

Supplemental Table 1. Iterative injury protocols. Development of the model involved the iterative increase in LPS dose, changes in dose timing, and manipulating the delay between LPS doses and hypoxia. Initially, LPS alone was used. Subsequently a period of hypoxia was added, then a period of hyperoxia after hypoxia and, finally, a second period of hypoxia with continuous rectal temperature monitoring during the insult period. Data from the penultimate row is depicted in Supplemental Figure 1. The current protocol is detailed in the final row.

Age at insult	LPS	No. of LPs doses	Delay (last LPS dose to hypoxia)	Hypoxia	Hyperoxia	2nd Hypoxia	Waterbath temp during insult (°C)	Rectal temp during insult (°C)
P9-P12	10mcg/kg/24h	4	N/A	N/A	N/A	N/A	N/A	N/A
P9-P12	50-100mcg/kg/24h	3	24h	8% for 4 h	N/A	N/A	35.7-37.5	N/A
P9-P11	100mcg/kg/24h	3	None	6% for 6h	N/A	N/A	35.7-37.5	N/A
P10	0.5-1mg/kg	1	None	6% for 4h	N/A	N/A	37	N/A
P9-P10	1mg/kg/12h	1 vs 3	None	6% for 45min	100% for 6h	N/A	37	N/A
P9-P10	2 vs 5 vs 10mg/kg/12h	3	None	10% for 2h	80% for 2h	10% for 2h	40	37 (hypoxia) 38 (hyperoxia)
P10-P11	5mg/kg	1	4h	9% for 2h	60% for 2h	9% for 2h	40	37 (hypoxia) 38 (hyperoxia)