**Experiment Number:** S0629

Route: Gavage, IV

Species/Strain: Rat/Sprague-Dawley

# **Toxicokinetics Data Summary**

Test Compound: Wyeth-14643

CAS Number: 50892-23-4

Date Report Requested: 11/09/2016 Time Report Requested: 14:05:25

Lab: Research Triangle Institute

		Male			
	Treatment Groups (mg/kg)				
	1 a	2 a	2 b	5 a	2 IV a
			Plasma		
C <sub>max(obs)</sub> (ug/mL)	0.961	1.42		5.13	
T <sub>max(obs)</sub> (minute)	30.0	30.0		15.0	
t <sub>1/2(Beta)</sub> (minute)	155.0	99.2		129.0	48.9
k <sub>01</sub> (min^-1)			0.0085		
k <sub>10</sub> (min^-1)			0.0471 ± 0.0051		
CI (mL/min/kg)					2.89
Cl <sub>1(F)</sub> (mL/min/kg)	6.35	8.48		5.48	
V1 (L/kg)			$0.0955 \pm 0.0086$		
MRT (minute)	221	185		246	38.9
AUCinf (ug/mL*min)	157.0	236.0		912.0	693.0
F (fraction)	0.45	0.34		0.53	

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## **LEGEND**

Data are displayed as mean ± SEM

### MODELING METHOD & BEST FIT MODEL

### **ANALYTE**

Wyeth-14643

### TK PARAMETERS

C<sub>max(obs)</sub> = Observed or Predicted Maximum plasma (or tissue) concentration

 $T_{max(obs)}$  = Time at which  $C_{max}$  predicted or observed occurs

 $t_{\frac{1}{2}(beta)}$  = Half-life for the beta phase

 $k_{01}$  = Absorption rate constant,  $k_a$ 

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

CI = Clearance, includes total clearance

Cl<sub>1(F)</sub> = Apparent clearance of the central compartment, also Cl<sub>(F)</sub> for gavage groups in non-compartmental model

 $V_1$  = 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

MRT = Mean residence time

AUC<sub>inf</sub> = Area under the plasma concentration versus time curve, AUC, extrapolated to time equals infinity

F = Bioavailability, absolute bioavailability

\*\* END OF REPORT \*\*

<sup>&</sup>lt;sup>a</sup> Models 200 and 201, PCNONLIN software, Version 4.2, SCI Software, Lexington, KY; Noncompartmental model

<sup>&</sup>lt;sup>b</sup> PCNONLIN software, Version 4.2, SCI Software, Lexington, KY; Best fit is one compartmental which simultaneously solves iv and mid dose oral data sets. Simultaneous solution of Sprague-Dawley rat intravenous dose ( 2.0 mg/kg Study X) and mid oral gavage dose (2.0 mg/kg Study Z).