## Supplemental Figure 5



## Supplementary Figure S5. Combined KRASG12C and SHP2 or MAPK inhibition improves in

 vitro efficacy. (A) Cell lines treated with ARS-1620 (1 $\mu \mathrm{M})$, SHP099 (10 $\mu \mathrm{M})$, RMC-4550 ( $1 \mu \mathrm{M}$ ), trametinib (10 nM), BVD-523 (1 $\mu \mathrm{M}$ ) or combination for 10-14 days and then stained with crystal violet. (B) Individual quantification of assays in (A) (C) SW837 and MIA PaCa-2 cell lines were treated with ARS-1620 ( $1 \mu \mathrm{M}$ ) alone or combined with SHP099 (10 $\mu \mathrm{M}$ ) or RMC-4550 $(1 \mu \mathrm{M})$ for 0 , 4, 24, 48, and 72 h . Blot analysis was performed for pMEK, pERK, pRSK, pAKT, and total MYC with GAPDH as a loading control. (D) SW837 and MIA PaCa-2 cell lines were treated ARS-1620 (1 $\mu \mathrm{M})$, SHP099 $(10 \mu \mathrm{M})$, afatinib $(1 \mu \mathrm{M})$, BGJ398 $(1 \mu \mathrm{M})$ or BVD-523 $(1 \mu \mathrm{M})$ or a combination and lysates were subject to a RAF-RBD pulldown and blot analysis of KRAS and total RAS as well as pERK, pRSK and GAPDH for input samples