

Analysis of dose-response trend in mouse for hemangioma -hemangiosarcoma separately

Analysis of Carcinogenic Potential in Male Mouse
 Test of Dose-Response (Tumor) Positive Linear Trend
 Ted Guo, PH.D, CDER/FDA

Run Date & Time: March 19, 1997 (8:34)

Source: c:_data\hemice.txt

Note: Dose Levels Included: CTRL CTR2 LOW MED HIGH MAXI (0 0 25 375 750 1500)
 For missing Tumor-Caused-Death set INCIDENTAL
 IN: Incidental (nonfatal) to all, FA: Fatal to all, MX: Mixed (Fatal to some)
 Symbols, ^ and + indicate that p-values are <0.005 and <0.025 respectively

ORGAN NAME	ORGAN CODE	TUMOR NAME	TUMOR CODE	TUMOR TIME TYPE	INTERVAL	ROW TABLE	P VALUES												
							EXACT PERMU	ASYMP-TOTIC	CONTINU CORRECT										
ORGAN WITH(ORG/1602)HEMANGIOMA	(C(160212)	IN	IN	53-78	1	[REDACTED]	0.420	0.440	0.440									
					IN 53-78	2													
					IN 92-104	1													
					IN 92-104	2													
					IN 105-105	1													
					IN 105-105	2													
					Tumor rate: 4% in CTL - Total	-													
					ORGAN WITH(ORG/1800)HEMANGIOSARCO	(180003)	MX	IN	79-91	1	[REDACTED]	0.199	0.199	0.199
														IN 79-91	2				
														IN 92-104	1				
														IN 92-104	2				
IN 105-105	1																		
IN 105-105	2																		
FA 82	1																		
FA 82	2																		
FA 83	1																		
FA 83	2																		
FA 85	1																		
FA 85	2																		
FA 93	1																		
FA 93	2																		
FA 96	1																		
FA 96	2																		
FA 100	1																		
FA 100	2																		
FA 104	1																		
FA 104	2																		
OTHER ORGAN(OTHER)OTHER TUMOR (OTHER)))	MX	IN	0-52	1	[REDACTED]	0.889	0.888	0.887									
					IN 0-52	2													
					IN 53-78	1													
					IN 53-78	2													
					IN 79-91	1													
					IN 79-91	2													
					IN 92-104	1													
					IN 92-104	2													
					IN 105-105	1													
					IN 105-105	2													
					FA 41	1													
FA 41	2																		
FA 52	1																		
FA 52	2																		
FA 55	1																		
FA 55	2																		
FA 62	1																		
FA 62	2																		
FA 65	1																		

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ORGAN NAME	ORGAN CODE	TUMOR NAME	TUMOR CODE	TUMOR TIME TYPE	INTERVAL	ROW TABLE	P VALUES		
							EXACT PERMU	ASYMP-TOTIC	CONTINU CORRECT
				FA 65		2			
				FA 66		1			
				FA 66		2			
				FA 70		1			
				FA 70		2			
				FA 71		1			
				FA 71		2			
				FA 72		1			
				FA 72		2			
				FA 74		1			
				FA 74		2			
				FA 78		1			
				FA 78		2			
				FA 80		1			
				FA 80		2			
				FA 81		1			
				FA 81		2			
				FA 82		1			
				FA 82		2			
				FA 83		1			
				FA 83		2			
				FA 84		1			
				FA 84		2			
				FA 85		1			
				FA 85		2			
				FA 88		1			
				FA 88		2			
				FA 89		1			
				FA 89		2			
				FA 90		1			
				FA 90		2			
				FA 91		1			
				FA 91		2			
				FA 93		1			
				FA 93		2			
				FA 94		1			
				FA 94		2			
				FA 96		1			
				FA 96		2			
				FA 98		1			
				FA 98		2			
				FA 99		1			
				FA 99		2			
				FA 100		1			
				FA 100		2			
				FA 101		1			
				FA 101		2			
				FA 102		1			
				FA 102		2			

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Analysis of Carcinogenic Potential in Male Mouse
 Test of Dose-Response (Tumor) Positive Linear Trend
 Run Date & Time: March 19, 1997 (8:34)
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Note: Dose Levels Included: CTRL CTR2 LOW MED HIGH MAXI (0 0 25 375 750 1500)
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 IN: Incidental (nonfatal) to all, FA: Fatal to all, MX: Mixed (Fatal to some)
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ORGAN NAME	ORGAN CODE	TUMOR NAME	TUMOR CODE	TUMOR TIME TYPE INTERVAL	ROW TABLE	P VALUES		
						EXACT PERMU	ASYMP- TOTIC	CONTINU CORRECT
				FA 103	1			
				FA 103	2			
				FA 104	1			
				FA 104	2			
Page - 3 -	(End of File)			Tumor rate: 87% in CTL - Total	-			

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Analysis of Carcinogenic Potential in Female Mouse
 Test of Dose-Response (Tumor) Positive Linear Trend
 Ted Guo, PH.D, CDER/FDA

Run Date & Time: March 19, 1997 (8:40)

Source: c:_data\hemice.txt

Note: Dose Levels Included: CTR1 CTR2 LOW MED HIGH MAXI (0 0 25 375 750 1500)
 For missing Tumor-Caused-Death set INCIDENTAL
 IN: Incidental (nonfatal) to all, FA: Fatal to all, MX: Mixed (Fatal to some)
 Symbols, ^ and + indicate that p-values are <0.005 and <0.025 respectively

