NDA 20-895

Sildenafil

Appendix B-Female-Rats

Female Rats

NDA 20-895, Sildenafil
Appendix B-Female-Rats

Table I_B-Female-Rats

Test for Positive Dose-Response (Tumor) Linear Trend
Species: Rat
Sex: Female
Sorted by: Organ Name

18:08 Friday, January 23, 1998

Organ Code	Organ Name	Tumor Code	Tumor Name	Exact-P	Asymp-P	AsyCor-P
45	Abdomen	327	B-LIPOMA	0.4649	0.3375	0.3417
45	Abdomen	474	B-PARAGANGLIOMA	0.3485	0.6021	0.6113
6	Adrenal	122	B-CORTICAL ADENOMA	0.8376	0.8545	0.8555
6	Adrenal	134	B-PHEOCHROMOCYTOMA	0.6008	0.6710	0.6730
6	Adrenal	353	M-CORTICAL CARCINOMA	0.3228	0.4029	0.4070
27	Brain	476	B-GRANULAR CELL TUMOUR	0.1515	0.0092	0.0098
27	Brain	177	M-ASTROCYTOMA	0.9255	0.8288	0.8318
24	Colon	362	M-ADENOCARCINOMA	0.1919	0.0204	0.0215
22	Duodenum	463	B-LEIOMYOMA	1.0000	0.6903	0.6986
43	Jejunum	202	M-ADENOCARCINOMA	1.0000	0.6903	0.6986
1	Kidney	373	B-LIPOMA	1.0000	0.7123	0.7196
1	Kidney	187	B-RENAL TUBULE ADENOMA	0.3485	0.6021	0.6113
1	Kidney	447	M-NEPHROBLASTOMA	0.3880	0.6362	0.6442
3	Liver	218	B-HEPATOCELLULAR ADENOMA	0.8281	0.7748	0.7769
3	Liver	337	M-HEPATOCELLULAR CARCINOM	1.0000	0.7418	0.7479
38	Lymphoreticular	315	M-HISTIOCYTIC SARCOMA	0.3728	0.2818	0.2861
38	Lymphoreticular	11	M-LARGE GRANULAR CELL LYM	0.2609	0.2771	0.2811
38	Lymphoreticular	252	M-LYMPHOBLASTIC LYMPHOMA	0.3918	0.6353	0.6433
38	Lymphoreticular	397	M-SMALL LYMPHOCYTIC LYMPH	1.0000	0.7730	0.7777
15	Ovaries	459	B-HAEMANGIOMA	0.2708	0.0507	0.0527
15	Ovaries	433	B-SEX CORD/STROMAL TUMOUR	0.6838	0.8253	0.8283
15	Ovaries	372	M-SCHWANNOMA, MALIGNANT	0.3412	0.1422	0.1458
15	Ovaries	355	M-SEX CORD/STROMAL TUMOUR	1.0000	0.7745	0.7792
19	Pancreas	257	B-ACINAR ADENOMA	0.6177	0.4520	0.4562
19	Pancreas	144	B-ISLET CELL ADENOMA	0.3496	0.4509	0.4533
19	Pancreas	431	B-MIXED ACINAR-ISLET CELL	0.5938	0.6586	0.6676
9	Parathyroid	369	B-ADENOMA	0.4342	0.2113	0.2154
7	Pituitary	6	B-ADENOMA, PARS DISTALIS	0.4403	0.4473	0.4479
7	Pituitary	127	M-CARCINOMA, PARS DISTALI	0.5347	0.6505	0.6532
29	Skeletal muscle	388	M-CHONDROSARCOMA	1.0000	0.7432	0.7493
30	Skin and adnexa	328	B-ADENOMA, MAMMARY GLAND	0.6281	0.6508	0.6529
30	Skin and adnexa	146	B-FIBROADENOMA, MAMMARY G	0.8369	0.8374	0.8369
30	Skin and adnexa	243	B-FIBROUS HISTIOCYTOMA	0.1276	0.0611	0.0624
30	Skin and adnexa	181	B-HAIR FOLLICLE TUMOUR	0.3875	0.6363	0.6442
30	Skin and adnexa	427	B-LIPOMA	0.7835	0.8332	0.8358
30	Skin and adnexa	290	M-ADENOCARCINOMA, MAMMARY	0.8307	0.8408	0.8416
30	Skin and adnexa	291	M-CARCINOMA ARISING IN FI	0.9627	0.9616	0.9620
30	Skin and adnexa	296	M-CARCINOSARCOMA, MAMMARY	1.0000	0.7125	0.7197
30	Skin and adnexa	238	M-FIBROSARCOMA	0.3810	0.6306	0.6388
30	Skin and adnexa	267	M-FIBROUS HISTIOCYTOMA, M	0.8301	0.7436	0.7490
30	Skin and adnexa	470	M-LIPOSARCOMA	0.5909	0.6647	0.6733
30	Skin and adnexa	481	M-MYCOSIS FUNGOIDES	0.3962	0.3476	0.3503
30	Skin and adnexa	259	M-SEBACEOUS/SQUAMOUS CELL	0.3620	0.1540	0.1576
30	Skin and adnexa	223	M-SQUAMOUS CELL CARCINOMA	0.5784	0.7126	0.7181

Source: E:\RAT\f_rat.fil

NDA 20-895, Sildenafil
Appendix B-Female-Rats

Table I_A-Male-Rats
(Continued)

