ATTACHMENT B.2

INHIBITION STUDY: DATA TABLES

Table 1: CYP3A4 Activity in Human Pooled Human Microsomes Following Incubation with 1 µM Ketoconazole

Sample Identification	6β-Hydroxytestosterone formation Raw (µM)	Adjusted (µM) Individual	Adjusted (µM) Mean ± SD	Specific Activity (pmol/min/mg protein) Individual	Specific Activity (pmol/min/mg protein) Mean ± SD	Percent of VC
			Incubation 1			
VC	0.12662*	N/A	0.742 ± 0.00679	N/A	594 ± 5.43	100
	0.74589	0.746		597		
	0.74640	0.746		597		
	0.73440	0.734		588		
1 µM Ketoconazole	0.06270 [a]	<0.100	<0.100 ± 0.000	<80.0	<80.0 ± 0.000	<13.5
	0.06153 [a]	<0.100		<80.0		
	0.06413 [a]	<0.100		<80.0		
	0.06278 [a]	<0.100		<80.0		
			Incubation 3			
VC	0.85601	0.856	0.844 ± 0.0149	685	676 ± 11.9	100
	0.83718	0.837		670		
	0.85744	0.857		686		
	0.82690	0.827		662		
1 µM Ketoconazole	0.07141 [a]	<0.100	<0.100 ± 0.000	<80.0	<80.0 ± 0.000	<11.8
	0.06825 [a]	<0.100		<80.0		
	0.06742 [a]	<0.100		<80.0		
	0.06916 [a]	<0.100		<80.0		

Abbreviations: SD, standard deviation; VC, vehicle control (1% Methanol)
* Sample has been removed from all calculations due to the incorrect volume being added to the sample to stop the reaction.
[a] The observed analyzed value (µM) was below the lowest concentration on the standard curve (0.1 µM).
Note: For all calculations above, the resulting values are shown with at least three significant figures for display purposes only.

Table 2: CYP1A2 Activity in Human Pooled Human Microsomes Following Incubation with Metaxalone

Metaxalone (μM)	Acetaminophen formation Raw (μM)	Adjusted (μM) Individual	Adjusted (μM) Mean ± SD	Specific Activity (pmol/min/mg protein) Individual	Specific Activity (pmol/min/mg protein) Mean ± SD	Percent of VC
0	0.23653	0.237	0.216 ± 0.0138	31.5	28.8 ± 1.84	100
(VC)	0.21124	0.211		28.2		
	0.21156	0.212		28.2		
	0.20568	0.206		27.4		
0.3	0.21120	0.211	0.210 ± 0.00536	28.2	28.0 ± 0.715	97.2
	0.21487	0.215		28.6		
	0.20431	0.204		27.2		
1	0.19966	0.200	0.200 ± 0.00246	26.6	26.6 ± 0.327	92.3
	0.19709	0.197		26.3		
	0.20200	0.202		26.9		
3	0.19900	0.199	0.195 ± 0.00589	26.5	26.0 ± 0.785	90.3
	0.18839	0.188		25.1		
	0.19813	0.198		26.4		
30	0.18924	0.189	0.194 ± 0.00544	25.2	25.9 ± 0.725	89.8
	0.19323	0.193		25.8		
	0.20000	0.200		26.7		
100	0.17757	0.178	0.177 ± 0.000206	23.7	23.6 ± 0.0275	82.0
	0.17733	0.177		23.6		
	0.17716	0.177		23.6		

Abbreviations: SD, standard deviation; VC, vehicle control (1% Methanol)
Note: For all calculations above, the resulting values are shown with at least three significant figures for display purposes only.

Table 3: CYP2A6 Activity in Human Pooled Human Microsomes Following Incubation with Metaxalone

