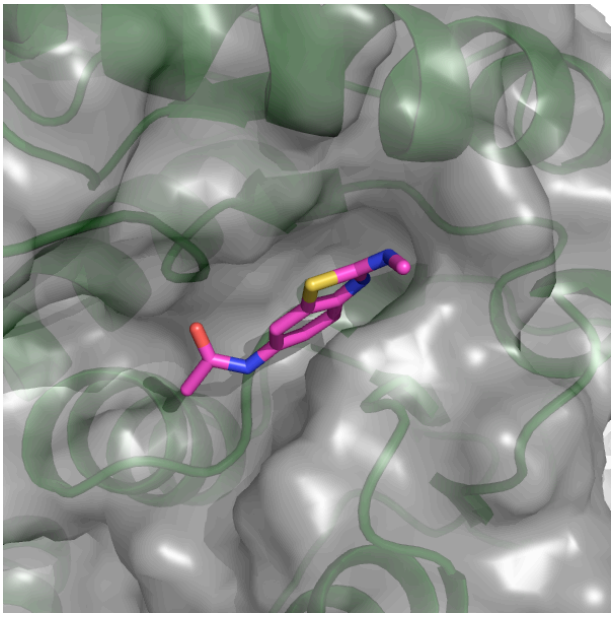
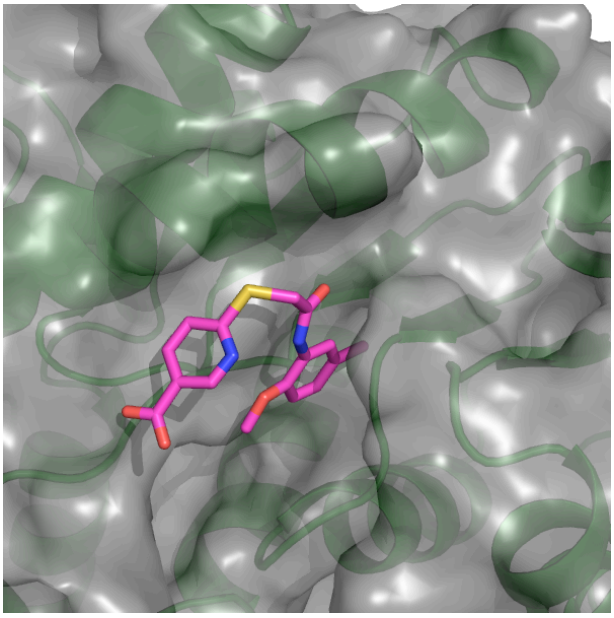
Text S5. Initial structures for steered MD simulations.

The inhibitor was pulled towards you (the viewer). LDHA structure is labeled by green cartoon (backbone) and grey surface, whereas inhibitors are represented by thick sticks with atoms colored as: carbon, magenta; oxygen, red; nitrogen, blue; phosphate, orange; sulfur, yellow; chlorine, green; fluorine, pale cyan. The mobile loop is also colored in magenta for S-site binders.

