

This week in therapeutics

Indication	Target/marker/pathway	Summary	Licensing status	Publication and contact information
Cancer				
Cancer	Colony-stimulating factor 1 receptor (CSF1R; C-FMS; CD115); inhibitor of κ light polypeptide gene enhancer in B cells kinase- β (IKBKB; IKK2); rap guanine nucleotide exchange factor 4 (RAPGEF4; EPAC2); toll-like receptor 4 (TLR4)	<p><i>In vitro</i> studies suggest inhibiting a subset of kinases could improve the efficacy of the chemotherapeutic paclitaxel. In human ovarian cancer cell lines, RNAi knockdown of multiple kinases—including <i>CSF1R</i>, <i>IKBKB</i>, <i>RAPGEF4</i> and <i>TLR4</i>—increased paclitaxel-induced cytotoxicity compared with that seen using paclitaxel alone. In a paclitaxel-resistant lung carcinoma cell line, <i>RAPGEF4</i> knockdown plus paclitaxel decreased cell growth compared with paclitaxel alone. Ongoing work includes simultaneous knockdown or pharmacological inhibition of multiple kinases in paclitaxel-treated cancer cell lines and animal models of cancer. Eritoran (E5564), a synthetic lipid A analog that blocks TLR4 activation from Eisai Co. Ltd., is in Phase III testing to treat sepsis. PLX5622, a CSF1R inhibitor from Daiichi Sankyo Co. Ltd., is in Phase I testing to treat rheumatoid arthritis (RA). SAR113945, an IKBKB inhibitor from Sanofi, is in Phase I testing to treat osteoarthritis (OA).</p> <p>SciBX 4(36); doi:10.1038/scibx.2011.1013 Published online Sept. 15, 2011</p>	Unpatented; available for licensing or partnering	<p>Ahmed, A.A. <i>et al. Cancer Res.</i>; published online July 20, 2011; doi:10.1158/0008-5472.CAN-11-0025 Contact: Robert C. Bast Jr., The University of Texas M.D. Anderson Cancer Center, Houston, Texas e-mail: rbast@mdanderson.org Contact: Ahmed Ashour Ahmed, University of Oxford, Oxford, U.K. e-mail: ahmed.ahmed@obs-gyn.ox.ac.uk</p>