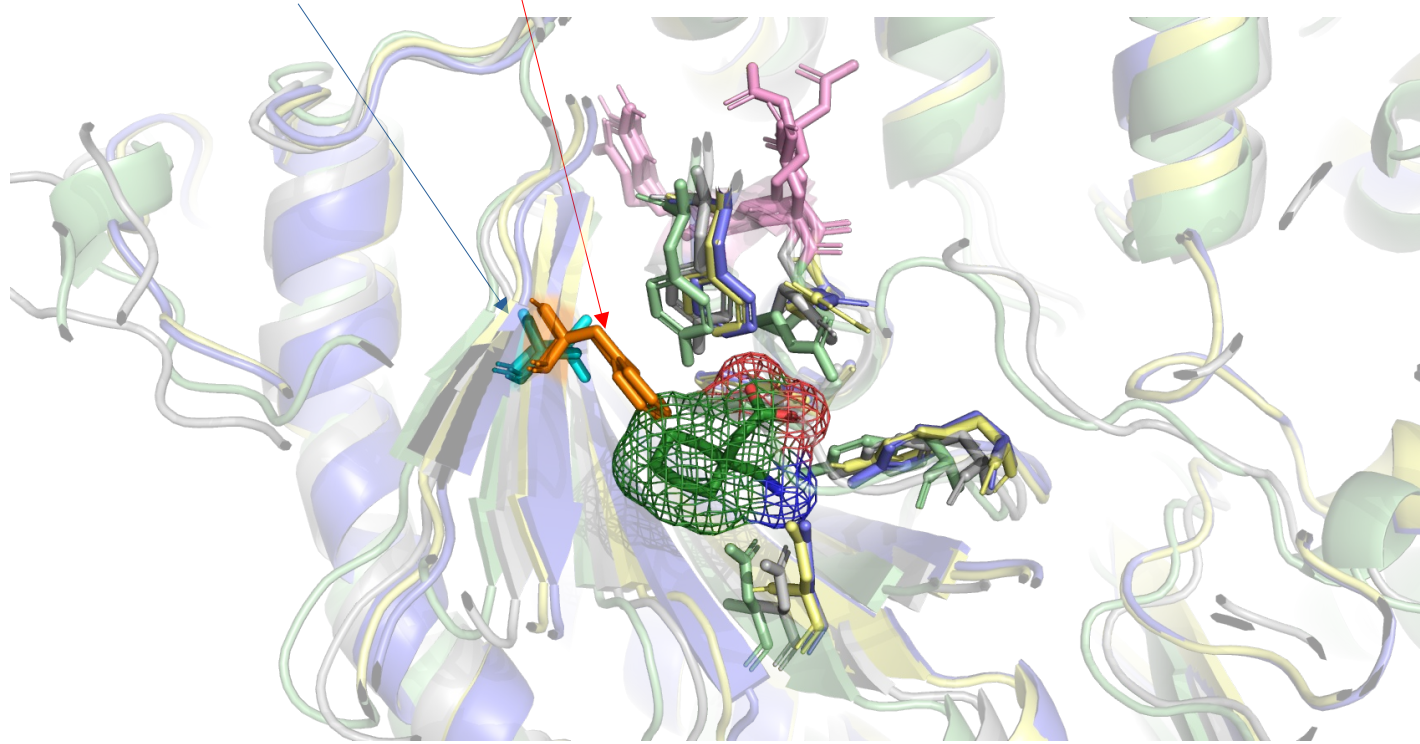


A217 (**A2D-1**) and T257 (**A2D-2**) F221 (**A2D-3**) and F256 (**A2D-4**)



Superimposed ligand-binding pockets of $\alpha_2\delta_1$ to $\alpha_2\delta_4$ proteins. Gabapentin was docked to $\alpha_2\delta_1$. $\alpha_2\delta_1$ (gray) is from rabbit, $\alpha_2\delta_2$ (light green), $\alpha_2\delta_3$ (light yellow), and $\alpha_2\delta_4$ (light blue) are AlphaFold models of human proteins. An amino acid causing steric hindrance is indicated by the red arrow. In pink the first two residue of the RRR (in $\alpha_2\delta_1$ and $\alpha_2\delta_2$)/RNR (in $\alpha_2\delta_3$ and $\alpha_2\delta_4$) sequence are shown; the last R of the RRR/RNR sequence is part of the amino acid binding motif. The amino acid binding motif (Gumerov et al., 2022) residues are also shown.