



**Supplementary Fig. 1** Stereoview of the positions of CnA residues Tyr341, Leu343 and Met347 in the full-length calcineurin structure (CnA ,yellow and grey; CnB, blue). These hydrophobic residues are surface-exposed in Cncat and therefore the mutations Y341S, L343A and M347D increased the solubility of Cncat. This figure was prepared using the full-length calcineurin structure (PDB entry 1aui).