## SUPPLEMENTARY DATA

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Format As	Select Summ	mary (protein	hits) 🗸					He	alp		
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605		919.4403	919.4684	-0.0281		55 0.00035	1	U	K.LSEAELMK.K 604 609 610		
684	468.7319	935.4493	935.4633	-0.0141	0 (	52) 0.00064	1	U	K.LSEAELMK.K 682 683		
801	482.7584	963.5022	963.5237	-0.0215	0	35 0.037	1	U	R.TVSTGTALSK.Y		
				-0.0109	0	52 0.0006	1	U			
<u>996</u>	497.7584	993.5022	993.5131	-0.0109	•	52 0.0000		•	K.QELVYTNK.K		
<u>996</u> 1312		993.5022 1025.5090		-0.0091		38 0.017	1	U	K.QELVYTNK.K R.QFFEIQSK.E <u>1292</u> <u>1298</u> <u>1306</u> <u>1307</u> <u>1310</u> <u>1311</u>		
	513.7618	1025.5090	1025.5182		0						
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1312 2548 2595 2733 2748	513.7618 590.3232 396.5168 601.3069 601.7944	1025.5090 1178.6317 1186.5286 1200.5993 1201.5742	1025.5182 1178.6407 1186.5652 1200.5986 1201.5826	-0.0091 -0.0090 -0.0366 0.0007 -0.0084	0 0 0 0 0	38 0.017   35 0.043   38 0.019   77 2.6e-006   47) 0.0026	1 1 1 1 1	U U U U	R.OFFEIQSK.E 1292 1298 1306 1307 1310 1311 K.LLQEHNNALK.T R.LWALSHENEK.L K.NSATPSELNEK.Q 2723 2724 2725 2727 2730 2734 2736 2737 2738 274 K.NSATPSELNEK.Q 2750		
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Figure S1: Confirmation of purified Optineurin by peptide mass fingerprinting followed by database search.

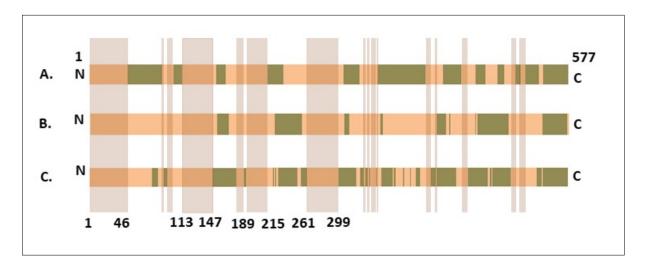


Figure S2: Schematic diagram of intrinsically disordered regions of OPTN; Disordered regions (peach) and ordered regions (olive) were predicted by online software A.PONDR, B.PONDR-FIT and C.IUPred2A. The shaded portions show the overlapping stretches of disordered regions by all three softwares.

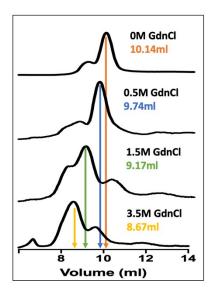


Figure S3: Size exclusion chromatography of Ni-NTA purified OPTN in presence of various concentrations of Guanidine hydrochloride

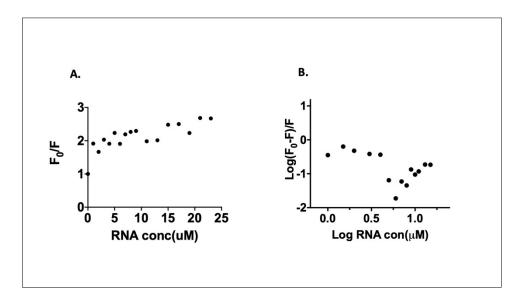


Figure S4: Fluorescence quenching of OPTN at 60 °C in the presence RNA; (A) Stern-Volmer plot of OPTN in the presence of various concentrations of RNA at 60 °C. (B) The plot of  $\log(F_0 - F)/F$  vs  $\log[Q]$  of OPTN at 60 °C as a function of RNA concentration.