Metaxalone (μM)	7-Hydrxoycoumarin formation Raw (μM)	Adjusted (μM) Individual	Adjusted (μM) Mean ± SD	Specific Activity (pmol/min/mg protein) Individual	Specific Activity (pmol/min/mg protein) Mean ± SD	Percent of VC
0 (VC)	1.03214	1.03	1.06 ± 0.0356	413	426 ± 14.2	100
	1.04464	1.04		418		
	1.06891	1.07		428		
	1.11282	1.11		445		
0.3	1.07439	1.07	1.03 ± 0.0399	430	413 ± 16.0	96.9
	0.99553	0.996		398		
	1.02457	1.02		410		
1	0.99854	0.999	1.02 ± 0.0184	399	407 ± 7.36	95.7
	1.02269	1.02		409		
	1.03468	1.03		414		
3	1.05100	1.05	1.09 ± 0.0402	420	436 ± 16.1	102
	1.13132	1.13		453		
	1.08822	1.09		435		
30	1.08205	1.08	1.14 ± 0.0493	433	455 ± 19.7	107
	1.15129	1.15		461		
	1.17736	1.18		471		
100	0.98864	0.989	1.01 ± 0.0416	395	404 ± 16.6	94.8
	0.98209	0.982		393		
	1.05713	1.06		423		

Abbreviations: SD, standard deviation; VC, vehicle control (1% Methanol)
Note: For all calculations above, the resulting values are shown with at least three significant figures for display purposes only.

Table 4: CYP2B6 Activity in Human Pooled Human Microsomes Following Incubation with Metaxalone

Metaxalone (μM)	Nirvanol formation Raw (μM)	Adjusted (μM) Individual	Adjusted (μM) Mean ± SD	Specific Activity (pmol/min/mg protein) Individual	Specific Activity (pmol/min/mg protein) Mean ± SD	Percent of VC
0 (VC)	0.23500	0.235	0.225 ± 0.0120	31.3	29.9 ± 1.60	100
	0.23266	0.233		31.0		
	0.22199	0.222		29.6		
	0.20877	0.209		27.8		
0.3	0.20942	0.209	0.203 ± 0.00904	27.9	27.0 ± 1.21	90.2
	0.19234	0.192		25.6		
	0.20601	0.206		27.5		
1	0.20438	0.204	0.223 ± 0.0201	27.3	29.8 ± 2.68	99.5
	0.22144	0.221		29.5		
	0.24442	0.244		32.6		
3	0.19695	0.197	0.203 ± 0.00751	26.3	27.1 ± 1.00	90.6
	0.20166	0.202		26.9		
	0.21166	0.212		28.2		
30	0.21681	0.217	0.217 ± 0.00162	28.9	28.9 ± 0.216	96.6
	0.21548	0.215		28.7		
	0.21871	0.219		29.2		
100	0.18648	0.186	0.188 ± 0.00436	24.9	25.1 ± 0.581	83.7
	0.18463	0.185		24.6		
	0.19293	0.193		25.7		

Abbreviations: SD, standard deviation; VC, vehicle control (1% Methanol)
Note: For all calculations above, the resulting values are shown with at least three significant figures for display purposes only.

Table 5: CYP2C8 Activity in Human Pooled Human Microsomes Following Incubation with Metaxalone

Metaxalone (μM)	6-Hydroxypaclitaxel formation Raw (μM)	Adjusted (μM) Individual	Adjusted (μM) Mean ± SD	Specific Activity (pmol/min/mg protein) Individual	Specific Activity (pmol/min/mg protein) Mean ± SD	Percent of VC
0	0.13462	0.135	0.136 ± 0.00522	17.9	18.2 ± 0.696	100
(VC)	0.14017	0.140		18.7		
	0.14074	0.141		18.8		
	0.12965	0.130		17.3		
0.3	0.14476	0.145	0.126 ± 0.0163	19.3	16.8 ± 2.18	92.7
	0.11377	0.114		15.2		
	0.12042	0.120		16.1		
1	0.13927	0.139	0.140 ± 0.00305	18.6	18.7 ± 0.406	103
	0.13749	0.137		18.3		
	0.14343	0.143		19.1		
3	0.15034	0.150	0.149 ± 0.00174	20.0	19.9 ± 0.232	109
	0.14945	0.149		19.9		
	0.14698	0.147		19.6		
30	0.14949	0.149	0.138 ± 0.0114	19.9	18.4 ± 1.52	101
	0.13724	0.137		18.3		
	0.12667	0.127		16.9		
100	0.13170	0.132	0.133 ± 0.0207	17.6	17.8 ± 2.76	97.8
	0.15467	0.155		20.6		
	0.11340	0.113		15.1		

Abbreviations: SD, standard deviation; VC, vehicle control (1% Methanol)
Note: For all calculations above, the resulting values are shown with at least three significant figures for display purposes only.