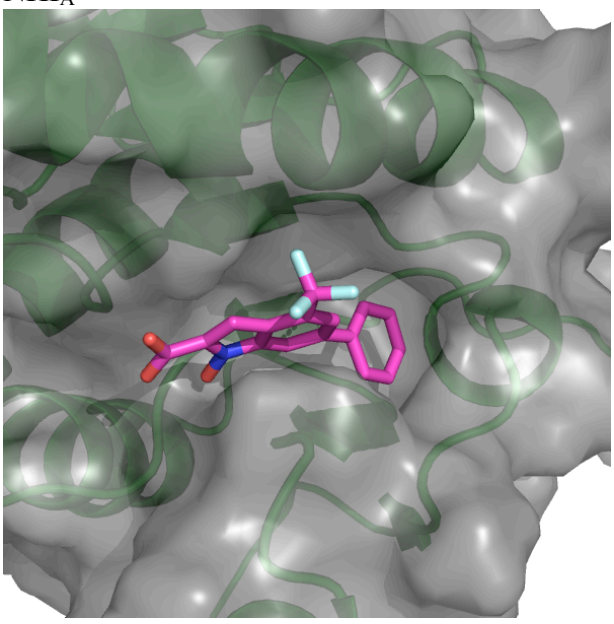
AJ1



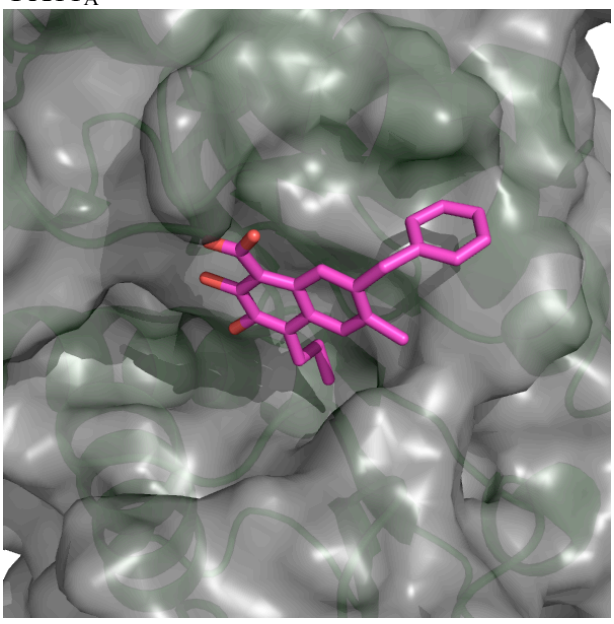
1E7



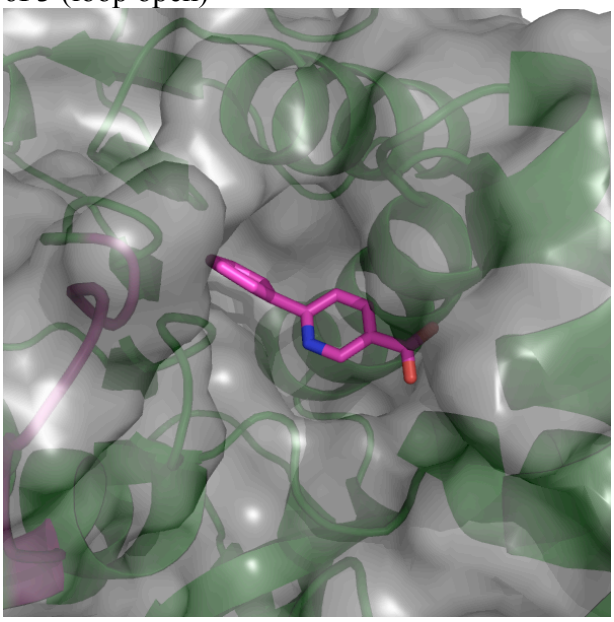
NHI_A



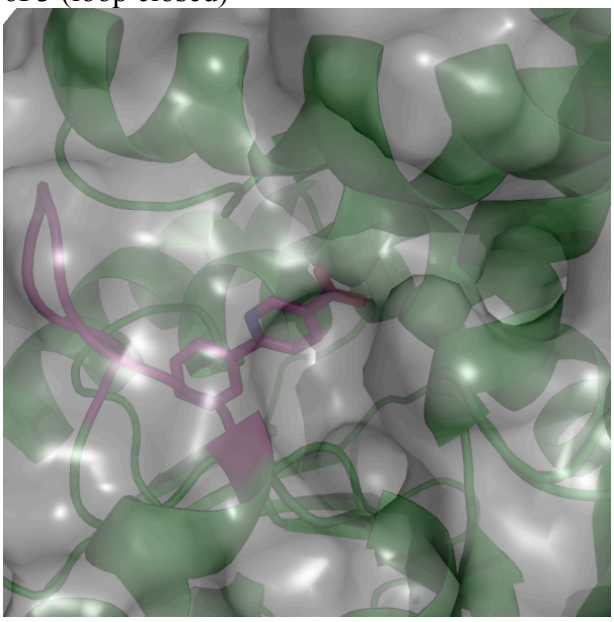
