

Supplementary Table 3. Measures of Cold-Induced Pain

<i>PAIN MEASURE</i>	<i>TRAMADOL VERSUS PLACEBO</i>	<i>DESMETRAMADOL VERSUS PLACEBO</i>	<i>DESMETRAMADOL VERSUS TRAMADOL</i>
Trial A (<i>n</i> = 43, males)			
Pain intensity, 0-10, MPD (SE)			
At 30 seconds	-0.46 (0.19) <i>P</i> = 0.015	-0.60 (0.15) <i>P</i> = 0.0019	-0.12 (0.16) <i>P</i> = 0.5
At first perception	-0.21 (0.10) NS	-0.15 (0.09) NS	0.07 (0.10) NS
Time, seconds, MD (SE)			
Withdrawal	12.4 (4.4) <i>P</i> = 0.0076	12.6 (3.8) <i>P</i> = 0.006	0.03 (4.4) <i>P</i> = 1.0
First pain	2.1 (0.85) <i>P</i> = 0.029	2.6 (0.73) <i>P</i> = 0.0057	0.42 (0.93) <i>P</i> = 0.7
Trial B (<i>n</i> = 42, males)			
Pain intensity, 0-10, MPD (SE)			
At 30 seconds	-0.25 (0.26) <i>P</i> = NS	-0.75 (0.28) <i>P</i> = 0.036	-0.62 (0.20) <i>P</i> = 0.003
At first perception	-0.08 (0.15) NS	-0.17 (0.14) NS	-0.09 (0.13) NS
Time, seconds, MPD (SE)			
Withdrawal	6.9 (1.6) <i>P</i> = 0.001	9.9 (1.9) <i>P</i> < 0.001	2.8 (2.4) <i>P</i> = 0.126
First pain	1.5 (0.5) <i>P</i> = 0.013	2.1 (0.5) <i>P</i> = 0.001	0.6 (0.6) <i>P</i> = 0.193
Trial B (<i>n</i> = 18, females)			
Pain intensity, 0-10, MPD (SE)			
At 30 seconds	0.03 (0.78) NS	0.28 (0.63) NS	0.09 (0.35) NS
At first perception	0.01 (0.21) NS	-0.24 (0.20) NS	-0.26 (0.26) NS
Time, seconds, MPD (SE)			
Withdrawal	7.5 (4.2) NS	8.2 (5.3) NS	0.61 (8.0) NS
First pain	2.6 (1.2) NS	1.9 (0.8) NS	-0.7 (1.5) NS

Abbreviations: MPD, mean paired difference; SE, standard error of the mean; NS, no significant treatment effect.