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Supplemental information

**Employing non-targeted interactomics approach
and subcellular fractionation to increase our
understanding of the ghost proteome**

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Supplemental Figures

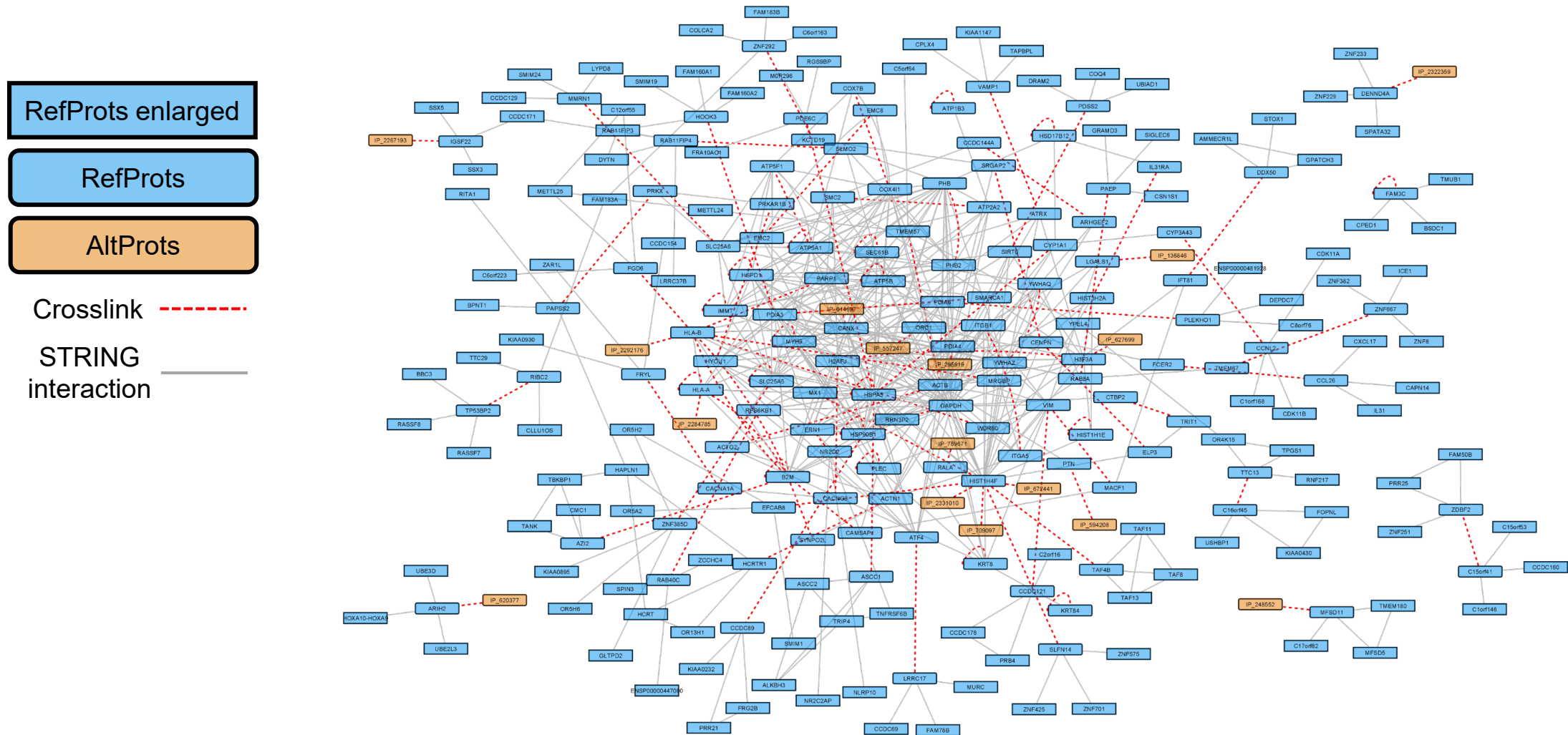
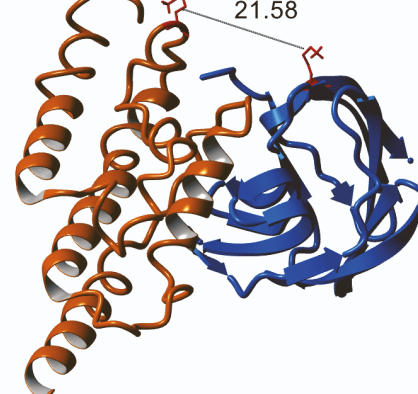
