## **Supplementary Information**



**Figure S1.** Melatonin-induced changes in mitochondrial morphology. Alterations in mitochondrial mass (NAO) (**A–C**) and mtDNA (**D–F**) in Cal-27 and SCC-9 cells. n = 6 per group. Data are presented as mean  $\pm$  SEM. \*p < 0.05, \*\*p < 0.01, \*\*\*p < 0.001 vs. control; #p < 0.05, ##p < 0.01, ###p < .001 vs. aMT 100 µM group; \$p < 0.05.



**Figure S2.** Modulation of melatonin receptor expression by melatonin in Cal-27 cells. Western Blot analysis of MT1 (A–C, J), MT2 (D–F, J) and ROR $\alpha$  (G–J). n = 6 per group. Data are presented as mean ± SEM. \*\*p < 0.01, \*\*\*p < 0.001 vs. control; #p < 0.05, ###p < 0.001 vs. aMT 100  $\mu$ M group.

## Full western data



**Figure S3.** Full scanned Western blots shown in Figure 3. Protein content of OXPHOS proteins (Complex I-V) in figure 3 A, B (**A**); 3 C, D (**B**) and 3 E, F (**C**). Experiments were performed with Cal-27 cells in Control, aMT 100, aMT 500 and aMT 1500 groups.



**Figure S4.** Full scanned Western blots shown in Figure 5 J–L. Protein content of hexokinase II in figure 5. Experiments were performed with Cal-27 cells in Control, aMT 100, aMT 500 and aMT 1500 groups.



**Figure S5.** Full scanned Western blots shown in Figure 7. Protein content of OPA-1, LETM, MFN2 and Drp1 in Figure 7 A–C, M (A); D–F, M (B); G–I, M (C) and J–M (D). Experiments were performed with Cal-27 cells in Control, aMT 100, aMT 500 and aMT 1500 groups.



**Figure S6.** Full scanned Western blots shown in Figure 8 A–J. Protein content of Bax and Bcl-2 in Figure 8 A–C, J (**A**); D–F, J (**B**). Experiments were performed with Cal-27 cells in Control, aMT 100, aMT 500 and aMT 1500 groups.



**Figure S7.** Full scanned Western blots shown in Figure 8 K-T. Protein content of Nix, LC3-II and ATG12-ATG5 in Figure 8 K–M, T (**A**); N–P, T (**B**) and Q–S, T (**C**). Experiments were performed with Cal-27 cells in Control, aMT 100, aMT 500 and aMT 1500 groups.



**Figure S8.** Full scanned Western blots shown in Figure S2. Protein content MT1, MT2 and RORα in Figure S2 A–C, J (**A**); D–F, J (**B**); K–M, T (**C**). Experiments were performed with Cal-27 cells in Control, aMT 100, aMT 500 and aMT 1500 groups.