Table S1. Results from DEXA analysis.

|  | Mirabegron $^{2}$ |  |  |
| :---: | :---: | :---: | :--- |
| Region | Pre |  | Post |

${ }^{\text {a }}$ Data are presented as mean $\pm$ SEM ( $n=13$ ) and were analyzed by paired, two-tailed Student's t-tests. There were no significant changes in the leg, gynoid, trunk, or android regions in fat, lean, or bone mass.

Table S2. Results from indirect calorimetry.

|  | Mirabegron $^{\mathbf{a}}$ |  |  |
| :---: | :---: | :---: | :--- |
| Measure | Pre | Post | P-value |
| $\mathbf{V O}_{2}$ (L/min) | $0.221 \pm 0.008$ | $0.221 \pm 0.008$ | 0.99 |
| $\mathbf{V C O}_{2}$ (L/min) | $0.183 \pm 0.009$ | $0.181 \pm 0.008$ | 0.66 |
| RQ | $0.827 \pm 0.014$ | $0.819 \pm 0.013$ | 0.49 |
| REE (kcal/day) | $1532 \pm 61$ | $1538 \pm 56$ | 0.8 |

${ }^{\text {a }}$ Data are presented as mean $\pm$ SEM ( $n=13$ ) and were analyzed by paired, two-tailed Student's t-tests.

Table S3. Results from PET-CT scans.

|  | Mirabegron $^{\mathbf{a}}$ |  |  |
| :---: | :---: | :---: | :---: |
| Measure | Pre | Post | P-value |
| SUV mean | $0.48 \pm 0.21$ | $0.53 \pm 0.23$ | 0.53 |
| SUV | max | $1.45 \pm 0.74$ | $1.87 \pm 1.02$ |
| SUV | 0.55 |  |  |
| Glycolysis | $1.21 \pm 0.62$ | $1.35 \pm 0.7$ | 0.77 |
|  | $80 \pm 51$ | $108 \pm 67$ | 0.71 |

${ }^{\text {a }}$ Standardized uptake value (SUV) and glycolysis data are presented as mean $\pm$ SEM and were analyzed by paired, two-tailed Student's t-tests ( $n=12$ ).

Table S4. Measurements of systemic inflammation and adipokines.

|  | Mirabegron ${ }^{\mathbf{a}}$ |  |  |
| :---: | :---: | :---: | :---: |
| Marker | Pre | Post | P-value |
| TNF $\boldsymbol{\alpha}$ (pg /ml) | $3.00 \pm 0.23$ | $2.93 \pm 0.20$ | 0.48 |
| MCP1 (pg /ml) | $138 \pm 5$ | $138 \pm 6$ | 0.98 |
| Adiponectin (ug /ml) | $6.09 \pm 0.65$ | $6.13 \pm 0.53$ | 0.90 |
| HMW Adiponectin (ug / ml) | $2.53 \pm 0.37$ | $2.49 \pm 0.36$ | 0.66 |
| Resistin (pg /ml) | $1518 \pm 35$ | $1492 \pm 33$ | 0.49 |
| FGF21 | $1299 \pm 60$ | $1578 \pm 83$ | 0.25 |
| Adipsin (ng/ ml) | $2300 \pm 34$ | $2393 \pm 29$ | 0.23 |
| Leptin (ng/ ml) | $52 \pm 2$ | $52 \pm 3$ | 0.85 |

${ }^{\text {a }}$ Data are presented as mean $\pm$ SEM ( $n=12$ ) and were analyzed by paired, two-tailed Student's t-tests.

Table S6. Primer sequences.

| Gene <br> symbol | Forward | Reverse |
| :--- | :--- | :--- |
| ACTB | GAGCACAGAGCCTCGCCTTT | CGCGGCGATATCATCATCCAT |
| COXIV | AGC CAG AAG GCA CTG AAG GA | AGC CCC TGT TCA TCT CAG CA |
| PGC1A | TCC TTC CTC CAT GCC TGA CG | TTA GCT GAG TGT TGG CTG GTG |
| PLIN5 | TGT CTG AAG CCG CTC GC | ACA CTG GAT CTG GGG ATC TGA |
| PPIA | CCCACCGTGTTCTTCGACAT | GCTGTCTTTGGGACCTTGTCT |
| PPIB | AAGTCACCGTCAAGGTGTATTTT | TGCTGTTTTTGTAGCCAAATCCT |
| TBP | CCCGAAACGCCGAATATAATCC | AATCAGTGCCGTGGTTCGTG |
| TFAM | GTT TCT CCG AAG CAT GTG GG | GAC AGA TGA AAA CCA CCT CGG |
| TUBB | ACCAACCTACGGGGATCTGAA | TTGACTGCCAACTTGCGGA |
| UBC9 | CTGGAAGATGGTCGTACCCTG | GGTCTTGCCAGTGAGTGTCT |



B


Figure S1. Insulin levels during the OGTT and euglycemic clamp. A) Insulin levels are shown during the OGTT. B) Insulin levels are shown during the final 30 min of insulin infusion during the the euglycemic clamp. Data are presented as mean $\pm$ SEM.


Figure S2. $\beta$ AR mRNA expression before and after mirabegron treatment. A-C) the mRNA expression of the indicated $\beta A R$ was determined using the nanostring nCounter system as described in "Methods." Data are presented as mean $\pm$ SEM.