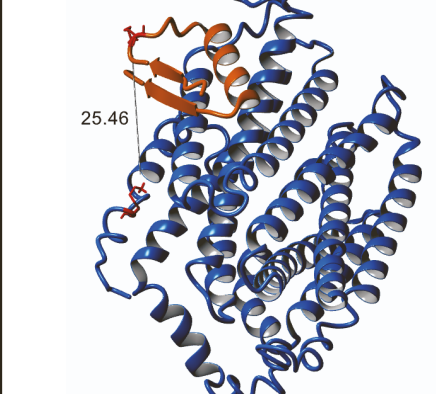
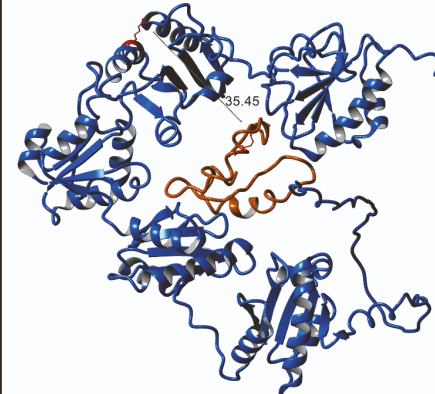
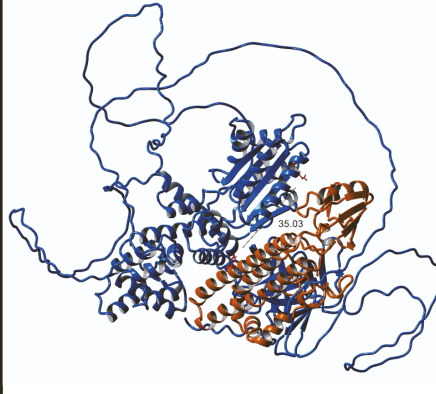
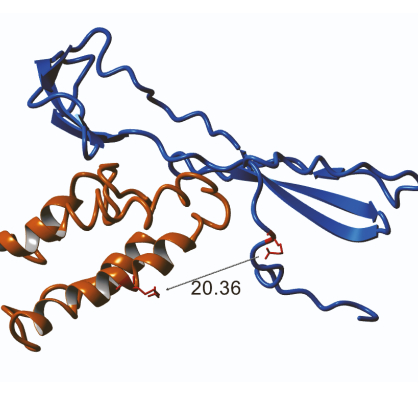
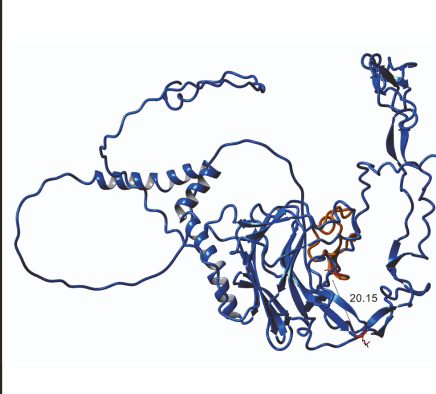
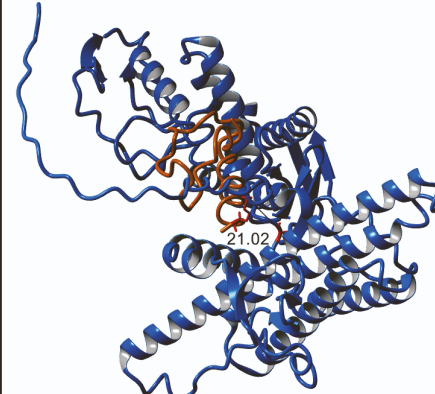
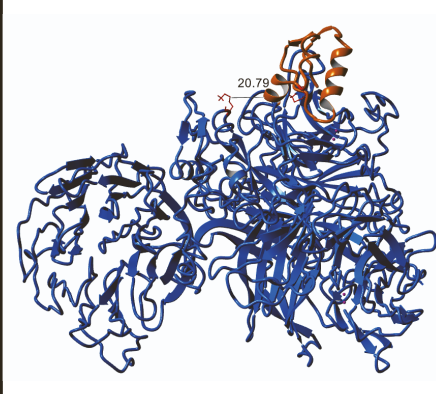
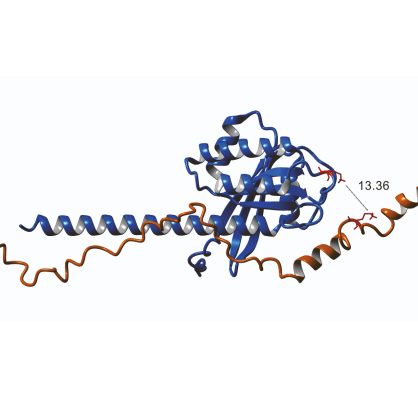
