**Experiment Number:** S0546

Species/Strain: Hamster/Syrian Golden

Route: Gavage, IV

# **Toxicokinetics Data Summary**

Test Compound: 2,4-Dichlorophenoxyacetic Acid

**CAS Number:** 94-75-7

Date Report Requested: 11/09/2016 Time Report Requested: 13:59:32

Lab: Research Triangle Institute

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IVI	a	ıe

	Treatment Groups (mg/kg)						
	<b>2</b> a	8 b	8 a	40 <sup>a</sup>	8 IV <sup>a</sup>		
	Plasma						
C <sub>0min(pred)</sub> (ug/mL)					223.0		
C <sub>max</sub> (ug/mL)	6.86		18.2	70.1			
T <sub>max</sub> (minute)	15		5	5			
t1/2(Beta) (minute)	36.9		24.6	210.0	21.0		
ko1 (min^-1)		0.0467 ± 0.012					
K10 (min^-1)		0.116 ± 0.010					
CI (mL/min/kg)					5.30		
Cl <sub>1(F)</sub> (mL/min/kg)	7.85		14.8	3.38			
√₁ (L/kg)		$0.0605 \pm 0.0055$					
MRT (minute)	103.0		33.0	306.0	12.8		
AUCinf (ug/mL*min)	255.0		541	11830	1510		
F (fraction)	0.67		0.36	1.57			

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# **LEGEND**

Route: Gavage, IV

Data are displayed as mean ± SEM

# MODELING METHOD & BEST FIT MODEL

<sup>a</sup> Modeling Method: Models 200 and 201, PCNONLIN software, SCI Software, Lexington, KY; noncompartmental model (not best fit)

# **ANALYTE**

2,4-Dichlorophenoxyacetic acid

# TK PARAMETERS

 $C_{0min(pred)}$  = Fitted plasma concentration at time zero (IV only)

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

 $T_{max}$  = 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<sub>10</sub> = Elimination rate constant from the central compartment also k<sub>e</sub> or k<sub>elim</sub>

CI = 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 \*\*

<sup>&</sup>lt;sup>b</sup> Analyzed using compartmental modeling techniques with established models or models written to simultaneously solve iv and oral data sets (PCNONLIN software, SCI Software, Lexington, KY); The hamster data were best fit using a 1-compartment model with simultaneous solution of the iv (Study T) and mid oral (Study V) data. The model underpredicted terminal concentrations for both the iv and oral studies.