Experiment Number: S0629
Route: Dosed Feed, Gavage, IV
Species/Strain: Mouse/B6C3F1

Toxicokinetics Data Summary
Test Compound: Wyeth-14643
CAS Number: 50892-23-4

Date Report Requested: 12/02/2016
Time Report Requested: 11:54:15
Lab: Research Triangle Institute

Male

	Treatment Groups (mg/kg)						
	2 [a]	**2 [b]**	**4 [a]**	**8 [a]**	**50 [c]**	**500 [c]**	**2 IV [a]**
			Plasma				
$C_{max(obs)}$ (ug/mL)	6.94		14.5	20.5	1.26	8.42	
$T_{max(obs)}$	30.0 m		15.0 m	15.0 m	1800.0 h	0200 h	
$t_{1/2(Beta)}$ (minute) k_{01}	64.6		61.6	67.0			243.0
(min^-1)		0.0374					
k_{10} (min^-1)		0.0230 ± 0.0027					
Cl (mL/min/kg)							1.48
$Cl_{1(F)}$ (mL/min/kg)	1.51		3.30	3.37			
V_1 (L/kg)		0.1310 ± 0.0132					
MRT (minute)	415		83.9	100			242
AUC_{inf} (ug/mL*min)	1325.0		1211.0	2376.0	777.0	7060.0	1353
F (fraction)	0.98		0.45	0.44			

LEGEND

Data are displayed as mean ± SEM

m = minutes; h = hours

MODELING METHOD & BEST FIT MODEL

[a] Models 200 and 201, PCNONLIN software, Version 4.2, SCI Software, Lexington, KY; Noncompartmental model

[b] PCNONLIN software, Version 4.2, SCI Software, Lexington, KY; Best fit is one compartmental which simultaneously solves iv and low dose oral data sets. Simultaneous solution of mouse intravenous dose (2.0 mg/kg Study P) and low oral gavage dose (2.0 mg/kg Study Q) fits the data well from time zero to approximately 240 minutes.

[c] PCNONLIN software, Version 4.2, SCI Software, Lexington, KY; Noncompartmental model

ANALYTE

Wyeth-14643

TK PARAMETERS

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

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

$t_{½(beta)}$ = 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}

Cl = Clearance, includes total clearance

$Cl_{1(F)}$ = Apparent clearance of the central compartment, also $Cl_{(F)}$ 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_{inf} = Area under the plasma concentration versus time curve, AUC, extrapolated to time equals infinity

F = Bioavailability, absolute bioavailability

** END OF REPORT **