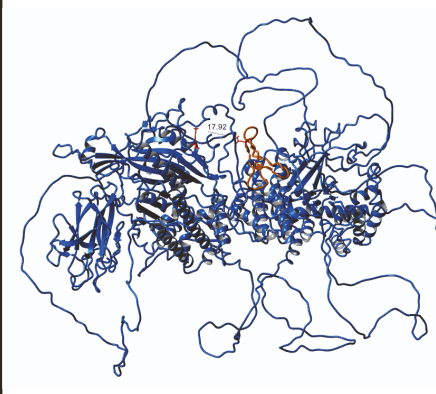
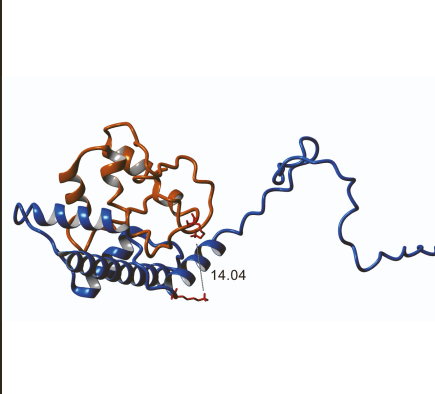
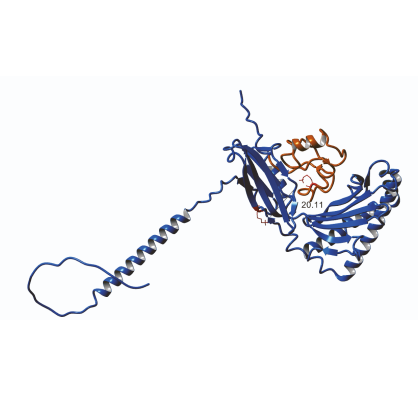
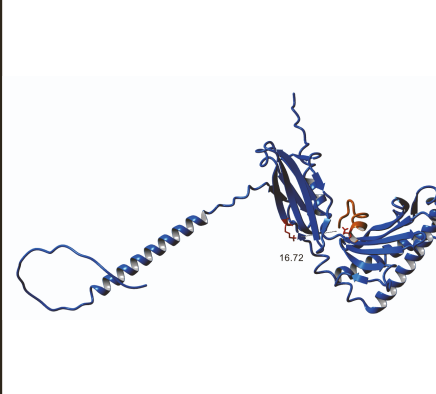
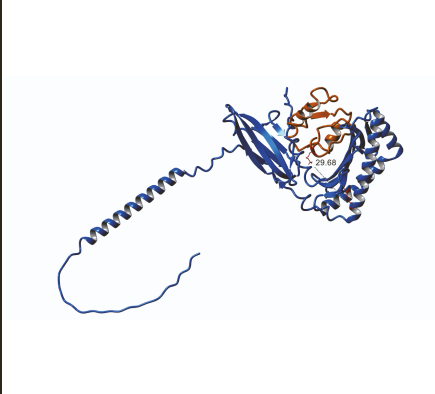
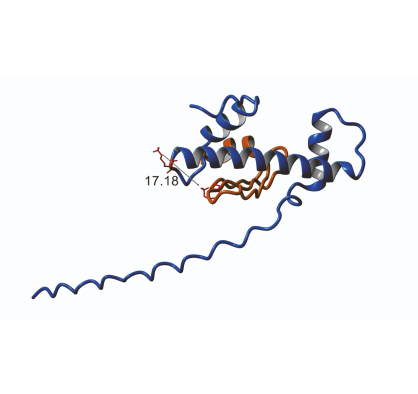
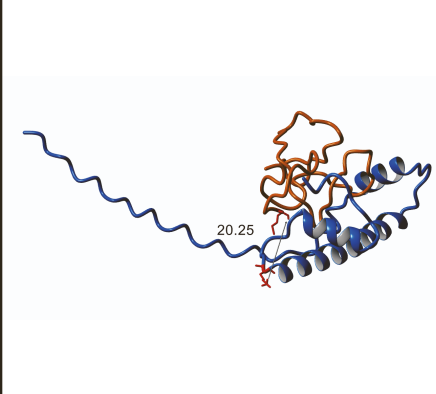
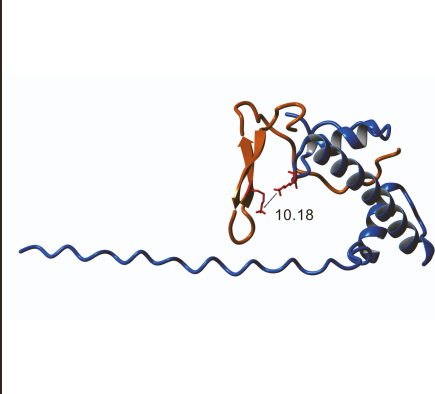
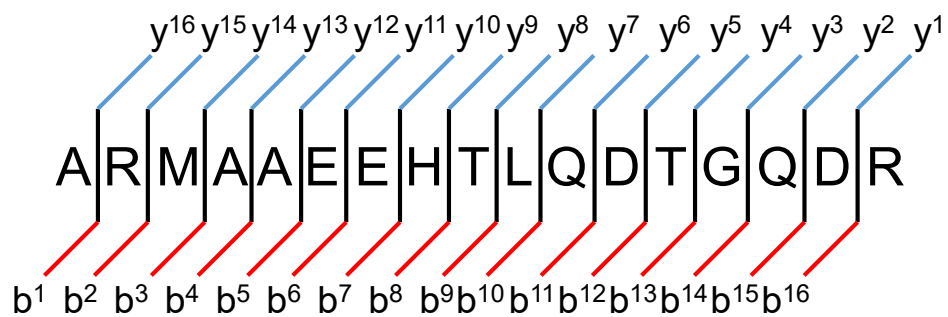