Table 6: CYP2C9 Activity in Human Pooled Human Microsomes Following Incubation with Metaxalone

Metaxalone (μM)	4'-Methylhydroxytolbutamide formation Raw (μM)	Adjusted (μM) Individual	Adjusted (μM) Mean ± SD	Specific Activity (pmol/min/mg protein) Individual	Specific Activity (pmol/min/mg protein) Mean ± SD	Percent of VC
0 (VC)	0.17476	0.175	0.166 ± 0.0208	46.6	44.3 ± 5.54	100
	0.14904	0.149		39.7		
	0.14954	0.150		39.9		
	0.19164	0.192		51.1		
0.3	0.13620	0.136	0.135 ± 0.00106	36.3	36.1 ± 0.283	81.4
	0.13415	0.134		35.8		
	0.13565	0.136		36.2		
1	0.15107	0.151	0.136 ± 0.0187	40.3	36.2 ± 4.98	81.6
	0.14080	0.141		37.5		
	0.11485	0.115		30.6		
3	0.13051	0.131	0.135 ± 0.0103	34.8	36.0 ± 2.75	81.2
	0.12759	0.128		34.0		
	0.14670	0.147		39.1		
30	0.14975	0.150	0.151 ± 0.00841	39.9	40.3 ± 2.24	91.0
	0.14376	0.144		38.3		
	0.16037	0.160		42.8		
100	0.16269	0.163	0.145 ± 0.0150	43.4	38.8 ± 4.00	87.4
	0.13711	0.137		36.6		
	0.13627	0.136		36.3		

Abbreviations: SD, standard deviation; VC, vehicle control (1% Methanol)
Note: For all calculations above, the resulting values are shown with at least three significant figures for display purposes only.

Table 7: CYP2C19 Activity in Human Pooled Human Microsomes Following Incubation with Metaxalone

Metaxalone (μM)	4'-Hydroxymephenytoin formation Raw (μM)	Adjusted (μM) Individual	Adjusted (μM) Mean ± SD	Specific Activity (pmol/min/mg protein) Individual	Specific Activity (pmol/min/mg protein) Mean ± SD	Percent of VC
0 (VC)	0.16904	0.169	0.168 ± 0.00550	22.5	22.4 ± 0.733	100
	0.17373	0.174		23.2		
	0.16915	0.169		22.6		
	0.16055	0.161		21.4		
0.3	0.13971	0.140	0.142 ± 0.00299	18.6	19.0 ± 0.399	84.6
	0.14558	0.146		19.4		
	0.14164	0.142		18.9		
1	0.11367	0.114	0.113 ± 0.00140	15.2	15.0 ± 0.186	67.0
	0.11336	0.113		15.1		
	0.11111	0.111		14.8		
3	0.11597	0.116	0.114 ± 0.00238	15.5	15.2 ± 0.317	67.7
	0.11127	0.111		14.8		
	0.11423	0.114		15.2		
30	0.08336	0.0834	0.107 ± 0.0211	11.1	14.3 ± 2.82	63.8
	0.12339	0.123		16.5		
	0.11502	0.115		15.3		
100	0.10857	0.109	0.109 ± 0.00205	14.5	14.5 ± 0.274	64.9
	0.11132	0.111		14.8		
	0.10730	0.107		14.3		

Abbreviations: SD, standard deviation; VC, vehicle control (1% Methanol)
Note: For all calculations above, the resulting values are shown with at least three significant figures for display purposes only.

Table 8: CYP2D6 Activity in Human Pooled Human Microsomes Following Incubation with Metaxalone

