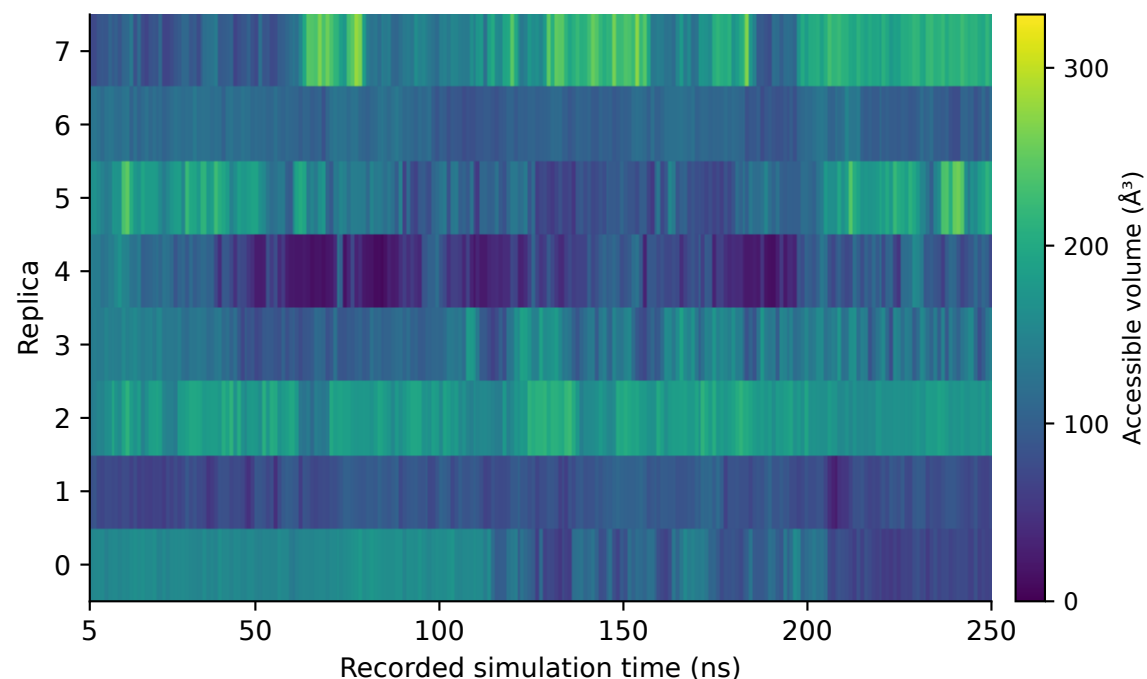


Pocket opening checks across eight simulation runs

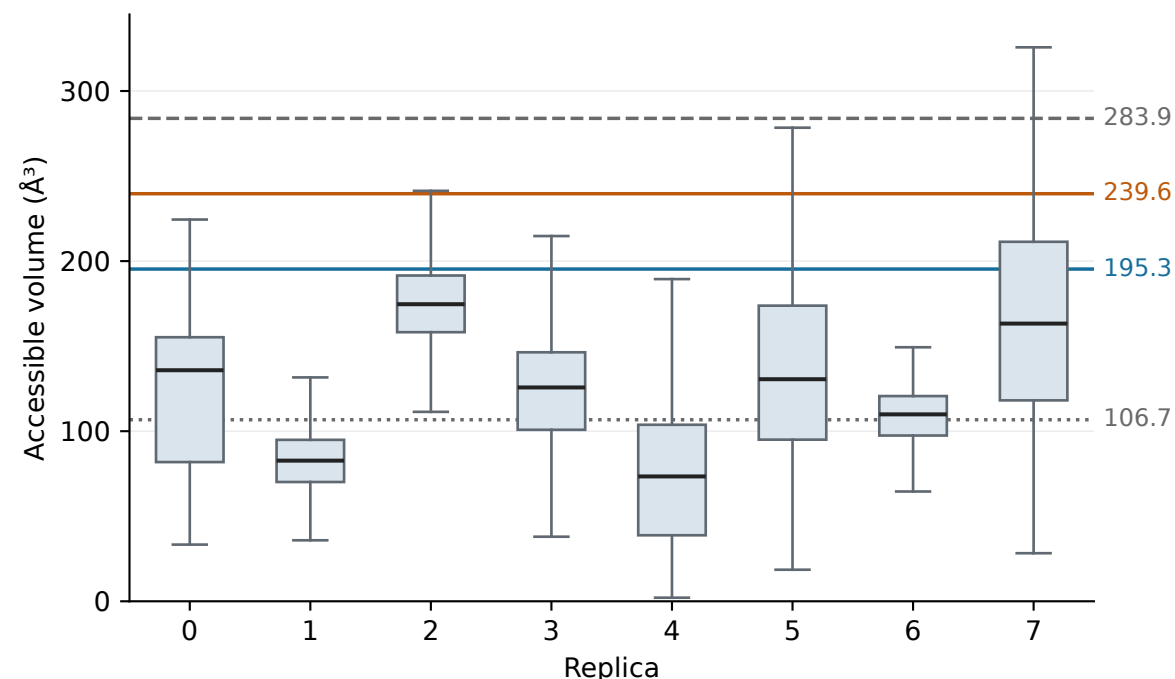
7,773 snapshots after the first 5 ns | Percentages describe the accelerated simulations

