### Supplemental Digital Content 2. Results of Mixed Effects Model from 4.5 to 6.5 Hours

<table>
<thead>
<tr>
<th>Outcome</th>
<th>Parameter</th>
<th>Coefficient</th>
<th>95% CI</th>
<th>P Value†</th>
</tr>
</thead>
<tbody>
<tr>
<td>NRS</td>
<td>Intercept</td>
<td>11.148</td>
<td>10.174 to 12.121</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td></td>
<td>Dose, low</td>
<td>0.825</td>
<td>0.632 to 1.019</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td></td>
<td>Time, per minute after 4.5h</td>
<td>-0.020</td>
<td>-0.023 to -0.018</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td></td>
<td>Session, 2nd</td>
<td>-0.394</td>
<td>-0.587 to -0.200</td>
<td>&lt;0.001</td>
</tr>
<tr>
<td></td>
<td>Order, low 1st</td>
<td>0.082</td>
<td>-0.660 to 0.824</td>
<td>0.821</td>
</tr>
</tbody>
</table>

| Log Hyperalgesia Area    | Intercept       | 3.866       | 3.065 to 4.666 | <0.001   |
|                          | Dose, low       | -0.305      | -0.444 to -0.166 | <0.001   |
|                          | Time, per minute after 4.5h | -0.001 | -0.003 to 0.001 | 0.432    |
|                          | Session, 2nd    | -0.427      | -0.566 to -0.288 | <0.001   |
|                          | Order, low 1st  | 0.169       | -0.345 to 0.683 | 0.502    |

| Log Allodynia Area       | Intercept       | 3.097       | 1.950 to 4.243 |         |
|                          | Dose, low       | -0.040      | -0.235 to 0.154 | 0.682    |
|                          | Time, per minute after 4.5h | -0.004 | -0.007 to -0.001 | 0.010    |
|                          | Session, 2nd    | -0.310      | -0.505 to -0.116 | 0.002    |
|                          | Order, low 1st  | 0.513       | -0.282 to 1.307 | 0.194    |

| Log CPP Threshold        | Intercept       | 3.412       | 2.518 to 4.306 |         |
|                          | Dose, low       | -0.430      | -0.563 to -0.297 | <0.001   |
|                          | Time, per minute after 4.5h | 0.000 | -0.001 to 0.001 | 0.876    |
|                          | Session, 2nd    | 0.366       | 0.233 to 0.498  | <0.001   |
|                          | Order, low 1st  | -0.110      | -0.976 to 0.756 | 0.795    |

| Log CPP Tolerance        | Intercept       | 4.687       | 3.887 to 5.488 |         |
|                          | Dose, low       | -0.325      | -0.434 to -0.217 | <0.001   |
|                          | Time, per minute after 4.5h | -0.001 | -0.002 to 0.001 | 0.283    |
|                          | Session, 2nd    | 0.161       | 0.053 to 0.270  | 0.004    |
|                          | Order, low 1st  | -0.208      | -1.051 to 0.635 | 0.614    |

† P values have been a topic of debate in mixed effects models. While we recommend using the 95% CI as the sole assessment, for those who would like P values, these have been included by ANOVA using a likelihood ratio test comparing a model with vs. without the examined parameter.

CI = confidence interval, CPP = cold pressor pain; NRS = numerical rating scale.