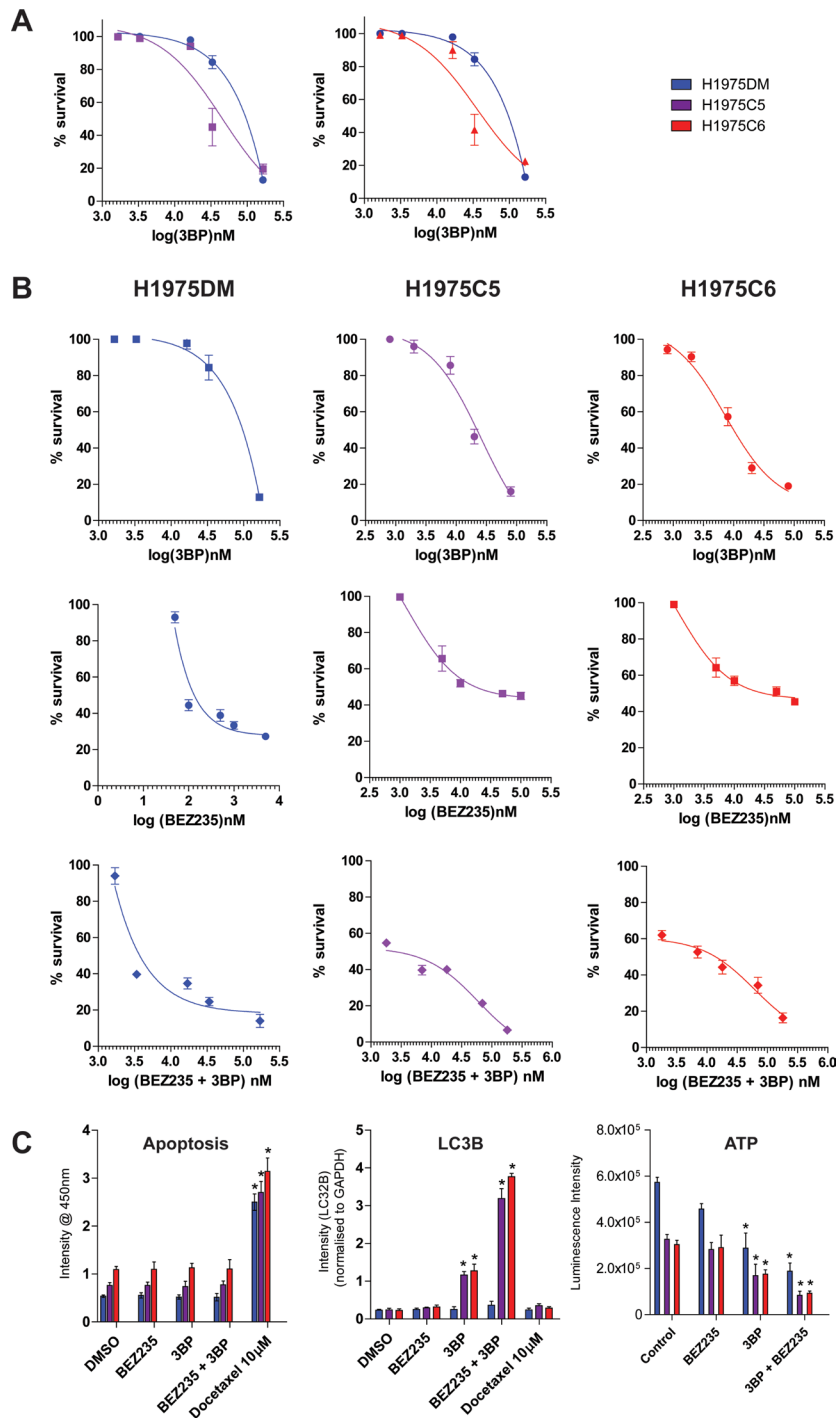


Acquired resistance to PI3K/mTOR inhibition is associated with mitochondrial DNA mutation and glycolysis

SUPPLEMENTARY MATERIALS



Supplementary Figure 1: Effect of treatment with BEZ235 alone, 3-bromopyruvate (3BP) alone, and 3BP and BEZ235 in combination in H1975DM, H1975C5, and H1975C6 cell lines. (A) Concentration response curves of 3BP. **(B)** Concentration response curves for 3BP (upper panels), BEZ235 (middle panels) and the fixed-ratio combination of BEZ235 and 3BP (lower panels). **(C)** Apoptosis, LC3B and ATP levels 24h post-exposure to IC₅₀ concentrations of BEZ235, 3BP or their combination. **p* < 0.05 to levels in control-treated cells for each cell line, paired *t*-test.

Supplementary Table 1: Gene expression levels and differences according to genome-wide gene expression array analysis. See Supplementary_Table_1

Supplementary Table 2: Mitochondrial DNA variants detected by next generation sequencing of mitochondrial DNA, according to their reported and predicted effects, and presence in H1975DM, H1975C5, and H1975C6 cells. See Supplementary_Table_2