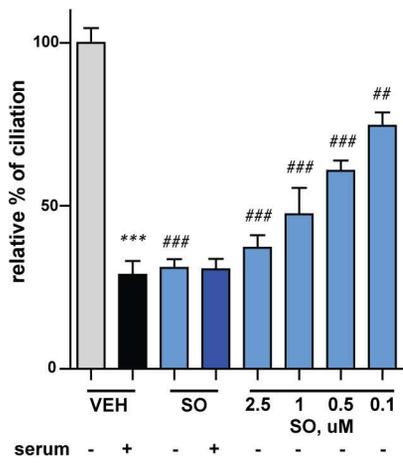
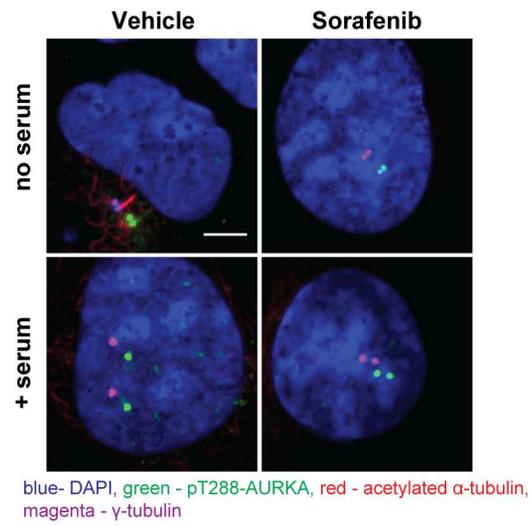
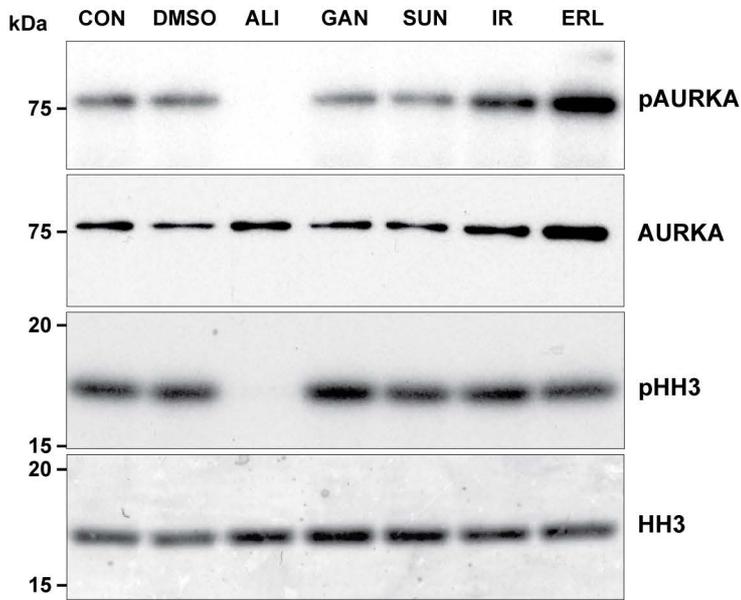
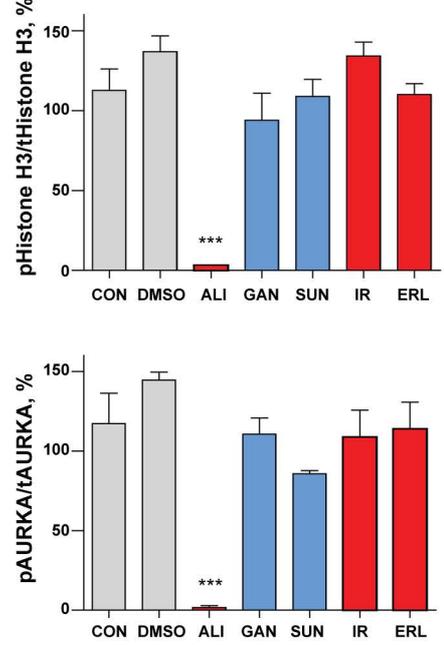