Figure S1. Crosslinked network enriched by the STRING interactions (gray lines) retrieved between these crosslinked (red dash lines) RefProts. Related to Figure 5. For the RefProts that did not present referenced STRING interaction, an enrichment has been performed to expand the network.

			
IP_136846-LGALS1 crosslink position = 21.58Å center energy = -487.3	IP_248552-MFSD11 crosslink position = 25.46Å center energy = -908.5	IP_295919-PDIA4 crosslink position = 35.45Å center energy = -825.2	IP_557247-ORC1 crosslink position = 35.03Å center energy = -951.6
			
IP_594208-PTN crosslink position = 20.36Å center energy = -569.2	IP_614697-CANX crosslink position = 20.15Å center energy = -664.4	IP_620377-ARIH2 crosslink position = 21.02Å center energy = -622	IP_2267193-IGSF22 crosslink position = 20.79Å center energy = -858.1
			<p>Figure S2. Predicted interaction models docked in ClusPro for the RefProts (blue) and AltProts (orange). Related to Figures 6 and 7. The distance between the residues crosslinked is displayed for each interaction.</p>
IP_789671-RALA crosslink position = 13.36Å center energy = -732.2	IP_2322359-DENND4A crosslink position = 17.92Å center energy = -977.7	IP_627699-H3F3A crosslink position = 14.04Å center energy = -860.9	
			
