

## **Supplemental Information**

### **The S1/S2 site containing PRRA affects cellular tropism of SARS-CoV-2 and ACE2 usage by the closely related Bat RaTG13**

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## Supplemental Methods

### Protein sequences used in this study

#### SARS-CoV-2 Spike Protein

MFVFLVLLPLVSSQCVNLTRTQLPPAYTNSFRGVYYPDKVFRSSVLHSTQDLFLPFFS  
NVTWFHAIHVSGTNGTKRFDNPVLPFNDGVYFASTEKSNIIRGWIFGTTLDSKTQSLLIV  
NNATNVVIKVEFQFCNDPFLGVYYHKNNKSWMESEFRVYSSANNCTFEYVSQPFLMD  
LEGKQGNFKNLREFVFKNIDGYFKIYSKHTPINLVRDLPQGFSALEPLVDLPIGINITRFQT  
LLALHRSYLTGDSSSGWTAGAAAYVGYLQPRFLLKYNENGTTDAVDCALDPLSET  
KCTLKSFTVEKGIYQTSNFRVQPTESIVRFPNITNLCPFGEVFNA TRFASVYAWNRKRISN  
CVADYSVLYNSASFSTFKCYGSPTKLNDLCFTNVYADSFVIRGDEV RQIAPGQTGKIA  
DYNYKLPPDDFTGC VIAWNSNNLDSKVGGNNYLYRLFRKSNLKPFERDISTEYQAGST  
PCNGVEGFNCYFPLQSYGFQPTNGVGYQPYRVVVL SFELLHAPATVC GPKNSTLVKN  
KCVNFNFNGLTGTGVLTESNKKFLPFQQFGRDIADTTDAVRDPQTLEILDITPCSFGGVS  
VITPGTN TSNQVA VLYQDV NCTEV PVAIHADQLTPTWRVY STGSNVFQTRAGCLIGAEH  
VNNSYECDIPIGAGICASYQTQTNSPRRAR SVASQSI IAYTMSLGAENS VAYSNN SIAI PT  
NFTISVTTEILPVSM TKTSDCTMYICGDSTECSNLLQYGSFCTQLNRA LTGIAVEQDK  
NTQEVAQVKQIYKTPPIKDFGGFNFSQILPDPSKPSKRSFIEDLLFNKVT LADAGFIKQY  
GDCLG DIAARDLICAQKFNGLTVL PPLTDE MIAQY TSALLA GTITSGWTFGAGA ALQIP  
FAMQMAYRFNGIGVTQNVLYENQKLIANQFNSAIGKIQDSSL TASALGKLQDV VNQN  
AQALNTLVQLSSNFGAISSV LNDILSRLDKVEAEVQIDRLITGRLQSLQTYVTQLIRAA  
EIRASANLAATKMSECVLGQSKR VDFCGKG YHLMSPQSAHGVVFLHVTYVPAQEKN  
FTTAPAICHDGKAHFREGV FV SNGTHWFVTQRNF YEPQI ITTDNTFVSGNC DVVIGIVN  
NTVYDPLQPELDSFKEELDKYFKNHTSPDVLGDISGINASVVNIQKEIDRLNEVAKNLN  
ESLIDLQELGKYEQYIKWPWYIWLGFIA GLIAIVMVTIMLCCMTSCSCLKGCCSCGSCC  
KFDED DSEPVLKGVKLHYT

