GRX-917 – Identical Metabolite Profile to Etifoxine

Identical Metabolite Profile

Importantly, GRX-917 showed an identical metabolite profile to etifoxine in human liver microsomes. The same metabolites were produced in the same ratios by both molecules but in lower overall amounts for deuterated etifoxine. The Company's GRX-917 is therefore expected to have the same, or superior safety and side-effect profile as etifoxine.

Major metabolites (Mass shift only) of Etifoxine and GRX-917 incubated with human liver microsomes in the presence of NADPH (0 min or 60 min)

A) Etifoxine

Peak ID	Mass Shift	Found m/z	R.T. (min)	UV absorption*	MS peak area
				HLM (60 min)	
Parent (0min)	0	301	4.98	100.00%	3.45E+05
Parent (60 min)	0	301	4.98	13.55%	9.86E+04
M1	-18	283	4.37	+	2.03E+03
M2	-12	289	4.66	+	5.37E+03
M3	+16	317	4.72	+	3.08E+04
M4	-28	273	4.74	50.81%	2.42E+05
M5	+16	317	4.99	+	2.09E+03

B) GRX-917

Peak ID	Mass Shift	Found m/z	R.T. (min)	UV absorption*	MS peak area
				HLM (60 min)	
Parent (0min)	0	306	4.98	100.00%	3.12E+05
Parent (60 min)	0	306	4.98	31.31%	1.41E+05
M1	-18	288	4.37	+	4.47E+03
M2	-17	289	4.66	+	2.33E+03
M3	+15	321	4.72	+	1.15E+04
M4	-33	273	4.74	35.69%	1.60E+05
M5	+16	322	4.99	+	7.34E+03

*All percentages were calculated based on the detected UV absorption relative to that of parent in T_{0min} samples (normalized as 100%)
+: only detected in MS