IP_2292176-HLA-B crosslink position = 20.11Å center energy = -847.7	[PEP]IP_2292176-HLA-B crosslink position = 16.72Å center energy = -587.1	IP_2284785-HLA-A crosslink position = 29.68Å center energy = -761.2	
			
IP_2331010-H4C1 crosslink position = 17.18Å center energy = -914.1	IP_709097-H4C1 crosslink position = 20.25Å center energy = -629.5	IP_672441-H4C1 crosslink position = 10.18Å center energy = -858.8	

IP_756980

MEKQLLEELE RQRQAELAAQ KARERKLARM
AAEEHTLQDT GQDRGRTCKT



Protein BLAST: No significant similarity was found
NextProt Peptide uniqueness checker: Peptide not found

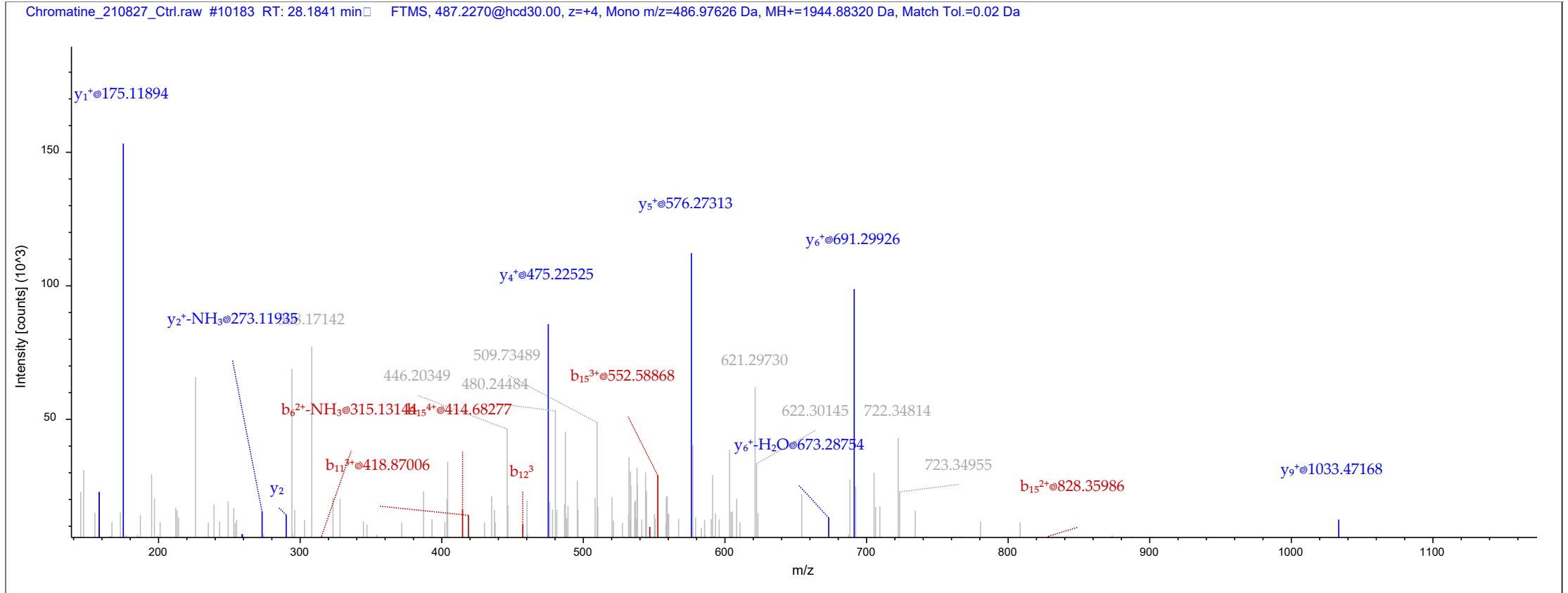


Figure S3. Tandem mass spectrum for the peptide ARMAAEEHTLQDT GQDRGRTCKT which was unique for the AltProt IP_756980. Related to STAR Methods. In red (b ions) and in blue (y ions) are observed in the spectra. No significant similarity was found for this AltProt in protein BLAST. Additionally, the peptide was not identified at NextProt Peptide uniqueness checker.