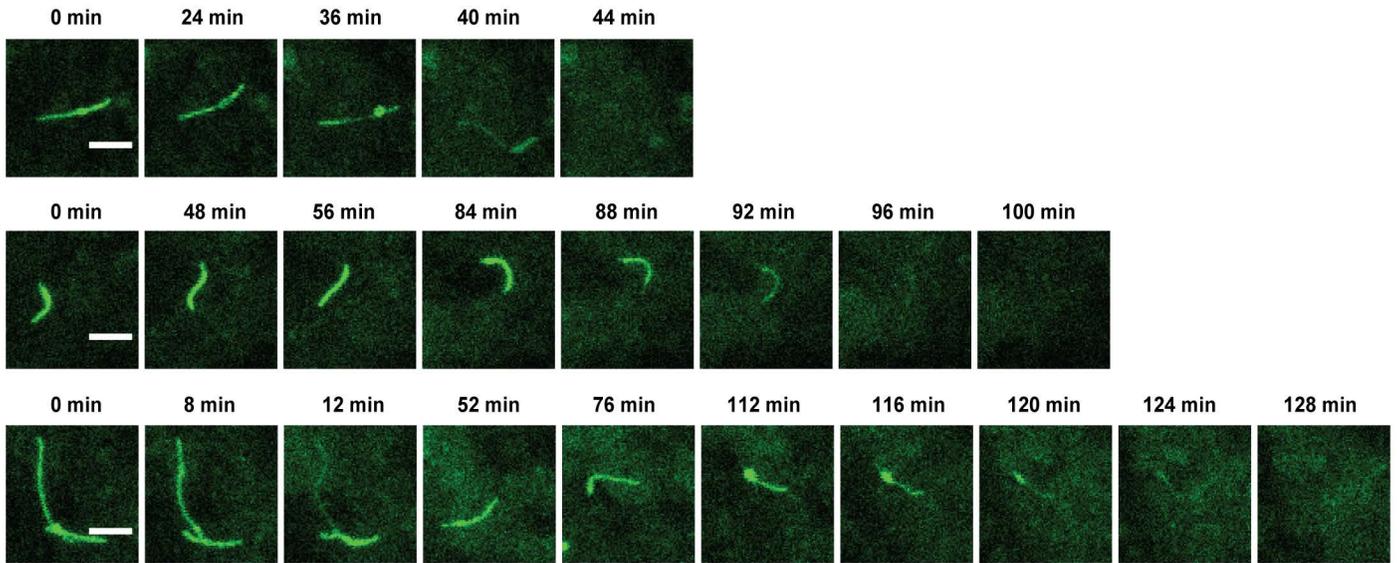
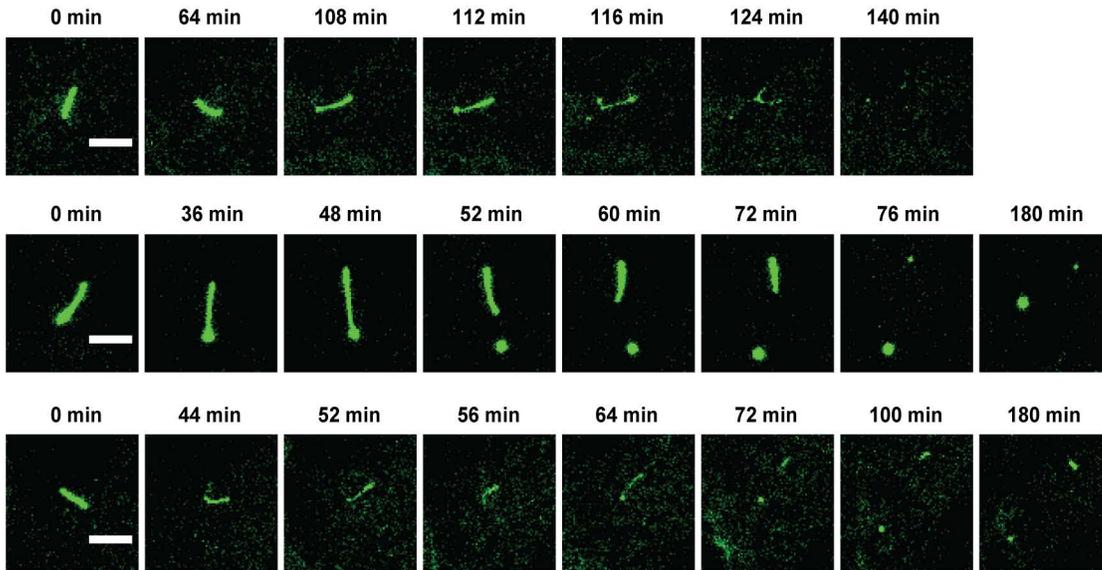
Supplementary Figure S1. Additional analysis of siRNA effects on ciliation and target depletion. **A.** Candidate protein kinase inhibitor targets that did not affect ciliation. For each datapoint, $n \geq 500$ cells (250 from each of two biological repetitions, visualized by confocal), with data analyzed by Tukey test. *, $p < 0.5$; **, $p < 0.01$; ***, $p < 0.001$, for each siRNA in serum versus no serum conditions. #, $p < 0.5$; ##, $p < 0.01$; ###, $p < 0.001$, relative to vehicle, no serum. +, $p < 0.5$; ++, $p < 0.01$; +++, $p < 0.001$, relative to vehicle plus serum. **B.** Quantitative RT-PCR to determine degree of target depletion 48 hours following transfection with the indicated siRNAs; all data normalized to mRNA encoding the ribosomal protein 36B4.

