

S6 Fig. Tumor 5.

Quantification of (A) relative total spheroid area, (B) relative spheroid number, and (C) relative average spheroid size, with error bars indicating standard error of the mean. Quantified mutant subpopulations rare plotted (D), with error bars indicating standard deviation. Although subpopulation of all four mutants were present, no erlotinib-treated culture had a significantly larger subpopulation than the 0  $\mu$ M erlotinib culture and all cultures had *PIK3CA* H1047R mutant subpopulations greater than the Tumor 5 TR (one-tailed Mann Whitney test, P = 0.0500). An example of spheroid culture appearance is provided (E), in which the scale bar = 200  $\mu$ m.