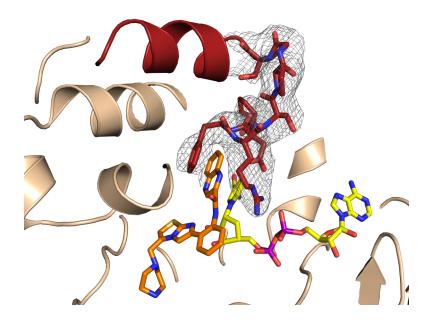
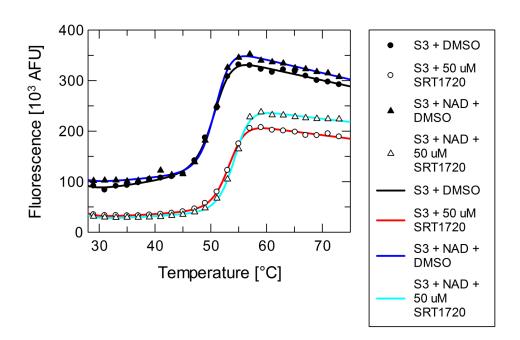
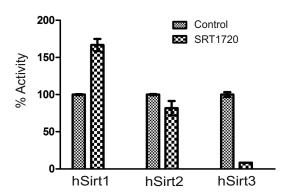
## **Supplementary material**



**Supplementary Figure S1:** Cofactor binding loop region of the refined Sirt3/carba-NAD $^{+}$ /SRT1720 complex. 2Fo-Fc electron density contoured at  $1\sigma$  is shown for residues 155-162 (2 Å distance cut-off).



**Supplementary Figure S2:** Thermal denaturation shift assays in presence and absence of 50  $\mu$ M SRT1720 and/or 500  $\mu$ M NAD+. The lines shown are nonlinear fits for a two state transition.



Supplementary Figure S3: Effects of 20  $\mu$ M SRT1720 on Fluor-de-Lys-1-deacetylation activity of different Sirtuin isoforms.