Table S1. Biochemical and cellular potency of encorafenib, dabrafenib, and vemurafenib in biochemical and cell-based assays. Purified BRAF V600E kinase domain was used in the biochemical experiments, using kinase-dead MEK1 as a substrate and a phosphorylated MEK1 (Ser217/221) alpha-screen readout. A375 cells were incubated with inhibitors for 3 hours to determine the phosphorylated ERK (pERK) half maximal effective concentration (EC₅₀) and 72 hours to determine the proliferation EC₅₀.

	Biochemical		A375 Cells		
Inhibitor	BRAF V600E IC ₅₀ , μΜ	Dissociation T _{1/2} , h	pERK EC _{50,} μM	Proliferation EC _{50,} μM	Fold Change pERK EC ₅₀ 3 Hours After Washout
Encorafenib	0.0004	> 30	0.003	0.004	2
Dabrafenib	0.00007	2	0.002	0.017	14
Vemurafenib	0.0007	0.5	0.017	0.060	23

EC₅₀, half maximal effective concentration; T_{1/2}, half-life.