

Experiment Number: C92012B
 Route: Gavage, IV
 Species/Strain: Mouse/B6C3F1

Toxicokinetics Data Summary
 Test Compound: 2-Methylimidazole
 CAS Number: 693-98-1

Date Report Requested: 11/09/2016
 Time Report Requested: 14:00:18
 Lab: Battelle Columbus

	Male							
	Treatment Groups (mg/kg)							
	25 ^a		50 ^a		100 ^a		10 IV ^b	
	Plasma							
C _{max} (ug/mL)	4.53	± 0.87	10.0	± 1.2	25.4	± 2.2	5.33	± 0.98
T _{max} (hour)	0.127	± 0.076	0.099	± 0.038	0.166	± 0.034		
k ₀₁ (hour ⁻¹)	19.9	± 18.7	26.0	± 15.2	12.9	± 4.6		
t _{1/2(k01)} (hour)	0.035	± 0.033	0.027	± 0.016	0.0537	± 0.0189		
k ₁₀ (hour ⁻¹)	2.07	± 0.34	2.53	± 0.15	2.18	± 0.20	2.66	± 0.32
t _{1/2(k10)} (hour)	0.334	± 0.054	0.274	± 0.016	0.318	± 0.028	0.260	± 0.032
Cl (mL/hr/kg)							5000	± 550
Cl _{1(F)} (mL/hr/kg)	8800	± 1100	9830	± 630	5990	± 410		
V _{ss} (mL/kg)							1880	± 340
MRT (hour)							0.376	± 0.046
AUC _{0-t} (ug/mL*hr)	2.84	± 0.35	5.09	± 0.32	16.7	± 1.1		
AUC _{inf} (ug/mL*hr)							2.00	± 0.22
F (percent)	51.2		49.7		85.0			

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	Female			
	Treatment Groups (mg/kg)			
	25 ^a	50 ^a	100 ^a	10 IV ^b
	Plasma			
C _{max} (ug/mL)	5.89 ± 0.53	13.0 ± 0.4	27.1 ± 1.2	6.27 ± 0.57
T _{max} (hour)	0.147 ± 0.027	0.165 ± 0.010	0.203 ± 0.018	
k ₀₁ (hour ⁻¹)	13.7 ± 4.4	12.2 ± 1.3	9.77 ± 1.66	
t _{1/2(k01)} (hour)	0.051 ± 0.016	0.057 ± 0.006	0.071 ± 0.012	
k ₁₀ (hour ⁻¹)	2.75 ± 0.19	2.44 ± 0.07	2.04 ± 0.12	3.01 ± 0.16
t _{1/2(k10)} (hour)	0.252 ± 0.018	0.284 ± 0.008	0.339 ± 0.020	0.231 ± 0.012
Cl (mL/hr/kg)				4800 ± 270
Cl _{1(F)} (mL/hr/kg)	7790 ± 510	6310 ± 140	4990 ± 180	
V _{ss} (mL/kg)				1600 ± 150
MRT (hour)				0.333 ± 0.018
AUC _{0-t} (ug/mL*hr)	3.21 ± 0.21	7.93 ± 0.17	20.0 ± 0.7	
AUC _{inf} (ug/mL*hr)				2.08 ± 0.12
F (percent)	59.9	74.4	93.9	

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LEGEND

Data are displayed as mean \pm SEM

MODELING METHOD & BEST FIT MODEL

^a WinNonlin V01.5A, using Gauss-Newton (Levenberg and Hartley) method; one-compartment model with no lag phase and first order absorption and elimination. The concentration values were weighted $1/y^2$ (predicted).

^b WinNonlin V01.5A, using Gauss-Newton (Levenberg and Hartley) method; one-compartment model with first order elimination. The concentration values were weighted $1/y^2$.

ANALYTE

2-Methylimidazole

TK PARAMETERS

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

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

k_{01} = Absorption rate constant, k_a

$t_{1/2(k01)}$ = Half-life of the absorption process to the central compartment

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

$t_{1/2(k10)}$ = Half-life for the elimination process from the central compartment

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_{ss} = Volume of distribution at steady state

MRT = Mean residence time

AUC_{0-t} = Area under the plasma concentration versus time curve, AUC, from time t_i (initial) to t_f (final), AUC_{last}

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

F = Bioavailability, absolute bioavailability

**** END OF REPORT ****