

## THE DISTILLERY

## This week in therapeutics

| Indication  | Target/marker/pathway   | Summary   | Licensing status                                       | Publication and contact<br>information   |
|-------------|---|---|--|--|
| Cancer      |   |   |  |  |
| Lung cancer | Anaplastic lymphoma<br>kinase (ALK); epidermal<br>growth factor receptor<br>(EGFR); stem cell factor<br>receptor tyrosine kinase<br>(c-Kit; KIT; CD117) | Patient and cell culture studies suggest targeting<br>EGFR and KIT signaling pathways could help<br>treat lung cancers resistant to ALK inhibitors.<br>In samples from 18 patients resistant to Xalkori<br>crizotinib, 17 showed EGFR activation and 2<br>samples showed KIT amplification. In Xalkori-<br>resistant human lung cancer cell lines, Xalkori<br>plus an EGFR or KIT inhibitor decreased<br>proliferation compared with Xalkori alone. Next<br>steps include identifying additional mechanisms<br>of Xalkori resistance and determining how they<br>influence resistance to next-generation ALK<br>inhibitors.<br>Pfizer Inc. markets Xalkori, a dual c-Met receptor<br>tyrosine kinase and ALK inhibitor, to treat non-<br>small cell lung cancer (NSCLC).<br>At least four other companies have ALK<br>inhibitors in Phase I/II testing or earlier to treat<br>cancer. | Work unpatented;<br>licensing status not<br>applicable | Katayama, R. <i>et al. Sci. Transl. Med.</i> ;<br>published online Jan. 25, 2012;<br>doi:10.1126/scitranslmed.3003316<br><b>Contact:</b> Jeffrey A. Engelman,<br>Massachusetts General Hospital<br>Cancer Center, Boston, Mass.<br>e-mail:<br>jengelman@partners.org |
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