Metaxalone (µM)	Dextrorphan formation Raw (µM)	Adjusted (µM) Individual	Adjusted (µM) Mean ± SD	Specific Activity (pmol/min/mg protein) Individual	Specific Activity (pmol/min/mg protein) Mean ± SD	Percent of VC
0 (VC)	0.18550	0.186	0.183 ± 0.00342	49.5	48.9 ± 0.911	100
	0.18569	0.186		49.5		
	0.18424	0.184		49.1		
	0.17843	0.178		47.6		
0.3	0.14820	0.148	0.149 ± 0.00258	39.5	39.8 ± 0.688	81.3
	0.14716	0.147		39.2		
	0.15206	0.152		40.5		
1	0.15910	0.159	0.154 ± 0.00482	42.4	41.2 ± 1.28	84.2
	0.14949	0.149		39.9		
	0.15485	0.155		41.3		
3	0.16116	0.161	0.164 ± 0.00353	43.0	43.7 ± 0.940	89.3
	0.16267	0.163		43.4		
	0.16788	0.168		44.8		
30	0.15533	0.155	0.156 ± 0.00335	41.4	41.6 ± 0.893	85.1
	0.15983	0.160		42.6		
	0.15328	0.153		40.9		
100	0.15992	0.160	0.158 ± 0.00255	42.6	42.0 ± 0.680	85.9
	0.15489	0.155		41.3		
	0.15813	0.158		42.2		

Abbreviations: SD, standard deviation; VC, vehicle control (1% Methanol)
Note: For all calculations above, the resulting values are shown with at least three significant figures for display purposes only.

Table 9: CYP2E1 Activity in Human Pooled Human Microsomes Following Incubation with Metaxalone

Metaxalone (µM)	6-Hydroxychlorzoxazone formation Raw (µM)	Adjusted (µM) Individual	Adjusted (µM) Mean ± SD	Specific Activity (pmol/min/mg protein) Individual	Specific Activity (pmol/min/mg protein) Mean ± SD	Percent of VC
0 (VC)	0.85346	0.853	0.852 ± 0.0127	228	227 ± 3.39	100
	0.86925	0.869		232		
	0.84615	0.846		226		
	0.83969	0.840		224		
0.3	0.73634	0.736	0.710 ± 0.0228	196	189 ± 6.08	83.3
	0.69947	0.699		187		
	0.69469	0.695		185		
1	0.72701	0.727	0.716 ± 0.0194	194	191 ± 5.18	84.0
	0.72685	0.727		194		
	0.69326	0.693		185		
3	0.76089	0.761	0.755 ± 0.0110	203	201 ± 2.94	88.6
	0.74221	0.742		198		
	0.76169	0.762		203		
30	0.71716	0.717	0.733 ± 0.0145	191	196 ± 3.88	86.1
	0.74538	0.745		199		
	0.73733	0.737		197		
100	0.74969	0.750	0.743 ± 0.0175	200	198 ± 4.66	87.2
	0.75620	0.756		202		
	0.72321	0.723		193		

Abbreviations: SD, standard deviation; VC, vehicle control (1% Methanol)
Note: For all calculations above, the resulting values are shown with at least three significant figures for display purposes only.

Table 10: CYP3A4 Activity in Human Pooled Human Microsomes Following Incubation with Metaxalone

Metaxalone (μM)	6β-Hydroxytestosterone formation Raw (μM)	Adjusted (μM) Individual	Adjusted (μM) Mean ± SD	Specific Activity (pmol/min/mg protein) Individual	Specific Activity (pmol/min/mg protein) Mean ± SD	Percent of VC
0	0.12662*	N/A	0.742 ± 0.00679	N/A	594 ± 5.43	100
(VC)	0.74589	0.746		597		
	0.74640	0.746		597		
	0.73440	0.734		588		
0.3	0.64318	0.643	0.647 ± 0.0130	515	517 ± 10.4	87.1
	0.66083	0.661		529		
	0.63550	0.636		508		
1	0.65762	0.658	0.654 ± 0.00353	526	523 ± 2.83	88.1
	0.65446	0.654		524		
	0.65057	0.651		520		
3	0.67154	0.672	0.668 ± 0.00420	537	534 ± 3.36	90.0
	0.66336	0.663		531		
	0.66907	0.669		535		
30	0.62513	0.625	0.633 ± 0.0370	500	506 ± 29.6	85.2
	0.67282	0.673		538		
	0.59996	0.600		480		
100	0.63960	0.640	0.596 ± 0.0454	512	477 ± 36.3	80.3
	0.59940	0.599		480		
	0.54904	0.549		439		

Abbreviations: SD, standard deviation; VC, vehicle control (1% Methanol)
* Sample has been removed from all calculations due to the incorrect volume being added to the sample to stop the reaction.
Note: For all calculations above, the resulting values are shown with at least three significant figures for display purposes only.