## SUPPLEMENTAL MATERIAL

**Figures**



n.s.

**Supplemental Figure 1.** Blood glucose (left) and insulin (right) concentrations before and after an oral glucose load of patients on verum (n=4) in the on drug (on) and off drug (off) phase. Data are given as mean ± SEM. Data of insulin concentrations are only available for the on drug phase, because samples of the off drug phase were lost prior to analysis.

**Table S1.** Body weight data as provided by the primary investigator of the CRAD001ADE12 trial

|  |  |  |  |  |  |  |  |  |  |  |  |  |
| --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- | --- |
|  | Total | Verum | Placebo | Total | Male | Verum | Placebo | Total | Female | Verum | Placebo | Total |
|  |  |   |   |   |  |   |   |   |  |   |   |   |
| Baseline (week 0) | **Pat.(n)** | 213 | 216 | 429 | **Pat.(n)** | 104 | 116 | 220 | **Pat.(n)** | 109 | 100 | 209 |
|  | **Mean** | 78,4 | 79,6 | 79,0 | **Mean** | 86,7 | 86,8 | 86,8 | **Mean** | 70,4 | 71,2 | 70,8 |
|  | **SEM** | 1,1 | 1,1 | 0,8 | **SEM** | 1,3 | 1,2 | 0,9 | **SEM** | 1,3 | 1,6 | 1,0 |
|  | **SD** | 15,6 | 16,2 | 15,9 | **SD** | 13,2 | 13,0 | 13,1 | **SD** | 13,4 | 15,5 | 14,4 |
|  | **Minimum** | 50,0 | 47,0 | 47,0 | **Minimum** | 59,0 | 60,0 | 59,0 | **Minimum** | 50,0 | 47,0 | 47,0 |
|  | **1st quartile** | 66,0 | 68,0 | 67,0 | **1st quartile** | 78,6 | 79,0 | 78,9 | **1st quartile** | 62,0 | 61,0 | 62,0 |
|  | **Median** | 78,0 | 79,0 | 78,0 | **Median** | 84,5 | 85,0 | 85,0 | **Median** | 67,2 | 68,8 | 68,0 |
|  | **3rd quartile** | 87,0 | 89,1 | 88,3 | **3rd quartile** | 94,5 | 94,2 | 94,2 | **3rd quartile** | 75,0 | 78,5 | 76,4 |
|  | **Maximum** | 132,0 | 152,0 | 152,0 | **Maximum** | 132,0 | 125,0 | 132,0 | **Maximum** | 124,0 | 152,0 | 152,0 |
| Month 6 | **Pat.(n)** | 181 | 203 | 384 | **Pat.(n)** | 90 | 111 | 201 | **Pat.(n)** | 91 | 92 | 183 |
|  | **Mean** | 77,6 | 80,2 | 78,9 | **Mean** | 86,4 | 87,6 | 87,1 | **Mean** | 68,9 | 71,2 | 70,0 |
|  | **SEM** | 1,1 | 1,2 | 0,8 | **SEM** | 1,3 | 1,2 | 0,9 | **SEM** | 1,3 | 1,6 | 1,0 |
|  | **SD** | 15,2 | 16,4 | 15,9 | **SD** | 12,6 | 13,0 | 12,8 | **SD** | 12,2 | 15,7 | 14,1 |
|  | **Minimum** | 49,5 | 46,6 | 46,6 | **Minimum** | 57,0 | 60,0 | 57,0 | **Minimum** | 49,5 | 46,6 | 46,6 |
|  | **1st quartile** | 65,8 | 69,0 | 67,0 | **1st quartile** | 78,0 | 78,0 | 78,0 | **1st quartile** | 60,1 | 61,0 | 61,0 |
|  | **Median** | 77,0 | 78,0 | 77,0 | **Median** | 84,5 | 86,3 | 85,6 | **Median** | 66,0 | 69,0 | 67,0 |