#### Bat RaTG13 Spike Protein

MFVFLVLLPLVSSQCVNLTRTQLPPAYTNSSTRGVYYPDKVFRSSVLHLTQDLFLPFFS  
NVTWFHAIHVSGTNGIKRFDNPVLPFNDGVYFASTEKSNIIRGWIFGTTLDSKTQSLLIV  
NATNVVIKVEFQFCNDPFLGVYYHKNNKSWMESEFRVYSSANNCTFEYVSQPFLMDL  
EGKQGNFKNLREFVFKNIDGYFKIYSKHTPINLVRDLPQGFSALEPLVDLPIGINITRFQTL  
LALHRSYLTGDSSSGWTAGAAAYVGYLQPRFLLKYNENGTTDAVDCALDPLSETK  
CTLKSFTVEKGIYQTSNFRVQPTDSIVRFPNITNLCPFGEVFNA TT FASVYAWNRKRISNC  
VADYSVLYNSASFSTFKCYGSPTKLNDLCFTNVYADSFVITGDEV RQIAPGQTGKIA DY  
NYKLPPDDFTGC VIAWNSKHIDAEGGNF NYLYRLFRKANLKPFERDISTEYQAGSKPC  
NGQTGLNCYYPLYRYGFYPTDGVGHQPYRVVVL SFELLNAPATVC GPKNSTLVKNKC  
VNFNFNGLTGTGVLTESNKKFLPFQQFGRDIADTTDAVRDPQTLEILDITPCSFGGVS VIT  
PGTN ASNQVA VLYQDV NCTEV PVAIHADQLTPTWRVY STGSNVFQTRAGCLIGAEHVN  
NSYECDIPIGAGICASYQTQTNSRVVASQSI IAYTMSLGAENS VAYSNN SIAI PTNFTISVTT  
EILPVSM TKTSDCTMYICGDSTECSNLLQYGSFCTQLNRA LTGIAVEQDKNTQEVAQ

VKQIYKTPPIKDFGGFNFSQILPDPSKPSKRSFIEDLLFNKVTLADAGFIKQYGDCLGDI  
AARDLICAQKFNGLTVPPLLTDEMIAQYTSALLAGTITSGWTFGAGAAALQIPFAMQMAYR  
FNGIGVTQNVLYENQKLIANQFNSAIGKIQDSSLTASALGKLQDVVNQNAQALNTLV  
QLSSNFGAISSVLDILSRLDKVEAEVQIDRLITGRLQSLQTYVTQQLIRAAEIRASANLA  
ATKMSECVLGQSKRVDFCGKGYHLMSPQSAPGVFLHVTYVPAQEKNFTTAPAI  
CHDGKAHFREGVFVSNGTHWFVTQRNFYEPQIITDNTFVSGCDVVIGIVNNNTVYDPLQP  
ELDSFKEELDKYFKNHTSPDVLGDISGINASVVNIQKEIDRLNEVAKNLNESLIDLQELG  
KYEQYIKWPWYIWLGFIAGLIAIMVTIMLCCMTSCSCLKGCCSCCKFDEDDSEPV  
LKGVKLHYT

### Pangolin Spike Protein

MFVFLFVLPLVSSQCVNLTRTGIPPGYTNSSTRGVYYPDKVFRSSILHLTQDLFLPFFSN  
VTWFNTINYQGGFKKFDNPVLPFNDGVYFASTEKSNIIRGWIFGTTLDARTQSLLIVNNA  
TNVVIKVCEFQFCDFPLGVYYHNNNKTWVENEFRVYSSANNCTFEYISQPFLMDLEGK  
QGNFKNLREFVFKNVDGYFKIYSKHTPIDLVRDLPGRGFAALEPLVLDLPIGINITRFQTLLA  
LHRSYLTGPNLESGWTTGAAAYVGYLQQRTFLLSYNQNQNTITDAVDCSLDPLSETKCT  
LKSLTVEKGIYQTSNFRVQPTISIVRFPNITNLCPGEVFNAASKFASVYAWNRKRISNCVA  
DYSVLYNSTSFSTFKCYGVSPKLNLCFTNVYADSFVVKGDEVRIQIAPGQTGVIADYN  
YKLPDDFTGCVIAWNSVKQDALTGNYGYLYRLFRKSCLKPFERDISTEIQAGSTPCN  
GQVGLNCYYPLERYGFHPTGVNYQPFRVVLSFELLNGPATVCGPKLSTTLVKDKCV  
NFNFNGLTGTGVLTTSKKQFLPFQQFGRDISDTDAVRDPQTLEILDITPCSFGGVSITP  
GTNTSNQAVLYQDVNCTEVPMAIHAEQLTPAWRVYASAGANVFQTRAGCLVGAEHVN  
NSYECDIPVGAGICASYHSMSSLRSVNQRSIIAYTMSLGAENSVA  
YSNNIAIPTNFTISVT  
TEILPVSMTKTSVDCTMYICGDSIECSNLLQYGSFCTQLNRALTGIAVEQDKNTQE  
VFA  
QVKQIYKTPPIKDFGGFNFSQILPDPSKPSKRSFIEDLLFNKVTLADAGFIKQYGDCLGDI  
AARDLICAQKFNGLTVPPLLTDEMIAQYTSALLAGTITSGWTFGAGAAALQIPFAMQMAY  
RFNGIGVTQNVLYENQKLIANQFNSAIGKIQDSSLTASALGKLQDVVNQNAQALNTLV  
KQLSSNFGAISSVLDILSRLDKVEAEVQIDRLITGRLQSLQTYVTQQLIRAAEIRASANL  
AAATKMSECVLGQSKRVDFCGKGYHLMSPQSAPGVFLHVTYVPAQEKNFTTAPAI  
CHEGKAHFREGVFVSNGTHWFITQRNFYEPQIITDNTFVSGCDVVIGIVNNNTVYDPLQ  
PELDSFKEELDKYFKNHTSPDVLGDISGINASVVNIQKEIDRLNEVAKNLNESPIDLQEL  
GKYEQYIKWPWYIWLGFIAGLIAIMVTIMLCCMTSCSCLKGCCSCCKFDEDDSEP  
VLKGVKLHYT

