

Discovery of potent dual ligands for Dopamine D4 and σ 1 receptors

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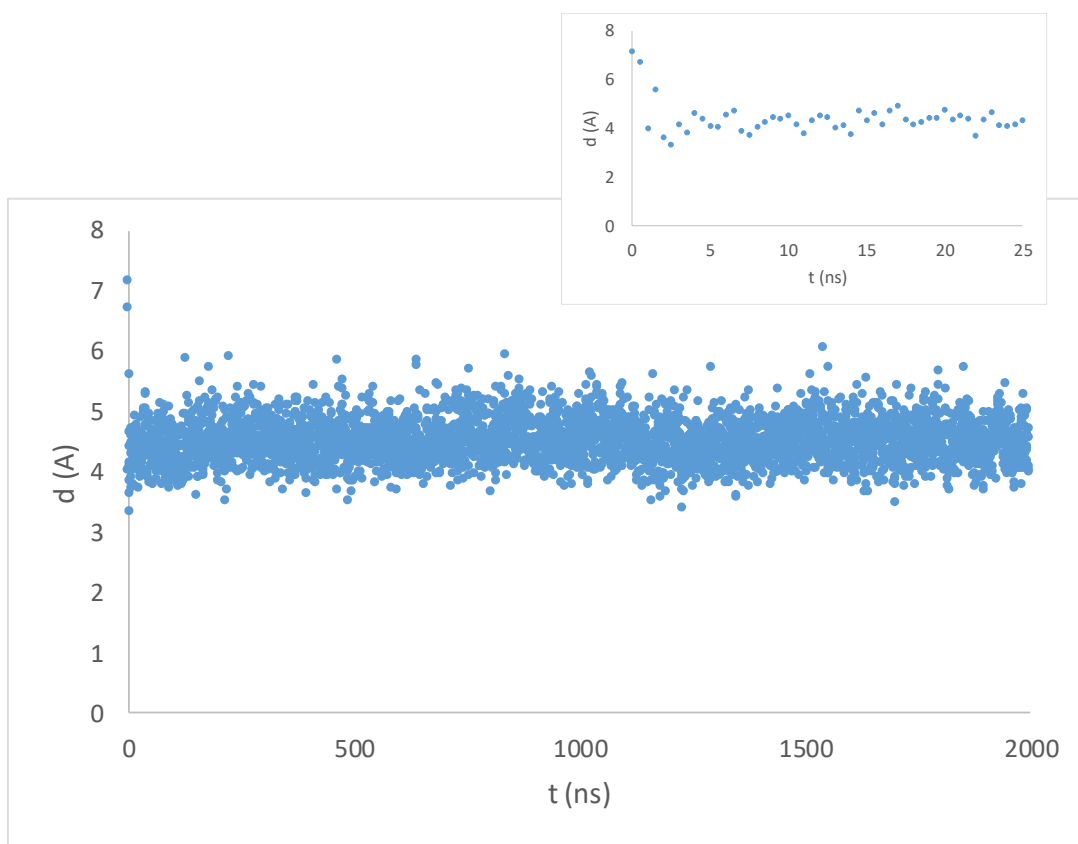
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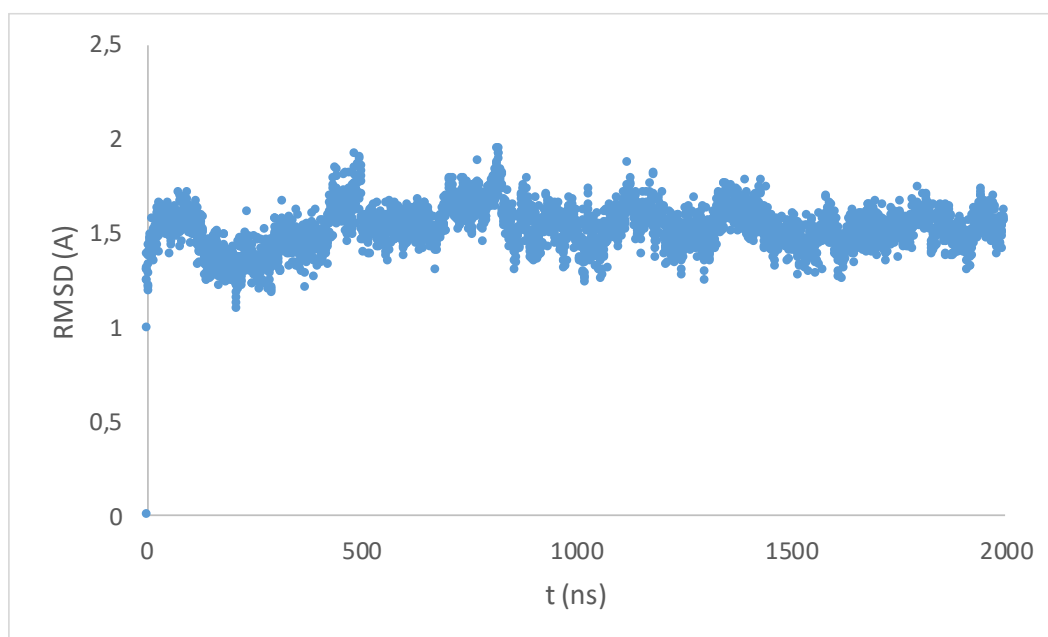
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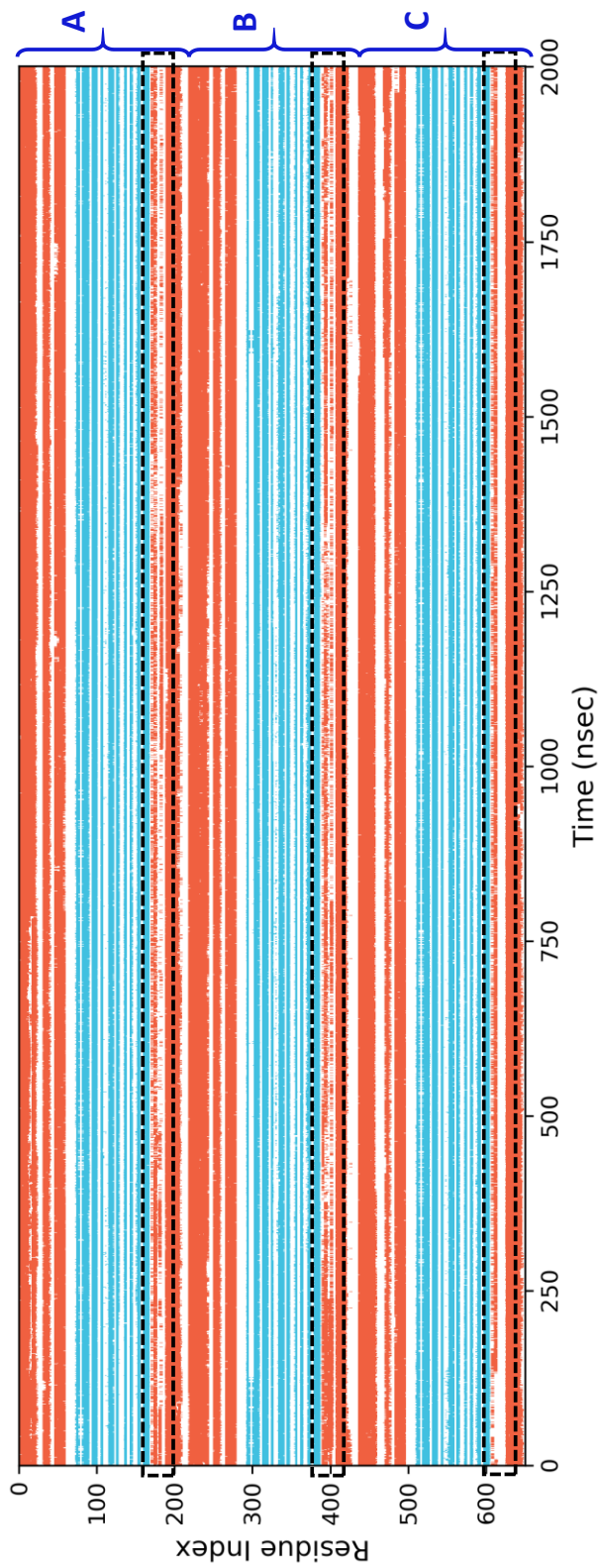
- **Suppl. Fig. 1** Distance between the sulfur atom of M93 and the centroid of the ligand triazole during the molecular dynamics experiment **page S2**
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Suppl. Fig. 1 Distance between the sulfur atom of M93 and the centroid of the ligand triazole during the molecular dynamics experiment. Note that the distance dropped very early, then stabilized till the end of the simulation at a mean distance of 4.5 Å.