OPTN HS71A HS90B CRYAA CRYBA1 CRYGC consensus>50	MAKAAAIGIDLGTTYSCVGVFQHGKVEIIANDQGNRTTPSYV MPEEVHHGEEEVETFAFQAEIAQLMSLIINTFYSNKEIFLRELISNASDALDKIRYE	
OPTN HS71A HS90B CRYAA CRYBA1 CRYGC consensus>50	AFTDTERL.IGDAAKNQVALNPQN.TVFDAKRLIGRKFGDPVVQ.SDMKHWPFQVINDGD SLTDPSKLDSGKELKIDIIPNPQERTLTLVDTGIGMTKADLINNLGTIAKSGTKAFMEAL	
OPTN HS71A HS90B CRYAA CRYBA1 CRYGC consensus>50	1 10 20 MSHQPLSCLTEKEDSPSESTGN KPKVQVSYKGETKA.FYPEEISSMVLTKMKEIAEAYLGYPVTNAVITVPAYFNDSORQ QAGADISMIGQFGVGFYSAYLVAEKVVVITKHNDDEQYAWESSAGGSFTVRADHG PIGR	
OPTN HS71A HS90B CRYAA CRYBA1 CRYBA1 CRYGC consensus>50	30 40 50 60 70 GPPHLAH. PNLDTFTPEELLQQMKELLTE. NHQLKEAMKLNNQAMKGRFEELSAWTEK ATK DAGVIAGLNVLRIINEPTAAAIAYGLDRIGKGERNVLIFDLGGGTFDVSI GTKVILHLKEDQTEYLEERRVKEVVKKHSQFIGYPITLYLEKEREKEISD.DEAEEEKGE  d	
OPTN HS71A HS90B CRYAA CRYBA1 CRYGC consensus>50	80 90 100 110 120 130 QKE RQFFEIQSKEAKERLMALSHENEKLKEELGKLKGKSERSSEDPTDDSRLPRAEAE LTIDDGIFEVKATAGDTHLGGEDFDNRLVNHFVEEFKRKHKKDISQNKRAVRRLATACE KEE OKDDEEKPKIEDVGSDEEDDSGKDKKKKKKKIKEKYIDQEELNKTKPIWTRNPDD 	I
OPTN HS71A HS90B CRYAA CRYBA1 CRYGC consensus>50	140 150 160 170 180 190 EKDQLRTQVVRLQAEKADLLGIVSELQLKLNSSGSSEDSFVEIFMAEGEABGSVKEIKH AKRILSSSTQASLEIDSLFEGL.DFYTSITRARFEELCSDLFFSTLEPVCKALRDAKL TQEEYGEFYKSLTNDWEDHLAV.KHFSVEGQLEFRALLFIPRFAPFDLFENKKKKNNI	SDK · · ·
OPTN HS71A HS90B CRYAA CRYBA1 CRYGC consensus>50	200 210 220 230 240 250 PGPTRTVSTGTALSKYRSRSADGAKNYFEHED TVSQLLLCLREGNQKVERLEVALKEAN KAQIHDLVLVGG.STRIPKVQKLLQDFFNGRO-NKSINPDEAVAYGAAVQAN LYVRRVFIMDSC.DELIPEYLNFIRGVVDSEDLPLNISREMLQQSKILKVIRKNIVKKC 	
OPTN HS71A HS90B CRYAA CRYBA1 CRYGC consensus>50	260 270 280 290 300 310 ERVSDFEKKTSDRSEIETQTEGSTEKENDEEKGPETVGSEVEALNLQV.ISLFKELQEA ILMG XSENVQLLLLDVAPLSLGLETAGGVMTALIKRNSTIPTKQTQIFTTY ELFSELAEDKENYKKFYEAFSKNIKLGIHEDSINRRLSELLRYHTSQSGDEMISLSEY 	s v w

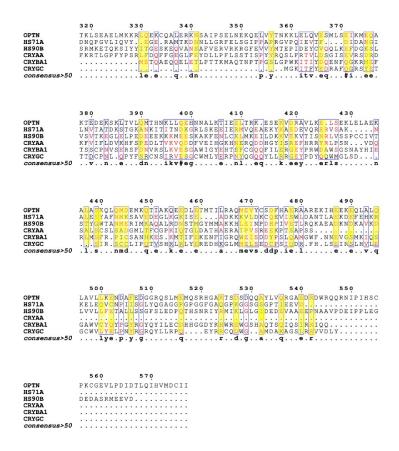


Figure S5: Multiple sequence alignment of human OPTN with HSPs (A) Optineurin (OPTN), (B) heat shock protein 70 (HS71A), (C) heat shock protein 90 (HS90B), (D)  $\alpha$ -crystallin (CRYAA), (E)  $\beta$ -crystallin (CRYBA1) and (F)  $\gamma$ -crystallin (CRYGC). The redcoloured letters depict the similar amino acid residues at the aligned position, while the yellow highlighted portion shows OPTN-specific similarity. The red line spanning the N terminal of OPTN represents the IDR residues in OPTN sequence.

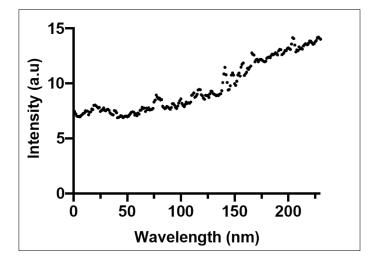


Figure S6: Light scattering profile of only OPTN (3mM) heated at 65 °C.