 mg/kg /day	501 mg/kg /day	2001 mg/kg /day	(n)
DIGESTIVE SYSTEM:-													
ESOPHAGUS,													
HISTIOCYTIC SARCOMA (M)	(70)	(61)	(61)	(57)	(69)		(70)	(46)	(46)	(42)	(70)		
STOMACH,													
HISTIOCYTIC SARCOMA (M)	(70)	(70)	(70)	(70)	(70)		(70)	(70)	(70)	(70)	(70)		
NEUROENDOCRINE CELL TUMOR (B), fundus													
NEUROENDOCRINE CELL TUMOR (M), fundus													
ADENOCARCINOMA (M), chief cell, fundus													
ADENOMATOUS POLYP (B), fundus													
ADENOCARCINOMA (M), pyloric region													
SQUAMOUS CELL PAPILLOMA (B)													
SQUAMOUS CELL CARCINOMA (M)													
MYELOMONOCYTIC LEUKEMIA (M)	1												
HISTIOCYTIC SARCOMA (M)	(70)	(61)	(62)	(56)	(69)		(70)	(52)	(49)	(42)	(70)		
DIVERGENT,													
HISTIOCYTIC SARCOMA (M)													

Figures in parentheses represent the number of animals from which this tissue was examined.

The absence of a numeral indicates that the lesion specified was not identified

(B) = benign, (M) = malignant

CG9024 (Study Completed)

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PLACES V6.206

Table 5 (Cont'd)
Summary of Primary Neoplasms

TUMORS	NUMERIC INCIDENCE OF NEOPLASTIC LESIONS														
	MALES						FEMALES								
GROUP	01	0.51	51	501	2001	01	0.51	51	501	2001	01	0.51	51	501	2001
	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day
DIGESTIVE SYSTEM:															
JEJUNUM:															
LEIOMYOSARCOMA (M), metastasizing	(68)	(61)	(62)	(58)	(70)	(69)	(48)	(49)	(62)	(69)					
MYELOMONOCYTIC LEUKEMIA (M)			1					1							
LEUKEMIA:						(70)	(48)	(48)	(42)	(70)					
LEIOMYOSARCOMA (M)															1
MYELOMONOCYTIC LEUKEMIA (M)								1							
COLON:						(69)	(61)	(62)	(57)	(70)	(70)	(48)	(48)	(42)	(70)
MYELOMONOCYTIC LEUKEMIA (M)															
HISTIOCYTIC SARCOMA (M)											1				
RECTUM:						(1)							(1)		
ADENOMA (B); hamorrhagic						1									
MESENTERY:						(2)	(3)	(2)	(2)	(2)	(1)	(1)	(2)	(2)	(2)
LIPOMA (B)															
MYELOMONOCYTIC LEUKEMIA (M)						1							1		

Figures in parentheses represent the number of animals from which this tissue was examined.

The absence of a numeral indicates that the lesion specified was not identified

(B) = benign, (M) = malignant

CS9024 (Study Completed)

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PLACES VP. 204

Table 5 (Cont'd)
Summary of Primary Neoplasms

TUMORS	NUMERIC INCIDENCE OF NEOPLASTIC LESIONS												
	MALES						FEMALES						
	0	0.51	51	501	2001		0	0.51	51	501	2001		
mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	(70)	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	mg/kg /day	(70)	mg/kg /day	
UROGENITAL SYSTEMS:-													
KIDNEY:													
TRANSITIONAL CELL CARCINOMA (H)													
SQUAMOUS CELL CARCINOMA (H)													
TUBULAR CELL ADENOMA (B)													
TUBULAR CELL ADENOCARCINOMA (H)													
LIPOMA (B)													
LIPOSARCOMA (H)													
LEUKEMIA (H), Lymphoid													
MYELOCHOCYTIC LEUKEMIA (H)													
HISTIOCYTIC SARCOMA (H)													
URINARY BLADDER:													
TRANSITIONAL CELL PAPILLOMA (B)													
TRANSITIONAL CELL CARCINOMA (H)													
LIPOMA (B)													
HISTIOCYTIC SARCOMA (H)													

Figures in parentheses represent the number of animals from which this tissue was examined.

The absence of a numeral indicates that the lesion specified was not identified

(B) = benign, (H) = malignant

CS9028 (Study Completed)