A.**B.**

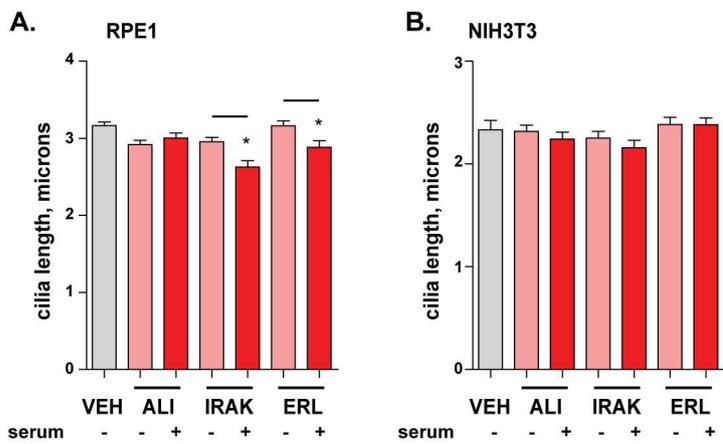
Supplementary Figure S2. Sorafenib is similar to sunitinib in inducing ciliary resorption. A., B. Ciliation rates (A) and T288ph-AURKA (B) images of hTERT-RPE1 cell line upon treatment with sorafenib. Green, T288ph -AURKA; red, acetylated α -tubulin; magenta, γ -tubulin; signal for T288ph-AURKA and basal body are offset for clarity; ($n \geq 750$ cells). Significance calculated by Tukey test, from three independent experiments; *, $p < 0.05$; **, $p < 0.01$; ***, $p < 0.001$, for each drug or siRNA in serum versus no serum conditions. #, $p < 0.05$; ##, $p < 0.01$; ###, $p < 0.001$, relative to vehicle, no serum.

