Supplementary data. Sequence alignments used for homology modelling. Conserved residues are labelled in dark green, same residues in lighter green and residues with similar properties in yellow.

SARS-CoV-2 S protein – close state

Sequence identity: 66%

SARS_CoV_2 6VXX_A 5X58_B	14 27 18	QCVNLTTRTQLPPAYTNSFTRGVYYPDKVFRSSVLHSTQDLFLPFFSNVTWFHAIHVSGTNGTKRFDNFVLPFNDGVYFASTEKSNIIRGNIFGTTLDSKTQSLLIVNNATNV
SARS_CoV_2 6VXX_A 5X58_B	127 127 124	VIKVCEFQFCNDPFLGVYYHKNNKSWMESEFRVYSSANNCTFEYVSQPFLMDLEGKQGNFKNLREFVFKNIDGYFKIYSKHTFINLVRDLPQGFSALEFLVDLFIGINITROT VIKVCEFQFCNDPFLGV
SARS_CoV_2	241	LLALHRSYLTPGDSSSGWTAGAAAYYVGYLOPRTFLLKYNENGTITDAVDCALDPLSETKCTLKSFTVEKGIYOTSNFRVOPTESIVRFPNITNLCPFGEVFNATRFASVYAWN
6VXX_A	241	LLALHAAYYVGYLOPRTFLLKYNENGTITDAVDCALDPLSETKCTLKSFTVEKGIYOTSNFRVOPTESIVRFPNITNLCPFGEVFNATRFASVYAWN
5X58_B	234	ILTAFSTWGTSAAAYFVGYLKPTTFMLKYDENGTITDAVDCSONPLAELKCSVKSFEIDKGIYOTSNFRVVPSGDVVRFPNITNLCPFGEVFNATKFPSVYAWE
SARS_CoV_2	355	RKRISNCVADYSVLYNSASFSTFKCYGVSPTKLNDLCFTNVYADSFVIRGDEVRQIAFGQTGKIADYNYKLPDDFTGGVIAWNSNNLDSKVGGNYNYLYRLFRKSNLKPFERDI
6vxx_a	355	RKRISNCVADYSVLYNSASFSTFKCYGVSPTKLNDLCFTNVYADSFVIRGDEVRQIAFGQTGKIADYNYKLPDDFTGGVIAWNSNNLDSKGNYNYLYRKPFERDI
5x58_b	342	RKKISNCVADYSVLYNSTFFSTFKCYGVSATKLNDLCFSNVYADSFVYKGDDVRQIAFGQTGVIADYNYKLPDDFMGGVLAWNTRNIDATSTGNYNYKYRYLRHGKLRPFERDI
SARS_CoV_2 6VXX_A 5X58_B	469 489 456	STEIYQAGSTPCNGVEGFNCYFPLQSYGFQPTNGVGYQPYRVVVLSFELLHAFATVCGFKKSTNLVKNKCVNFNFNGLTGTGVLTESNKKFLPFQQFGRDIADTTDAVRDFQT
SARS_CoV_2 6VXX_A 5X58_B	583 583 569	EILDITPCSFGGVSVITEGTNTŠNOVAVLYODVNCTEVPVAIHADQLTPTWRVYSTGSNVFQTRAGCLIGAEHVNNSYECDIPIGAGICASYQTQTNSPRRARSVASQŠIIAYT EILDITPCSFGGVSVITEGTNTŠNOVAVLYODVNCTEV
SARS_CoV_2	697	MSLGAENSVAYSNNSIAIFTNFTISVTTEILPVSMTKTSVDCTMYICGDSTECSNLLLQYGSFCTQLNRALTGIAVEQDKNTQEVFAQVKQIXKTPPIKDFGGFNFSQILPDPS
6VXX_A	697	MSLGAENSVAYSNNSIAIPTNFTISVTTEILPVSMTKTSVDCTMYICGDSTECSNLLLQYGSFCTQLNRALTGIAVEQDKNTQEVFAQVKQIXKTPPIKDFGGFNFSQILPDPS
5X58_B	679	MSLGADSSIAYSNNTIAIPTNFSISITTEVMPVSMAKTSVDCNMYICGDSTECANLLLQYGSFCTQLNRALSGIAAEQDRNTREVFAQVKQMYKTPTLKYFGGFNFSQILPDPL
SARS_CoV_2 6VXX_A 5X58_B	811 811 793	KESKRSFIEDLLENKVTLADAGFIKQYGDCLGDIAARDLICAQKENGLTVLPPLLTDEMIAQYTSALLAGTITSGWTEGAGAALQIPFAMQMAYRENGIGVTQNVLYENQKLIA KESKRSFIEDLLENKVT
SARS_COV_2	925	NQENSAIGKIQDSLSSTASALGKLQDVVNQNAQALNTLVKQLSSNEGAISSVLNDILSRLDKVEAEVQIDRLITGRLQSLQTYVTQQLIRAAEIRASANLAATKMSECVLGQSK
6VXX_A	925	NQENSAIGKIQDSLSSTASALGKLQDVVNQNAQALNTLVKQLSSNEGAISSVLNDILSRLDEPEAEVQIDRLITGRLQSLQTYVTQQLIRAAEIRASANLAATKMSECVLGQSK
5X58_B	907	NQENKAISQIQESLTTTSTALGKLQDVVNQNAQALNTLVKQLSSNEGAISSVLNDILSRLDKVEAEVQIDRLITGRLQSLQTYVTQQLIRAAEIRASANLAATKMSECVLGQSK
SARS_COV_2	1039	RVDFCGKGYHIMSFPQSAPHGVVFLHVTYVPAQEKNFTTAPAICHDGKAHFPREGVFVSNGTHWFVTQRNFYEPQIITTDNTFVSGNCDVVIGIVNNTVYDPLQPELDS
6VXX_A	1039	RVDFCGKGYHIMSFPQSAPHGVVFLHVTYVPAQEKNFTTAPAICHDGKAHFPREGVFVSNGTHWFVTQRNFYEPQIITTDNTFVSGNCDVVIGIVNNTVYDPLQPELDS
5X58_B	102	RVDFCGKGYHIMSFPQAAPHGVVFLHVTYVESCERNFTTAPAICHEGKAYFPREGVFVSNGTWFTSWIITORNFFSPQIITTDNTFV

