**RUN1397** 0.8 binding binding (all GENS)  $\Delta\Delta G$  / kcal mol $^{-1}$ 0.6 0.4 0.2 0.0 -0.2solvent 49.8 complex  $\Delta G$  / kcal mol<sup>-1</sup> 49.6 49.4 49.2 49.0 0

**GEN**