A.**B.**

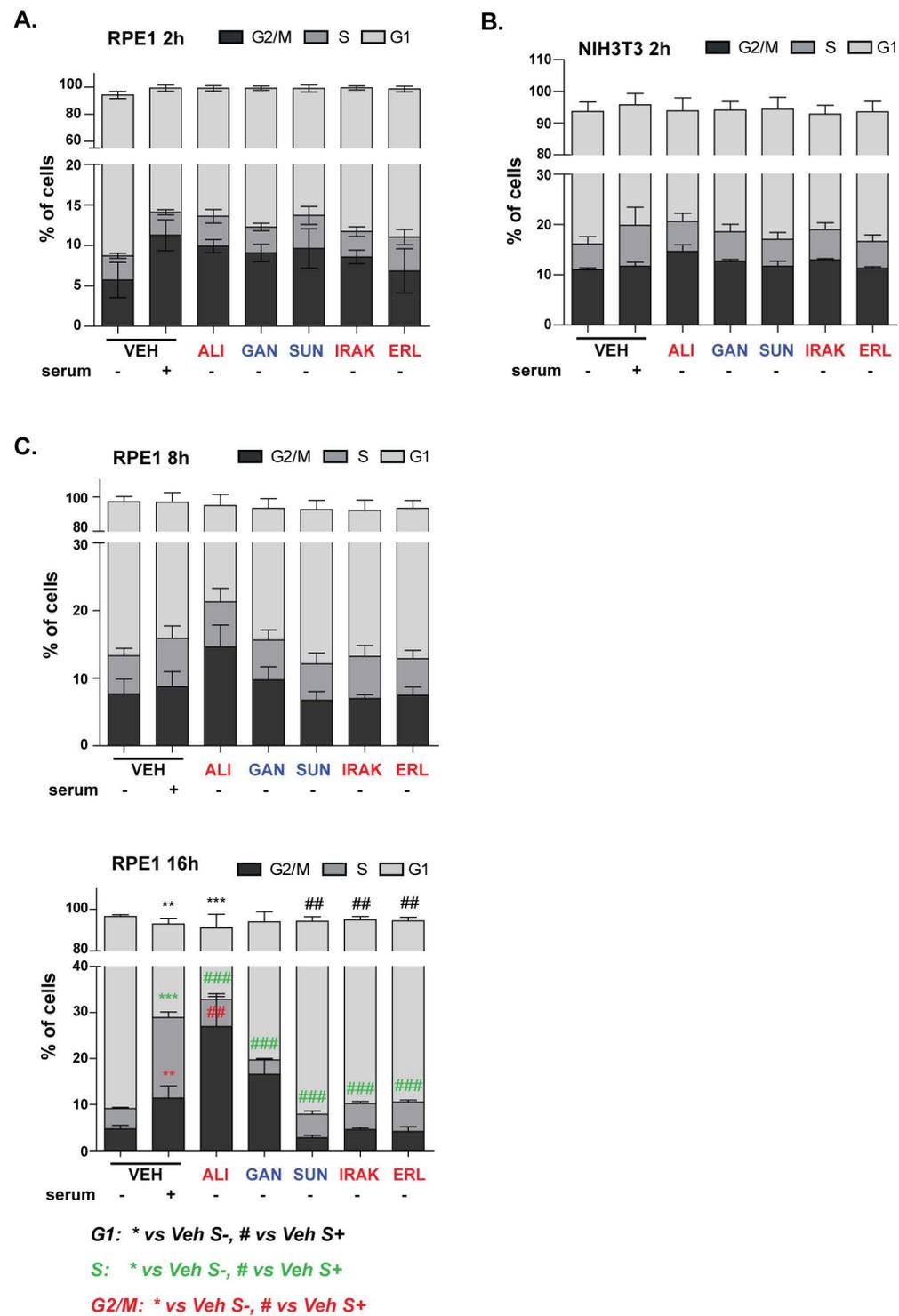
Supplementary Figure S3. Effect of compounds on AURKA catalytic activity. Representative WB images (A) and quantification of *in vitro* kinase assay (B) demonstrating alisertib inhibits AURKA autophosphorylation and phosphorylation of histone H3 (HH3), but other drugs tested do not, gauged by *in vitro* kinase assay with ^{32}P -ATP. Total AURKA is visualized by antibody, and HH3 by Coomassie blue. CON, no drug diluent; GAN, ganetespib; SUN, sunitinib; IR, IRAK1/4i; ERL, erlotinib. Significance calculated by Tukey test, from three independent experiments; ***, $p < 0.001$.

A.**Sunitinib treatment****B.****Ganetespiib treatment**

Supplementary Figure S4. Representative images demonstrating the effect of sunitinib (A) and ganetespiib (B) on cilia resorption in hTERT-RPE1-Arl13b-GFP cells. Three examples are shown; green stain reflects Arl13b-GFP signal. Scale bar, 5 μ m.

