

**Supplemental Table S2.** Adverse side effect profiling of VU0152100 using the Modified Irwin Neurological Test Battery

	<i>Vehicle</i>					<i>1 mg/kg Oxotremorine</i>					<i>56.6 mg/kg VU0152100</i>				
<i>Time</i>	<i>5</i>	<i>15</i>	<i>60</i>	<i>120</i>	<i>240</i>	<i>5</i>	<i>15</i>	<i>60</i>	<i>120</i>	<i>240</i>	<i>5</i>	<i>15</i>	<i>60</i>	<i>120</i>	<i>240</i>
<b><i>Autonomic nervous system</i></b>															
<i>Ptosis</i>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
<i>Exopthalmus</i>	-	-	-	-	-	++ **	++ **	-	-	-	-	-	-	-	-
<i>Miosis</i>	++	++++	+++	+++	+++	- **	- **	- **	- **	- **	++	++ **	(+)	++	++
<i>Mydriasis</i>	-	-	-	-	-	++++ **	+++ **	++ **	-	-	-	(+)	(+)	-	-
<i>Corneal reflex loss</i>	-	-	-	-	-	(+)	-	-	-	-	-	-	-	-	-
<i>Pinna reflex loss</i>	-	-	-	-	(+)	++	++ **	(+)	-	-	-	-	-	-	-
<i>Piloerection</i>	-	-	-	-	-	+++ **	++ **	(+)	-	-	-	-	-	-	-
<i>Respiratory rate</i>	-	-	-	-	-	++ **	(+)	(+)	-	-	-	-	-	-	-
<i>Writhing</i>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
<i>Tail erection</i>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
<i>Lacrimation</i>	-	-	-	-	-	+++ **	++ **	-	-	-	-	-	-	-	-
<i>Salivation</i>	-	-	-	-	-	+++ **	++ **	-	-	-	-	-	-	-	-
<i>Vasodilation</i>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
<i>Skin colour</i>	-	-	-	-	-	+ **	-	-	-	-	-	-	-	-	-
<i>Irritability</i>	(+)	-	-	(+)	(+)	+++ **	++ **	++ **	+	(+)	-	-	-	-	-
<i>Rectal temp. (°C)</i>	36.5	36.9	37.0	36.9	36.9	35.7	35.5 **	36.0 *	36.7	36.9	37.1	37.2	37.0	37.3	36.7
<b><i>Somatomotor systems</i></b>															
<i>Motor activity</i>	-	-	-	-	-	++++ **	++++ **	+++ **	+ **	(+)	-	-	-	-	-
<i>Convulsions</i>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
<i>Arch/roll</i>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
<i>Tremors</i>	-	-	-	-	-	++ **	(+)	(+)	-	-	-	-	-	-	-
<i>Leg weakness</i>	-	-	-	-	-	++++ **	++++ **	+++ **	++ **	(+)	-	-	-	-	-
<i>Rigid stance</i>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
<i>Spraddle</i>	-	-	-	-	-	++++ **	+++ **	+	-	-	-	-	-	-	-
<i>Placing loss</i>	-	-	-	-	-	(+)	-	-	-	-	-	-	-	-	-
<i>Grasping loss</i>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
<i>Righting loss</i>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
<i>Catalepsy</i>	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-
<i>Tail pinch</i>	-	-	-	-	-	++++ **	++++ **	++++ **	-	-	-	-	-	-	-
<i>Escape loss</i>	-	-	-	-	-	++++ **	++++ **	++ **	(+)	-	-	-	-	-	-

The effects of VU0152100 (56.6 mg/kg, ip) or vehicle (10 % Tween 80) on autonomic nervous system and somatomotor function were compared to oxotremorine (1 mg/kg, sc). The mean scores of five animals per treatment are classified as follows: -, = no effect; (+), 0.01 – 0.25; +, 0.251 – 0.50; ++, 0.51 – 1.0; +++, 1.01 – 1.50; +++++, 1.51 – 2.0; \* p<0.05, \*\* p<0.01 vs. Vehicle (Dunnett's test)