SARS-CoV-2 S protein – open state

Sequence identity: 10%

In this case, the sequence identity is very low because the template structures used to model SARS-CoV-2 S protein in the open conformation with PDB ID 6LZG and 6M0J are the structures of the S protein RBD domain. Therefore, when the sequences of these structures are aligned with the sequences of the template structures with PDB ID 6VSB, 6VYB and 5X5B – which are the structures of an entire S protein monomer – the only region of the alignment in which the sequences of all the template structures align is in the section of the sequences corresponding the RBD.

SARS_COV_2 6VSB_A 6VYB_B 6LZG_B	27 27 27 333	AYTNSFTRGVYY PDKVFRSSVLHSTQDLFLPFFSNVTWFHAIHVSGTNGTKRFDNPULPFNDGVYFASTEKSNIIGAUTFGTTLDSKTQSLLIVNAINVVIKVCEFQFCNDPF AYTNSFTRGVYY PDKVFRSSVLHSTQDLFLPFFSNVTWFHFDNPULPFNDGVYFASTNIIGAUTFGTTLDSKTQSLLIVNAINVVIKVCEFQFCNDPF AYTNSFTRGVYY PDKVFRSSVLHSTQDLFLPFFSNVTWFHPDNPULPFNDGVYFASTEKSNIIGAUTFGTTLDSKTQSLLIVNAINVVIKVCEFQFCNDPF
5X5B_A	333 15	QHTSSM-RGVYYPDEIFRSDTLYLTQDLFLPFYSNVTGFHTINHTFDNPVIPFKDGIYFAATEKSNVVRGWVFGSTMNNKSQSVIIINNSTNVVIRACNFELCDNPF
SARS_CoV_2 6VSB_A	141 141	LGVYYHKNNKSWMESEFRVYSSANNCTFEYVSQPPLMDLEGKQGNFKNLREFVFKNIDGYFKIYSKHTPINLVRDLPQGFSALEPLVDLPIGINITRPQTLLALHRSYLTPGDSS LGERVYSSANNCTFEYVSQPFLKNLREFVFKNIDGYFKIYSKHTPINLVRDLPQGFSALEPLVDLPIGINITRFQTLLALH
6VYB_B	164	YFKIYSKHTPINDLPQGFSALEPLVDLPIGINITRFQTLLYFKIYSKHTPINDLPQGFSALEPLVDLPIGINITRFQTLL
6MOJ E	333	
5x5B_A	121	${\tt Fav}{\tt SkpmgtQthtmifdnafnctfeyisdafsldvseksgnfkhlrefvfknkdgflyvykgyQpidvvrdlpsgfntlkpifklplginitnfrailtafs}$
SARS_CoV_2	256	SCWTAGAAAYYVCYLQPRTFLLKYNENGTITDAVDCALDPLSETKCTLKSFTVEKGIYQTSNFRVQPTESIVRFPNITNLCPFGEVFNATRASVYAWNRKRISNCVADYSVLYN
6VYB B	263	
6LZG B	333	TNLCPFGEVFNATRFASVYAWNRKRISNCVADYSVLYN
6M0J_E	333	
5x5b_a	223	-TWGTSAAAYFYGYLKPTTFMLKYDENGTITDAVDCSQNPLAELKCSVKSFEIDKGIYQTSNFRVVPSGDVVRL <mark>CPFGEVFNATK</mark> PSVYAHERKKISNCVADYSVLYN