### Intermediate horseshoe bat (*Rhinolophus affinis*) ACE2

MSGSSWLLSLVAVTTAQSTTEDEAKMFLDKFNTKAEDLSHQSSLASWDYNTNINDEN  
VQKMDEAGAKWSAFYEEQSKLAKNYSLEQIQNVTVKLQLQILQQSGSPVLS  
EDKS  
KRL  
NSILNAMSTIYSTGKVCKPNKPQECLLLEPGLDNIMGTSKDYN  
ERLWA  
WE  
GWRAEV  
GK  
QLRPLYEEYVVLKNEMARGYHYEDYGDYWR  
RDYETEESP  
PG  
GYSR  
DQL  
MKD  
VER  
IFT  
EIKPL  
YEHL  
HAYVRA  
KLM  
DTYP  
FH  
ISPT  
GCL  
PA  
HLL  
GDM  
WGR  
FW  
TN  
LY  
PL  
TV  
PFG  
QK  
P

NIDVTDEMLKQGWDADRIFKEAEKFFVSVGLPNMTEGFVNNSMLTEPGDGRKVVCHEP  
TAWDLGKGDFRIKMCTKVTMEDFLTAHHEMGHIQYDMAIASQPYLLRNGANEGFHEA  
VGEVMSLSVATPKHLKTMGLLSPDFREDNETEINFLLKQALNIVGTLPTYMLEDKWRW  
MVFKGEIPKEEWMKWWEMKRKIVGVVEPVPHDETYCDPASLFHVANDYSFIRYYTR  
TIEFQFHEALCRIAQHDGPLHKCDISNSTDAGKKLHQMLSVGKSQAWTKTLEDIVDSR  
NMDVGPLKYFEPLYTWLQEQRKSYVGWNTDWSPYSDQSIKVRISLKSALGENAYE  
WNDNEMYLFRSSVAYAMREYFLKEKHQTILFGAENVWVSNLKPRISFNHFVTSPGNLS  
DIIPRPEVEGAIRMSRSRINDAFLDDNSLEFLGIQPTLGPPYQPPVTIWLIVFGVVMAVV  
VVGIVVLIITGIRDERRKTDQARSEENPYSSVDLSKGENNPGFQNGDDVQTSF