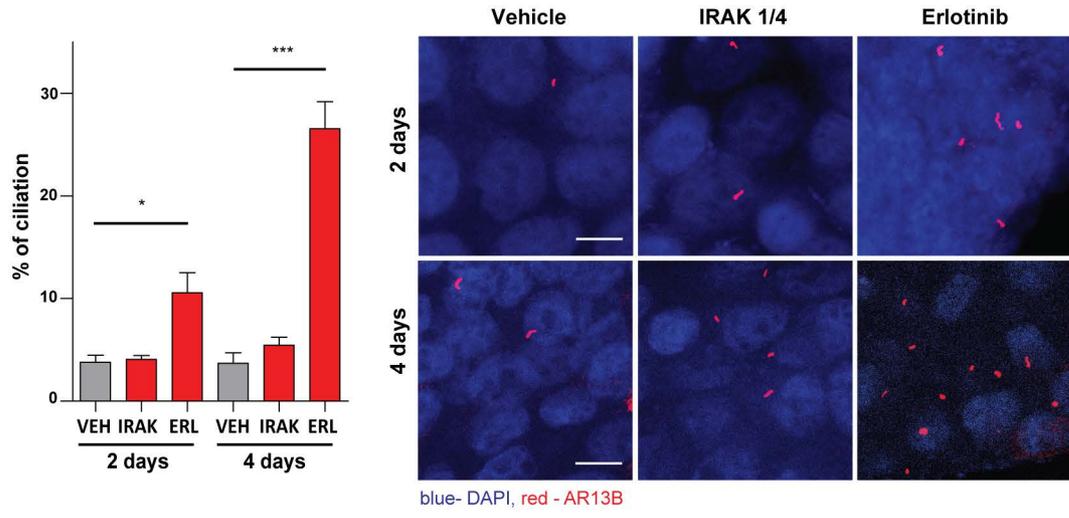


Supplementary Figure S5. Cilia length upon treatment with indicated cilia-stabilizing compounds in hTERT-RPE1 (**A**) and NIH3T3 (**B**) cell lines. $n \geq 150$ cells (in each of 2 independent repetitions), based on Tukey test. *, $p < 0.5$; **, $p < 0.01$; ***, $p < 0.001$, for each drug in serum versus no serum conditions.

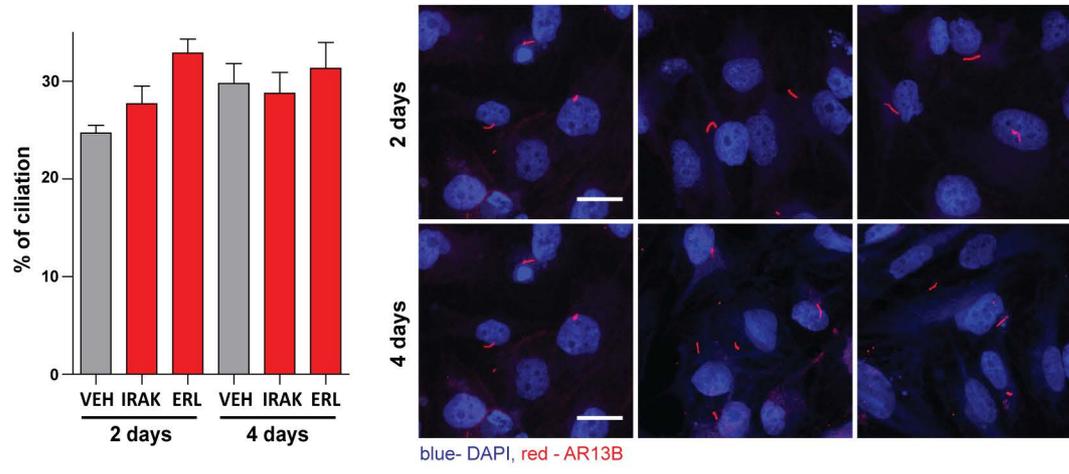


Supplementary Figure S6. Drug effects on ciliation correspond to changes in cell cycle over time. Cell cycle distribution of hTERT-RPE1 (A) and NIH3T3 (B) cells treated for 2.5 hours with the indicated drugs, based on FACS. (C) Cell cycle distribution of hTERT-RPE1 cells treated for 8 and 16 hours with the indicated drugs. Significance calculated by Tukey test in all cases, from three independent experiments. * or #, $p < 0.5$; ** or ##, $p < 0.01$; *** or ###, $p < 0.001$.

