

THE DISTILLERY

This week in therapeutics

| Indication | Target/marker/ pathway | Summary | Licensing status | Publication and contact information |
|----------------------|---|---|---|--|
| Metabolic disease | | | | |
| Hypercholesterolemia | Niemann-Pick C1-like protein (NPC1L1) | In vitro studies of the ezetimibe (EZE) binding site of NPC1L1 could help guide the design of improved therapies for hypercholesterolemia. NPC1L1 is an intestinal protein that mediates the absorption of dietary cholesterol, and it is inhibited by EZE. Biochemical and proteomics analysis of the EZE- NPC1L1 complex identified the loop C region of NPC1L1 as the binding site for EZE. Next steps should include design of next-generation NPC1L1 inhibitors. Zetia ezetimibe is marketed by Schering- Plough Corp. and Merck & Co. Inc. to treat hypercholesterolemia. | Patent and licensing status unavailable | Weinglass, A. <i>et al. Proc. Natl.</i> <i>Acad. Sci. USA</i> ; published online Aug 4, 2008; doi:10.1073/pnas.0800936105 Contact: Adam B. Weinglass, Merck Research Laboratories, Rahway, N.J. e-mail: adam_weinglass@merck.com |

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