Test for Positive Dose-Response (Tumor) Linear Trend
Species: Mouse 15:05 Monday, February 2, 1998
Sex: Female
Sorted by: Organ Name

Organ Code	Organ Name	Tumor Code	Tumor Name	Exact-P	Asymp-P	AsyCor-P
56	Abdomen	348	B-HAEMANGIOMA	0.1458	0.0118	0.0170
7	Adrenal	285	B-SUBCAPSULAR CELL ADENOM	1.0000	0.7319	0.7771
37	Harderian gland	258	B-ADENOMA	0.9535	0.9104	0.9156
1	Kidney	266	B-RENAL TUBULE ADENOMA	1.0000	0.7319	0.7771
3	Liver	332	B-HAEMANGIOMA	0.4710	0.4597	0.5145
33	Lungs	154	B-BRONCHIOLAR-ALVEOLAR AD	0.3570	0.3467	0.3542
33	Lungs	149	M-BRONCHIOLAR-ALVEOLAR CA	0.3670	0.3767	0.3872
58	Lymphoreticular	244	M-HISTIOCYTIC SARCOMA	0.3001	0.3299	0.3445
58	Lymphoreticular	234	M-IMMUNOBLASTIC LYMPHOMA	0.5625	0.2620	0.2759
58	Lymphoreticular	183	M-LYMPHOBLASTIC LYMPHOMA	0.2908	0.2812	0.2875
16	Ovaries	209	B-CYSTADENOMA	0.4794	0.4707	0.4818
16	Ovaries	246	B-SEX CORD/STROMAL TUMOUR	0.7605	0.7184	0.7461
16	Ovaries	407	B-TERATOMA	1.0000	0.8073	0.8438
31	Skin and adnexa	259	M-ADENOCARCINOMA, MAMMARY	1.0000	0.7310	0.7783
31	Skin and adnexa	204	M-FIBROUS HISTIOCYTOMA, M	1.0000	0.8073	0.8438
19	Spleen	330	B-HAEMANGIOMA	0.3750	0.4028	0.4594
22	Stomach	272	B-SQUAMOUS CELL PAPILLOMA	1.0000	0.7319	0.7771
22	Stomach	270	M-SQUAMOUS CELL CARCINOMA	1.0000	0.8114	0.8381
32	Thymus	359	M-THYMOMA, EPITHELIAL PRE	1.0000	0.8517	0.8710
9	Thyroid	331	M-FOLLICULAR CELL CARCINO	0.6047	0.7032	0.7181
17	Uterus	264	B-HAEMANGIOMA	1.0000	0.7310	0.7783
17	Uterus	287	B-LEIOMYOMA	0.2585	0.1731	0.1951
17	Uterus	284	M-STROMAL CELL SARCOMA	0.8448	0.7974	0.8086
18	Vagina	282	B-LEIOMYOMA	0.8400	0.7765	0.7890

Appendix B-Female Rats

Table II_B_Female_Rats

Analysis of Carcinogenic Potential in Female Rats
Test of Dose-Response (Tumor) Positive Linear Trend
Ted Guo, PH.D, CDER/FDA
Run Date & Time: January 29, 1998 (17:55)
Source: E:\RAT\f_rat.fil

Note: Dose Levels Included: CTRL1 CTRL2 LOW MED HIGH (0 0 1.5 5 60)
Missing value in Tumor-Caused Death is treated as tumor not causing death
Tumor Type: IN: Incidental (nonfatal) tumor, FA: Fatal tumor.

ORGAN/TISSUE NAME AND TUMOR NAME	(ORG#) (TMR#)	TUMOR TYPES	TIME STRATA	ROW NO.	2xC_CONTINGENCY -----TABLE-----	EXACT PROB =PR(STATISTIC.GE.OBSERVED)	ASYMP TOTIC	ASYMP(CONTI NUITY CORR)
Kidney	(1)	IN	93-93	1	0 0 0 1 0	0.3485	0.6021	0.6113
B-RENAL TUBULE ADENOMA	(187)	IN	93-93	2	14 13 16 12 10			
Spontaneous tumor rate LE 1% in ctrl.		-	Total	-	0 0 0 1 0			
Kidney	(1)	IN	65-80	1	0 1 0 0 0	1.0000	0.7123	0.7196
B-LIPOMA	(373)	IN	65-80	2	17 18 13 16 16			
Spontaneous tumor rate LE 1% in ctrl.		-	Total	-	0 1 0 0 0			
Kidney	(1)	FA	72	1	0 0 0 1 0	0.3880	0.6362	0.6442
M-NEPHROBLASTOMA	(447)	FA	72	2	51 45 46 43 46			
Spontaneous tumor rate LE 1% in ctrl.		-	Total	-	0 0 0 1 0			
Ovaries	(15)	IN	93-93	1	1 0 0 0 0	1.0000	0.7745	0.7792
M-SEX CORD/STROMAL TUMOUR	(355)	IN	93-93	2	13 13 16 13 10			
		FA	91	1	1 0 0 0 0			
		FA	91	2	27 24 25 19 25			
Spontaneous tumor rate 2%	in ctrl.	-	Total	-	2 0 0 0 0			
Ovaries	(15)	IN	41-64	1	0 1 0 0 0	0.3412	0.1422	0.1458
M-SCHWANNOMA, MALIGNANT	(372)	IN	41-64	2	17 13 18 21 15			
		FA	57	1	0 0 0 0 1			
		FA	57	2	57 55 54 54 54			
Spontaneous tumor rate LE 1% in ctrl.		-	Total	-	0 1 0 0 1			
Ovaries	(15)	IN	65-80	1	0 0 0 1 0	0.6838	0.8253	0.8283
B-SEX CORD/STROMAL TUMOUR	(433)	IN	65-80	2	17 19 12 15 16			
		IN	81-92	1	0 1 1 0 0			
		IN	81-92	2	11 10 7 5 13			
Spontaneous tumor rate LE 1% in ctrl.		-	Total	-	0 1 1 1 0			
Ovaries	(15)	IN	81-92	1	0 0 0 0 1	0.2708	0.0507	0.0527
B-HAEMANGIOMA	(459)	IN	81-92	2	11 11 8 5 12			
Spontaneous tumor rate LE 1% in ctrl.		-	Total	-	0 0 0 0 1			
Uterus	(16)	IN	41-64	1	1 0 0 0 1	0.3357	0.1361	0.1395
M-STROMAL CELL SARCOMA	(303)	IN	41-64	2	16 15 18 21 15			
Spontaneous tumor rate LE 1% in ctrl.		-	Total	-	1 0 0 0 1			
Uterus	(16)	IN	41-64	1	0 0 0 0 1	0.1084	0.0438	0.0448
B-GRANULAR CELL TUMOUR	(321)	IN	41-64	2	17 15 18 21 15			
		IN	65-80	1	0 0 0 0 1			
		IN	65-80	2	17 19 13 16 15			
		IN	93-93	1	1 0 1 0 0			
		IN	93-93	2	13 13 15 13 10			
Spontaneous tumor rate LE 1% in ctrl.		-	Total	-	1 0 1 0 2			

