

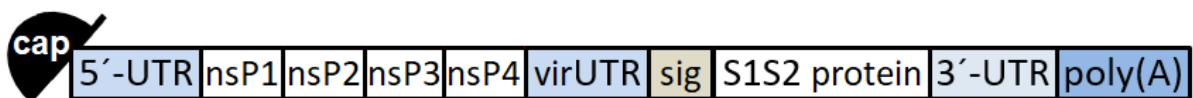


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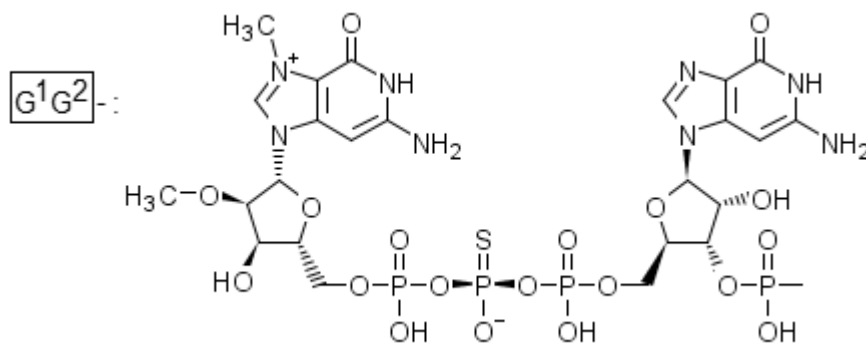
Description

Self-replicating messenger RNA encoding the full-length SARS-CoV-2 spike glycoprotein.

Schematic



UTR = Untranslated region; nsP1, nsP2, nsP3 and nsP4 = Venezuelan equine encephalitis virus (VEEV) RNA-dependent RNA polymerase; virUTR = 5'-UTR derived from VEEV; sig = extended signal sequence of the S glycoprotein. S1S2 protein = S glycoprotein sequence containing mutations K986P and V987P.



5'- capping structure

The 5'- capping structure is = $m^7G^+m^2-5'-p-(R)p^s-p-5'-G$

Table of features

Element	Description	Position
cap	cap: $m^7G^+m^2-5'-p-(R)p^s-p-5'-G$	1-2
5'-UTR	5'-untranslated region derived from the Venezuelan equine encephalitis virus (VEEV)	3-46



nsP1	Venezuelan equine encephalitis virus (VEEV) RNA-dependent RNA polymerase (RNA replicase) with a subgenomic promotor starting in nsP4. - In conjunction with the virUTR, these sequences introduce a self-amplifying function.	47-1651
nsP2		1652-4033
nsP3		4034-5704
nsP4		5705-7528
virUTR	VEEV untranslated region encoding 5 nucleotides of the subgenomic promotor as well as replication and translation supporting sequence elements.	7529-7569
sig	S glycoprotein signal peptide (extended leader sequence), which guides translocation of the nascent polypeptide chain into the endoplasmic reticulum.	7570 - 7617
S1S2 protein	Codon-optimized sequences encoding full-length SARS-CoV-2 S glycoprotein; two mutations, K986P and V987, ensure that the S glycoprotein remains in an antigenically optimal pre-fusion conformation.	7618 - 11394
3'-UTR	3'-untranslated region comprising sequence elements derived from the amino-terminal enhancer of split (AES) mRNA and the mitochondrial encoded 12S ribosomal RNA, to confer RNA stability and highest total protein expression, followed by a further Venezuelan equine encephalitis virus (VEEV) derived sequence of the 26S mRNA.	11395-11808
poly(A)	A 110-nucleotide poly(A)-tail consisting of a stretch of 30 adenosine residues, followed by a 10-nucleotide linker sequence and another 70 adenosine residues.	11809-11918

Sequence / Séquence / Secuencia

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GGAUGGGCGG CGAUGAGAG AAGCCAGAC CAUUACCUA CCCAAAUGG 50
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UGACCAUGCU AAUGCCAGAG CGUUUUCGCA UCUGGCUUCA AAACUGAUCG 200
AAACGGAGGU GGACCCAUCC GACACGAUCC UUGACAUUGG AAGUGCGCCC 250
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AUGUGCGGAA GAUCCGGACA GAUUGUAUAA GUAUGCAACU AAGCUGAAGA 350
AAAACUGUAA GGAAAUAAU GAUAAGGAAU UGGACAAGAA AAUGAAGGAG 400
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CCACGACGAC GAGUCGUGUC GCUACGAAGG GCAAGUCGCU GUUUACCAGG 500

