Experiment Number: S0609
Route: Oral followed by IV
Species/Strain: Mouse/B6C3F1

Toxicokinetics Data Summary
Test Compound: AZT + TMP/SMX (mixture) combination
CAS Number: AZTTMPSMX

Date Report Requested: 02/01/2017
Time Report Requested: 12:59:21
Lab: Research Triangle Institute International

	Male							
	Treatment Groups (mg/kg)							
	100/250 #	100/250 °	100/250 *	100/250 ~	100/1000 #	100/1000 °	100/1000 *	100/1000 ~
	Plasma							
C_{max} (ug/mL)	179	1.08	2.04	0.442	184	0.846	1.45	0.590
T_{max} (hour)		0.250	6.00	0.500		0.250	0.750	0.750
$Lambda_z$ (hour^-1)	2.15				1.45			
$t_{1/2}$ (hour)	0.322				0.480			
Cl (mL/min/kg)	20.7				18.9			
V_1 (L/kg)	0.577				0.784			
MRT (hour)	0.488				0.519			
AUC_{0-t} (ug*hr/mL)		0.580	7.34	0.183		0.557	1.33	0.345

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	Female							
	Treatment Groups (mg/kg)							
	100/250 °	100/250 ~	100/250 *	100/250 #	100/1000 °	100/1000 #	100/1000 ~	100/1000 *
	Plasma							
C_{max} (ug/mL)	0.724	0.105	1.15	164	0.629	105	0.365	0.716
T_{max} (hour)	0.250	1.00	6.00		0.250		1.00	0.500
Lambda$_z$ (hour^-1)				1.87		1.78		
$t_{1/2}$ (hour)				0.370		0.391		
Cl (mL/min/kg)				18.6		21.3		
V_1 (L/kg)				0.597		0.721		
MRT (hour)				1.07		0.548		
AUC_{0-t} (ug*hr/mL)	0.340	0.0222	3.05		0.415		0.0834	0.535

LEGEND

Data are displayed as mean values

MODELING METHOD & BEST FIT MODEL

WinNonlin (Model 200 and 201, WinNonlin Ver. 1.5A, Scientific Consulting, Inc. now Pharsight Corporation, Apex, NC); Non compartmental analysis

ANALYTE

3'-Azido-3'-deoxythymidine

* 3'-Amino-3'-deoxythymidine

~ 3'-amino-3'-deoxythymidine glucuronide

° Beta-D-glucuronide

DOSING

Mice were given a single oral dose of Trimethoprim-Sulfamethoxazole (TMP/SMX) followed one hour later by a single intravenous dose of 3'-Azido-3'-deoxythymidine (AZT) on Study Day 1.

TK PARAMETERS

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

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

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

$t_{½}$ = $Lambda_z$ half-life, $t_{1/2}$, the terminal elimination half-life based on non-compartmental analysis

Cl = Clearance, includes total clearance

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

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

** END OF REPORT **