Uterus	(16)	IN 81-92	1	0	0	0	0	1	0.2708	0.0507	0.0527
B-HAEMANGIOMA	(389)	IN 81-92	2	11	11	8	5	12			
Spontaneous tumor rate LE 1% in ctrl. -		Total	-	0	0	0	0	1			
Uterus	(16)	IN 41-64	1	1	0	0	0	0	0.7537	0.7110	0.7136
B-ENDOMETRIAL STROMAL POL	(393)	IN 41-64	2	16	15	18	21	16			
		IN 65-80	1	0	1	0	1	0			
		IN 65-80	2	17	18	13	15	16			
		IN 81-92	1	3	0	0	0	0			
		IN 81-92	2	8	11	8	5	13			
		IN 93-93	1	0	0	0	0	1			
		IN 93-93	2	14	13	16	13	9			
Spontaneous tumor rate 4% in ctrl. -		Total	-	4	1	0	1	1			
Uterus	(16)	IN 81-92	1	0	0	0	1	0	0.3750	0.6766	0.6834
M-SARCOMA, N.O.S.	(443)	IN 81-92	2	11	11	8	4	13			
Spontaneous tumor rate LE 1% in ctrl. -		Total	-	0	0	0	1	0			
Uterus	(16)	IN 93-93	1	0	0	1	0	0	0.5909	0.6647	0.6733
M-CARCINOMA	(468)	IN 93-93	2	14	13	15	13	10			
Spontaneous tumor rate LE 1% in ctrl. -		Total	-	0	0	1	0	0			
Uterus	(16)	IN 93-93	1	0	1	0	0	0	1.0000	0.6903	0.6986
B-ADENOMA	(479)	IN 93-93	2	14	12	16	13	10			
Spontaneous tumor rate LE 1% in ctrl. -		Total	-	0	1	0	0	0			
Uterus	(16)	IN 93-93	1	1	0	0	0	0	1.0000	0.6903	0.6986
B-LEIOMYOMA	(486)	IN 93-93	2	13	13	16	13	10			
Spontaneous tumor rate LE 1% in ctrl. -		Total	-	1	0	0	0	0			
Vagina	(17)	FA 61	1	0	0	1	0	0	0.5837	0.6887	0.6963
M-LEIOMYOSARCOMA	(437)	FA 61	2	56	56	50	54	52			
Spontaneous tumor rate LE 1% in ctrl. -		Total	-	0	0	1	0	0			
Vagina	(17)	FA 76	1	0	0	0	1	0	0.4563	0.7216	0.7267
M-SCHWANNOMA, MALIGNANT	(440)	FA 76	2	43	44	42	38	40			
		FA 79	1	0	0	1	0	0			
		FA 79	2	41	43	35	34	38			
Spontaneous tumor rate LE 1% in ctrl. -		Total	-	0	0	1	1	0			
Vagina	(17)	IN 93-93	1	0	0	0	0	1	0.1515	0.0092	0.0098
B-SQUAMOUS CELL PAPILLOMA	(475)	IN 93-93	2	14	13	16	13	9			
Spontaneous tumor rate LE 1% in ctrl. -		Total	-	0	0	0	0	1			
Pancreas	(19)	IN 41-64	1	0	0	0	1	0	0.3496	0.4509	0.4533
B-ISLET CELL ADENOMA	(144)	IN 41-64	2	17	15	19	20	16			
		IN 65-80	1	0	1	1	0	0			
		IN 65-80	2	17	18	11	16	16			
		IN 81-92	1	0	0	0	1	1			
		IN 81-92	2	11	11	7	4	12			
		IN 93-93	1	0	3	0	2	1			
		IN 93-93	2	13	10	16	11	8			
Spontaneous tumor rate 3% in ctrl. -		Total	-	0	4	1	4	2			
Pancreas	(19)	IN 81-92	1	0	1	0	0	1	0.6177	0.4520	0.4562
B-ACINAR ADENOMA	(257)	IN 81-92	2	11	10	7	5	12			
		IN 93-93	1	1	1	0	0	0			
		IN 93-93	2	12	12	16	13	9			
Spontaneous tumor rate 3% in ctrl. -		Total	-	1	2	0	0	1			
Pancreas	(19)	IN 93-93	1	0	0	1	0	0	0.5938	0.6586	0.6676
B-MIXED ACINAR-ISLET CELL	(431)	IN 93-93	2	13	13	15	13	9			
Spontaneous tumor rate LE 1% in ctrl. -		Total	-	0	0	1	0	0			
Duodenum	(22)	IN 93-93	1	1	0	0	0	0	1.0000	0.6903	0.6986
B-LEIOMYOMA	(463)	IN 93-93	2	13	13	16	13	10			
Spontaneous tumor rate LE 1% in ctrl. -		Total	-	1	0	0	0	0			

