

Supporting information:

Table S1: amino acid sequences of peptides that mediate the oligomerization of STIL IDR as found in the peptide array screening

Residues	peptide sequence
367-381	FSKASKNF SIKRSSQ
376-390	IKRSS SQKLSS SGKMPI
382-396	KLSS SGKMPI IHDH DSG
406-420	PIPS HPV SQKISKI
451-465	ENPPLIN HLEHLKPL
457-471	NHLEHLKPL QPQLY D
487-501	GIPNQLN QDKPALLR
496-510	KPALLR HCKV RQPPA
502-516	HCKV RQPPAYKKGNP
511-525	YKKGNP HTRNSIKPS
517-531	HTRNSIKPSS HNGPS
547-561	QNE EYPIRPS TLNSR
571-585	PHDFV SPHNS GRPM
592-606	PPLPSY CSTNVC RC CC
601-615	NV CRCCQH HS HIQYS
607-621	QHHS HIQY SPLNS WQ
631-645	DVQSEAL LQKHS LFHP
652-666	YCNAFC SSSSPIALR

* Colored residues (His, Cys, Glu, Asp) are common residues for binding Zn²⁺ ions.

** The motif that was detected using the Block Maker alignment tool ¹ is colored yellow.

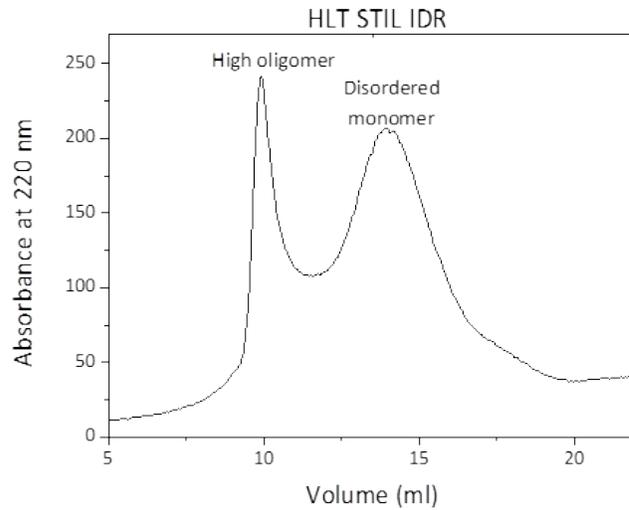


Figure S1: Analytical SEC of HLT STIL IDR: the protein exists as a mixture of monomer and high order oligomer.

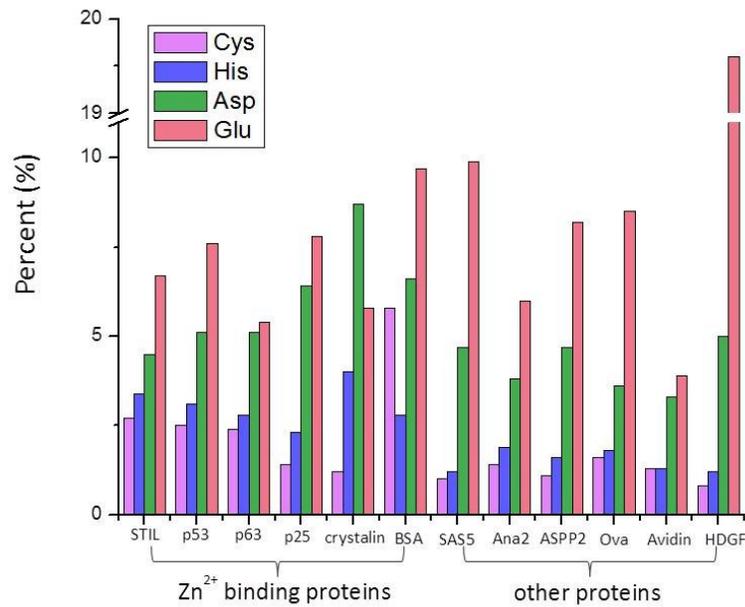


Figure S2: STIL is composed of a high portion of Cys and His residues. Shown is the amino acid composition of several Zn²⁺ and non-Zn²⁺ binding proteins, as was calculated by the Bioinformatics Resource Portal ProtParam tool ².

