

Supplementary Note. Intra-domain and inter-domain linkages in TcdB identified by XL-MS.

Intra-Domain Linkages (60)						
Subunit a	Subunit b	Position a	Position b	Sum Ids	Bio Rep IDs	Distance(Å)
TcdB-CPD	TcdB-CPD	K572	K601	26	3*	8.1
TcdB-CPD	TcdB-CPD	K572	K646	16	3	22.2
TcdB-CPD	TcdB-CPD	K572	K648	18	3	20.7
TcdB-CPD	TcdB-CPD	K572	K765	16	3*	20.4
TcdB-CPD	TcdB-CPD	K572	K771	11	3	23.3
TcdB-CPD	TcdB-CPD	K601	K646	14	3*	15.6
TcdB-CPD	TcdB-CPD	K601	K648	14	3*	12.9
TcdB-CPD	TcdB-CPD	K601	K771	23	3*	20.8
TcdB-CPD	TcdB-CPD	K601	K776	17	3*	18.1
TcdB-CPD	TcdB-CPD	K646	K771	7	3	26.4
TcdB-CPD	TcdB-CPD	K646	K776	16	3*	18.7
TcdB-CPD	TcdB-CPD	K646	K784	11	3	24.9
TcdB-CPD	TcdB-CPD	K646	K787	22	3	24.9
TcdB-CPD	TcdB-CPD	K646	K791	78	3*	15.8
TcdB-CPD	TcdB-CPD	K646	K793	12	3*	16.7
TcdB-CPD	TcdB-CPD	K648	K771	12	3*	22.3
TcdB-CPD	TcdB-CPD	K648	K791	12	3*	15.1
TcdB-CPD	TcdB-CPD	K765	K776	34	3*	17.4
TcdB-CPD	TcdB-CPD	K771	K784	8	3	32.7
TcdB-CPD	TcdB-CPD	K771	K787	7	3	26.2
TcdB-CPD	TcdB-CPD	K771	K791	7	3	18.5
TcdB-CPD	TcdB-CPD	K771	K793	8	3	13.6
TcdB-CPD	TcdB-CPD	K776	K787	7	3	19.1
TcdB-CPD	TcdB-CPD	K776	K791	8	3	10.7
TcdB-CPD	TcdB-CPD	K776	K793	22	3*	7.6
TcdB-CPD	TcdB-CPD	K784	K791	12	3	16.2
TcdB-CROPS	TcdB-CROPS	K1977	K1997	9	3*	14
TcdB-CROPS	TcdB-CROPS	K2230	K2234	5	3	12
TcdB-CROPS	TcdB-CROPS	K2230	K2249	11	3	14.3
TcdB-CROPS	TcdB-CROPS	K2234	K2249	20	3*	6.8
TcdB-GTD	TcdB-GTD	K11	K73	19	3	15.1
TcdB-GTD	TcdB-GTD	K11	K74	19	3	13.5
TcdB-GTD	TcdB-GTD	K142	K536	25	3*	20.9
TcdB-GTD	TcdB-GTD	K143	K216	288	3*	12.6
TcdB-GTD	TcdB-GTD	K143	K517	21	3*	23.9
TcdB-GTD	TcdB-GTD	K143	K536	53	3*	18
TcdB-GTD	TcdB-GTD	K143	K537	13	3	19.1
TcdB-GTD	TcdB-GTD	K194	K247	97	3*	17.3
TcdB-GTD	TcdB-GTD	K299	K506	6	3*	16.8
TcdB-GTD	TcdB-GTD	K303	K380	22	3*	21.4
TcdB-GTD	TcdB-GTD	K303	K506	38	3*	20.5
TcdB-GTD	TcdB-GTD	K334	K517	5	3*	13
TcdB-GTD	TcdB-GTD	K380	K506	22	3*	11.3
TcdB-GTD	TcdB-GTD	K50	K402	70	3*	17.2
TcdB-GTD	TcdB-GTD	K7	K70	8	3	17.4
TcdB-GTD	TcdB-GTD	K7	K73	8	3	17.5
TcdB-GTD	TcdB-GTD	K7	K74	8	3	16.2
TcdB-DRBD	TcdB-DRBD	K1126	K1250	31	3*	8.9
TcdB-DRBD	TcdB-DRBD	K1427	K1459	18	3*	10.3
TcdB-DRBD	TcdB-DRBD	K1459	K1522	12	3*	12.3
TcdB-DRBD	TcdB-DRBD	K1471	K1518	28	3*	11.6
TcdB-DRBD	TcdB-DRBD	K1481	K1504	12	3*	12.1
TcdB-DRBD	TcdB-DRBD	K1504	K1549	121	3*	12.5
TcdB-DRBD	TcdB-DRBD	K1504	K1552	9	3	17.3
TcdB-DRBD	TcdB-DRBD	K1522	K1552	37	3	13.9
TcdB-DRBD	TcdB-DRBD	K1534	K1568	49	3*	15.4
TcdB-DRBD	TcdB-DRBD	K1773	K1780	18	3	13.3
TcdB-DRBD	TcdB-DRBD	K846	K1765	21	3*	16
TcdB-DRBD	TcdB-DRBD	K846	K1773	40	3*	15.5
TcdB-DRBD	TcdB-DRBD	K857	K1773	26	3	15