Appendix B-Female Rats: Table II_Female Rats (Continued)

Colon	(24)	FA 95	1	0	0	0	0	1	0.1919	0.0204	0.0215
M-ADENOCARCINOMA	(362)	FA 95	2	21	20	22	17	18			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	0	0	0	1			
Brain	(27)	FA 66	1	0	1	0	0	0	0.9255	0.8288	0.8318
M-ASTROCYTOMA	(177)	FA 66	2	55	50	50	49	48			
		FA 75	1	1	0	0	0	0			
		FA 75	2	45	44	44	40	42			
		FA 87	1	0	0	1	0	0			
Spontaneous tumor rate 2% in ctrl. - Total			-	1	1	1	0	0			
Brain	(27)	IN 93-93	1	0	0	0	0	1	0.1515	0.0092	0.0098
B-GRANULAR CELL TUMOUR	(476)	IN 93-93	2	14	13	16	13	9			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	0	0	0	1			
Skeletal muscle	(29)	IN 81-92	1	1	0	0	0	0	1.0000	0.7432	0.7493
M-CHONDROSARCOMA	(388)	IN 81-92	2	10	11	8	4	13			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	1	0	0	0	0			
Liver	(3)	IN 41-64	1	0	0	0	1	0	0.8281	0.7748	0.7769
B-HEPATOCELLULAR ADENOMA	(218)	IN 41-64	2	17	15	19	20	16			
		IN 65-80	1	0	1	0	0	0			
		IN 65-80	2	17	18	13	16	16			
		IN 81-92	1	2	1	0	0	0			
		IN 81-92	2	9	10	8	5	13			
		IN 93-93	1	0	2	1	0	1			
		IN 93-93	2	14	11	15	13	9			
Spontaneous tumor rate 5% in ctrl. - Total			-	2	4	1	1	1			
Liver	(3)	IN 81-92	1	0	1	0	0	0	1.0000	0.7418	0.7479
M-HEPATOCELLULAR CARCINOM	(337)	IN 81-92	2	11	10	8	5	13			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	1	0	0	0			
Skin and adnexa	(30)	IN 0-40	1	0	1	0	0	0	0.8369	0.8374	0.8369
B-FIBROADENOMA, MAMMARY G	(146)	IN 0-40	2	1	1	4	5	5			
		IN 41-64	1	2	1	7	5	3			
		IN 41-64	2	14	13	11	16	13			
		IN 65-80	1	4	6	2	3	5			
		IN 65-80	2	10	6	9	11	10			
		IN 81-92	1	4	2	3	2	4			
		IN 81-92	2	5	8	5	2	7			
		IN 93-93	1	7	8	11	11	5			
		IN 93-93	2	7	5	4	2	5			
		FA 67	1	0	0	1	0	0			
		FA 67	2	54	50	49	47	48			
		FA 69	1	0	1	0	0	0			
		FA 69	2	53	48	47	46	48			
		FA 74	1	1	0	0	0	0			
		FA 74	2	49	45	45	39	42			
		FA 79	1	0	1	0	0	0			
		FA 79	2	42	42	37	33	39			
		FA 80	1	1	2	0	0	0			
		FA 80	2	40	38	35	31	38			
		FA 81	1	0	2	0	0	0			
		FA 81	2	40	36	34	28	37			
		FA 83	1	0	0	1	0	0			
		FA 83	2	37	35	31	25	36			
		FA 84	1	1	1	1	0	0			
		FA 84	2	36	33	30	24	35			
		FA 86	1	0	1	0	0	0			
		FA 86	2	34	28	29	23	33			
		FA 87	1	1	0	0	0	0			
		FA 87	2	33	27	28	21	33			
		FA 89	1	0	0	0	1	0			
		FA 89	2	31	25	27	20	30			
		FA 90	1	0	0	0	0	1			
		FA 90	2	30	24	26	19	27			
		FA 93	1	0	0	0	0	1			

Appendix B-Female Rats: Table II_Female Rats (Continued)