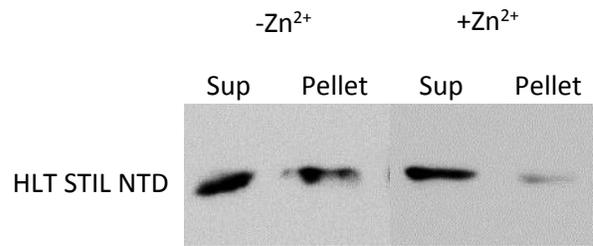


Figure S3: the presence of ZnCl₂ in the medium improved the expression of HLT STIL NTD. Western blot analysis of supernatant and pellet fractions from bacterial expression of HLT STIL NTD with or without addition of ZnCl₂ to the medium.

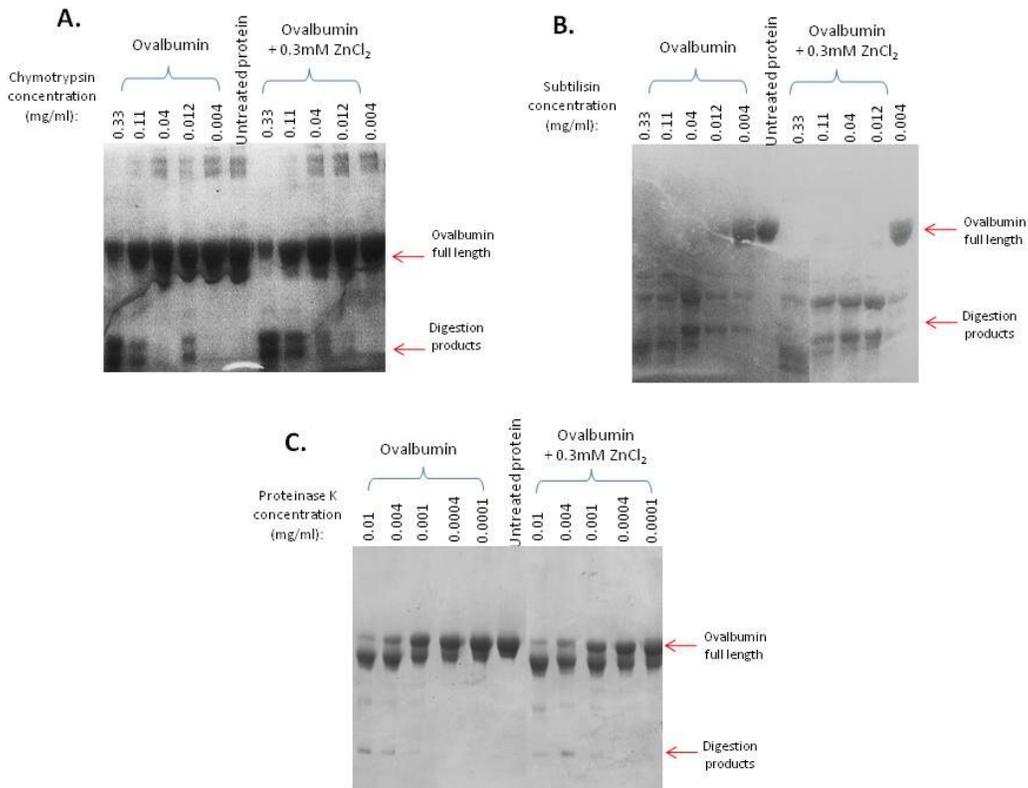


Figure S4: The presence of ZnCl₂ does not affect protease digestion of OvAlbumin. Shown is SDS-PAGE gel of OvAlbumin with and without ZnCl₂ at several concentrations of Chymotrypsin (A), Subtilisin (B) and Proteinase K (C).