|  | **3rd quartile** | 86,4 | 90,4 | 89,0 | **3rd quartile** | 92,0 | 94,5 | 93,5 | **3rd quartile** | 73,0 | 76,0 | 75,0 |
|  | **Maximum** | 132,0 | 152,0 | 152,0 | **Maximum** | 132,0 | 125,0 | 132,0 | **Maximum** | 110,0 | 152,0 | 152,0 |
| Month 12 | **Pat.(n)** | 162 | 194 | 356 | **Pat.(n)** | 81 | 109 | 190 | **Pat.(n)** | 81 | 85 | 166 |
|  | **Mean** | 77,5 | 80,7 | 79,2 | **Mean** | 86,4 | 87,4 | 87,0 | **Mean** | 68,6 | 72,1 | 70,4 |
|  | **SEM** | 1,2 | 1,2 | 0,8 | **SEM** | 1,3 | 1,3 | 0,9 | **SEM** | 1,3 | 1,8 | 1,1 |
|  | **SD** | 14,8 | 16,8 | 16,0 | **SD** | 12,1 | 13,7 | 13,0 | **SD** | 11,6 | 16,5 | 14,4 |
|  | **Minimum** | 49,0 | 48,0 | 48,0 | **Minimum** | 57,0 | 59,0 | 57,0 | **Minimum** | 49,0 | 48,0 | 48,0 |
|  | **1st quartile** | 66,0 | 68,0 | 67,0 | **1st quartile** | 78,0 | 78,0 | 78,0 | **1st quartile** | 62,0 | 61,0 | 62,0 |
|  | **Median** | 76,0 | 80,0 | 78,0 | **Median** | 84,0 | 86,0 | 84,7 | **Median** | 66,0 | 69,0 | 67,0 |
|  | **3rd quartile** | 86,0 | 89,6 | 89,0 | **3rd quartile** | 93,0 | 95,0 | 94,0 | **3rd quartile** | 72,0 | 79,0 | 75,0 |
|  | **Maximum** | 130,0 | 149,0 | 149,0 | **Maximum** | 130,0 | 125,0 | 130,0 | **Maximum** | 112,0 | 149,0 | 149,0 |
| Month 18 | **Pat.(n)** | 154 | 192 | 346 | **Pat.(n)** | 76 | 106 | 182 | **Pat.(n)** | 78 | 86 | 164 |
|  | **Mean** | 77,6 | 80,5 | 79,2 | **Mean** | 86,8 | 87,7 | 87,3 | **Mean** | 68,7 | 71,7 | 70,2 |
|  | **SEM** | 1,2 | 1,2 | 0,9 | **SEM** | 1,4 | 1,3 | 1,0 | **SEM** | 1,3 | 1,8 | 1,1 |
|  | **SD** | 14,9 | 17,0 | 16,1 | **SD** | 12,4 | 13,5 | 13,0 | **SD** | 11,3 | 16,8 | 14,5 |
|  | **Minimum** | 50,0 | 49,0 | 49,0 | **Minimum** | 58,0 | 61,0 | 58,0 | **Minimum** | 50,0 | 49,0 | 49,0 |
|  | **1st quartile** | 65,0 | 68,0 | 67,0 | **1st quartile** | 78,0 | 80,0 | 78,0 | **1st quartile** | 60,9 | 60,0 | 60,4 |
|  | **Median** | 77,8 | 80,0 | 78,0 | **Median** | 84,0 | 85,8 | 85,0 | **Median** | 65,5 | 68,5 | 68,0 |
|  | **3rd quartile** | 88,0 | 90,1 | 89,0 | **3rd quartile** | 93,8 | 95,0 | 95,0 | **3rd quartile** | 74,0 | 78,6 | 75,8 |
|  | **Maximum** | 132,3 | 152,0 | 152,0 | **Maximum** | 132,3 | 127,0 | 132,3 | **Maximum** | 111,0 | 152,0 | 152,0 |
| Month 24 | **Pat.(n)** | 167 | 194 | 361 | **Pat.(n)** | 82 | 110 | 192 | **Pat.(n)** | 85 | 84 | 169 |
|  | **Mean** | 77,2 | 81,0 | 79,3 | **Mean** | 85,8 | 87,6 | 86,8 | **Mean** | 68,9 | 72,5 | 70,7 |
|  | **SEM** | 1,1 | 1,2 | 0,8 | **SEM** | 1,3 | 1,3 | 0,9 | **SEM** | 1,3 | 1,9 | 1,2 |
|  | **SD** | 14,5 | 17,1 | 16,0 | **SD** | 11,8 | 13,6 | 12,8 | **SD** | 11,8 | 17,6 | 15,0 |