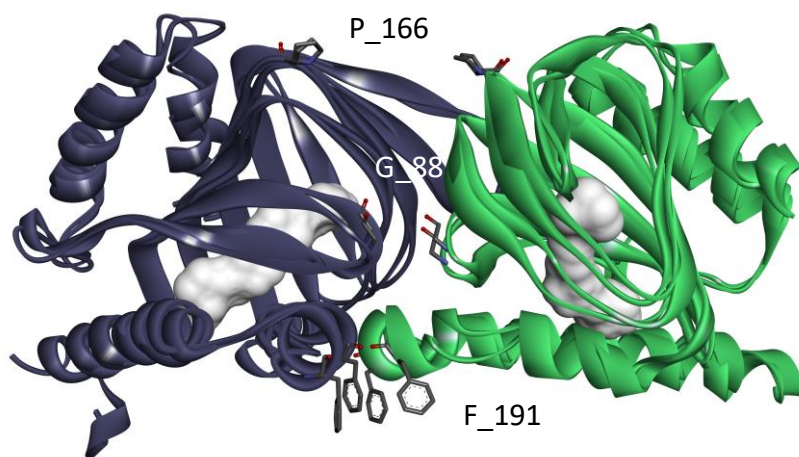


Suppl. Fig. 2 RMSD of the aromatic amino-acids (W89, F107, Y120, F133, H154, F184, W164) located at the cytosolic end of the ligand binding site during the molecular dynamics experiment.



Helix α_4 of the 3 protomers A, B, C

Suppl. Fig. 3 Secondary structures of σ_1 receptor during molecular dynamics. α -helices are in red, β -strands in blue. Note the partial unfolding of helix α_4 .



	F191		G88		P166	
	d0 (Å)	df (Å)	d0 (Å)	df (Å)	d0 (Å)	df (Å)
d_{AB}	6,4	8,7	5,5	6,8	15,4	15,5
d_{BC}	6,1	10,3	5,7	7,1	14,4	14,8
d_{AC}	5,6	5,3	6,1	7,3	14,2	14,9

Suppl. Table 1 Distance between protomers during the simulation, from the bottom (F191), middle (G88) and top of the interface (P166). The initial (d0) and final distance (df) are the mean of the 10 first/last frames of the molecular dynamics.