		FA 93	2	25	21	24	18	21			
		FA 94	1	1	0	0	0	0			
		FA 95	1	0	0	0	1	0			
		FA 95	2	21	20	22	16	20			
		FA 96	1	0	1	0	0	0			
		FA 96	2	21	19	22	15	18			
		FA 100	1	1	0	0	0	0			
		FA 100	2	19	15	18	14	14			
		FA 102	1	0	0	0	0	1			
		FA 102	2	14	15	18	14	12			
		FA 104	1	0	0	1	0	0			
		FA 104	2	14	13	15	13	10			
Spontaneous tumor rate 42% in ctrl. - Total			-	23	27	27	23	20			
Skin and adnexa	(30)	IN 65-80	1	0	0	0	1	0	0.3875	0.6363	0.6442
B-HAIR FOLLICLE TUMOUR	(181)	IN 65-80	2	17	19	13	14	16			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	0	0	1	0			
Skin and adnexa	(30)	IN 93-93	1	0	0	0	1	0	0.5784	0.7126	0.7181
M-SQUAMOUS CELL CARCINOMA	(223)	IN 93-93	2	14	13	16	12	10			
		FA 99	1	0	1	0	0	0			
		FA 99	2	20	16	18	15	15			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	1	0	1	0			
Skin and adnexa	(30)	FA 66	1	0	0	0	1	0	0.3810	0.6306	0.6388
M-FIBROSARCOMA	(238)	FA 66	2	55	51	50	47	48			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	0	0	1	0			
Skin and adnexa	(30)	IN 93-93	1	0	0	0	0	1	0.1276	0.0611	0.0624
B-FIBROUS HISTIOCYTOMA	(243)	IN 93-93	2	14	13	16	13	9			
		FA 64	1	1	0	0	0	0			
		FA 64	2	55	54	51	50	49			
		FA 90	1	0	0	0	0	1			
		FA 90	2	30	24	26	19	27			
		FA 93	1	0	0	1	0	0			
		FA 93	2	25	21	23	18	22			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	1	0	1	0	2			
Skin and adnexa	(30)	IN 65-80	1	0	1	0	0	1	0.3620	0.1540	0.1576
M-SEBACEOUS/SQUAMOUS CELL	(259)	IN 65-80	2	17	18	13	15	15			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	1	0	0	1			
Skin and adnexa	(30)	IN 93-93	1	1	0	0	0	0	0.8301	0.7436	0.7490
M-FIBROUS HISTIOCYTOMA, M	(267)	IN 93-93	2	13	13	15	13	10			
		FA 104	1	0	0	1	0	0			
		FA 104	2	14	13	15	13	10			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	1	0	1	0	0			
Skin and adnexa	(30)	IN 0-40	1	0	0	0	0	1	0.8307	0.8408	0.8416
M-ADENOCARCINOMA, MAMMARY	(290)	IN 0-40	2	1	2	4	4	3			
		IN 41-64	1	5	0	1	1	1			
		IN 41-64	2	11	14	17	18	14			
		IN 65-80	1	5	1	0	5	1			
		IN 65-80	2	11	16	13	10	15			
		IN 81-92	1	1	1	2	0	3			
		IN 81-92	2	9	7	6	5	10			
		IN 93-93	1	2	2	5	8	1			
		IN 93-93	2	12	10	11	5	9			
		FA 37	1	0	0	0	1	0			
		FA 37	2	60	60	60	57	58			
		FA 51	1	0	0	0	0	1			
		FA 51	2	59	58	59	55	55			
		FA 61	1	0	0	0	0	1			
		FA 64	1	0	0	1	0	0			
		FA 64	2	56	54	50	50	49			
		FA 67	1	1	1	0	0	0			
		FA 67	2	53	49	50	47	48			
		FA 70	1	0	0	0	1	0			

Appendix B-Female Rats: Table II_Female Rats (Continued)

		FA 70	2	53	46	46	45	47			
		FA 72	1	0	0	0	1	0			
		FA 72	2	51	45	46	42	46			
		FA 79	1	0	1	0	0	0			
		FA 79	2	42	42	37	33	39			
		FA 87	1	0	1	0	0	0			
		FA 87	2	34	26	28	21	33			
		FA 90	1	1	0	0	0	0			
		FA 90	2	29	24	26	19	28			
		FA 92	1	0	1	0	0	0			
		FA 92	2	25	23	24	18	23			
		FA 96	1	0	1	0	0	0			
		FA 96	2	21	19	22	15	18			
		FA 101	1	1	0	0	0	0			
		FA 101	2	14	15	18	14	14			
		FA 102	1	0	1	0	0	0			
		FA 102	2	14	14	18	14	13			
		FA 104	1	0	1	0	0	0			
		FA 104	2	14	12	16	13	10			
Spontaneous tumor rate 23%	in ctrl.	- Total	-	16	11	9	17	9			
Skin and adnexa	(30)	IN 41-64	1	0	0	1	0	0	0.9627	0.9616	0.9620
M-CARCINOMA ARISING IN FI	(291)	IN 41-64	2	17	15	18	21	16			
		IN 65-80	1	3	1	2	1	0			
		IN 65-80	2	13	17	10	13	16			
		IN 81-92	1	0	1	1	0	1			
		IN 81-92	2	11	10	7	5	12			
		IN 93-93	1	2	0	1	3	0			
		IN 93-93	2	12	13	15	10	10			
		FA 80	1	0	1	0	0	0			
		FA 80	2	41	39	35	31	38			
		FA 82	1	0	0	0	1	0			
		FA 82	2	39	35	33	26	36			
		FA 87	1	1	0	0	0	0			
		FA 87	2	33	27	28	21	33			
		FA 89	1	0	0	1	0	0			
		FA 89	2	31	25	26	21	30			
Spontaneous tumor rate 8%	in ctrl.	- Total	-	6	3	6	5	1			
Skin and adnexa	(30)	FA 54	1	1	0	0	0	0	1.0000	0.7125	0.7197
M-CARCINOSARCOMA, MAMMARY	(296)	FA 54	2	57	57	56	54	55			
Spontaneous tumor rate LE 1%	in ctrl.	- Total	-	1	0	0	0	0			
Skin and adnexa	(30)	IN 65-80	1	1	0	0	0	0	1.0000	0.7128	0.7199
M-SQUAMOUS CELL CARCINOMA	(319)	IN 65-80	2	16	19	13	15	16			
Spontaneous tumor rate LE 1%	in ctrl.	- Total	-	1	0	0	0	0			
Skin and adnexa	(30)	IN 41-64	1	0	1	0	0	0	0.6281	0.6508	0.6529
B-ADENOMA, MAMMARY GLAND	(328)	IN 41-64	2	17	14	19	21	16			
		IN 65-80	1	0	0	1	1	1			
		IN 65-80	2	17	19	12	14	15			
		IN 81-92	1	0	1	0	0	1			
		IN 81-92	2	11	10	8	5	12			
		IN 93-93	1	2	3	1	2	0			
		IN 93-93	2	12	10	15	11	10			
Spontaneous tumor rate 6%	in ctrl.	- Total	-	2	5	2	3	2			
Skin and adnexa	(30)	IN 81-92	1	0	0	0	1	0	0.7835	0.8332	0.8358
B-LIPOMA	(427)	IN 81-92	2	11	11	8	4	13			
		IN 93-93	1	2	0	0	0	0			
		IN 93-93	2	12	13	16	13	10			
		FA 76	1	0	0	1	0	0			
		FA 76	2	44	44	43	38	41			
Spontaneous tumor rate 2%	in ctrl.	- Total	-	2	0	1	1	0			

