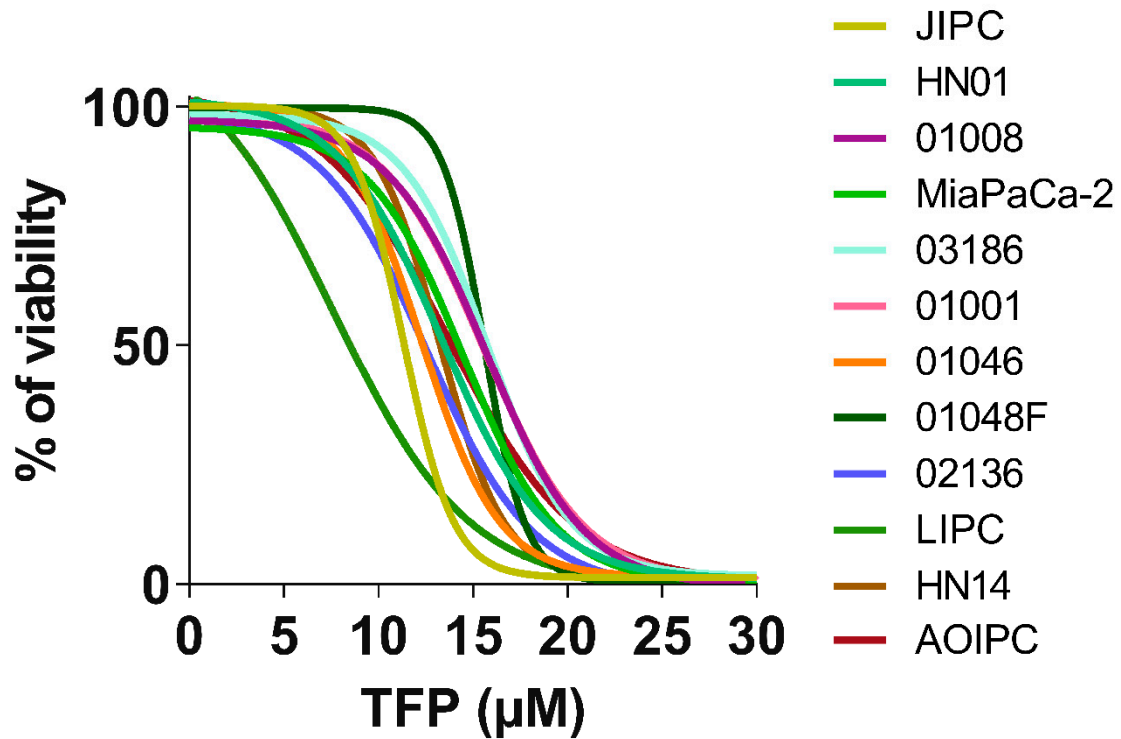


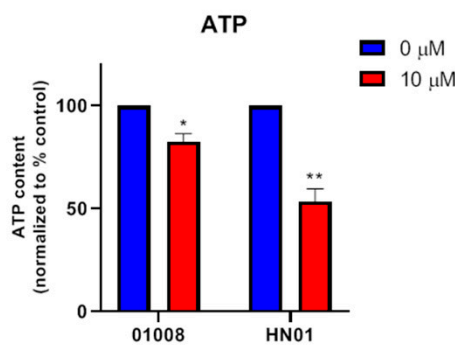
Supplementary Materials

# Dissecting the Anticancer Mechanism of Trifluoperazine on Pancreatic Ductal Adenocarcinoma

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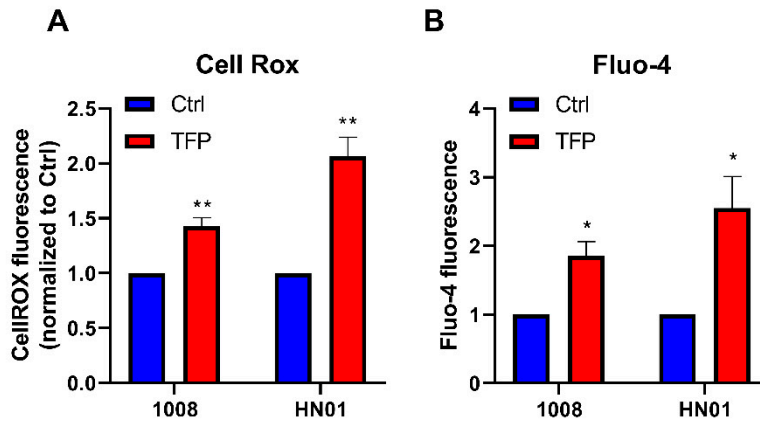


**Figure S1.** Trifluoperazine induces cell death via apoptosis and necroptosis in PDAC cells. Viability of MiaPaCa-2 cells and 11 PDX-derived cell lines upon a 24-h treatment with TFP at increasing concentrations of trifluoperazine was measured.



**Figure S2.** Trifluoperazine decreases ATP production in PDAC cells. Cells were incubated with TFP at 10 μM, and ATP content was measured after 24 h of treatment. For each treatment, statistical sig-

nificance is \*  $p < 0.05$ , \*\*  $p < 0.01$ , compared with untreated cells (Student's 2-tailed unpaired  $t$ -Test). Data represent mean  $\pm$  SEM,  $n = 3$  (with technical triplicates).



**Figure S3.** Trifluoperazine promotes mitochondrial and ER coupled stress (A) ROS production detected using MitoSOX Red by flow cytometry analysis in TFP and non-treated cells. (B) Flow cytometry analysis with Fluo-4-AM performed to determine cytosolic calcium concentration. For each treatment, statistical significance is \*  $p < 0.05$ , \*\*  $p < 0.01$ , compared with untreated cells (Student's 2-tailed unpaired  $t$ -Test). Data represent mean  $\pm$  SEM,  $n = 3$  (with technical triplicates).

