

## THE DISTILLERY

## This week in therapeutics

Indication	Target/marker/pathway	Summary	Licensing status	Publication and contact information
Cancer				
Cancer	HER2 (EGFR2; ERBB2; neu); neuregulin 1 (NRG1)	In vitro, mouse and patient studies suggest HER2 inhibitors could help treat EGFR inhibitor-resistant cancer. In primary and metastatic colorectal cancer patients, greater levels of HER2 or NRG1 in tumors were associated with resistance to Erbitux cetuximab and correlated with poor survival. In a panel of Erbitux-resistant human cancer cell lines, HER2 and NRG1 levels were higher than those in Erbitux-sensitive cell lines. In xenograft mice with Erbitux-resistant cancer cells, Erbitux plus Tykerb lapatinib or pertuzumab decreased tumor growth compared with Erbitux alone. Ongoing work includes additional studies in xenograft models and primary tumors. Eli Lilly and Co., Bristol-Myers Squibb Co. and Merck KGaA market Erbitux, a mAb targeting EGFR, to treat colorectal cancer and head and neck cancer. GlaxoSmithKline plc markets Tykerb/Tyverb, an inhibitor of HER2 and epidermal growth factor receptor 1 (EGFR1; HER1; ERBB1) to treat breast cancer. Pertuzumab (2C4; R1273; RG1273), a mAb HER dimerization inhibitor that prevents HER2 from binding to HER1, HER3 (EGFR3; ERBB3) and HER4 (EGFR4; ERBB4) from Roche's Genentech Inc. unit and Chugai Pharmaceutical Co. Ltd., is in Phase III testing to treat breast cancer, Phase II testing to treat breast cancer, Phase II testing to treat non-small cell lung cancer (NSCLC) and Phase I testing to treat ovarian cancer.	Patent and licensing status for findings in first study unavailable Findings in second study unpatented; available for partnering	Yonesaka, K. <i>et al. Sci. Transl. Med.</i> ; published online Sept. 7, 2011; doi:10.1126/scitranslmed.3002442 <b>Contact:</b> Kazuhiko Nakagawa, Kinki University School of Medicine, Osaka, Japan e-mail: <b>nakagawa@med.kindai.ac.jp</b> <b>Contact:</b> Pasi A. Jänne, Dana-Farber Cancer Institute, Brigham and Women's Hospital and Harvard Medical School, Boston, Mass. e-mail: <b>pjanne@partners.org</b> Bertotti, A. <i>et al. Cancer Discov.</i> ; published online Sept. 2, 2011; doi:10.1158/2159-8290.CD-11-0109 <b>Contact:</b> Livio Trusolino, Institute for Cancer Research and Treatment, Torino, Italy e-mail: livio.trusolino@ircc.it

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