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<u>AAACCGUGUU</u>	<u>AACGGCUCGU</u>	<u>AACAUAGGCC</u>	<u>UAUGCAGCUC</u>	<u>UGACGUUAUG</u>	700
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<u>GGGACUUACU</u>	<u>GAGGAGCUGG</u>	<u>CACCUGCCGU</u>	<u>CUGUAUUUCA</u>	<u>CUUACGUGGC</u>	850
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<u>CAGGUACCUG</u>	<u>CACCAUUAUG</u>	<u>CCACACAUGG</u>	<u>AGGAGCGCUG</u>	<u>AACACUGAUG</u>	2000
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UAAGAACAUC	GACGGCUACU	UCAAGAUJUA	CAGCAAGCAC	ACCCCUAUCA	8200
ACCUCJUGCG	GGAUUCJCCU	CAGGGCUUCU	CUGCUCUGGA	ACCCUJGGUG	8250
GAUCUGCCCA	UCGGCAUCA	CAUCACCCGG	UUUCAGACAC	UGCUGGCCU	8300



GCACAGAAGC	UACCUGACAC	CUGGCGAUAG	CAGCAGCGGA	UGGACAGCUG	8350
GUGCCGCCGC	UUACUAUGUG	GGCUACCUGC	AGCCUAGAAC	CUUCCUGCUG	8400
AAGUACAACG	AGAACGGCAC	CAUCACCGAC	GCCGUGGAUU	GUGCUCUGGA	8450
UCCUCUGAGC	GAGACAAAGU	GCACCCUGAA	GUCCUUCACC	GUGGAAAAGG	8500
GCAUCUACCA	GACCAGCAAC	UUCCGGGUGC	AGCCCACCGA	AUCCAUCGUG	8550
CGGUUCCCCA	AUAUCACCAA	UCUGUGCCCC	UUCGGCGAGG	UGUUCAAUUC	8600
CACCAGAUUC	GCCUCUGUGU	ACGCCUGGAA	CCGGAAGCGG	AUCAGCAAUU	8650
GCGUGGCCGA	CUACUCCGUG	CUGUACAACU	CCGCCAGCUU	CAGCACCUUC	8700
AAGUGCUACG	GCGUGUCCCC	UACCAAGCUG	AACGACCUGU	GCUUCACAAA	8750
CGUGUACGCC	GACAGCUUCG	UGAUCCGGGG	AGAUGAAGUG	CGGCAGAUUG	8800
CCCCUGGACA	GACAGGCAAG	AUCGCCGACU	ACAACUACAA	GCUGCCCGAC	8850
GACUUCACCG	GCUGUGUGAU	UGCCUGGAAC	AGCAACAACC	UGGACUCCAA	8900
AGUCGGCGGC	AACUACAAUU	ACCUGUACCG	GCUGUCCCGG	AAGUCCAAUC	8950
UGAAGCCCUU	CGAGCGGGAC	AUCUCCACCG	AGAUCUAUCA	GGCCGGCAGC	9000
ACCCUUGUA	ACGGCGUGGA	AGGCUUCAAC	UGCUCUUC	CACUGCAGUC	9050
CUACGGCUUU	CAGCCCACAA	AUGGCGUGGG	CUAUCAGCCC	UACAGAGUGG	9100
UGGUGCUGAG	CUUCGAACUG	CUGCAUGCCC	CUGCCACAGU	GUGCGGCCCU	9150
AAGAAAAGCA	CCAUCUCGU	GAAGAACAAA	UGCUGAACU	UCAACUCAA	9200
CGGCCUGACC	GGCACCGGCG	UGCUGACAGA	GAGCAACAAG	AAGUCCUGC	9250
CAUCCAGCA	GUUUGGCCGG	GAUAUCGCCG	AUACCACAGA	CGCCGUUAGA	9300
GAUCCCAGA	CACUGGAAAU	CCUGGACAUC	ACCCUUGCA	GCUUCGGCGG	9350
AGUGUCUGUG	AUCACCCUG	GCACCAACAC	CAGCAAUCAG	GUGGCAGUGC	9400
UGUACCAGGA	CGUGAACUGU	ACCGAAGUGC	CCGUGGCCAU	UCACGCCGAU	9450
CAGCUGACAC	CUACAUGGCG	GGUGUACUCC	ACCGGCAGCA	AUGUGUUUCA	9500
GACCAGAGCC	GGCUGUCUGA	UCGGAGCCGA	GCACGUGAAC	AAUAGCUACG	9550
AGUGCGACAU	CCCCAUCGGC	GCUGGAAUCU	GCGCCAGCUA	CCAGACACAG	9600
ACAAACAGCC	CUCGGAGAGC	CAGAAGCGUG	GCCAGCCAGA	GCAUCAUUGC	9650
CUACACAAUG	UCUCUGGGCG	CCGAGAACAG	CGUGGCCUAC	UCCAACAACU	9700
CUAUCGCUAU	CCCCACCAAC	UUCACCAUCA	GCGUGACCAC	AGAGAUCUG	9750
CCUGUGUCCA	UGACCAAGAC	CAGCGUGGAC	UGCACCAUGU	ACAUCUGCGG	9800
CGAUUCCACC	GAGUGCUGCA	ACCUGCUGCU	GCAGUACGGC	AGCUUCUGCA	9850
CCCAGCUGAA	UAGAGCCUG	ACAGGGAUCC	CCGUGGAACA	GGACAAGAAC	9900
ACCCAAGAGG	UGUUCGCCCA	AGUGAAGCAG	AUCUACAAGA	CCCCUCCUUA	9950
CAAGGACUUC	GGCGGCUUCA	AUUUCAGCCA	GAUUCUGCCC	GAUCCUAGCA	10000
AGCCAGCAA	GCGGAGCUUC	AUCGAGGACC	UGCUGUCAA	CAAAGUGACA	10050
CUGGCCGACG	CCGGCUUCAU	CAAGCAGUUA	GGCGAUUGUC	UGGGCGACAU	10100
UGCCGCCAGG	GAUCUGAUUU	GCGCCAGAA	GUUUAACGGA	CUGACAGUGC	10150
UGCCUCCUCU	GCUGACCGAU	GAGAUGAUCG	CCCAGUACAC	AUCUGCCCUG	10200
CUGGCCGGCA	CAAUCACAAG	CGGCUGGACA	UUUGGAGCAG	GCGCCGCUCU	10250
GCAGAUCCCC	UUUGCUAUGC	AGAUGCCUA	CCGGUUCAAC	GGCAUCGGAG	10300
UGACCCAGAA	UGUGCUGUAC	GAGAACCAGA	AGCUGAUCGC	CAACCAGUUC	10350
AACAGCGCCA	UCGGCAAGAU	CCAGGACAGC	CUGAGCAGCA	CAGCAAGCGC	10400
CCUGGGAAAG	CUGCAGGACG	UGGUCAACCA	GAAUGCCAG	GCACUGAACA	10450
CCCUGGUCAA	GCAGCUGUCC	UCCAACUUCG	GCGCCAUCAG	CUCUGUGCUG	10500
AACGAUAUCC	UGAGCAGACU	GGACCCUCCU	GAGGCCGAGG	UGCAGAUCCA	10550
CAGACUGAUC	ACAGGCAGAC	UGCAGAGCCU	CCAGACAUAC	GUGACCCAGC	10600
AGCUGAUCAG	AGCCGCCGAG	AUUAGAGCCU	CUGCCAAUCU	GGCCGCCACC	10650
AAGAUGUCUG	AGUGUGUGCU	GGGCCAGAGC	AAGAGAGUGG	ACUUUUGCGG	10700
CAAGGGCUAC	CACCUGAUGA	GCUUCCCUCA	GUCUGCCCU	CACGGCGUGG	10750
UGUUUCUGCA	CGUGACAUUA	GUGCCCGCUC	AAGAGAAGAA	UUUACCACC	10800
GCUCCAGCCA	UCUGCCACGA	CGGCAAAGCC	CACUUCCUA	GAGAAGGCGU	10850
GUUCGUGUCC	AACGGCACCC	AUUGGUUCGU	GACACAGCGG	AACUUCUACG	10900