$FX11_A$



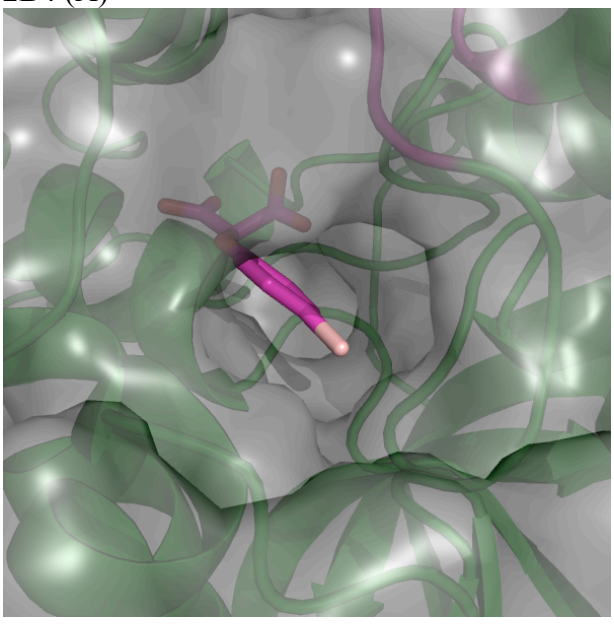
6P3 (loop open)



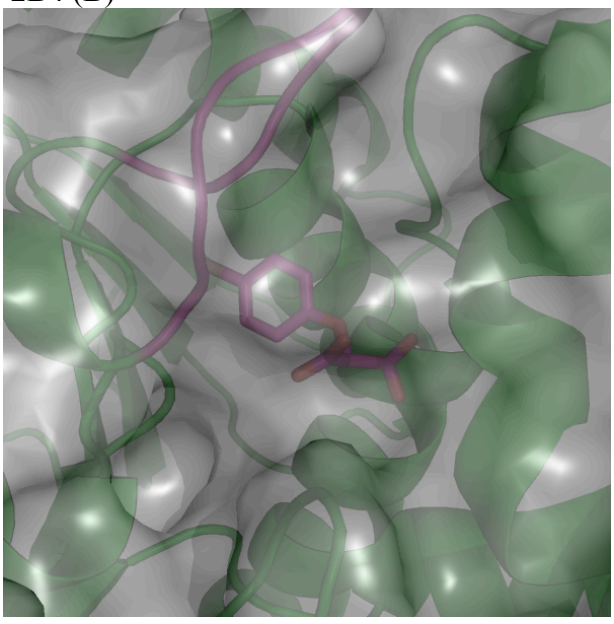
6P3 (loop closed)



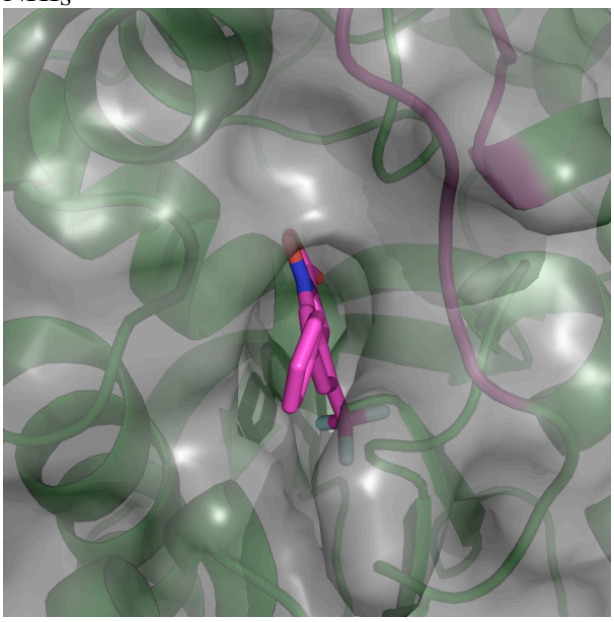
2B4 (A)



