Electronic Supplementary Material (ESI) for Toxicology Research.

SUPPLEMENTARY INFORMATION

Table S1. P450 inhibition results for seven methylindole (MI) analogs for major CYP isoforms. The two CYP3A probes were testosterone (T) and midazolam (M).

Compound	CYP1A2	CYP2C9	CYP2C19	CYP2D6	CYP3A (T)	CYP3A (M)
1MI	>20	>20	>20	>20	4.8	3.3
2MI	>20	>20	>20	>20	>20	>20
3MI	>20	>20	>20	>20	>20	>20
4MI	>20	>20	>20	>20	>20	>20
5MI	>20	>20	>20	>20	>20	>20
6MI	>20	>20	>20	>20	>20	>20
7MI	>20	>20	>20	>20	>20	>20

Table S2. Time-dependent inhibition IC_{50} (μM) and AUC shift for seven methylindole (MI) analogs. The assay's positive results are with AUC shift of greater than 15% with or without NADPH preincubation for 30 min.

Compound	P450 TDI IC_{50} in μM (% change in AUC)					
	CYP1A2	CYP2C9	CYP2C19	CYP2D6	CYP3A-M	CYP3A-T
1MI	N	N	N	N	0.63 (28)	1.76 (38)
2MI	N	N	N	N	N	N
3MI	N	N	N	N	N	N
4MI	N	N	N	N	N	7.23 (21)
5MI	N	N	N	N	>10.0 (22)	5.40 (15)
6MI	N	N	N	N	>10 (33)	2.06 (34)
7MI	>10 (15)	N	>10 (16)	7.42 (26)	>10 (23)	6.60 (32)

N = no time-dependent inhibition with an IC_{50} greater than 10 μM and an AUC change of less than 15%.

Table S3.

Analyte	**RT** (min)	**Conjugate** $[M+H]^+$	**Modification**	**Fragmentation** *(m/z)*
1MI	4.37	322.0863	+O-2H+GSH-Glu	304.0763, 258.0884, 203.0645
2MI	4.32	453.1443	+O+GS	435.1338, 378.1126, 324.1026, 178.0327
3MI	4.97	453.1445	+O+GS	435.1350, 378.1134, 324.1025, 274.1047, 146.0602
	5.17 (A)	437.1495	+GS	308.0922, , 179.0489, 130.0648
	5.87 (B)	437.1495	+GS	362.1185, 308.1076, 274.1042
4MI	3.94	338.0814	+2O-2H+GS-Glu	320.0712, 274.0834, 219.0596
	4.28	453.1443	+O+GS	324.1020, 308.0923, , 179.0488
	5.00	322.0861	+O-2H+GS-Glu	304.0759, 258.0882, 229.0434, 203.0643
5MI	4.38	322.0863	+O-2H+GS-Glu	304.0762, 270.0869, 258.0883, 229.0437, 219.0788, 203.0645
6MI	3.40	338.0813	+2O-2H+GS-Glu	320.0709, 274.0831, 219.0592,
	3.83	338.0811	+2O-2H+GS-Glu	320.0710, 274.0832, 235.0549, 219.0594,
	5.14	322.0862	+O-2H+GS-Glu	258.0884, 229.0441, 219.0767, , 203.0646
7MI	3.56	338.0814	+2O-2H+GS-Glu	320.0709, 274.0830, 219.0594, ,
	4.94	322.0861	+O-2H+GS-Glu	304.0760, 258.0880, 229.0433, 203.0643,