SARS_COV_2	371	SASFSTFKCYGVSPTKLNDLCFTNVYADSFVIRGDEVRQIAPGQTGKIADYNYKLPDDFTGCVIAWNSNNLDSKVGGNYNYLYRLFRKSNLKPFERDISTEIYQAGSTPCNGVEG
6VYB B	371	SASPSTFRCYGVSPTKINDLCFTNVYADSFVIRGDEVRGIAPQOTGRIADYNYKLPDDFTGCVIAWNSNNLDSKVGGNYNYLYRKPFER
6LZG_B	371	SASFSTFKCYGVSPTKLNDLCFTNVYADSFVIRGDEVRQIAPGOTGKIADYNYKLPDDFTGCVIAWNSNNLDSKVGGNYNYLYRLFRKSNLKPFERDISTEIYQAGSTPCNGVEG
6M0J_E 5X5B_A	371 331	SASFSTFKCYGVSPTKLNDLCTNVYADSFVIRGDEVRQIAFGQTGXIADYNYKLPDDFTGCVIANSNNLDSKVGGNYNYLYRLFRKSNLKPFERDISTEIYQAGSTFCNGVEG STFFSTFKCYGVSATKLNDLCFSNVYADSFVVKGDDVRQIAFGQTGVIADYNYKLPDDFMGCVLANNTRNIDATSTGNYNYKYRYLRHGKLRPFERDISNVPFSPDGKFCTP-PA
SARS_CoV_2	486	FNCYF PLOS YGF OPTNGVGY OPYRVVVLSFELHAPATVCGPKKSTNLVKNKCVNFNFNGLTGTGVLTESNKKFLPFQOFGRUIADTTDAVRDPOTLEILDITPCSFGGVSVITP
6VYB B	491	PLOSYGFOOTNGYGYOOYRVVVLSTATVCGPKKSTNLVKNKCVNFNFNGLTGTGVLTESNKKFLPFOOFGRDIADTTDAVRDPOTLEILDITFCSFGGVSVITP
6LZG B	486	FNCYFPLOSYGFOPTNGVGYOPYRVVVLSFELLHAPATVCGP
6M0J_E	486	FNCYFPLQSYGFQPTNGVGYQPYRVVVLSFELLHAPATVCG
5X5B_A	445	LNCYWPLNDYGFYTTTGIGYQPYRVVVLSFEDLIKNQCVNFNFNGLTGTGVLTPSSKRFQPFQQFGRDVSDFTDSVRDPKTSEILDISPCSFGGVSVITP
SARS_CoV_2	601	${\tt GTNTSNQVAVLYQDVNCTEVPVAIHADQLTPTWRVYSTGSNVFQTRAGCLIGAEHVNNSYECDIPIGAGICASYQTQTNSPRRARSVASQSIIAYTMSLGAENSVAYSNNSIAIPIGAENSVAYSNNSIAIPIGAENSVAYSNNSIAIPIGAENSVAYSNNSIAIPIGAENSVAYSNNSIAIPIGAENSVAYSNNSIAIPIGAENSVAYSNNSIAIPIGAENSVAYSNNSIAIPIGAENSVAYSNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIANPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSYNNSIAIPIGAENSVASYNNSIAIPIGAENSVASYNNSYNNSIAIPIGAENSVASYNNSYNNSYNNSYNNSYNNSYNT$
6VSB_A	601	GTNTSNQVAVLYQDVNCTEVSQSIIAYTMSLGAENSVAYSNNSIAIP
6VYB_B	601	GTNTSNEVAVLIQDVNCTEVSQSIIAYTMSLGAENSVAYSNNSIAIP
6M0J E		
5X5B_A	545	GTNASSEVAVLYQDVNCTDVSTAIHADQLTPAWRIYSTGNNVFQTQAGCLIGAEHVDTSYECDIPIGAGICASYHTKSIVAYTMSLGADSSIAYSNNTIAIP