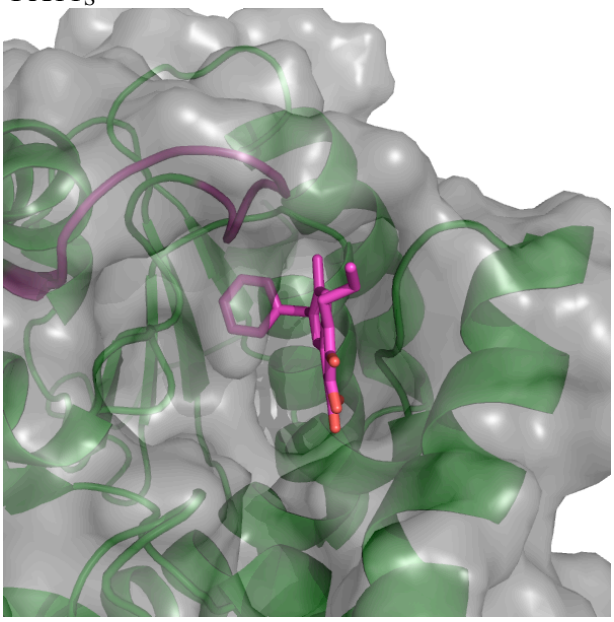
2B4 (B)



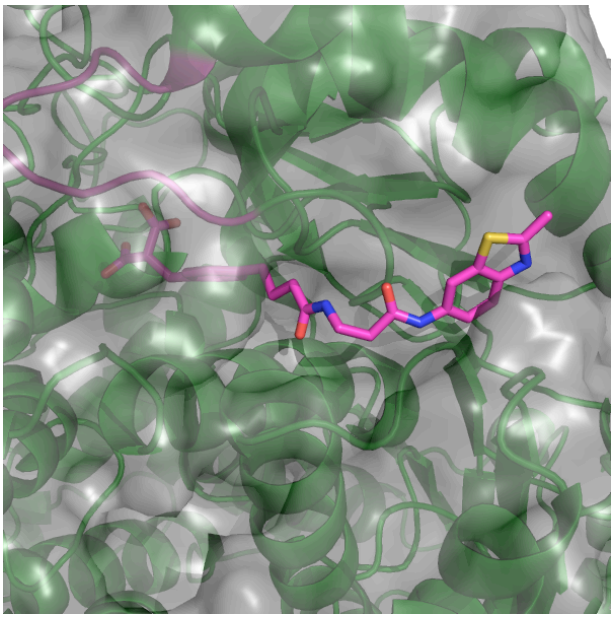
NHI_S



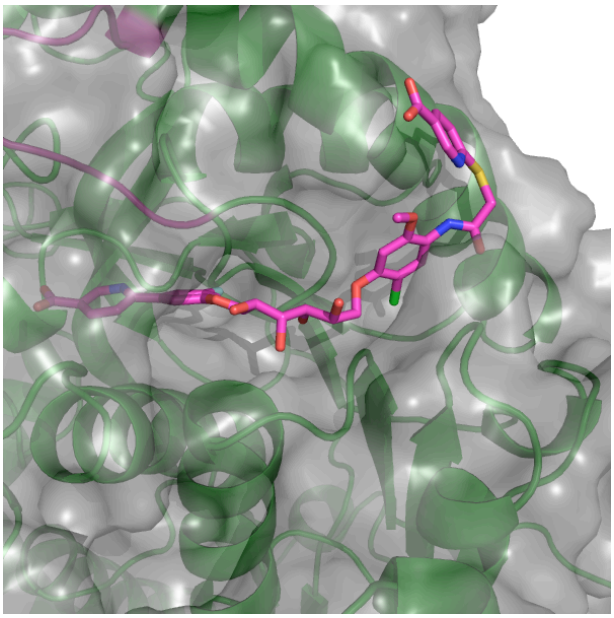
$FX11_S$



0SN



1E4



Apparently, the pulling of 6P3 (loop closed), 2B4 (A and B), and NHI_S, $FX11_S$, 0SN, and 1E4 is complicated by steric clashes of different extents.