

**Supplemental information**

**Binding and molecular basis of the bat coronavirus**

**RaTG13 virus to ACE2 in humans and other species**

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**Supplementary tables:**

**Table S1. The coding genes of 25 ACE2 orthologs, Related to Figure 2**

Species of ACE2	Organism	Database	Code
Human	<i>Homo sapiens</i>	NCBI Reference Sequence	NP_001358344.1
Macaca	<i>Macaca fascicularis</i>	NCBI Reference Sequence	XP_005593094.1
Rabbit	<i>Oryctolagus cuniculus</i>	NCBI Reference Sequence	XP_002719891.1
Mouse	<i>Mus musculus</i>	NCBI Reference Sequence	NP_001123985.1
Rat	<i>Rattus norvegicus</i>	NCBI Reference Sequence	NP_001012006.1
Malayan pangolin	<i>Manis javanica</i>	NCBI Reference Sequence	XP_017505746.1
Cat	<i>Lynx canadensis</i>	NCBI Reference Sequence	XP_030160839.1
Civet	<i>Paguma larvata</i>	NCBI Reference Sequence	Q56NL1.1
Fox	<i>Vulpes vulpes</i>	NCBI Reference Sequence	XP_025842512.1
Dog	<i>Canis lupus familiaris</i>	NCBI Reference Sequence	XP_005641049.1
Raccoon Dog	<i>Nyctereutes procyonoides</i>	GenBank	ABW16956.1
Horse	<i>Equus caballus</i>	NCBI Reference Sequence	XP_001490241.1
Pig	<i>Sus scrofa</i>	NCBI Reference Sequence	XP_020935033.1
Wild bactrian camel	<i>Camelus ferus</i>	NCBI Reference Sequence	XP_006194263.1
Alpaca	<i>Vicugna pacos</i>	NCBI Reference Sequence	XP_006212709.1
Bovine	<i>Bos taurus</i>	NCBI Reference Sequence	XP_005228486.1
Goat	<i>Capra hircus</i>	NCBI Reference Sequence	XP_005701129.2
Sheep	<i>Ovis aries</i>	NCBI Reference Sequence	XP_011961657.1
Intermediate horseshoe bat	<i>Rhinolophus affinis</i>	GenBank	QMQ39222.1
Little brown bat	<i>Myotis lucifugus</i>	NCBI Reference Sequence	XP_023609439.1
Fulvous fruit bat	<i>Rousettus leschenaultii</i>	GenBank	ADJ19219.1
Greater horseshoe bat	<i>Rhinolophus ferrumequinum</i>	NCBI Reference Sequence	BAH02663.1
Least horseshoe bat	<i>Rhinolophus pusillus</i>	GenBank	ADN93477.1
Big-eared horseshoe bat	<i>Rhinolophus macrotis</i>	GenBank	ADN93471.1
Lesser hedgehog tenrec	<i>Echinops telfairi</i>	NCBI Reference Sequence	XP_004710002.1

**Table S3. The binding affinities between ACE2s and SARS-CoV-2 RBD, RaTG13 RBD or RaTG13 RBD mutants, Related to Figure 3**

Species	Binding affinity ( $\mu\text{M}$ )								
	SARS-CoV-2 RBD	RaTG13-CoV RBD	F449Y	L486F	Y493Q	Y498Q	D501N	H505Y	6-mutant
Human	$0.015 \pm 0.001^1$	$3.86 \pm 1.1$	$1.43 \pm 0.36$	$0.88 \pm 0.16$	$9.78 \pm 0.46$	-- <sup>2</sup>	$0.34 \pm 0.03$	$0.7 \pm 0.03$	$0.028 \pm 0.001$
Mouse	--	$2.55 \pm 0.01$	$1.02 \pm 0.41$	$6.65 \pm 3.7$	--	--	$35.6 \pm 10.5$	$0.6 \pm 0.05$	--
Horse	$0.056 \pm 0.006$	$0.69 \pm 0.1$	$0.48 \pm 0.15$	$1.21 \pm 0.43$	$0.33 \pm 0.07$	$9.39 \pm 3.1_4$	$0.25 \pm 0.04$	$0.15 \pm 0.009$	$0.111 \pm 0.001$
Intermediate horseshoe bat	$0.382 \pm 0.054$	$42.3 \pm 15.4$	--	$21.4 \pm 9.9$	--	--	$0.59 \pm 0.11$	$5.82 \pm 0.74$	$0.179 \pm 0.04$

<sup>1</sup> The binding affinities between ACE2s and RaTG13 RBD or RaTG13 RBD mutants are shown with the means  $\pm$  SD of three independent experiments.

<sup>2</sup> "--" means no detectable binding.

**Table S4. The capture level of the indicated MAb and concentrations of the indicated RBD, together with the calculated  $K_D$  in the SPR assay were listed, Related to Figure 5.**

MAbs	Capture level (RU)	Solution	Concentrations of Solutions (nM)						$K_D$
H4	662.3 ± 4.97	SARS-CoV-2 RBD	6.25 12.5 25 50 100						1.72 ± 0.11nM
B38	776.3 ± 12.55	SARS-CoV-2 RBD	50 100 200 400 800						10.9 ± 0.85nM
CB6	744.1 ± 4.01	SARS-CoV-2 RBD	6.25 12.5 25 50 100						1.95 ± 0.17nM
S2H14	575.1 ± 2.80	SARS-CoV-2 RBD	50 100 200 400 800						3.98 ± 0.462nM
REGN10933	495.7 ± 4.69	SARS-CoV-2 RBD	1.5625 3.125 6.25 12.5 25						0.223 ± 0.0018nM
REGN10987	701.2 ± 9.03	SARS-CoV-2 RBD	6.25 12.5 25 50 100						1.66 ± 0.13nM
C110	398.8 ± 9.90	SARS-CoV-2 RBD	6.25 12.5 25 50 100						0.122 ± 0.007nM
H4	662.2 ± 4.88	RaTG13 RBD	200 400 800 1600 3200						--
B38	757.1 ± 3.95	RaTG13 RBD	200 400 800 1600 3200						963 ± 16.9nM
CB6	744.2 ± 5.49	RaTG13 RBD	50 100 200 400 800						39.0 ± 0.45nM
S2H14	571.7 ± 2.17	RaTG13 RBD	200 400 800 1600 3200						--
REGN10933	492.7 ± 5.02	RaTG13 RBD	200 400 800 1600 3200						551 ± 20.2nM
REGN10987	696.5 ± 11.82	RaTG13 RBD	200 400 800 1600 3200						5930 ± 381nM
C110	391.2 ± 24.53	RaTG13 RBD	200 400 800 1600 3200						1100 ± 59.3nM

<sup>1</sup> The binding affinities between ACE2s and RaTG13 RBD or RaTG13 RBD mutants are shown with the means ± SD of three independent experiments.

2 “--” means no detectable binding.

**Table S5. The information of donors, Related to Figure 5**

Donors	Gender	Age	Developmental stage
1	Male	50	Convalescent
2	Male	51	Convalescent
3	Male	56	Convalescent
4	Male	20	Convalescent
5	Male	28	Convalescent
6	Female	61	Convalescent
7	Male	30	Healthy
8	Female	27	Healthy
9	Male	24	Healthy