(α) IP_295919 – (β) PDIA4

Max. XlinkX Score: 46.35

CSM:1

NextProt Peptide uniqueness checker: Peptide not found

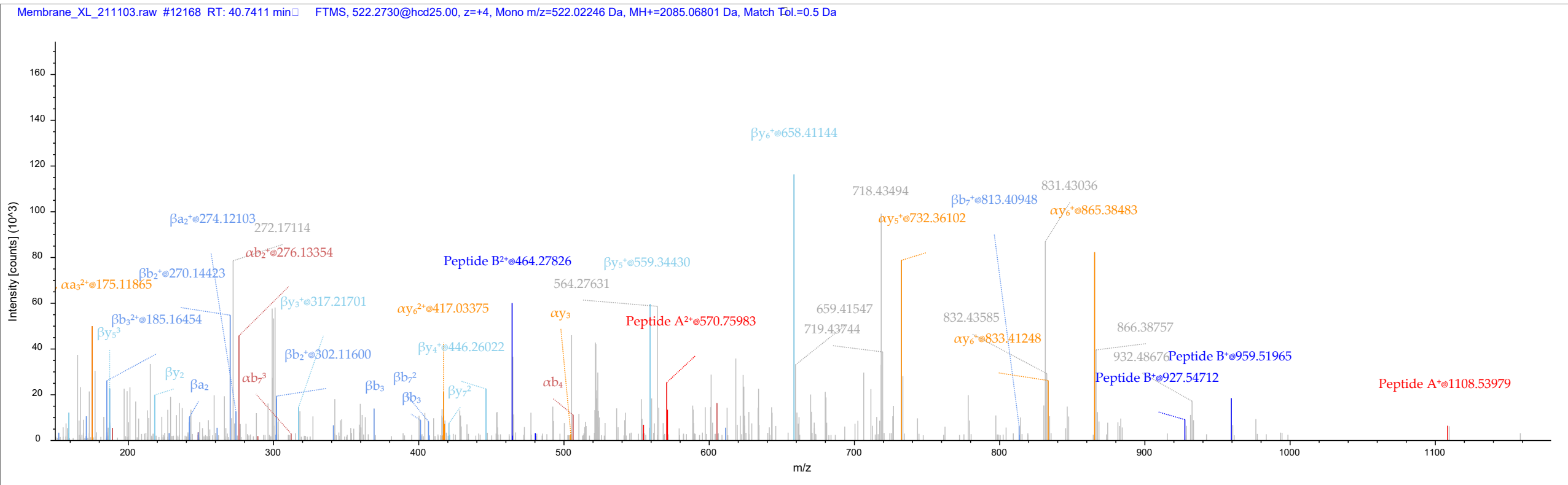
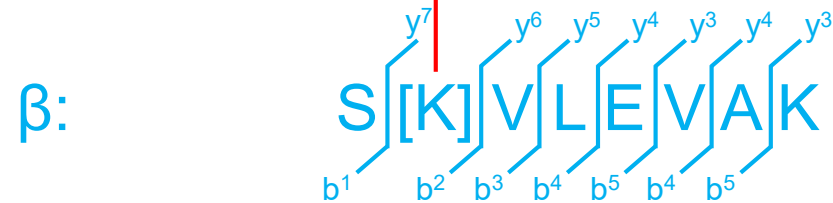
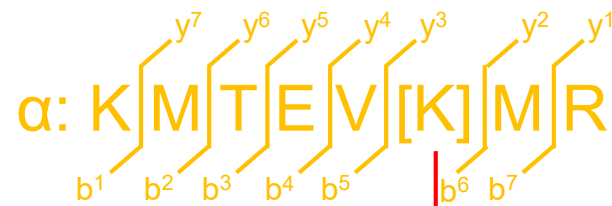


Figure S4. Crosslinking tandem mass spectrum which identified the interaction between IP_295919 (orange) and PDIA4 (blue). Related to STAR Methods. The b and y ions for the individual and crosslinking peptides are observed. The unique peptide of IP_295919 was not identified at NextProt.

