

Figure S1. Viability in H295R cells treated with rosuvastatin. Viability of rosuvastatin-treated H295R cells. Cell viability was measured by WST1 assay. Results are the means \pm SEM of 8 to 32 independent determinations from 3 different experiments. (A) H295R cells were treated for 24, 48 and 72 h with rosuvastatin (100 μ M; R100). (B) H295R cells were treated with increasing concentrations of rosuvastatin. Data are the means \pm SEM of 8 to 32 independent determinations from 3 different experiments. ***P<0.0001 vs. control (DMSO).

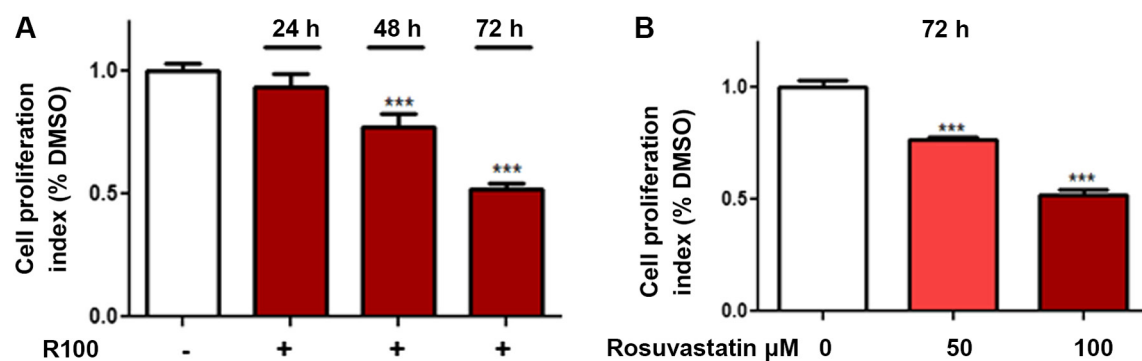


Figure S2. Viability of H295R cells treated with mitotane and/or rosuvastatin at 72 h. Cell viability in mitotane and/or rosuvastatin-treated H295R cells. H295R cells were treated for 72 h with mitotane (50 μ M or 100 μ M), rosuvastatin (100 μ M), alone or in combination. Cell viability was measured by WST1 assay. Results are the means \pm SEM of 8 to 16 independent determinations from 2 different experiments. Data are the means \pm SEM of 8 to 32 independent determinations from 3 different experiments. * P <0.05, ** P <0.01 and *** P <0.001 vs. control (DMSO); ns, not significant.

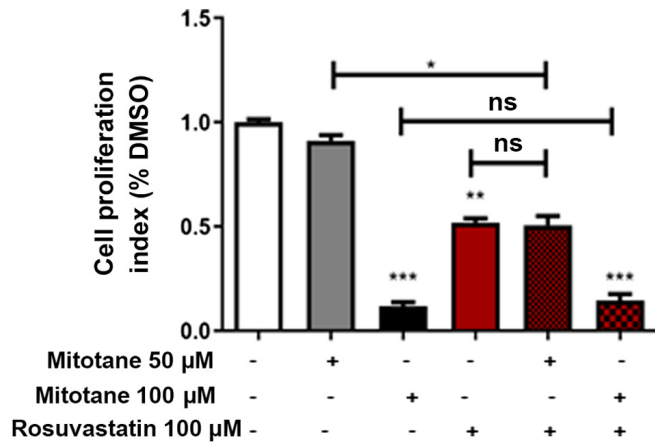


Figure S3. Steroidogenesis in H295R cells treated with mitotane and/or rosuvastatin. H295R cells were treated for 48 h with mitotane (25 μ M; M25), rosuvastatin (50 μ M; R50) alone or in combination (R50M25). Steroid concentrations in cell supernatants were measured by LC-MS/MS and are expressed in ng/ml. (A) Cortisol concentration; (B) corticosterone concentration. Data are the means \pm SEM of 4 independent determinations. * P <0.05 vs. control (FC or BC); NS, not significant. FC, cortisol control; FM25, cortisol mitotane 25 μ M; FR50, cortisol rosuvastatin 50 μ M; FR50M25, cortisol rosuvastatin 50 μ M and mitotane 25 μ M; BC, corticosterone control; BM25, corticosterone mitotane 50 μ M; BR50, corticosterone rosuvastatin 50 μ M; BR50M25, corticosterone rosuvastatin 50 μ M and mitotane 25 μ M.

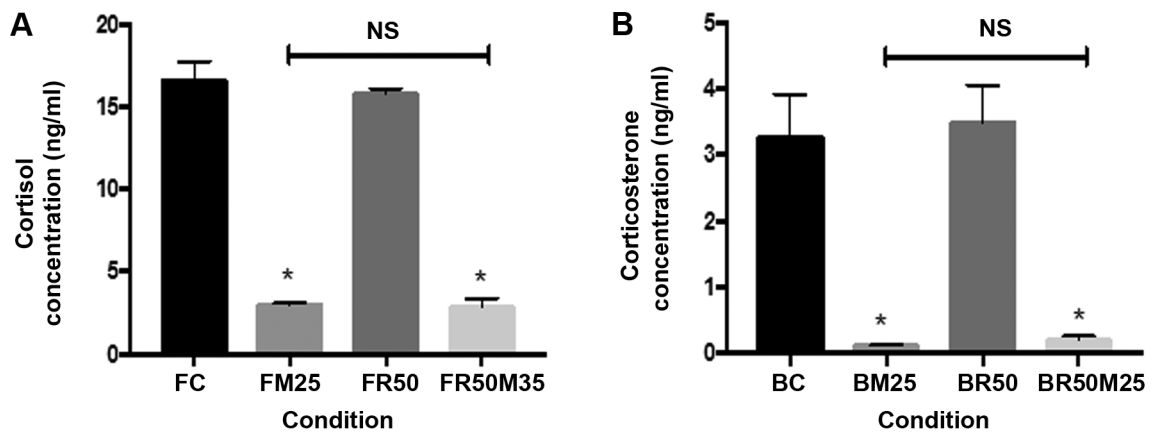


Table SI. Primer sequences.

| Gene | Amplicon size (bp) | Sense primer | Antisense primer |
|----------------|--------------------|-------------------------------|-----------------------------------|
| <i>HMGCR</i> | 119 | 5'-GGACCAACCTACTACCTCAGCAA-3' | 5'-CCATTACGGTCCCACACACA-3' |
| <i>ABCA1</i> | 67 | 5'-CCCTCATTCCAAGCACTTTACG-3' | 5'-CAATTCTCAGATATTCCAGTGCAAA-3' |
| <i>CYP11A1</i> | 149 | 5'-CGATTACCGTGGCATCCTCTA-3' | 5'-AGGTTGCGTGCCATCTCATA-3' |
| <i>StAR</i> | 121 | 5'-GCCACAGACTTCGGGAACAT-3' | 5'-AGTAGCCACGTAAGTTTGGTCTTAGAG-3' |
| <i>18S</i> | 71 | 5'-GTGCATGGCCGTTCTTAGTTG-3' | 5'-CATGCCAGAGTCTCGTTCGTT-3' |
| <i>36B4</i> | 75 | 5'-AGCGCGTCCTGGCATTGTCTGT-3' | 5'-GGGCAGCAGTGGTGGCAGCAGC-3' |

bp, base pairs.