Appendix B-Female Rats: Table II_Female Rats (Continued)

Skin and adnexa	(30)	IN 93-93	1	0	0	1	0	0	0.5909	0.6647	0.6733
M-LIPOSARCOMA	(470)	IN 93-93	2	14	13	15	13	10			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	0	1	0	0			
Skin and adnexa	(30)	IN 41-64	1	1	0	0	0	0	0.3962	0.3476	0.3503
M-MYCOSIS FUNGOIDES	(481)	IN 41-64	2	16	15	19	21	16			
		IN 81-92	1	0	0	0	0	2			
		IN 81-92	2	11	11	8	5	11			
		IN 93-93	1	2	1	1	0	0			
		IN 93-93	2	12	12	15	13	10			
		FA 79	1	0	0	0	1	0			
		FA 79	2	42	43	37	32	39			
Spontaneous tumor rate 3% in ctrl. - Total			-	3	1	1	1	2			
Thymus	(31)	IN 41-64	1	1	0	0	1	1	0.7410	0.6854	0.6883
M-THYMOMA, LYMPHOCYTIC PR	(209)	IN 41-64	2	16	15	19	20	15			
		IN 65-80	1	1	0	0	0	0			
		IN 65-80	2	16	19	13	16	16			
		IN 81-92	1	1	1	0	0	0			
		IN 81-92	2	10	10	8	5	13			
		IN 93-93	1	1	0	0	0	0			
		IN 93-93	2	13	13	16	13	10			
Spontaneous tumor rate 4% in ctrl. - Total			-	4	1	0	1	1			
Thymus	(31)	FA 81	1	0	1	0	0	0	1.0000	0.7154	0.7224
M-SCHWANNOMA, MALIGNANT	(374)	FA 81	2	40	37	34	29	37			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	1	0	0	0			
Thymus	(31)	IN 81-92	1	0	0	0	0	1	0.2708	0.0507	0.0527
M-CARCINOMA, NOS	(391)	IN 81-92	2	11	11	8	5	12			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	0	0	0	1			
Thymus	(31)	IN 41-64	1	0	0	1	0	0	0.8513	0.7561	0.7612
M-THYMOMA, EPITHELIAL PRE	(426)	IN 41-64	2	17	15	18	21	16			
		IN 93-93	1	1	0	0	0	0			
		IN 93-93	2	13	13	16	13	10			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	1	0	1	0	0			
Lymphoreticular	(38)	IN 81-92	1	0	0	0	1	0	0.2609	0.2771	0.2811
M-LARGE GRANULAR CELL LYM	(11)	IN 81-92	2	11	11	8	4	13			
		IN 93-93	1	0	1	0	0	0			
		IN 93-93	2	14	12	16	13	10			
		FA 21	1	0	0	0	0	1			
		FA 21	2	60	60	60	60	59			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	1	0	1	1			
Lymphoreticular	(38)	FA 38	1	0	0	0	1	0	0.3918	0.6353	0.6433
M-LYMPHOBLASTIC LYMPHOMA	(252)	FA 38	2	60	60	60	57	58			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	0	0	1	0			
Lymphoreticular	(38)	FA 61	1	1	0	0	0	0	0.3728	0.2818	0.2861
M-HISTIOCYTIC SARCOMA	(315)	FA 61	2	56	56	53	54	53			
		FA 96	1	0	0	1	0	0			
		FA 96	2	21	20	21	15	18			
		FA 98	1	0	0	0	0	1			
		FA 98	2	20	18	20	15	17			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	1	0	1	0	1			
Lymphoreticular	(38)	IN 65-80	1	0	1	0	0	0	1.0000	0.7730	0.7777
M-SMALL LYMPHOCYTIC LYMPH	(397)	IN 65-80	2	17	18	13	16	16			
		IN 93-93	1	0	1	0	0	0			
		IN 93-93	2	14	12	16	13	10			
Spontaneous tumor rate 2% in ctrl. - Total			-	0	2	0	0	0			
Jejunum	(43)	IN 93-93	1	0	1	0	0	0	1.0000	0.6903	0.6986
M-ADENOCARCINOMA	(202)	IN 93-93	2	14	12	16	13	10			
Spontaneous tumor rate LE 1% in ctrl. - Total			-	0	1	0	0	0			