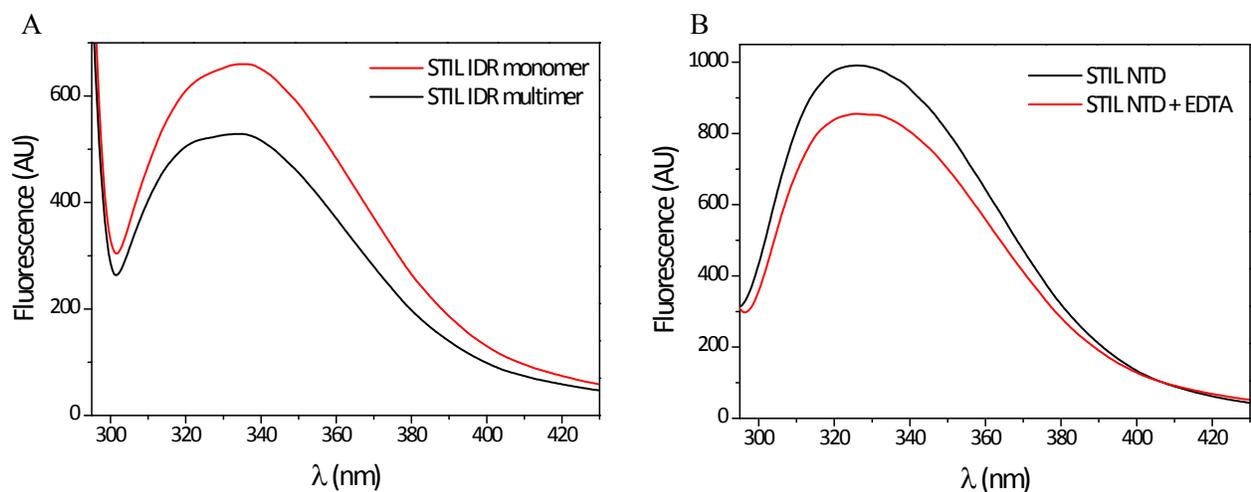


Figure S5: The presence of Zn^{2+} affects the structure of STIL IDR and NTD. A) Tryptophan fluorescence spectra of apo-STIL IDR (monomer - red) and holo-STIL IDR (multimer - black). B) Tryptophan fluorescence spectra of holo-STIL NTD (without EDTA addition - black) and apo-STIL NTD (with EDTA - red).

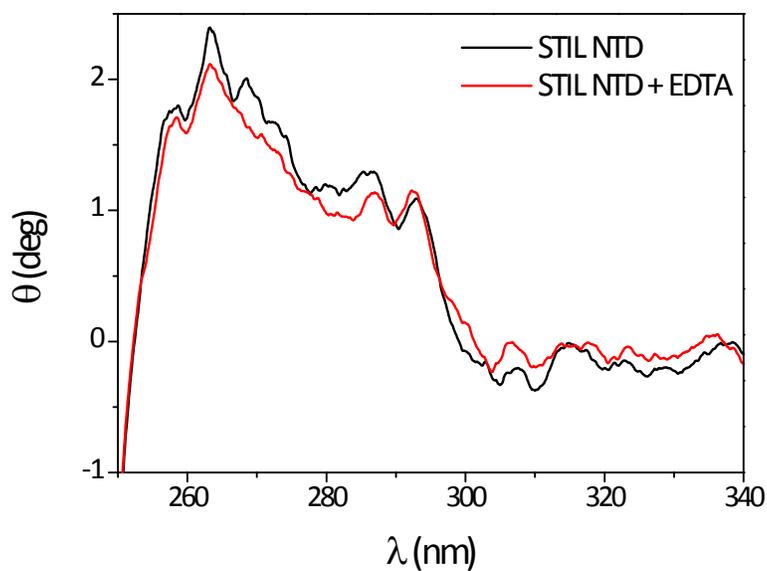


Figure S6: Near UV CD studies show that the structure of STIL NTD changes in presence of Zn²⁺. Near UV CD spectra of Zn²⁺ pre-bound STIL NTD (black) and without Zn²⁺, in the presence of EDTA (red).

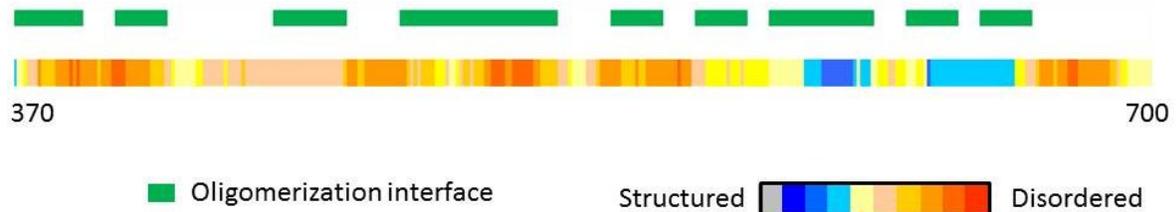


Figure S7: Oligomerization of the STIL IDR: The STIL IDR peptides that mediate its oligomerization were revealed by peptide array screening and are colored green on the STIL disordered region.



Figure S8: STIL oligomerization motifs on IDR sequence. The oligomerization motif SxKxS/SxHxS/SxLxS - colored orange and the Zn binding residues found in the oligomerization sites identified by peptide array screening: Cys residues - colored green and His residues -colored pink.

