*Supplemental table 1. Comparisons between the changes in nighttime and 36-hour blood pressure after NWT-03 and placebo*

|  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
|   | 1 gr  |   |   | 2 gr  |   |   | 5 gr  |   |   |
|   | NWT-03a | Placeboa | P-valueb | NWT-03a | Placeboa | P-valueb | NWT-03a | Placeboa | P-valueb |
| Nighttime SBP |  |  |  |  |  |  |  |  |  |
|  Normotensive | n = 9 |  | n = 9  |  | n = 9 |  |
|  Before treatment | 112.4 ± 9.2 | 111.7 ± 5.7 |  | 116.5 ± 6.8 | 120.0 ± 10.0 |  | 109.0 ± 6.6 | 111.3 ± 5.2 |  |
|  After treatment | 112.0 ± 9.1  | 114.9 ± 8.7 |  | 119.5 ± 10.1 | 118.3 ± 7.5 |  | 109.6 ± 7.8 | 107.3 ± 3.4 |  |
|  Effect |  -0.4 ± 6.4 |  3.1 ± 8.3 | 0.317 |  3.0 ± 9.9 |  -1.8 ± 7.3 | 0.193 |  0.6 ± 10.9 |  -4.0 ± 3.7 | 0.375 |
|  High normotensive | n = 12 |  | n = 11 |  | n = 11 |  |
|  Before treatment | 119.9 ± 11.1 | 123.4 ± 12.8 |  | 120.8 ± 7.5  | 124.2 ± 10.8 |  | 118.0 ± 12.3 | 118.3 ± 10.6 |  |
|  After treatment | 119.8 ± 7.9 | 119.4 ± 10.4 |  | 117.0 ± 12.1 | 125.0 ± 8.3 |  | 116.1 ± 6.4 | 116.6 ± 6.9 |  |
|  Effect |  -0.2 ± 9.3 |  -4.0 ± 8.4 | 0.186 |  -3.8 ± 8.5 |  0.8 ± 7.4 | 0.203 |  -1.9 ± 11.8 |  -1.7 ± 6.4 | 0.965 |
|  Mild hypertensive | n = 10 |  | n = 10 |  | n = 9 |  |
|  Before treatment | 131.6 ± 11.3 | 128.1 ± 10.7 |  | 126.5 ± 9.7 | 125.1 ± 12.9 |  | 130.6 ± 13.5 | 121.6 ± 8.9 |  |
|  After treatment | 132.8 ± 17.2 | 128.0 ± 11.8 |  | 126.5 ± 9.7 | 128.4 ± 12.9 |  | 118.6 ± 10.7 | 124.3 ± 13.1 |  |
|  Effect |  1.2 ± 14.6 |  -0.2 ± 8.0 | 0.782 |  -0.0 ± 6.8 |  3.3 ± 8.8 | 0.348 |  -12.0 ± 12.0 |  2.8 ± 7.3 | 0.008 |
| Nighttime DBP |   |   |   |   |   |   |   |   |   |
|  Normotensive | n = 9 |  | n = 9  |  | n = 9 |  |
|  Before treatment | 70.0 ± 5.6 | 69.9 ± 4.6 |  | 71.2 ± 7.6 | 75.6 ± 6.1 |  | 66.4 ± 5.6 | 67.4 ± 6.6 |  |
|  After treatment | 69.8 ± 6.9 | 70.9 ± 5.7 |  | 74.0 ± 6.0 | 71.6 ± 5.3 |  | 65.8 ± 5.0 | 63.8 ± 4.1 |  |
|  Effect | -0.2 ± 3.9 |  1.0 ± 4.8 | 0.483 |  2.8 ± 9.9 | -4.0 ± 4.6 | 0.076 | -0.7 ± 5.1 | -3.6 ± 6.3 | 0.311 |
|  High normotensive | n = 12 |  | n = 11 |  | n = 11 |  |
|  Before treatment | 73.1 ± 11.0 | 74.0 ± 11.2 |  | 72.5 ± 8.3 | 72.6 ± 7.0 |  | 72.0 ± 9.7 | 73.7 ± 6.9 |  |
|  After treatment | 71.5 ± 9.0 | 73.3 ± 1278 |  | 71.8 ± 9.8 | 75.2 ± 10.9 |  | 73.6 ± 8.0 | 72.6 ± 6.8 |  |
|  Effect | -1.6 ± 7.1 |  -0.7 ± 8.6 | 0.372 | -0.7 ± 7.4 |  2.6 ± 6.2 | 0.258 |  1.6 ± 10.0 | -1.1 ± 3.9 | 0.388 |
|  Mild hypertensive | n = 10 |  | n = 10 |  | n = 9 |  |
|  Before treatment | 81.7 ± 8.4 | 80.2 ± 6.0 |  | 71.6 ± 6.0 | 71.7 ± 9.9 |  | 77.6 ± 10.4 | 71.5 ± 6.3 |  |
|  After treatment | 83.5 ± 9.5 | 78.0 ± 7.6 |  | 73.6 ± 8.7 | 75.4 ± 7.3 |  | 72.6 ± 5.5 | 75.0 ± 6.9 |  |
|  Effect |  1.8 ± 7.6 |  -2.1 ± 4.1 | 0.147 |  2.0 ± 5.9 |  3.7 ± 7.6 | 0.395 | -5.0 ± 7.9 |  3.4 ± 5.4 | 0.020 |