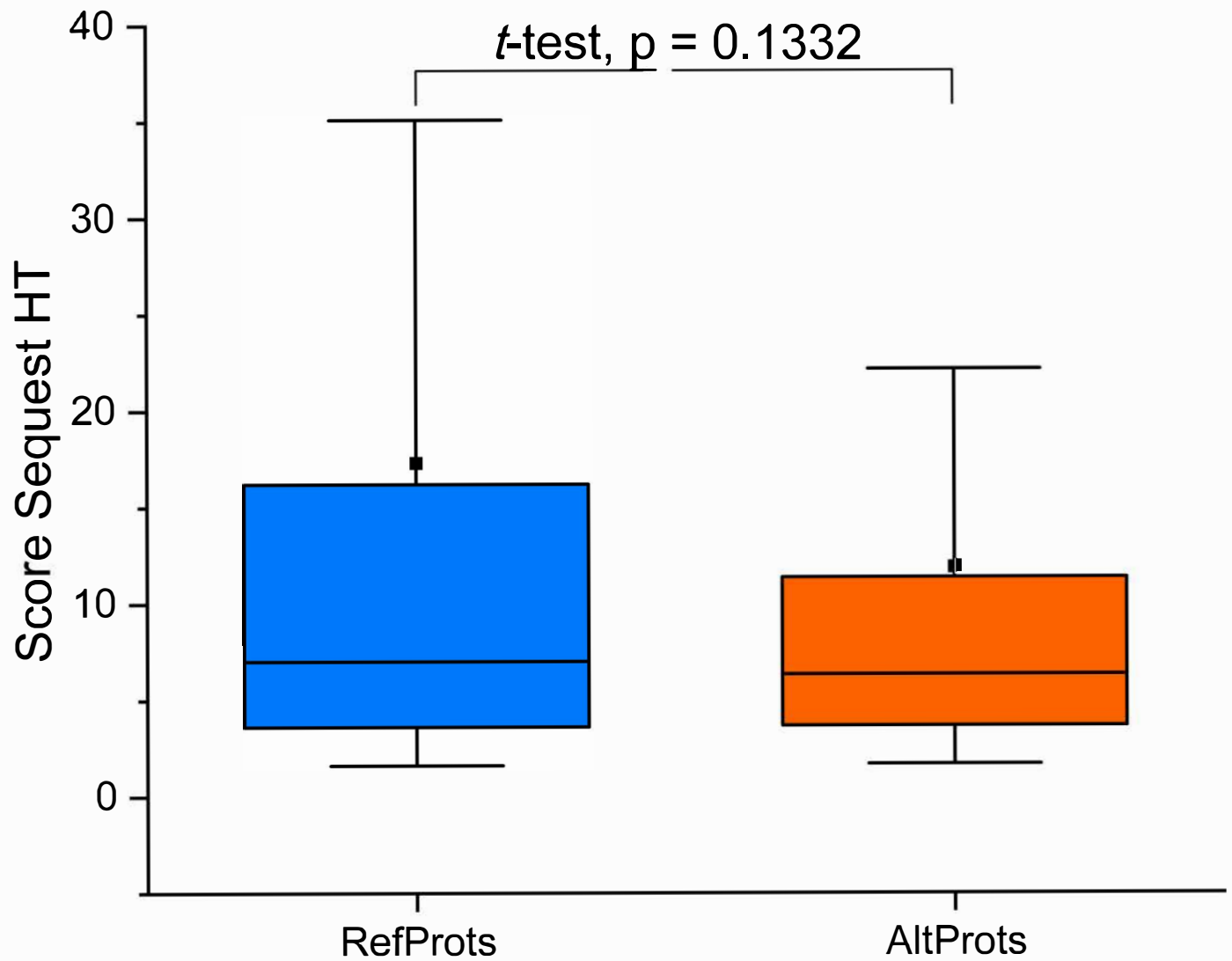


Figure S5. Boxplot of the Sequest HT Scores from the RefProts (blue) and AltProts (orange) identified using at least one peptide and OpenProt from the nuclear fraction. Related to STAR Methods. The Sequest HT Scores are not significantly different between RefProts (n=5764) and AltProts (n=151) (t-test, average Sequest HT Score 17.29 vs 11.91, $p = 0.1332$).