■ 195.3 Å³ cutoff ■ 239.6 Å³ cutoff ···· Start 106.7 Å³ - - - 7RPZ reference 283.9 Å³

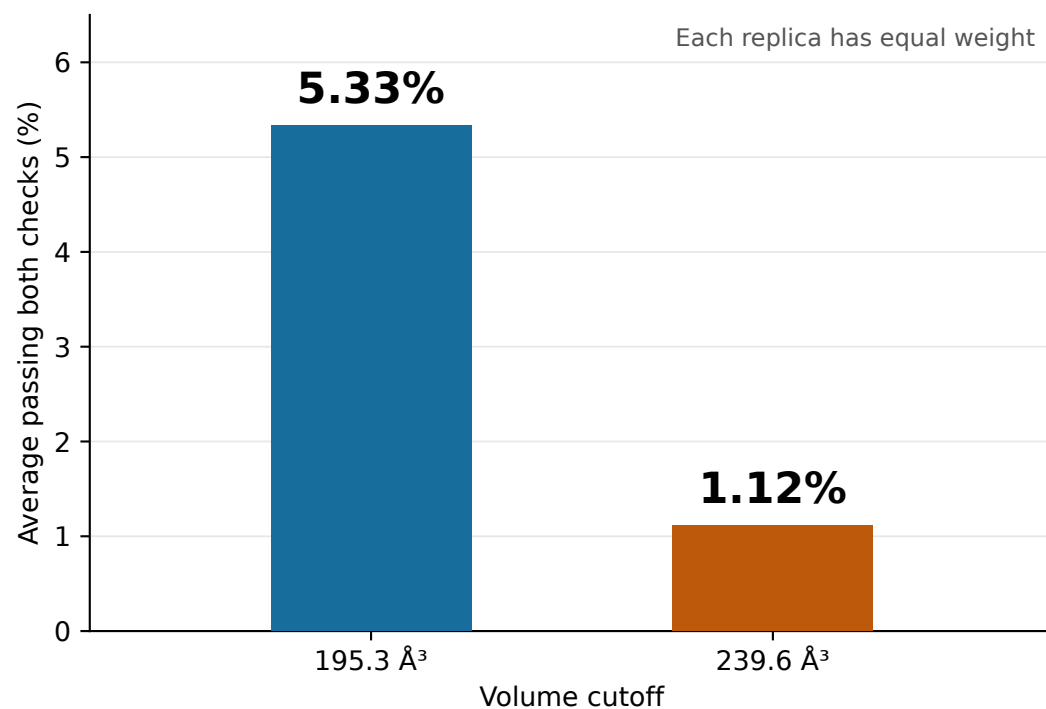
A Space at the binding site over time



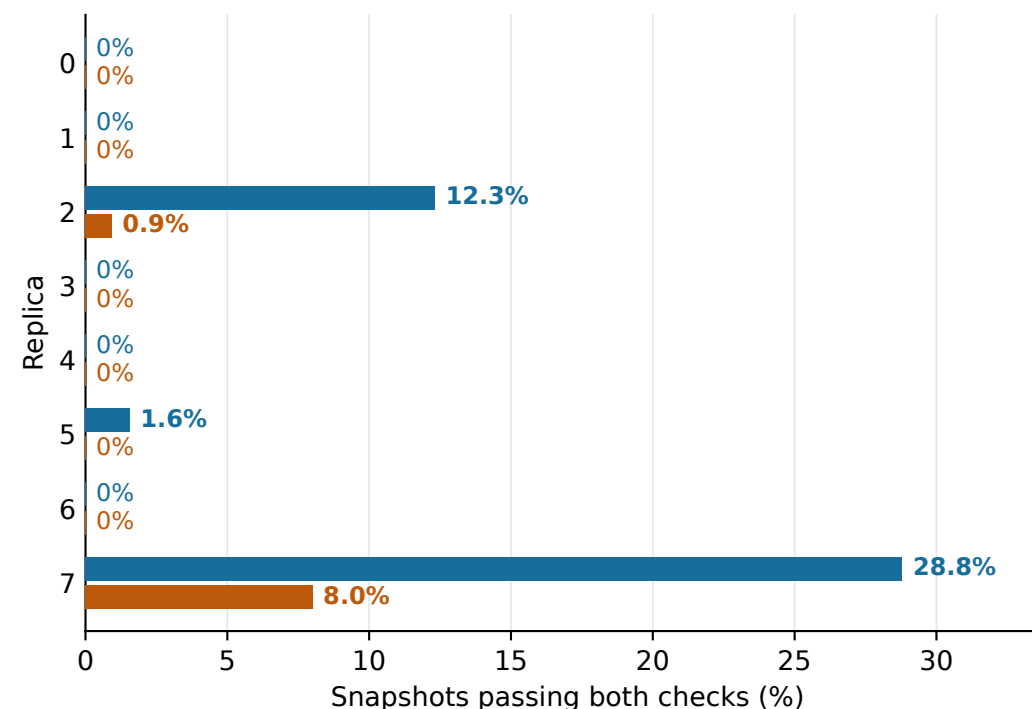
B Volume distributions



C Average across all eight replicas



D Percentages for each replica



Passing requires BOTH the volume cutoff and a switch II shape closer to the reference than to the start.

C shows the mean of the eight percentages in D. Neither panel estimates how often the pocket opens in nature.