## Supplementary Figure



Supplementary Fig. 1 ADR- or VCR-resistant SGC7901 cell lines were truly resistant to ADR or VCR. (A) SGC7901 cells were treated with different concentrations of ADR (0, 0.2, 0.4 , $0.8,1.6,3.2,6.4 \mu \mathrm{~g} / \mathrm{ml}$ ) for 48 h and SGC7901/ADR cells were stimulated with different concentrations of $\operatorname{ADR}(0,1,2,4,8,16 \mu \mathrm{~g} / \mathrm{ml})$ for 48 h . Then, the $\mathrm{IC}_{50}$ value of $\operatorname{ADR}$ was determined by CCK-8 assay. (B) SGC7901 cells were stimulated with different doses of VCR (0, $0.4,0.8,1.6,3.2,6.4,12.8 \mu \mathrm{~g} / \mathrm{ml})$ for 48 h and $\mathrm{SGC7901/VCR}$ cells were treated with various doses of VCR $(0,2,4,8,16,32 \mu \mathrm{~g} / \mathrm{ml})$ for 48 h . Then, the $\mathrm{IC}_{50}$ value of VCR was determined by CCK-8 assay. $\quad * P \quad 0.05$.


Supplementary Fig. 2 GACAT1 knockdown enhanced the resistance of GC cells to ADR and VCR. (A) SGC7901/ADR or SGC7901/VCR cells were infected with shRNA-NC or shRNAGACAT1 for 24 h . Then, SGC7901/ADR cells were stimulated with different concentrations of $\operatorname{ADR}(0,1,2,4,8,16 \mu \mathrm{~g} / \mathrm{ml})$ and SGC7901/VCR cells were treated with various doses of VCR ( 0 , $2,4,8,16,32 \mu \mathrm{~g} / \mathrm{ml})$ for additional 48 h . Subsequently, cell viability was determined by CCK-8 assay. (B) SGC7901/ADR or SGC7901/VCR cells were infected with shRNA-NC or shRNAGACAT1 for 24 h and then treated with $\operatorname{ADR}(2 \mu \mathrm{~g} / \mathrm{ml})$ or VCR $(4 \mu \mathrm{~g} / \mathrm{ml})$ for another 48 h . Then, cell apoptotic percentage was determined through flow cytometry. $* P<0.05$.


Supplementary Fig. 3 PTEN loss enhanced ADR and VCR resistance by regulating PTEN/AKT/mTOR/S6K1 signaling pathway in GC cells. (A) SGC7901/ADR or SGC7901/VCR cells were infected with shRNA-NC or shRNA-PTEN for 24 h and then treated with different concentrations of ADR or VCR for additional 48 h . Then, cell viability was measured by CCK-8 assay. (B-D) SGC7901/ADR or SGC7901/VCR cells were infected with shRNA-NC or shRNA-PTEN for 24 h and then treated with ADR $(2 \mu \mathrm{~g} / \mathrm{ml})$ or VCR $(4 \mu \mathrm{~g} / \mathrm{ml})$ for another 48 h . (B) Cell apoptotic percentage was examined through flow cytometry. (C and D) Protein levels of AKT, p-AKT, mTOR, p-mTOR, p-S6K1 and S6K1 were determined by western blot assay. ${ }^{*} P<0.05$.