### Syrian Hamster ACE2

MSSSWLLLLSLVAVTTAQSIIEEQAKTFLDKFNQEAEDLSYQSALASWNYNNTNITEENAQM  
KMNEAAAKWSAFYEEQSKLAKNYSLQEVLQNLTIKRLQLQQSGSSALSADKNKQLN  
TILNTMSTIYSTGVCNPKNPQECLLLEPGLDDIMATSTDYNERLWAEGWRAEVGKQ  
LRPLYEEYVVLKNEMARANNYEDYGDYWRGDYEAEAGADGYNNGNQLIEDVERTFK  
EIKPLYEQLHAYVRTKLMNTYPSYISPTGCLPAHLLGDMWGRFWTNLYPLTVFGQKPN  
IDVTDAMVNQGWNAERIFKEAEKFFVSVGLPYMTQGFWENSMLTDPGDDRKVVCHEP  
AWDLGKGDFRIKMCTKVTMDNFTAHHEMGHIQYDMAyatQPFLRNGANEGFHEA  
VGEIMSLSAATPEHLKSIGLLPSDFQEDNETEINFLLKQALTIVGTLPTYMLEDKWRWMV  
FKGDIPKEQWMEKWWEWKREIVGVVEPLPHDETYCDPAALFHVSNDYSFIRYYTRTIY  
QFQFQEALCQAACKHDGPLHKCDISNSTEAGQKLLNMLRLGKSEPWTLALENVVGARN  
MDVRPLLNYFEPLSVWLKEQNKNFVGWNTDWSPYADQSIKVRISLKSALGENAYEWD  
DNEMYLFRASVAYAMRVYFAKNKTQTPFGVEDIRVSDLKPRVSFNFVTSPQNVSDII  
PRNEVEEAVRLSRGRINDVFGDDNSLEFLGINPTLSPPYQPPVTIWLIIFGVVVMGIVVVG  
IILIFTGIKGRKKKNETKREENPYDSVDIGKGESNAGFLSNDDAQTSF

### Mouse ACE2

MSSSWLLLLSLVAVTTAQSLTEENAKTFLNNFNQEAEDLSYQSSLASWNYNNTNITEENAQM  
QKMSEAAAKWSAFYEEQSKTAQSFSLQEIQTPIIKRQLQLQQSGSSALSADKNKQLNTI  
LNTMSTIYSTGVCNPKNPQECLLLEPGLDEIMATSTDYNSRLWAEGWRAEVGKQLR  
PLYEEYVVLKNEMARANNYNDYGDYWRGDYEAEAGADGYNNGNQLIEDVERTFAEIK  
PLYEHLHAYVRRKLMDTYPSYISPTGCLPAHLLGDMWGRFWTNLYPLTVFAQKPNID  
VTDAMMNQGWDAERIFQEAEKFFVSVGLPHMTQGFWANSMLTEPADGRKVVCHEP  
WDLGHGDFRIKMCTKVTMDNFTAHHEMGHIQYDMAyarQPFLRNGANEGFHEAV  
GEIMSLSAATPKHLKSIGLLPSDFQEDSETEINFLLKQALTIVGTLPTYMLEDKWRWMVF  
RGEIPKEQWMKKWWEMKREIVGVVEPLPHDETYCDPASLFHVSDYFIRYYTRTIYQF  
QFQFQEALCQAACKYNGSLHKCDISNSTEAGQKLLKMLSLGNSEPWTKALENVGARNMD  
VKPLLNYFQPLFDWLKEQRNSFVGWNTEWSPYADQSIKVRISLKSALGANAYEWTNN  
EMFLFRSSVAYAMRKYFSIIKNQTVFLEEDVRVSDLKPRVSFYFFVTSPQNVSDVIPRSE

