eriment Number: S0548 te: IV, Gavage cies/Strain: Rats/FISCHER 344	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/k	g)	
	20 IV Plasma ^{a,b}	40 Gavage Plasma ^{a,b}	80 Gavage Plasm	a ^{a,b}
Cmax pred (percent)	26.3	18.8	40.1	
Tmax_pred (min)	45	60	120	
Half-life (min)	53	50	75	
MRT (min)	94	171	117	
AUC_0-T (percent min)	3260	10500	2620	

periment Number: S0548 pute: IV, Gavage ecies/Strain: Rats/FISCHER 344	Toxicokinetics Data Summary Compound: Sodium Nitrite / Analyte: Methemoglobin C AS Number: 7632-00-0			Request Date: 7/11/2023 Request Time: 10:03:16 Lab: Midwest Research Institute		
		Female				
	Treatment Group (mg/kg)					
	20 IV Plasma ^{a,b}	40 Gavage Plasma ^{a,b}	80 Gavage Plası	ma ^{a,b}		
Cmax pred (percent)	25.8	65.9	31.5			
Tmax_pred (min)	45	60	120			
Half-life (min)	82	73	81			
MRT (min)	130	125	186			
AUC_0-T (percent min)	4340	5480	16400			

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

TK PARAMETERS PROTOCOL

ANALYSIS METHOD

Blood collection time points for this group are 2, 5, 10, and 30 minutes, 1, 2, 4, 6, 8, and 10 hours 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, and 30 minutes, 1, 2, 4, 6, 8, and 10 hours post-dose.

TK_GAVAGE PLASMA

40 mg/kg Male and Female

A single oral gavage dose of 40 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.

80 mg/kg Male and Female

A single oral gavage dose of 80 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.