

Study Number: C11054B-01
Test Type: TOX
Route: Dosing in Water
Species/Strain: Rat/Harlan Sprague Dawley

R16: Pubertal Markers Summary (Interim and Chronic Animals)
Test Compound: Sulfolane
CAS Number: 126-33-0

Date Report Requested: 07/15/2021
Time Report Requested: 14:59:24
Lab: Battelle with CRL

Study Number: C11054B-01
Study Sex: Both
PWG Approval Date: See web page for date of PWG Approval
Version: v1.2.7
Stat Version: S

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			Male				
Generation	Cohort		Treatment Groups (mg/L)				
			0	30	100	300	1000
F1	All Males	No. Examined (litters)	120 (38)	120 (31)	120 (31)	120 (29)	120 (31)
		No. Removed (litters)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
		No. Not Attaining BPS (litters)	0 (0)	0 (0)	1 (1)	0 (0)	1 (1)
		Day of BPS					
		Mean Analysis					
		Litter Mean $\pm$ SE	44.6 $\pm$ 0.2 *	44.5 $\pm$ 0.4	44.5 $\pm$ 0.2	44.5 $\pm$ 0.3	45.4 $\pm$ 0.4
		Litter Mean of Adjusted $\pm$ SE	44.8 $\pm$ 0.2	44.5 $\pm$ 0.3	44.6 $\pm$ 0.2	44.5 $\pm$ 0.2	45.1 $\pm$ 0.4
		Proportional Hazards Analysis					
		Litter-based Model	p=0.035	p=0.724	p=0.853	p=0.853	p=0.853
		BW at Attainment (g)	192.2 $\pm$ 1.7	187.0 $\pm$ 1.9	189.6 $\pm$ 1.7	189.5 $\pm$ 1.8	186.2 $\pm$ 2.1 *
		BW at Weaning (g)	89.1 $\pm$ 1.4 **	86.2 $\pm$ 1.3	86.9 $\pm$ 1.3	86.7 $\pm$ 1.3	83.0 $\pm$ 1.2 **

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			Female				
Generation	Cohort		Treatment Groups (mg/L)				
			0	30	100	300	1000
F1	All Females	No. Examined (litters)	120 (33)	120 (31)	117 (31)	120 (32)	120 (32)
		No. Removed (litters)	23 (6)	2 (1)	0 (0)	9 (3)	12 (4)
		No. Not Attaining VO (litters)	0 (0)	0 (0)	0 (0)	0 (0)	0 (0)
		Day of VO					
		Mean Analysis					
		Litter Mean $\pm$ SE	36.2 $\pm$ 0.3 **	37.1 $\pm$ 0.3	36.5 $\pm$ 0.3	37.4 $\pm$ 0.3 *	38.4 $\pm$ 0.3 **
		Litter Mean of Adjusted $\pm$ SE	36.4 $\pm$ 0.3 **	37.1 $\pm$ 0.3	36.6 $\pm$ 0.3	37.4 $\pm$ 0.3 *	38.2 $\pm$ 0.3 **
		Proportional Hazards Analysis					
		Litter-based Model	p<0.001	p=0.145	p=0.758	p=0.074	p<0.001
		BW at Attainment (g)	115.7 $\pm$ 1.3	116.1 $\pm$ 1.4	115.0 $\pm$ 1.3	116.3 $\pm$ 1.6	116.4 $\pm$ 1.5
		BW at Weaning (g)	81.6 $\pm$ 1.2 **	78.7 $\pm$ 1.0	79.9 $\pm$ 1.0	78.3 $\pm$ 1.0	76.0 $\pm$ 0.8 **

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LEGEND

BPS = Balanopreputial separation; BW = Body weight; VO = Vaginal opening

No. Examined (litters) = the number of animals or pups examined (number of litters)

No. Removed (litters) is the number of animals (number of litters contributing) that died or were removed prior to the end of the observation period and did not attain. These animals were excluded from all analyses.

Summary statistics and mixed model results are presented for animals that attained during the observation period for Day of BPS and Day of VO Mean Analysis endpoint.

Means of litter means presented for Day of BPS and Day of VO Litter Mean $\pm$ SE. Trend and pairwise tests were based on mixed models for day of attainment with dose as a covariate. The Dunnett-Hsu adjustment was used for multiple comparisons.

Mean adjusted day of attainment was calculated from the mean of the litter means of the weaning weight-adjusted attainment days for individual pups. Trend and pairwise tests were based on mixed models for day of attainment with dose and weaning weight as covariates. The Dunnett-Hsu adjustment was used for multiple comparisons.

Animals that did not attain by the end of the observation period were included in the proportional hazards analysis.

P-values for trend and pairwise comparisons for the Litter-based Model of the Proportional Hazards Analysis were calculated from a Cox proportional hazards model with dose and weaning weight as covariates and a random effect for litter, and a Hommel adjustment for multiple comparisons.

Analysis of body weight at attainment and body weight at weaning were performed using mixed effects models with dose as covariate and a random effect for litter. The Dunnett-Hsu adjustment was used for multiple comparisons. Animals that attained during the observation period were used for analysis.

Statistical significance for the control group indicates a significant trend test

Statistical significance for a treatment group indicates a significant pairwise test compared to the vehicle control group

* Statistically significant at $P \leq 0.05$

** Statistically significant at $P \leq 0.01$

**** END OF REPORT ****