VEDAIRMSRGRINDVGLNDNSLEFLGIHPTLEPPYQPPVTIWLIIFGVVMALVVVGIIILIV  
TGIKGRKKNETKREENPYDSMDIGKGESNAGFQNSDDAQTSF

### Pig ACE2

MSGFWLLLIPVTAAQSTTEELAKTFLEKFNLAEADLAYQSSLASWTINTNITDENIQK  
MNDARAKWSAFYEEQSRIAKEYPLDEIQLTLIKRQLQALQQSGTSGSADKSKRLNTILN  
TMSTIYSSGKVLDPPNPQECLVLEPGLDEIMENSKDYSRRLWAWESWRAEVGKQLRPL  
YEEYVLENEMARANNYEDYGDYWRGDYEVGTGDYDYSRNQLMEDVERTFAEIKPL  
YEHLHAYVRAKLMAYPSRISPTGCLPAHLLGDMWGRFWTNLYPLTVPGKEKPSIDVT  
EAMVNQSWDAIRIFEEAKFFVSIGLPNMTQGFNWNNMLTEPGDGRKVVCHPTAWDLG  
KGDFRIKMCTKVTMDDFLTAHHEMGHIQYDMAYAIQPYLLRNGANEGFHEAVGEIMSL  
SAATPHYLKALGLLPPDFYEDSETEINFLLKQALTIVGTLPTYMLEKWRWMVFKEIPK  
EQWMQKWWEMKREIVGVVEPLPHDETYCDPACLFVAEDYSFIRYYTRTIYQFQFHEA  
LCRTAKHEGPLYKCDISNSTEAGQKLLQMLSLGKSEPWTIALENIVGKTMVDVKPLLSY  
FEPLLTLKAQNGNNSVGWNTDWTPYADQSIKVRISLKSALGEDAYEWNDNEMYLFRS  
SIAYAMRNYFSSAKNETIPFGAVDVWVSDLKPRISFNFFVTSPANMSDIIPRSDVEKAISM  
SRSRINDAFRLDDNTLEFLGIQPTLGPPDEPPVTWLIIFGVVMGLVVVGIVVLIFTGIRDR  
RKKKQASSEENPYGSMDLSKGESNSGFQNGDDIQTsf

### Chinese hamster ACE2

MSSSWLLLTLAVTTAQSIIEEQAKTFLDKFNQEAEDLSYQSLASWNYNTNITEENAQ  
KMNEAAAKWSAFYEEQSCLKAKNYSLQEVTQNLIIKRQLQALQQSGSSALSADKNKQLNT  
ILNTMSTIYSTGKVCNPKNPQECLLLEPGLDDIMATSTDYNERLWAWEGWRAEVGKQL  
RPLYEEYVVLKNEMARANNYKDYGDYWRGDYEAEGADGYNYNGNQLIEDVERTFKEI  
KPLYEQLHAYVRTKLMDTYPSFISPTGCLPAHLLGDMWGRFWTNLYPLTVPGQKPNID  
VTDAMVNQGWDAERIFKEAEKFFVSVGLPHMTQGFWGNSMLTDPGDDRKVVCPTA  
WDLGKGDFRIKMCTKVTMDNFTAHHEMGHIQYDMAYATQPFLLRNGANEGFHEAV  
GEIMSLSAATPKHLKSIGLLPSNFHEDNETEINFLLKQALTIVGTLPTYMLEKWRWMVF  
KGDIPEKEWMEKWEMKREIVGVVEPLPHDETYCDPAALHVSNDYSFIRYYTRTIYQ  
FQFQEALCQAALKHDGPLHKCDISNSTEAGQKLLNMLRLGKSEPWTIALENVGARNM  
DVRPLNYFEPLSVWLKEQNKNFVGWNTDWSPYADQSIKVRISLKSALGENAYEWND  
NEMYLFRTVAYAMRVYFAKNKTQTVLFGVEDIRVSDLKPRVSFNFFVTSPQNVSDIIP  
RNEVEEARFSRGRINDVGLDDNSLEFLGINPTLAPPYQPPVTIWLIIFGVVVMGIVVGI  
VILITGIRARKKNEAKREENPYDSVDIGKGESNAGFQSNDDVQTSF

