**Experiment Number:** S0555

Route: Dermal, IV

Species/Strain: Mouse/B6C3F1

# Toxicokinetics Data Summary Test Compound: DL-Camphor

**CAS Number:** 21368-68-3

Date Report Requested: 11/09/2016 Time Report Requested: 14:02:35

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							Male					
	Treatment Groups (mg/kg)											
	200 <sup>a, 2</sup>		500 c, 3		500 <sup>d, 4</sup>		<b>500</b> b, 2	500 <sup>a, 4</sup>	500 a, 2	500 a, 3	1000 <sup>a, 2</sup>	50 IV <sup>a, 1</sup>
	Plasma											
Beta (min^-1)	0.0105							0.0091	0.0076	0.0304	0.0062	0.0337
t <sub>1/2(Beta)</sub> (minute)	66.0							75.8	91.5	22.8	112	20.5
k <sub>01</sub> (min^-1)		0.057	1 ± 0.037	0.012	0 ± 0.0066	0.010	0 ± 0.0045					
k <sub>10</sub> (min^-1)		0.066	4 ± 0.0066	0.061	0 ± 0.0050	0.059	3 ± 0.0044					
CI (L/min/kg)												0.121
CI <sub>1(F)</sub> (L/min/kg)	15.1							22.2	17.7	23.7	15.6	
V <sub>1</sub> (L/kg)	1438	1.89	± 0.21	1.99	± 0.18	2.02	± 0.16	2434	2333	780	2522	3.59
MRT (minute)	104							115	140	38.7	188	16.6
AUC <sub>inf</sub> (ug*min/L)	11457							18878	23924	16944	59483	391079
F (fraction)	0.00803							0.00545	0.00686	0.00511	0.00777	

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# Female

		Treatment Groups (mg/kg)								
	200 a, 2	500 e, 2	500 <sup>a, 2</sup>	1000 <sup>a, 2</sup>	50 IV <sup>a, 1</sup>					
			Plasma							
Beta (min^-1)	0.0053		0.0062	0.0067	0.0335					
t <sub>1/2(Beta)</sub> (minute)	131		112	104	20.7					
k <sub>01</sub> (min^-1)		$0.100 \pm 0.0046$								
k <sub>10</sub> (min^-1)		$0.0571 \pm 0.0043$								
Cl (L/min/kg)					0.152					
CI <sub>1(F)</sub> (L/min/kg)	16.6		24.2	22.5						
V <sub>1</sub> (L/kg)	3129	2.59 ± 0.21	3933	3368	4.52					
MRT (minute)	226		190	142	17.4					
AUC <sub>inf</sub> (ug*min/L)	10782		17863	42010	309567					
F (fraction)	0.00913		0.00625	0.00674						

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## **LEGEND**

Data are displayed as mean ± SEM

#### MODELING METHOD & BEST FIT MODEL

<sup>a</sup> WinNonlin, Version 1 .0 (Scientific Consulting Inc., 1995); non-compartmental analysis (WinNonlin Models 200 or 201).

#### **ANALYTE**

**DL-Camphor** 

#### **ROUTE & DOSE FREQUENCY**

- <sup>1</sup> Intravenous, 1 per study
- <sup>2</sup> Dermal Protected, 1 per study
- <sup>3</sup> Dermal Unprotected, 1 per study
- <sup>4</sup> Dermal Unprotected, Repeated dose, 7 per study

### TK PARAMETERS

Beta = Hybrid rate constant of the beta phase

 $t_{\frac{1}{2}(beta)}$  = 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

AUCinf = Area under the plasma concentration versus time curve, AUC, extrapolated to time equals infinity

F = Bioavailability, absolute bioavailability

<sup>&</sup>lt;sup>b</sup> WinNonlin, Version 1 .0 (Scientific Consulting Inc ., 1995); Compartmental models were written to simultaneously solve iv and dermal data sets (WinNonlin) with 1/YHAT weighting, where YHAT is the predicted plasma d,l-camphor concentration at a given time. M and S simultaneously solved iv and single administration mid dose dermal protected, male mice.

<sup>&</sup>lt;sup>c</sup> WinNonlin, Version 1 .0 (Scientific Consulting Inc., 1995); Compartmental models were written to simultaneously solve iv and dermal data sets (WinNonlin) with 1/YHAT weighting, where YHAT is the predicted plasma d,l-camphor concentration at a given time. M and AC simultaneously solved iv and single administration mid dose dermal unprotected, male mice.

<sup>&</sup>lt;sup>d</sup> WinNonlin, Version 1 .0 (Scientific Consulting Inc ., 1995); Compartmental models were written to simultaneously solve iv and dermal data sets (WinNonlin) with 1/YHAT weighting, where YHAT is the predicted plasma d,l-camphor concentration at a given time. M and AF simultaneously solved iv and repeated administration mid dose dermal unprotected, male mice.

<sup>&</sup>lt;sup>e</sup> WinNonlin, Version 1 .0 (Scientific Consulting Inc., 1995); Compartmental models were written to simultaneously solve iv and dermal data sets (WinNonlin) with 1/YHAT weighting, where YHAT is the predicted plasma d,l-camphor concentration at a given time. N and T simultaneously solved iv and single administration mid dose dermal protected, female mice.