Route: Gavage

Species/Strain: Rat/Fischer 344

Toxicokinetics Data Summary

Test Compound: Primidone CAS Number: 125-33-7

Date Report Requested: 12/29/2016 Time Report Requested: 14:55:31

Lab: Battelle Columbus Laboratories

Male

_			Treatment Groups (mg/kg)	
_	30 a, #	30 b, *	80 a,#	80 b, *
			Plasma	
C _{max} (ug/mL)	14.4 ± 0.9	3.31	39.3 ± 1.2	7.12
T _{max} (minute)	60.9 ± 4.6	360	114 ± 5	540
t _{1/2} (minute)	42.2 ± 3.2		78.8 ± 3.1	
k ₁₀ (minute^-1)	0.0164 ± 0.0012		0.00880 ± 0.00035	
V ₁ (mL/kg)	0.767 ± 0.051		0.750 ± 0.023	
AUCinf (ug/mL/min)	2381 ± 207	2944	12127 ± 561	6276

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Male

	Treatment Groups (mg/kg)				
	130 ^{a,#} 130 ^{b, *} Plasma			130 b, *	
				sma	
C _{max} (ug/mL)	53.4	±	1.9	9.68	
T _{max} (minute)	131	±	6	540	
t _{1/2} (minute)	91.1	±	4.1		
k ₁₀ (minute^-1)	0.0076	1 ±	0.00034		
V ₁ (mL/kg)	0.896	±	0.032		
AUC _{inf} (ug/mL/min)	19067	± 10	034	10599	

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Date Report Requested: 12/29/2016 Time Report Requested: 14:55:31 Lab: Battelle Columbus Laboratories

Female

_	Treatment Groups (mg/kg)			
_	30 a, #	30 b, *	80 a, #	80 b, *
			Plasma	
C _{max} (ug/mL)	27.4 ± 2.6	1.02	70.7 ± 3.3	2.28
T _{max} (minute)	110 ± 13	360	152 ± 9	540
t _{1/2} (minute)	76.3 ± 9.3		105 ± 6	
k ₁₀ (minute^-1)	0.00908 ± 0.00111		0.00659 ± 0.00037	
V ₁ (mL/kg)	0.402 ± 0.038		0.416 ± 0.019	
AUCinf (ug/mL/min)	8213 ± 1098	1115	29157 ± 1921	2272

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Female

	Treatment Groups (mg/kg)				
	130 ^{a,#} 130 ^{b, *}			130 b, *	
				asma	
C _{max} (ug/mL)	102	±	5	2.74	
T _{max} (minute)	179	±	10	540	
t _{1/2} (minute)	124	±	7		
k ₁₀ (minute^-1)	0.0055	8 ±	0.00030		
V ₁ (mL/kg)	0.467	±	0.021		
AUC _{inf} (ug/mL/min)	49924	± 3	159	3735	

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LEGEND

Data are displayed as mean ± SEM

MODELING METHOD & BEST FIT MODEL

- ^a PCNONLIN (Statistical Consultants, Inc., Lexington, KY); one one-compartment with equal first-order absorption and first-order elimination with no lag time (Model 5).
- ^b Phenobarbital plasma concentration versus time data were not entered into the PCNONLIN program. Reported phenobarbital toxicokinetic parameters are observed values.

ANALYTE

- [†] Primidone
- Phenobarbital

TK PARAMETERS

C_{max} = Observed or Predicted Maximum plasma (or tissue) concentration

 T_{max} = Time at which C_{max} predicted or observed occurs

 $t_{1/2}$ = Lambda_z half-life, $t_{1/2}$, the terminal elimination half-life based on non-compartmental analysis

 k_{10} = Elimination rate constant from the central compartment also k_e or k_{elim}

 V_1 = Volume of distribution of the central compartment, includes V_d and V_{volume} of distribution, V_z apparent volume of distribution NCA, V_{app} apparent volume of distribution for intravenous studies

AUC_{inf} = Area under the plasma concentration versus time curve, AUC, extrapolated to time equals infinity

** END OF REPORT **