Appendix B-Female Rats: Table II_Female Rats (Continued)

Abdomen	(45	)	IN 41-64	1	0	0	0	0	1	0.4649	0.3375	0.3417
B-LIPOMA	(327	)	IN 41-64	2	17	15	19	21	15			
			IN 93-93	1	0	2	1	0	0			
			IN 93-93	2	14	11	15	13	10			
Spontaneous tumor rate 2%	in ctrl.		- Total	-	0	2	1	0	1			
Abdomen	(45	)	IN 93-93	1	0	0	0	1	0	0.3485	0.6021	0.6113
B-PARAGANGLIOMA	(474	)	IN 93-93	2	14	13	16	12	10			
Spontaneous tumor rate LE 1%	in ctrl.		- Total	-	0	0	0	1	0			
Adrenal	(6	)	IN 0-40	1	0	0	1	0	0	0.8376	0.8545	0.8555
B-CORTICAL ADENOMA	(122	)	IN 0-40	2	1	2	3	5	5			
			IN 41-64	1	1	0	2	0	0			
			IN 41-64	2	16	15	17	21	16			
			IN 65-80	1	1	0	2	0	2			
			IN 65-80	2	16	19	11	16	14			
			IN 81-92	1	2	3	2	1	1			
			IN 81-92	2	9	8	6	4	12			
			IN 93-93	1	0	1	1	3	0			
			IN 93-93	2	14	12	15	10	10			
Spontaneous tumor rate 7%	in ctrl.		- Total	-	4	4	8	4	3			
Adrenal	(6	)	IN 41-64	1	0	0	1	0	1	0.6008	0.6710	0.6730
B-PHEOCHROMOCYTOMA	(134	)	IN 41-64	2	17	15	18	21	15			
			IN 65-80	1	0	2	1	1	0			
			IN 65-80	2	17	17	12	15	16			
			IN 81-92	1	1	1	0	0	1			
			IN 81-92	2	10	10	8	5	12			
			IN 93-93	1	0	2	0	3	0			
			IN 93-93	2	14	11	16	10	10			
Spontaneous tumor rate 5%	in ctrl.		- Total	-	1	5	2	4	2			
Adrenal	(6	)	IN 81-92	1	0	1	1	0	0	0.3228	0.4029	0.4070
M-CORTICAL CARCINOMA	(353	)	IN 81-92	2	11	10	7	5	13			
			IN 93-93	1	0	0	0	1	1			
			IN 93-93	2	14	13	16	12	9			
Spontaneous tumor rate LE 1%	in ctrl.		- Total	-	0	1	1	1	1			
Pituitary	(7	)	IN 65-80	1	0	1	0	0	0	0.5347	0.6505	0.6532
M-CARCINOMA, PARS DISTALI	(127	)	IN 65-80	2	17	17	13	15	15			
			IN 93-93	1	0	0	0	1	0			
			IN 93-93	2	13	12	16	12	10			
			FA 63	1	0	0	0	0	1			
			FA 63	2	56	54	51	53	49			
			FA 78	1	0	0	0	1	0			
			FA 78	2	43	42	39	36	38			
			FA 80	1	0	0	0	1	0			
			FA 80	2	41	39	35	31	37			
			FA 85	1	0	1	0	0	0			
			FA 85	2	34	31	29	23	33			
			FA 94	1	0	1	0	0	0			
			FA 94	2	23	19	23	18	21			
			FA 104	1	1	0	0	0	0			
			FA 104	2	13	12	16	13	10			
Spontaneous tumor rate 3%	in ctrl.		- Total	-	1	3	0	3	1			
Pituitary	(7	)	IN 0-40	1	0	1	0	0	0	0.4403	0.4473	0.4479
B-ADENOMA, PARS DISTALIS	(6	)	IN 0-40	2	1	0	0	3	5			
			IN 41-64	1	6	1	3	1	2			
			IN 41-64	2	2	3	2	6	2			
			IN 65-80	1	7	7	4	5	1			
			IN 65-80	2	0	5	1	2	3			
			IN 81-92	1	4	5	2	1	5			
			IN 81-92	2	0	1	0	0	1			
			IN 93-93	1	11	11	16	12	8			
			IN 93-93	2	3	1	0	1	1			
			FA 39	1	0	0	1	1	0			
			FA 39	2	60	59	59	56	56			
			FA 50	1	0	1	0	0	0			

Appendix B-Female Rats: Table II_Female Rats (Continued)

