

Supplement Table S1. Comparison of the structural characteristics of SARS-CoV-2 wild and variants and the prediction of the effect of ASPNJ on S protein

		1	2	3	4	5	6	7	
WHO PANGO	Wild wild	Alpha B.1.1.7	Beta B.1.351	Gamma P.1	Delta B.1.617.2	Omicron BA.1	Lambda C.37	Mu B.1.621	
S	aa mutations of S protein	del69/70(HV), del144(Y), N501Y, A570D, D614G, P681H, T716I, S982A, D1118H	D80A, D215G, K417N, E484K, N501Y, D614G, A701V	L18F, T20N, P26S, D138Y, R190S, K417T, E484K, N501Y, D614G, H655Y, T1027I, V1176F	T19R, E156G, del157/158 (FR), L452R, T478K, D614G, P681R, D950N	A67V,del69/70(HV),T95I,G142D, del143/145(VYY),N211I,del212/212(L), ins214(EPE),G339D,S371L,S373P,S375F, Q493R,G496S,Q498R,N501Y,Y505H,T547K ,D614G,H655Y,N679K,P681H,N764K, D796Y,N856K,Q954H, N969K,L981F	G75V, T76I, R246N, del247/253 , (SYLTPGD) L452Q, F490S, D614G, T859N,	T95I, Y145N, R346K, E484K, N501Y, D614G, P681H, D950N	
	aa length/pI	1261/6.23	1258/6.35	1261/6.63	1261/6.38	1259/6.93	1258/6.79	1254/6.32	1261/6.66
	No. of K/R	61/42	61/42	61/42	61/41	62/44	66/44	61/41	63/41
	No. of CS/ No. of NCS	101/2	101/2	101/2	100/2	102/4	107/3	100/2	102/2
	PNCS	462(*450) KP , 811(*799) KP	462(*447) KP , 811(*796) KP	462(*450) KP , 811(*799) KP	462(*450) KP , 811(*799) KP	462(*448) KP , 478(*464) KP , 682(*668) RR 811(*797) KP	462(*447) KP , 498(*483) RP , 811(*796) KP	462(*443) KP , 811(*792) KP	462(*450) KP , 811(*799) KP
S1	aa length/pI	673/8.24	670 / 8.24	673/8.59	673/8.43	671/8.65	670/8.35	666/8.35	673/8.51
	No. of K/R	30/29	30/29	30/29	30/28	31/31	32/31	30/28	32/28
	No. of CS/ No. of NCS	58/1	58/1	58/1	57/1	59/2	61/2	57/1	59/1
	C terminal	669PRRAR	666HRRAR	669PRRAR	669PRRAR	667RRRAR	666HRRAR	662PRRAR	669HRRAR
RBD	aa length/pI	223/8.91	223/8.90	223/9.00	223/9.00	223/9.09	223/9.01	223/8.91	223/9.07
	No. of K/R	12/11	12/11	12/11	12/11	13/12	12/13	12/11	14/10
	No. of CS/ No. of NCS	22/1	22/1	22/1	22/1	23/2	23/2	22/1	23/1
S2	aa length/pI	588/5.25	588/5.39	588/5.25	588/5.25	588/5.32	588/5.58	588/5.25	588/5.32
	No. of K/R	31/13	31/13	31/13	31/13	31/13	34/13	31/13	31/13
	No. of CS/ No. of NCS	43/1	43/1	43/1	43/1	43/1	46/1	43/1	43/1

Columns 1-5 are Variants of Concern (VOC), and columns 6,7 are Variants of Interest (VOI) according to the WHO (World Health Organization) and PANGO (Phylogenetic assignment of the named global outbreak). S: Spike glycoprotein, S1: S1 protein, RBD: receptor binding domain. Amino acids (aa) in bold letters such as 462**KP** and 682**RR** indicate that they cannot be hydrolyzed by trypsin. No. of CS: theoretical No. of cleavage sites (CS) by trypsin; No. of NCS: theoretical No. of non-cleavage sites (NCS) by trypsin. PNCS: Trypsin's

theoretical position of NCS in S, S1 and RBD. *: The position of NCS in the mature sequence of S, S1 and RBD without signal peptide. No. of K/R: numbers of Lysine / number of Arginine. The c-terminal sequence RRAR of S1 protein, called CendR, can be hydrolyzed by trypsin, causing the c-terminal of S1 protein to fail to bind and activate neuropilin receptors (NRP1 and NRP2). The aa sequence data of S, S1 and RBD were quoted from <https://www.uniprot.org/>, <https://cov-lineages.org> and <https://outbreak.inf>. The pI, No. of CS and NCS by trypsin were calculated using https://web.expasy.org/compute_pi/ and https://web.expasy.org/peptide_cutter/. Trypsin preferentially cleaves the carbonyl side of arginine or lysine. Since this catalytic specificity is also shared by ASPNJ, this table predicts CS etc. using trypsin.