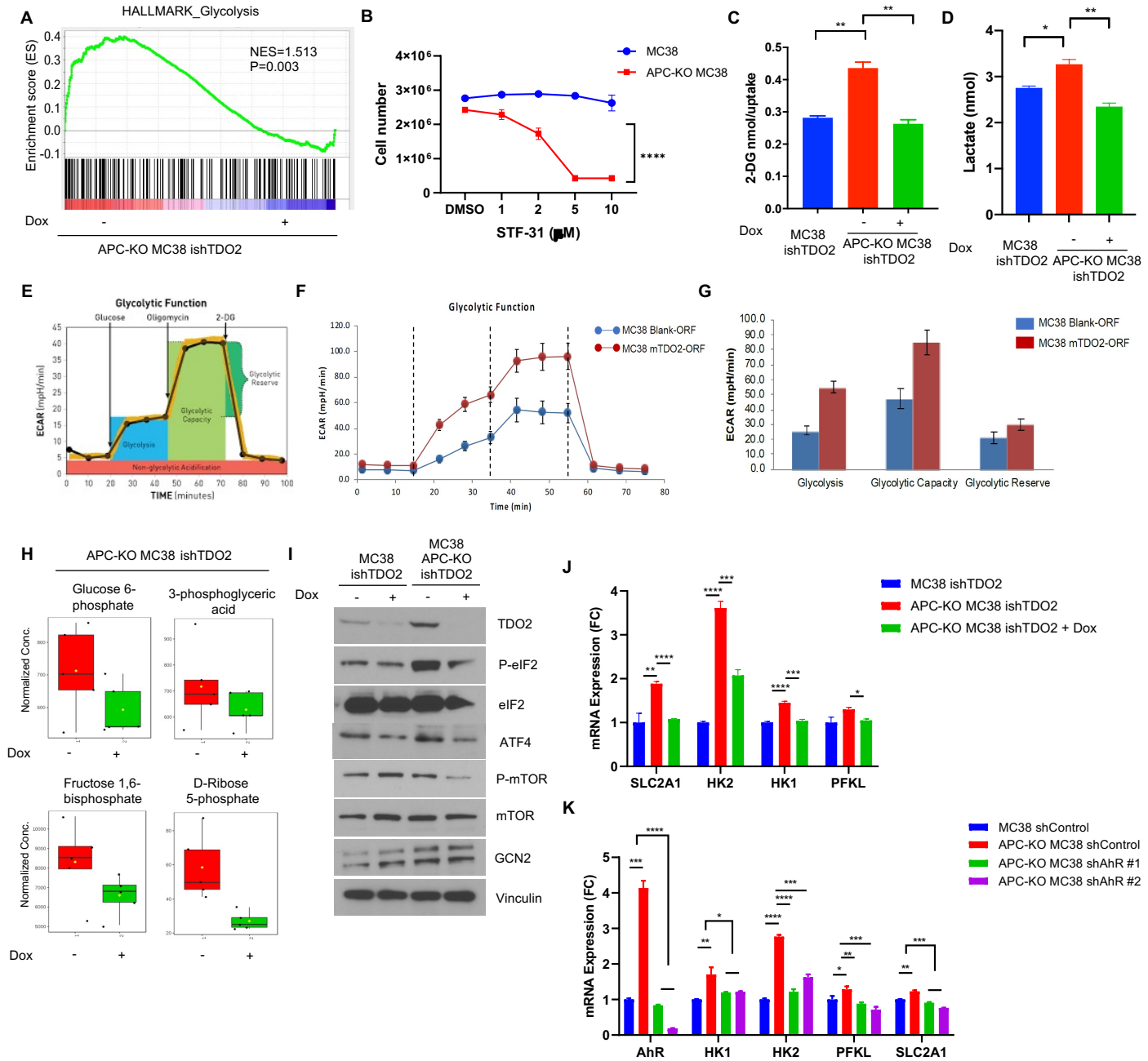


Supplementary Fig. S6.



Supplementary Fig. S6.

(A) GSEA correlation of glycolysis with altered gene expression in TDO2 depleted APC-KO MC38 cells. Normalized enrichment score (NES) and nominal P value are shown. **(B)** Cell viability assay of APC-WT and APC-KO MC38 cells treated with STF-31 for 24 hr in a dose-dependent manner. Six replicates per group. *** $P < 0.001$, Two-way ANOVA. Two independent experiments were performed. **(C)** 2-DG uptake assay with ishTDO2 APC-WT and APC-KO MC38 cell lines with and without dox treatment. ** $P < 0.01$, *** $P < 0.001$ **(D)** Measurement of secreted lactate with conditioned media from ishTDO2 APC-WT and APC-KO MC38 cell lines with and without dox treatment. * $P < 0.05$, ** $P < 0.01$ **(E)** Representation of key parameters of glycolytic flux measured by Seahorse experiment. **(F and G)** Measured extracellular acidification rate (ECAR) in MC38 Blank-ORF and mouse TDO2-ORF expressing cell lines. **(H)** Changes in concentration of key glycolysis metabolites in ishTDO2 APC-KO MC38 cell line lysates. The red bars indicate cells without dox treatment and the green bars indicate dox-treated cells. $n = 5$ per biological replicates. **(I)** Western blot of GCN2 and mTOR pathway components in ishTDO2 APC-WT and APC-KO MC38 cells. **(J)** RT-qPCR analysis of glycolysis pathway genes in ishTDO2 APC-WT and APC-KO MC38 cell lines. ** $P < 0.01$, *** $P < 0.001$, **** $P < 0.0001$ **(K)** RT-qPCR analysis of glycolysis pathway genes in APC-WT and APC-KO MC38 cell lines expressing shControl or two shAhR constructs. * $P < 0.05$, ** $P < 0.01$, *** $P < 0.001$, **** $P < 0.0001$