Experiment Number: S0548 Route: IV, Gavage Species/Strain: Mouse/B6C3F1	Toxicokinetics Data Summary Compound: Sodium Nitrite / Analyte: Methemoglobin CAS Number: 7632-00-0		Request Date: 7/11/2023 Request Time: 10:03:16 Lab: Midwest Research Institute	
		Male		
		Treatment Group (mg/kg)		
	20 IV Plasma ^{a,b}	20 IV Plasma ^{a,b} 62.5 Gavage Plasma ^{a,b} 125 Gavage Pla		sma ^{a,b}
Cmax_pred (percent)	4.2	14.9	35.1	
Tmax_pred (min)	5	15	15	
Half-life (min)	18	23	25	
MRT (min)	29	38	55	
AUC_0-T (percent min)	151	665	2720	

 Experiment Number: \$0548
 Toxicokinetics Data Summary
 Request Date: 7/11/2023

 Route: IV, Gavage
 Compound: Sodium Nitrite / Analyte: Methemoglobin
 Request Time: 10:03:16

 Species/Strain: Mouse/B6C3F1
 CAS Number: 7632-00-0
 Lab: Midwest Research Institute

 Female
 Treatment Group (mg/kg)
 120 Gavage Plasma^{a,b}

Cmax_pred (percent)	4.8	11.2	38.6
Tmax_pred (min)	2	15	60
Half-life (min)	15	18	14
MRT (min)	26	37	60
AUC_0-T (percent min)	141	669	3620

LEGEND

MODELING SOFTWARE PCNONLIN

MODELING METHOD & BEST FIT MODEL

^aPCNONLIN Statistical Consultants, Inc., Lexington, KY, Non compartmental (NCA) model

EXCEPTIONS

^bwhere Half-life is equal to Kc half-life

ANALYTE

Methemoglobin

TK PARAMETERS

Cmax_pred = Observed or Predicted Maximum plasma (or tissue) concentration Tmax_pred = Time at which Cmax predicted or observed occurs Half-Life = Lambda z Half life, t 1/2, the terminal elimination half-life based on non-compartmental analysis MRT = Mean residence time AUCinf = Area under the plasma concentration versus time curve, AUC, extrapolated to time equals infinity Toxicokinetics Data Summary Compound: Sodium Nitrite / Analyte: Methemoglobin CAS Number: 7632-00-0 Request Date: 7/11/2023 Request Time: 10:03:16 Lab: Midwest Research Institute

TK PARAMETERS PROTOCOL

ANALYSIS METHOD

Blood collection time points for this group are 2, 5, 10, 15, 20, 30, 45, 60, 75, and 90 minutes post-dose.

TK_INTRAVENOUS PLASMA

20 mg/kg Male and Female

A single intravenous dose of 20 mg/kg was given per study via lateral tail vein. Toxicokinetic analyses were performed using the average concentration for each time point. The data were modeled using nonlinear regression analysis (PCNONLIN, Statistical Consultants, Inc., Lexington, KY). The nitrite data was modeled using compartmental models.

ANALYSIS METHOD

Blood collection time points for this group are 2, 5, 10, 15, 30, 45, 60, 90, 120, and 150 minutes post-dose.

TK_GAVAGE PLASMA

62.5 mg/kg Male and Female

A single oral gavage dose of 62.5 mg/kg was given per study. Toxicokinetic analyses were performed using the average concentration for each time point. The data were modeled using nonlinear regression analysis (PCNONLIN, Statistical Consultants, Inc., Lexington, KY). The nitrite data was modeled using compartmental models.

125 mg/kg Male and Female

A single oral gavage dose of 125 mg/kg was given per study. Toxicokinetic analyses were performed using the average concentration for each time point. The data were modeled using nonlinear regression analysis (PCNONLIN, Statistical Consultants, Inc., Lexington, KY). The nitrite data was modeled using compartmental models.