

THE DISTILLERY

This week in techniques

Approach	Summary	Licensing status	Publication and contact information
Drug platforms			
Crystal structure of Selzentry maraviroc- bound HIV-1 co- receptor CC chemokine receptor 5 (CCR5; CD195)	The crystal structure of HIV-1 co-receptor CCR5 bound by the allosteric inhibitor Selzentry maraviroc could help to guide design of therapies for HIV-1 infection. The crystal structure of the complex was determined at 2.7 Å resolution and showed Selzentry bound at a site distinct from proposed recognition sites for chemokines and HIV gp120. Crystal structure–based modeling showed that different charge distributions and steric hindrances in the co-receptor ligand-binding pocket could be major determinants for HIV-1 co-receptor selectivity. Next steps include structural studies of CCR5 and CXC chemokine receptor 4 (CXCR4; NPY3R) in complex with the HIV envelope protein gp120 and CD4 to obtain more insight into the process of viral infection. Selzentry is marketed by Pfizer Inc. to treat HIV/AIDS. CytoDyn Inc.'s CCR5 inhibitor, PRO 140, is in Phase II trials. Tobira Therapeutics Inc. has the dual CCR5 and CCR2 (CD192) antagonist cenicriviroc in Phase II trials to treat HIV/AIDS.	Unpatented; licensing status not applicable	Tan, Q. <i>et al. Science</i> ; published online Sept. 12, 2013; doi:10.1126/science.1241475 Contact: Beili Wu, Shanghai Institute of Materia Medica, Chinese Academy of Sciences, Shanghai, China e-mail: beiliwu@simm.ac.cn

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