



This week in therapeutics

Indication	Target/marker/ pathway	Summary	Licensing status	Publication and contact information
Cancer				
Brain cancer	Hedgehog pathway; P glycoprotein (MDR1; ABCB1; P-gp; CD243)	Mouse studies suggest P-gp inhibitors could help reverse resistance to saridegib in medulloblastoma. In a mouse model of aggressive medulloblastoma, the hedgehog pathway inhibitor saridegib decreased tumor size and increased survival compared with vehicle. In the model, hedgehog pathway signaling rebounded and levels of P-gp, a transporter for which saridegib is a known substrate, increased after six weeks of treatment. In the same model, saridegib plus the P-gp inhibitor verapamil reversed the drug resistance seen at six weeks. Next steps could include testing a P-gp inhibitor plus saridegib in clinical trials for medulloblastoma. Infinity Pharmaceuticals Inc.'s saridegib is in Phase II trials to treat bone cancer and myeloproliferative disorder and is in preclinical development for non–small cell lung cancer (NSCLC). Verapamil is a generic calcium channel blocker approved for multiple cardiovascular indications.	Unpatented; licensing status not applicable	Lee, M.J. et al. Proc. Natl. Acad. Sci. USA; published online May 1, 2012; doi:10.1073/pnas.1114718109 Contact: James M. Olson, Fred Hutchinson Cancer Research Center, Seattle, Wash. e-mail: jolson@fhcrc.org
		multiple cardiovascular indications. SciBX 5(19); doi:10.1038/scibx.2012.489 Published online May 10, 2012		