ORGAN NAME	ORGAN CODE	TUMOR NAME	TUMOR CODE	TUMOR TIME TYPE	TUMOR TIME INTERVAL	ROW TABLE	P VALUES														
							EXACT PERMU	ASYMP-TOTIC	CONTINU CORRECT												
ORGAN WITH (ORG/1602) HEMANGIOMA (C(160212)				IN	IN 0-52	1	[REDACTED]	0.283	0.278	0.278											
						IN 0-52					2										
						IN 79-91					1										
						IN 79-91					2										
						IN 92-104					1										
						IN 92-104					2										
						- Total					-										
						ORGAN WITH (ORG/1800) HEMANGIOSARCO (180003)								MX	IN 79-91	1	[REDACTED]	0.481	0.494	0.494	
																IN 79-91					2
																IN 92-104					1
IN 92-104	2																				
FA 69	1																				
FA 69	2																				
FA 82	1																				
FA 82	2																				
FA 85	1																				
FA 85	2																				
OTHER ORGA (OTHER) OTHER TUMOR (OTHER)				MX	IN 0-52	1	[REDACTED]	0.188	0.189	0.190											
						IN 0-52					2										
						IN 53-78					1										
						IN 53-78					2										
						IN 79-91					1										
						IN 79-91					2										
						IN 92-104					1										
						IN 92-104					2										
						FA 28					1										
						FA 28					2										
FA 30	1																				
FA 30	2																				
FA 33	1																				
FA 33	2																				
FA 39	1																				
FA 39	2																				
FA 43	1																				
FA 43	2																				
FA 50	1																				
FA 50	2																				
FA 52	1																				
FA 52	2																				
FA 54	1																				
FA 54	2																				
FA 57	1																				

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Analysis of Carcinogenic Potential in Female Mouse
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IN: Incidental (nonfatal) to all, FA: Fatal to all, MX: Mixed (Fatal to some)
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ORGAN NAME	ORGAN CODE	TUMOR NAME	TUMOR CODE	TUMOR TIME TYPE	INTERVAL	ROW TABLE	P VALUES		
							EXACT PERMU	ASYMP-TOTIC	CONTINU CORRECT
				FA 57		2			
				FA 61		1			
				FA 61		2			
				FA 62		1			
				FA 62		2			
				FA 63		1			
				FA 63		2			
				FA 65		1			
				FA 65		2			
				FA 66		1			
				FA 66		2			
				FA 67		1			
				FA 67		2			
				FA 68		1			
				FA 68		2			
				FA 69		1			
				FA 69		2			
				FA 70		1			
				FA 70		2			
				FA 72		1			
				FA 72		2			
				FA 73		1			
				FA 73		2			
				FA 74		1			
				FA 74		2			
				FA 75		1			
				FA 75		2			
				FA 76		1			
				FA 76		2			
				FA 77		1			
				FA 77		2			
				FA 78		1			
				FA 78		2			
				FA 79		1			
				FA 79		2			
				FA 80		1			
				FA 80		2			
				FA 81		1			
				FA 81		2			
				FA 82		1			
				FA 82		2			
				FA 83		1			
				FA 83		2			
				FA 84		1			
				FA 84		2			
				FA 85		1			
				FA 85		2			
				FA 86		1			
				FA 86		2			

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ORGAN NAME	ORGAN CODE	TUMOR NAME	TUMOR CODE	TUMOR TIME TYPE INTERVAL	ROW TABLE	P VALUES		
						EXACT PERMU	ASYMP- TOTIC	CONTINU CORRECT
				FA 87	1			
				FA 87	2			
				FA 88	1			
				FA 88	2			
				FA 89	1			
				FA 89	2			
				FA 90	1			
				FA 90	2			
				FA 91	1			
				FA 91	2			
				FA 92	1			
				FA 92	2			
				FA 93	1			
				FA 93	2			
				FA 94	1			
				FA 94	2			
				FA 95	1			
				FA 95	2			
				Total				

Page - 3 - (End of File) Tumor rate: 90% in CTL - Total

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References

- 1 Lin et al (1994). "Statistical Review and Evaluation of Animal Tumorigenicity Studies." Statistics in the Pharmaceutical Industry. Marcel Dekker, Inc. pp. 19-57.
 - 2 Thomas et al (1976). "Trend and Homogeneity Analyses of Proportions and Life Table Data." Computers and Biomedical Research. pp. 373-81.
 - 3 Peto et al (1980). "Guidelines for Simple, Sensitive Significance Tests for Carcinogenic Effects in Long-Term Animal Experiments. In Long-Term and Short-Term Screening Assays for Carcinogens: An Critical Appraisal, International Agency for Research on Cancer, Lyon, France. IARC Monographs Supplement 2. pp. 311-426.
- Haseman, J. K. (1983). "A re-examination of false-positive rates for carcinogenesis studies." Fundamental and Applied Toxicology 3, pp. 334-9.
- Haseman, J. K. (1984). "Issues in carcinogenicity testing: Dose selection." Fundamental and Applied Toxicology, Vol. 5. pp. 66-78.

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