

**Experiment Number:** S0305\_1  
**Route:** Gavage, IV  
**Species/Strain:** Mouse/B6C3F1

**Toxicokinetics Data Summary**  
**Test Compound:** Methadone hydrochloride  
**CAS Number:** 1095-90-5

**Date Report Requested:** 01/11/2017  
**Time Report Requested:** 12:23:12  
**Lab:** Research Triangle Institute

	Male				
	Treatment Groups (mg/kg)				
	2.5 <sup>a</sup>	2.5 <sup>b</sup>	15 <sup>b</sup>	2.5 IV <sup>a</sup>	7.5 IV <sup>b</sup>
	Plasma				
C <sub>max</sub> (percent of dose*g/mL)	1.03	1.03	2.63		2.41
T <sub>max</sub> (minute)	15	15	15		45
Lambdaz (minute <sup>-1</sup> )	0.0058			0.0170	
t <sub>1/2</sub> (minute)	119.2			59.4	
k <sub>10</sub> (minute <sup>-1</sup> )		0.0107	0.0063		0.0112
t <sub>1/2(k10)</sub> (minute)		65.0	110.5		61.9
AUC <sub>0-t</sub> (percent of dose*g*min/mL)	73.9	73.9	556.4	910.4	239.1
F (percent of intravenous)	8.1	8.1	61	100	26

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**Female**

**Treatment Groups (mg/kg)**

**2.5<sup>a</sup>**

**15<sup>b</sup>**

**2.5 IV<sup>a</sup>**

**7.5 IV<sup>a</sup>**

**Plasma**

$C_{max}$ (percent of dose*g/mL)	0.83	3.06		2.22
$T_{max}$ (minute)	75	45		75
$\text{Lambdaz}$ (minute <sup>-1</sup> )	0.0181		0.0149	0.0316
$t_{1/2}$ (minute)	38.3		46.7	21.9
$k_{10}$ (minute <sup>-1</sup> )		0.0074		
$t_{1/2(k10)}$ (minute)		93.1		
$AUC_{0-t}$ (percent of dose*g*min/mL)	40.6	232.2	907.7	116.5
F (percent of intravenous)	4.5	26	100	12.8

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#### LEGEND

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Data are displayed as mean values

#### MODELING METHOD & BEST FIT MODEL

<sup>a</sup> ADAPT II (a pharmacokinetic modeling package) was used to perform the nonlinear curve fitting; non-model dependent analysis

<sup>b</sup> ADAPT II (a pharmacokinetic modeling package) was used to perform the nonlinear curve fitting; one compartment model

#### ANALYTE

Methadone hydrochloride

#### TK PARAMETERS

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

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

$\lambda_{dz}$  = Non-compartmental analysis (NCA) terminal elimination rate constant, NCA  $k_e$  or  $k_{elim}$

$t_{1/2}$  =  $\lambda_{dz}$  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  $k_{elim}$

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

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

F = Bioavailability, absolute bioavailability

**\*\* END OF REPORT \*\***