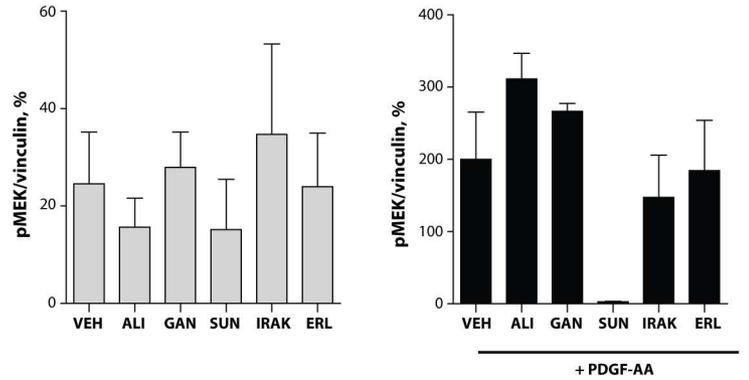
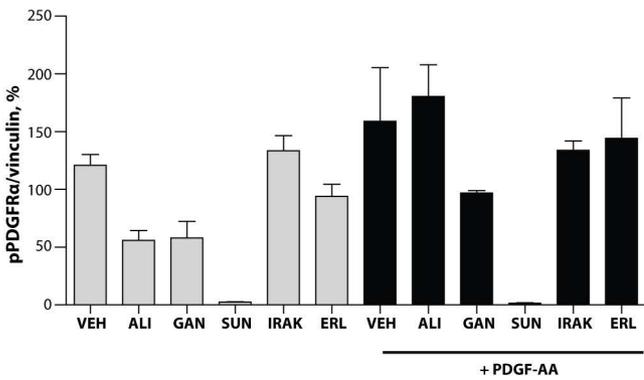
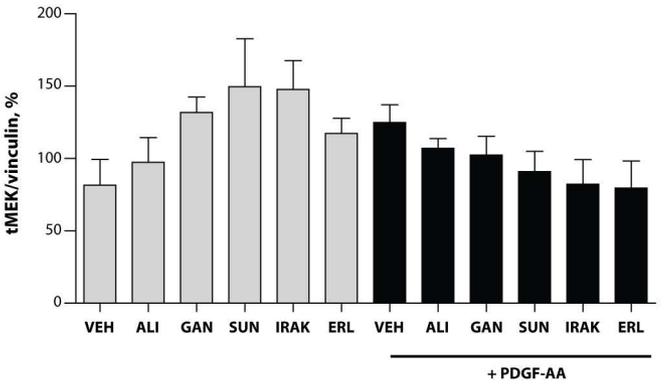
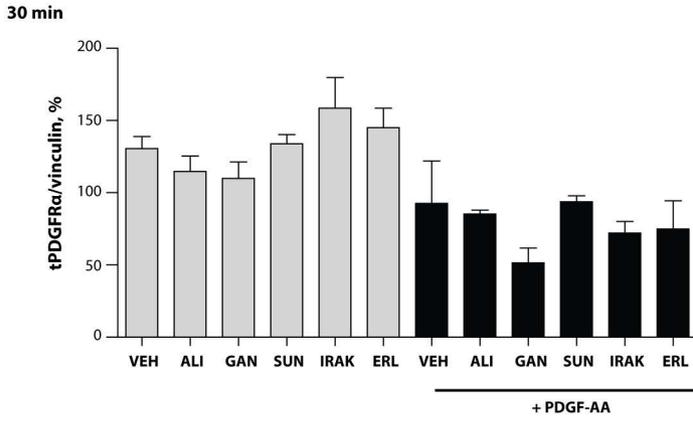
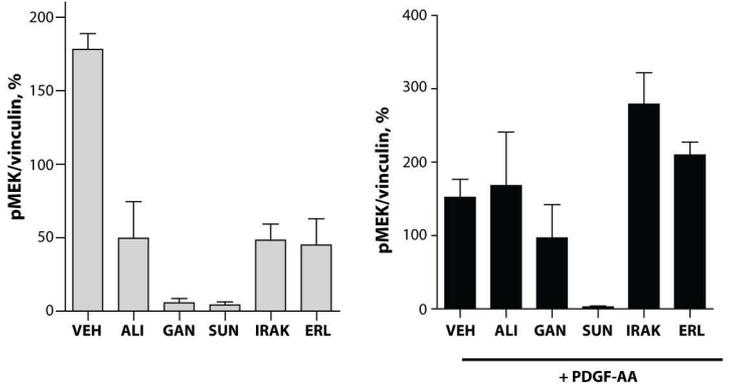
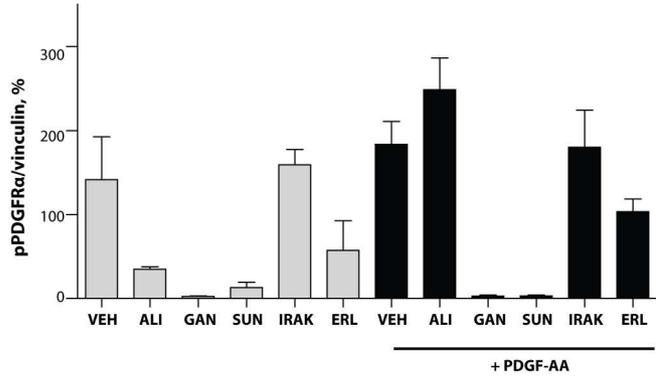
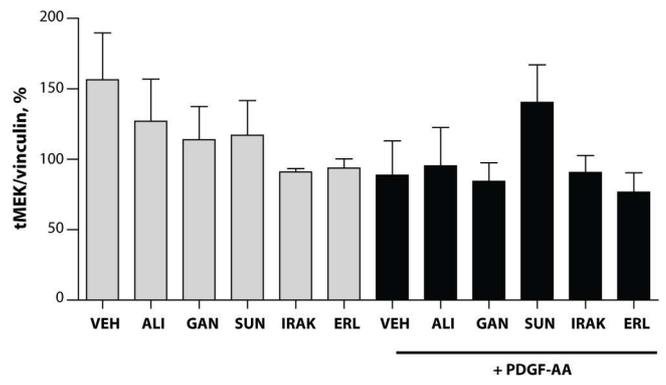
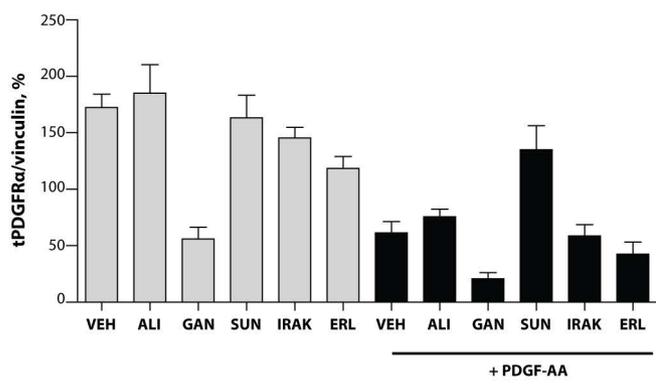
A. CFPAC-1