FA	50	2	59	57	59	56	56
FA	52	1	0	0	3	1	0
FA	52	2	59	57	56	55	54
FA	53	1	0	1	0	0	0
FA	53	2	59	56	56	55	54
FA	54	1	0	1	0	1	0
FA	54	2	58	55	56	54	54
FA	57	1	0	0	0	0	1
FA	57	2	57	55	56	54	53
FA	59	1	0	0	2	0	0
FA	59	2	57	55	54	54	52
FA	61	1	0	0	0	1	2
FA	61	2	57	55	53	53	50
FA	62	1	0	1	1	0	0
FA	62	2	56	54	51	53	50
FA	63	1	0	1	0	2	1
FA	63	2	56	53	51	51	49
FA	64	1	0	3	0	1	0
FA	64	2	56	50	51	50	48
FA	67	1	0	0	2	1	0
FA	67	2	54	49	48	47	47
FA	69	1	0	2	1	0	1
FA	69	2	53	46	46	47	46
FA	70	1	1	0	0	1	0
FA	70	2	52	45	46	46	46
FA	71	1	1	1	0	1	1
FA	71	2	51	44	46	44	45
FA	72	1	1	0	0	2	1
FA	72	2	50	44	46	42	44
FA	73	1	0	0	1	0	2
FA	73	2	50	44	45	40	41
FA	74	1	3	1	1	0	0
FA	74	2	47	43	44	40	41
FA	75	1	1	0	0	1	1
FA	75	2	45	43	44	39	40
FA	76	1	1	0	3	1	2
FA	76	2	43	43	41	38	38
FA	77	1	0	0	1	0	0
FA	77	2	43	42	39	37	38
FA	78	1	1	0	2	2	0
FA	78	2	42	42	37	35	38
FA	79	1	1	1	1	1	1
FA	79	2	41	41	36	33	37
FA	80	1	0	0	1	2	1
FA	80	2	41	39	34	30	36
FA	81	1	1	0	1	0	1
FA	81	2	39	37	33	29	35
FA	82	1	1	0	1	0	0
FA	82	2	38	34	32	28	35
FA	83	1	0	1	0	1	1
FA	83	2	37	33	32	25	34
FA	84	1	2	0	1	2	1
FA	84	2	35	33	30	23	33
FA	85	1	0	2	0	0	0
FA	85	2	34	30	29	23	33
FA	86	1	0	1	1	1	0
FA	86	2	34	27	28	22	32
FA	87	1	1	0	0	0	2
FA	87	2	33	26	28	21	30
FA	88	1	1	1	0	0	1
FA	88	2	31	24	27	21	29
FA	89	1	1	1	0	1	1
FA	89	2	30	23	27	20	28
FA	90	1	0	0	1	0	0
FA	90	2	30	23	25	19	27
FA	91	1	2	0	1	1	2
FA	91	2	26	23	24	18	23
FA	92	1	0	2	0	0	1
FA	92	2	25	21	24	18	22

Appendix B-Female Rats: Table II_Female Rats (Continued)

		FA 93	1	2	0	0	0	0			
		FA 93	2	23	20	24	18	22			
		FA 94	1	1	0	1	1	0			
		FA 94	2	22	20	22	17	21			
		FA 95	1	0	0	0	1	1			
		FA 95	2	21	19	22	16	19			
		FA 97	1	1	1	1	0	0			
		FA 97	2	20	17	20	15	18			
		FA 98	1	0	1	2	0	2			
		FA 98	2	20	16	18	15	16			
		FA 99	1	0	1	0	1	1			
		FA 99	2	20	15	18	14	14			
		FA 100	1	3	0	0	0	0			
		FA 100	2	17	14	18	14	14			
		FA 101	1	0	0	0	0	1			
		FA 101	2	15	14	18	14	13			
		FA 102	1	0	0	2	0	1			
		FA 102	2	14	14	16	14	12			
		FA 103	1	0	0	0	1	0			
		FA 103	2	14	12	16	13	10			
		FA 104	1	0	0	0	0	1			
		FA 104	2	14	12	16	13	9			
Spontaneous tumor rate 86%	in ctrl.	- Total	-	54	49	57	48	47			
Thyroid	(8)	IN 41-64	1	1	1	1	1	0	0.7008	0.6837	0.6855
B-C-CELL ADENOMA	(141)	IN 41-64	2	16	14	18	20	16			
		IN 65-80	1	0	1	1	1	0			
		IN 65-80	2	17	18	12	15	16			
		IN 81-92	1	1	1	0	0	0			
		IN 81-92	2	10	10	8	5	13			
		IN 93-93	1	1	3	4	1	3			
		IN 93-93	2	12	10	12	12	7			
Spontaneous tumor rate 8%	in ctrl.	- Total	-	3	6	6	3	3			
Thyroid	(8)	IN 41-64	1	0	0	0	1	1	0.2694	0.2457	0.2497
B-FOLLICULAR CELL ADENOMA	(170)	IN 41-64	2	17	15	19	20	15			
		IN 65-80	1	0	1	0	0	0			
		IN 65-80	2	17	18	13	16	16			
Spontaneous tumor rate LE 1%	in ctrl.	- Total	-	0	1	0	1	1			
Thyroid	(8)	IN 93-93	1	0	0	1	0	0	0.6000	0.6666	0.6751
M-C-CELL CARCINOMA	(254)	IN 93-93	2	13	13	15	13	10			
Spontaneous tumor rate LE 1%	in ctrl.	- Total	-	0	0	1	0	0			
Thyroid	(8)	IN 81-92	1	0	1	0	0	0	1.0000	0.7418	0.7479
M-FOLLICULAR CELL CARCINO	(375)	IN 81-92	2	11	10	8	5	13			
Spontaneous tumor rate LE 1%	in ctrl.	- Total	-	0	1	0	0	0			
Parathyroid	(9)	IN 65-80	1	0	1	0	0	1	0.4342	0.2113	0.2154
B-ADENOMA	(369)	IN 65-80	2	12	11	10	9	13			
Spontaneous tumor rate LE 1%	in ctrl.	- Total	-	0	1	0	0	1			

Appendix B-Female Rats: Table II_Female Rats (Continued)