SARS_CoV_2	716	$\label{eq:construct} TNFTISVTTEILPVSMTKTSVDCTMYICGDSTECSNLLLQYGSFCTQLNRALTGIAVEQDKNTQEVFAQVKQIYKTPPIKDFGGFNFSQILPDPSKPSKRSFIEDLLFNKVTLADFGFNFSQILPDPSKPSKRSFIEDLFNKVTLADFGFNFSQILPDPSKPSKRSFIEDLFNKVTLADFGFNFSQILPDPSKPSKRSFIEDLFNKVTLADFGFNFSQILPDSKPSKRSFIEDLFNKVTLADFGFNFSQILPDSKPSKRSFIEDLFNKVTLADFGFNFSQILPDSKPSKRSFIEDLFNKVTLADFGFNFSQILPDSKPSKRSFIEDLFNKVTLADFGFNFFF$
6VYB B	716	$\label{eq:construction} The Leventrisy double (gds) and (gds) an$
6LZG B	120	
6M0J_E		
5x5b_A	647	TNFSISITTEVMPVSMAKTSVDCNMYICGDSTECANLLLQYGSFCTQLNRALSGIAAEQDRNTREVFAQVKQMYKTPTLKYFGGFNFSQILPDPLKPTKRSFIEDLLFNKVT
SARS_COV_2	831	AGFIKQYGDCLGDIAARDLICAQKFNGLTVLPPLLTDEMIAQYTSALLAGTITSGWTFGAGAALQIPFAMQMAYRFNGIGVTQNVLYENQKLIANQFNSAIGKIQDSLSSTASAL
6VYB B	854	
6LZG B	004	
6M0J_E		
5X5B_A	759	DLICAQKFNGLTVLPPLLTDDMIAAYTAALVSGTATAGWTFGAGAALQIPFAMQMAYRFNGIGVTQNVLYENQKQIANQFNKAISQIQESLTTTSTAL
SARS_COV_2	946	GKLQDVVNQNAQALNTIVKQLSSNFGAISSVINDILSRLDKVEAEVQIDRLITGRLQSLQTYVTQQLIRAAEIRASANLAATKMSECVLGQSKRVDFCGKGYHLMSFPQSAPHGV
6VYB B	946	GALGUVVNUNAQALNTIVAULSSNEGATSSVLINDTISKLIPPEAEVULDKLITGKLUSLUTIVTUULIKAAELIKASALAATAMSEVILGUSKVUFUGAGIHLMSFEQASAHGV (skilonuvniadalnitvaulssnegatssvlindtisklippeaevuldkiltgkluslutivaulsenses)
6LZG_B	210	
6M0J_E		
5X5B_A	857	GKLQDVVNQNAQALNTLVKQLSSNFGAISSVLNDILSRLDKVEAEVQIDRLITGRLQSLQTYVTQQLIRAAEIRASANLAATKMSECVLGQSKRVDFCGKGYHLMSFPQAAPHGV
SARS_CoV_2 6VSB_A 6VYB_B 6UZG_B	106: 106: 106:	.VFLHVTYVPAQEKNFTTAPAICHDGKAHFPREGVFVSNGTHWFVTQRNFYEPQIITTDNTFVSGNCDVVIGIVNNTVYDPLQPELDS .VFLHVTYVPAQEKNFTTAPAICHDGKAHFPREGVFVSNGTHWFVTQRNFYEPQIITTDNTFVSGNCDVVIGIVNNTVYDPLQPELD - .VFLHVTYVPAQEKNFTTAPAICHDGKAHFPREGVFVSNGTHWFVTQRNFYEPQIITTDNTFVSGNCDVVIGIVNNTVYDPLQPELDS
6M0J E		
5X5B_A	972	VFLHVTYVPSQERNFTTAPAICHEGKAYFPREGVFVFNGTSWFITQRNFFSPQIITTDNTFV

SARS-CoV S protein – close state

Sequence identity: 100%



SARS-CoV S protein - open state

Sequence identity: 100%