B. PNX_0010



Supplementary Figure S7. Erlotinib increases ciliation in CFPAC-1 but not in PNX0010 cell line. Ciliation rates and representative images of CFPAC-1 (A) and PNX0010 (B) cell lines after 2 and 4 days incubation with IRAK1/4i or erlotinib. Red, ARL13B; blue, DAPI. Scale bar for (A) 10 μ m, for (B) 20 μ m. $n \geq 750$ cells, reflecting $n \geq 250$ from each of 3 independent experiments. Statistics determined by Tukey test in all cases; *, $p < 0.5$; **, $p < 0.01$; ***, $p < 0.001$, for each drug in serum versus no serum conditions.



Supplementary Figure S8. Sorafenib is similar to sunitinib in inducing ciliary resorption. Representative Western blot images and quantitation of PDGF-AA-induced activation of PDGFR and MEK1/2 in NIH3T3 cells, related to Figure 3F, G. Cells were treated with indicated drugs for 2 h (top 4 panels) or 30 min (bottom 4 panels); values represent data normalized to vinculin loading control for phosphorylated and total protein, (n=3, n, number of biological replicates). Statistics determined by Tukey test in all cases; *, p < 0.5; **, p < 0.01; ***, p < 0.001, relative to vehicle, no ligand; #, p < 0.5; ##, p < 0.01; ###, p < 0.001, relative to vehicle, plus