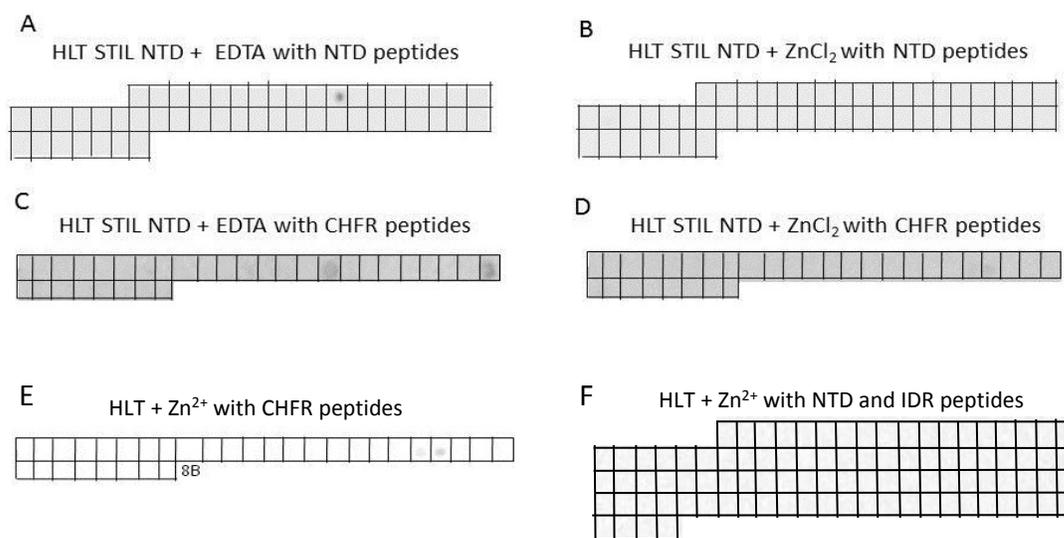


Figure S9: Testing the binding of STIL NTD and HLT to arrays of peptides derived from STIL and CHFR: Screening for binding of HLT STIL NTD to an array of peptides derived from STIL NTD derived peptides in the presence of EDTA (A) or ZnCl₂ (B), and to an array of CHFR derived peptides in the presence of EDTA (C) or ZnCl₂ (D). HLT was screened for binding CHFR derived peptide array (E) and STIL derived peptide array (F). Each black spot indicates binding between the protein and the corresponding peptide.

HHHHHS GAFEFKLPDIGEGIH EGEIVKWFVKPGDEVNEDDVLCE

VQNDKAVVEIPSPVKGKVL EILVPEGTVATVGQTLITLDAPGYENM

TTGSDTGENLYFQG

A

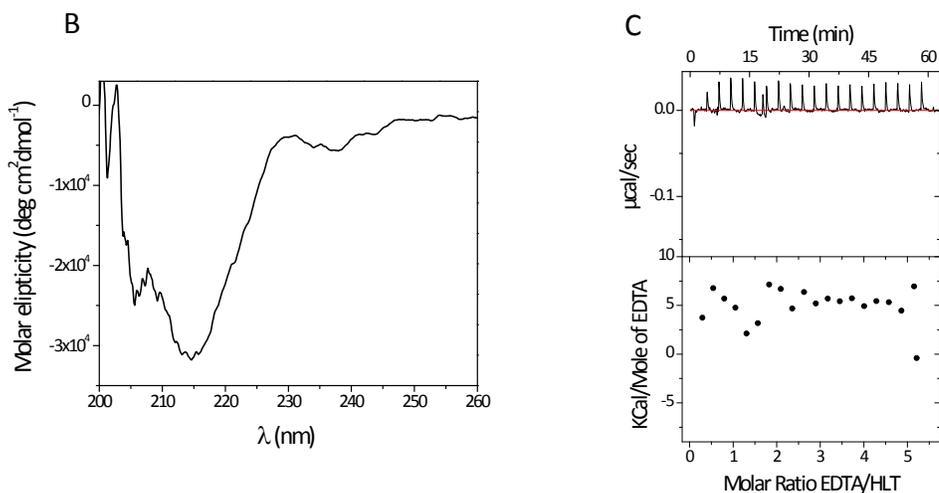


Figure S10: The HLT tag. (A) Primary sequence of HLT tag, composed of 6 His residues (pink), a Lipodomain (blue) and a Tev cleavage site (red). (B) CD spectrum of HLT tag. (C) ITC control - titration of EDTA into a HLT.

References

1. S. Henikoff, S. Pietrokovski, and J. G. Henikoff, *Nucleic Acids Res.*, 1998, **26**, 309–312.

2. E. Gasteiger, A. Gattiker, C. Hoogland, I. Ivanyi, R. D. Appel, and A. Bairoch, *Nucleic Acids Res.*, 2003, **31**, 3784–3788.