Inter-Domain Linkages (27)						
TcdB-CPD	TcdB-DRBD	K572	K1760	104	3*	28.3
TcdB-CPD	TcdB-DRBD	K646	K1775	7	3	29.1
TcdB-CPD	TcdB-DRBD	K646	K846	52	3*	11
TcdB-CPD	TcdB-DRBD	K784	K1773	8	3	21.2
TcdB-CPD	TcdB-DRBD	K784	K846	103	3	24.4
TcdB-CPD	TcdB-DRBD	K787	K846	25	3*	24.7
TcdB-CPD	TcdB-DRBD	K791	K1773	9	3	14.3
TcdB-CPD	TcdB-DRBD	K791	K846	41	3*	13.7
TcdB-GTD	TcdB-CPD	K303	K784	13	3	14.5
TcdB-GTD	TcdB-CPD	K303	K787	11	3	16.4
TcdB-GTD	TcdB-CPD	K380	K784	18	3*	13.8
TcdB-GTD	TcdB-CPD	K380	K787	17	3*	7.2
TcdB-GTD	TcdB-CPD	K506	K784	76	3*	17.3
TcdB-GTD	TcdB-CPD	K506	K787	108	3*	14.7
TcdB-GTD	TcdB-CPD	K529	K572	4	3	31.8
TcdB-GTD	TcdB-DRBD	K536	K765	7	3*	20.1
TcdB-GTD	TcdB-DRBD	K11	K904	288	3*	15.1
TcdB-GTD	TcdB-DRBD	K7	K904	48	3	20
TcdB-GTD	TcdB-DRBD	K70	K857	12	3	22.4
TcdB-GTD	TcdB-DRBD	NT	K916	24	3*	17.7
Novel Set						
						Distance (Å)
TcdB-DRBD	TcdB-CROPS	K1117	K2249	25	3	209.2
TcdB-DRBD	TcdB-CROPS	K1120	K2234	14	3	203.8
TcdB-DRBD	TcdB-CROPS	K1120	K2249	203	3*	207.2
TcdB-DRBD	TcdB-CROPS	K1126	K2249	72	3*	204.2
TcdB-DRBD	TcdB-CROPS	K1504	K1977	15	3*	117.1
TcdB-DRBD	TcdB-CROPS	K1549	K1965	141	3*	96.4
TcdB-DRBD	TcdB-CROPS	K1549	K1977	41	3*	108

* The stars indicate the cross-linked peptides that were confirmed by in-gel digestion LC-MSⁿ analysis.