|  | **Minimum** | 51,1 | 48,0 | 48,0 | **Minimum** | 56,0 | 60,0 | 56,0 | **Minimum** | 51,1 | 48,0 | 48,0 |
|  | **1st quartile** | 65,3 | 69,0 | 68,0 | **1st quartile** | 78,2 | 80,0 | 78,9 | **1st quartile** | 61,0 | 60,8 | 61,0 |
|  | **Median** | 77,0 | 80,5 | 79,0 | **Median** | 84,0 | 86,0 | 85,0 | **Median** | 66,6 | 69,0 | 68,0 |
|  | **3rd quartile** | 87,0 | 90,0 | 88,0 | **3rd quartile** | 92,0 | 95,3 | 94,0 | **3rd quartile** | 75,0 | 79,1 | 77,0 |
|  | **Maximum** | 114,4 | 152,0 | 152,0 | **Maximum** | 114,4 | 133,0 | 133,0 | **Maximum** | 110,0 | 152,0 | 152,0 |
| Month 36 | **Pat.(n)** | 151 | 161 | 312 | **Pat.(n)** | 73 | 89 | 162 | **Pat.(n)** | 78 | 72 | 150 |
|  | **Mean** | 78,3 | 81,2 | 79,8 | **Mean** | 86,0 | 88,8 | 87,5 | **Mean** | 71,0 | 71,9 | 71,4 |
|  | **SEM** | 1,2 | 1,4 | 0,9 | **SEM** | 1,3 | 1,5 | 1,0 | **SEM** | 1,4 | 2,1 | 1,2 |
|  | **SD** | 14,2 | 17,7 | 16,2 | **SD** | 11,4 | 14,0 | 12,9 | **SD** | 12,8 | 17,5 | 15,2 |
|  | **Minimum** | 53,0 | 49,0 | 49,0 | **Minimum** | 60,0 | 61,0 | 60,0 | **Minimum** | 53,0 | 49,0 | 49,0 |
|  | **1st quartile** | 67,0 | 67,0 | 67,0 | **1st quartile** | 79,9 | 79,0 | 79,0 | **1st quartile** | 63,0 | 61,5 | 62,0 |
|  | **Median** | 78,0 | 82,0 | 79,9 | **Median** | 85,0 | 88,0 | 86,0 | **Median** | 67,3 | 67,0 | 67,1 |
|  | **3rd quartile** | 88,0 | 90,6 | 89,0 | **3rd quartile** | 92,0 | 97,0 | 94,0 | **3rd quartile** | 76,5 | 80,0 | 78,0 |
|  | **Maximum** | 118,0 | 152,0 | 152,0 | **Maximum** | 118,0 | 141,0 | 141,0 | **Maximum** | 115,0 | 152,0 | 152,0 |
| Month 48 | **Pat.(n)** | 130 | 129 | 259 | **Pat.(n)** | 60 | 73 | 133 | **Pat.(n)** | 70 | 56 | 126 |
|  | **Mean** | 79,1 | 81,2 | 80,1 | **Mean** | 87,3 | 89,4 | 88,5 | **Mean** | 72,1 | 70,4 | 71,4 |
|  | **SEM** | 1,3 | 1,5 | 1,0 | **SEM** | 1,6 | 1,7 | 1,2 | **SEM** | 1,4 | 1,8 | 1,1 |
|  | **SD** | 14,3 | 16,8 | 15,6 | **SD** | 12,2 | 14,4 | 13,5 | **SD** | 12,1 | 13,2 | 12,6 |
|  | **Minimum** | 51,2 | 48,0 | 48,0 | **Minimum** | 59,0 | 63,0 | 59,0 | **Minimum** | 51,2 | 48,0 | 48,0 |
|  | **1st quartile** | 68,6 | 69,0 | 69,0 | **1st quartile** | 78,7 | 81,0 | 80,0 | **1st quartile** | 64,0 | 61,5 | 63,0 |
|  | **Median** | 77,5 | 81,0 | 79,5 | **Median** | 85,5 | 88,0 | 88,0 | **Median** | 69,0 | 69,6 | 69,0 |
|  | **3rd quartile** | 89,0 | 92,0 | 90,0 | **3rd quartile** | 93,0 | 97,0 | 95,5 | **3rd quartile** | 78,0 | 79,0 | 78,8 |
|  | **Maximum** | 120,0 | 136,5 | 136,5 | **Maximum** | 120,0 | 136,5 | 136,5 | **Maximum** | 107,0 | 106,0 | 107,0 |