

THE DISTILLERY

This week in therapeutics

| Indication | Target/marker/ pathway | Summary | Licensing status | Publication and contact information |
|------------|--|--|--|---|
| Cancer | | | | |
| Cancer | Heat shock protein 90 (HSP90AA1; Hsp90) | Studies in mice and cell culture suggest that gamitrinibs may be useful for treating cancer. Gamitrinibs consist of a 17-(allylamino)-17- demethoxygeldanamycin (17-AAG)-derived scaffold linked to a mitochondrial targeting moiety. In human leukemia, breast cancer and lung cancer xenograft mouse models, a gamitrinib analog decreased tumor proliferation compared with that seen using vehicle or 17-AAG. In cancer cell lines, gamitrinibs entered mitochondria, inhibited Hsp90 activity and induced cell death via mitochondrial apoptosis. Gamitrinibs also had broad-spectrum activity across a panel of 12 cancer cell lines, whereas 17-AAG did not. Next steps include evaluating gamitrinibs in long-term studies in animals. Tanespimycin (17-AAG), an Hsp90 inhibitor from Bristol-Myers Squibb Co., is in Phase III testing to treat multiple myeloma (MM). At least 11 other companies have Hsp90 inhibitors in Phase II or earlier to treat cancer. | Multiple patent applications filed covering structure and biology of the compounds; available for licensing from the University of Massachusetts Commercial Ventures and Intellectual Property Contact: James McNamara, University of Massachusetts Medical School, Worcester, Mass. phone: 508-856-4390 e-mail: james.mcnamara@umassmed.edu | Kang, B.H. <i>et al. J. Clin. Invest.</i> published online Feb. 23, 2009 doi:10.1172/JCI37613 Contact: Dario C. Altieri, University of Massachusetts Medical School, Worcester, Mass. e-mail: dario.altieri@umassmed.edu |
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