### Ferret ACE2

MLGSSWLLLTLAAQSTTEDLAKTFLEKFNYEAEEELSYQNSLASWNYNTNITDENI  
QKMNIAGAKWSAFYEEESQHAKTYPLEEIQDPIIKRQLRALQQSGSSVLSADKRERLNTI

LNAMSTIYSTGKACPNNPQECLLLEPGLDDIMENSKDYNERLWAWEGRSEVGKQLR  
PLYEEYVALKNEMARANNYEDYGDYWRGDYEEEWADGYSYSRNQLIEDVEHTFTQIK  
PLYEHLHAYVRAKMDAYPSRISPTGCLPAHLLGDMWGRFWTNLYPLMVPFRQKPNID  
VTDAMVNQSVDARRIFEEAETFFVSVGLPNMTEGFWQNSMLTEPGDNRKVVCPTAW  
DLGKRDRIKMCTKVTMDDFLTAHHEMGHIQYDMAYAEQPFLRNGANEGFHEAVGEI  
MSLSAATPNHLKNIGLLPPDFSEDSETDINFLLKQALTIVGTLPTYMLEKWRWMVFKG  
EIPKEQWMQKWEMKRDIVGVEPLPHDETYCDPAALFHVANDYSFIRYYTRTIYQFQ  
FQEALCQIAKHEGPLYKCDISNSSEAGQKLHEMLSLGRSKPWTFALEVVGAKTMDVR  
PLLNYFEPLFTWLKEQRNSFVGWNTDWSPYADQSIKVRISLKSALGENAYEWNDNEM  
YFFQSSIAYAMREYFSKVKNQTIPFGDKDVRVSDLKPRISFNFIITSPENMSDIIPRADVEE  
AIRKSRGRINDAFRLDDNSLEFLGIQPTLEPPYQPPVTIWLIVFGVVMGVVVVGIFLLIFSG  
IRNRRKNNQARSEENPYASVDSLKGENNPGFQNVDVQTSF

#### Bovine ACE2

MTGSFWLLSLVAVTAAQSTTEEQAKTFLEKFNHEAEDLSYQSSLASWNYNTNDENV  
QKMNEARAKWSAFYEEQSRMAKTYSEEIQNLTLRQLKALQHSGTSALSAEKSKRLN  
TILNMSTIYSTGKVLDPNTQECLALEPGLDDIMENS RDYNRRLWAWEGRRAEVGKQL  
RPLYEEYVLENEMARANNYEDYGDYWRGDYEVGTAGDYDYSRDQLMKDVERTFAE  
IKPLYEQLHAYVRAKLMHTPSYISPTGCLPAHLLGDMWGRFWTNLYSLTVPEHKPSI  
DVTEKMENQSWDAERIFKEAEKFFVSISLPYMTQGFWDNSMLTEPGDGRKVVCPTAW  
DLGKGDFRIKMCTKVTMDDFLTAHHEMGHIQYDMAYAAQPYLLRNGANEGFHEAVGE  
IMSLSAATPHYLKALGLAPDFHEDNETEINFLLKQALTIVGTLPTYMLEKWRWMVF  
GEIPKQQWMEKWWEMKREIVGVEPLPHDETYCDPACLFHVAEDYSFIRYYTRTIYQF  
QFHEALCKTAKHEGALFKCDISNSTEAGQRLLQMLRLGKSEPWTLALENIVGIKTMDVK  
PLLNYFEPLFTWLKEQRNSFVGWSTEWPYSDQSIKVRISLKSALGENAYEWNDNEM  
YLFQSSVAYAMRKYFSEARNETVLFGEDNVWVSDKPRISFKFFVTSPNNVSDIIPRT  
ENAIRLSRDRFNDVQLDDNSLEFLGIQPTLGPPYEPPVTIWLIVFGVVMGVVVIGIVVLIF  
TGIRNRRKKNQASSEENPYGSVDLNKGENNSGFQNIDDVQTSF