*Supplemental table 1. Continued*

|  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
| 36-hour SBP |   |   |   |   |   |   |   |   |   |
|  Normotensive | n = 9 |  | n = 9  |  | n = 9 |  |
|  Before treatment | 124.5 ± 10.8 | 127.4 ± 11.0 |  | 124.6 ± 6.9 | 127.5 ± 6.2 |  | 125.9 ± 8.1 | 125.9 ± 6.8 |  |
|  After treatment | 125.0 ± 12.1 | 128.6 ± 11.5 |  | 128.7 ± 10.1 | 131.6 ± 7.7 |  | 127.6 ± 7.3 | 126.3 ± 5.8 |  |
|  Effect |  0.4 ± 6.7 |  1.2 ± 6.0 | 0.706 |  4.1 ± 5.5 |  4.1 ± 5.0 | 1.000 |  1.8 ± 5.4 |  0.4 ± 6.0 | 0.688 |
|  High normotensive | n = 12 |  | n = 11 |  | n = 11 |  |
|  Before treatment | 132.5 ± 12.2 | 134.5 ± 11.9 |  | 134.6 ± 8.2 | 136.6 ± 7.2  |  | 130.4 ± 5.3 | 129.7 ± 6.3 |  |
|  After treatment | 133.1 ± 10.0 | 132.8 ± 10.3 |  | 135.2 ± 9.4 | 136.9 ± 9.5 |  | 130.7 ± 4.7 | 129.2 ± 5.1 |  |
|  Effect |  0.6 ± 5.2 |  -1.5 ± 4.8 | 0.323 |  0.6 ± 6.5 |  0.4 ± 5.4 | 0.945 |  0.3 ± 3.5 |  -0.5 ± 3.5 | 0.59 |
|  Mild hypertensive | n = 10 |  | n = 10 |  | n = 9 |  |
|  Before treatment | 141.9 ± 5.5 | 141.3 ± 9.5 |  | 145.1 ± 12.3 | 141.5 ± 8.4 |  | 137.0 ± 8.5 | 135.9 ± 6.7 |  |
|  After treatment | 140.5 ± 10.1 | 142.9 ± 10.2 |  | 142.2 ± 12.5 | 145.3 ± 13.7 |  | 133.8 ± 13.5 | 139.0 ± 11.5 |  |
|  Effect |  -1.5 ± 6.7 |  1.6 ± 3.1 | 0.285 |  -3.0 ± 5.5 |  3.9 ± 6.6 | 0.015 |  -3.2 ± 7.3 |  3.1 ± 10.7 | 0.202 |
| 36-hour DBP |   |   |   |   |   |   |   |   |   |
|  Normotensive | n = 9 |  | n = 9  |  | n = 9 |  |
|  Before treatment | 78.7 ± 7.2 | 81.17 ± 6.38 |  | 79.6 ± 5.3 | 81.9 ± 4.9 |  | 80.2 ± 7.4 | 78.7 ± 4.4 |  |
|  After treatment | 79.6 ± 6.0 | 81.84 ± 6.04 |  | 81.8 ± 4.7 | 83.0 ± 4.7 |  | 80.1 ± 8.3 | 80.4 ± 8.1 |  |
|  Effect |  0.9 ± 4.7 |  0.67 ± 4.28 | 0.863 |  2.2 ± 3.5 |  1.2 ± 3.6 | 0.519 | -0.1 ± 4.8 |  1.7 ± 6.3 | 0.523 |
|  High normotensive | n = 12 |  | n = 11 |  | n = 11 |  |
|  Before treatment | 83.1 ± 6.9 | 84.20 ± 7.27 |  | 84.0 ± 7.4 | 84.2 ± 7.6 |  | 84.0 ± 4.0 | 82.7 ± 2.5 |  |
|  After treatment | 83.1 ± 6.3 | 82.58 ± 7.06 |  | 84.1 ± 8.3 | 84.5 ± 6.7 |  | 83.2 ± 3.9 | 81.3 ± 4.5 |  |
|  Effect |  0.0 ± 3.2 |  -1.62 ± 3.51 | 0.304 |  0.2 ± 3.9 |  0.3 ± 4.4 | 0.964 | -0.8 ± 4.0 | -1.4 ± 3.3 | 0.613 |
|  Mild hypertensive | n = 10 |  | n = 10 |  | n = 9 |  |
|  Before treatment | 88.9 ± 7.4 | 88.06 ± 5.98 |  | 85.6 ± 9.3 | 83.8 ± 8.7 |  | 85.5 ± 7.4 | 83.9 ± 6.9  |  |
|  After treatment | 89.1 ± 7.67 | 89.31 ± 7.27 |  | 83.8 ± 7.1 | 85.5 ± 7.3 |  | 83.0 ± 9.4 | 85.4 ± 7.9 |  |
|  Effect |  0.2 ± 4.8 |  1.25 ± 1.79 | 0.543 | -1.8 ± 3.0 |  1.7 ± 3.2 | 0.035 | -2.5 ± 5.2 |  1.6 ± 5.6 | 0.189 |

a Values are expressed as mean ± SD

b P-value for the difference between the active and the placebo period are based on paired samples t-test