#### Grivet (African green monkey) ACE2

MSSSWLLSLVAVTAAQSTIEEQAKTFLDKFNHEAEDLFYQSSLASWNYNTNITEENV  
QNMNNAGEKWSAFLKEQSTLAQMYPLQAIQNLTVKLQLQALQQNGSSVLSEDKSKRL  
NTILNTMSTIYSTGKVCNPNNPQECLLDPLNEIMEKSLDYNERLWAWEGRSEVGK  
QLRPLYEEYVVLKNEMARANHYKDYGDYWRGDYEVNGVDGYDYNRDQLIEDVERTF  
EEIKPLYEHLHAYVRAKLMNAYPSYISPTGCLPAHLLGDMWGRFWTNLYSLTVPEFGQK  
PNIDVT DAMVNQAWNAQRIFKEAEKFFVSVGLPNMTQGFWENSMLTDPGNVQKVVC  
PTAWDLKGDFRIIMCTKVTMDDFLTAHHEMGHIQYDMAYAAQPYLLRNGANEGFHE  
AVGEIMSLAATPKHLKSIGLLSPDFQEDNETEINFLLKQALTIVGTLPTYMLEKWRWM  
VFKGEIPKDQWMKKWWEMKREIVGVEPVPHDETYCDPASLFHVSNDYSFIRYYTRTL  
YQFQFQEALCQAQAKHEGPLHKCDISNSTEAGQKLLNMLKLGKSEPWTLALENVVGAKN

MSVRPLLNYFEPLFTWLKDQNKNFVGWSTDWSPYADQSIKVRISLKSALGANAYKWN  
DNEMYLFRSSVAYAMRQYFLENKHQTILFGEEDVRVADLKPRISFNFYVTAPKNVSDIIP  
RTEVEEAIRFSRSRINDAFQLNDNSLEFLGIQSTLVPPYQSPITTWLIVFGVVMAVIVAGIV  
VLIFTGIRDRKKKNQARSEENPYASIDISKGENNPGFQNTDDVQTSF

Malayan pangolin ACE2

MSGSSWLLSLVAVTAAQSTSDEEAKTFLEKFNSEAEELSYQSSLASWNYNTNITDEN  
QKMNVAGAKWSTFYEEQSKIAKNYQLQNIQNDTIKRQLQALQLSGSSALSADKNQRLN  
TILNTMSTIYSTGVCNPQECISLLEPGLDNIMESSKDYNELWAEGWRSEVGKQL  
RPLYEEYVVLKNEMARANHYEDYGDYWRGDYEAE GANGYNYSRDHLIEDVEHIFTQI  
KPLYEHLHAYVRAKLMNDNYPHSIPTGCLPAHLLGDMWGRFWTNLYPLTVPFRQKPNI  
DVT DAMVNQTWDANRIFKEAEKFFVSGLPKMTQTFWENSMLTEPGDGRKVVC  
WDLGKHDFRIKMCTKVTMDDFLTAHH EMGHIQYDMA YAMQPYLLRNGA  
NEG FHEAV GEIMSLSAATPKHLKNIGLLPPDFYEDNETEINFLLKQALTIVGTL  
PFTYMLEKWRWMVF SGQIPKEQWMKKWWEMKREIVGVVEPVPHDETYCDP  
ASL FH VANDY SFIR YYTR TIY Q FQFQEALC  
QTAKHEGPLHKCDISNSAEAGQKLLQMLSLGKSKPWT  
LA LERVVGTKNMD VRPLLNYFEPLLTWLKEQNKNFVGWNTDWSPYAAQSIK  
VRISLKSALGEKAYEWNDS EMYLFRSSVAYAMREYFSKVKQTIPFEDEC  
VRVSDLKPRVSFIFFVTLPKNVSAVIPRA EVEEAIRISRSRINDA  
FRLDDNSLEFLGIQPTLQPPYQPPVTIWLIVFGVV  
MGVVVVGIVV LIFTGIRDRKKDQARSEQNPYASVDLSKG  
GENNPGFQNVDDVQTSF

## Supplemental Figures and Legends

**Figure S1.** Protease repertoires in Calu-3 and Caco-2. Calu-3 and Caco-2 cells were infected with SARS-CoV-2 USA/WA1-2020 at MOI of 0.1 for 4, 24, and 48 hours. RNA was isolated from uninfected and infected cells were subject to mRNA-Seq. Reads per kilobase per million mapped reads (RPKM) was calculated to show expression level of proteases that may cleave SARS-CoV-2 spike protein at the S1/S2 site. Also refer to Table S1.