AGCCCCAGAU	CAUCACCACC	GACAACACCU	UCGUGUCUGG	CAACUGCGAC	10950
GUCGUGAUCG	GCAUUGUGAA	CAAUACCGUG	UACGACCCUC	UGCAGCCCGA	11000
GCUGGACAGC	UUCAAAGAGG	AACUGGACAA	GUACUUUAAG	AACCACACAA	11050
GCCCCGACGU	GGACCUGGGC	GAUAUCAGCG	GAAUCAAUUG	CAGCGUCGUG	11100
AACAUCCAGA	AAGAGAUCGA	CCGGCUGAAC	GAGGUGGCCA	AGAAUCUGAA	11150
CGAGAGCCUG	AUCGACCUGC	AAGAACUGGG	GAAGUACGAG	CAGUACAUCA	11200
AGUGGCCCCUG	GUACAUCUGG	CUGGGCUUUA	UCGCCGGACU	GAUUGCCAUC	11250
GUGAUGGUCA	CAAUCAUGCU	GUGUUGCAUG	ACCAGCUGCU	GUAGCUGCCU	11300
GAAGGGCUGU	UGUAGCUGUG	GCAGCUGCUG	CAAGUUCGAC	GAGGACGAUU	11350
CUGAGCCCGU	GCUGAAGGGC	GUGAAACUGC	ACUACACAUG	AUGACUCGAG	11400
CUGGUACUGC	AUGCACGCAA	UGCUAGCUGC	CCCUUCCCG	UCCUGGGUAC	11450
CCCGAGUCUC	CCCCGACCUC	GGGUCCCAGG	UAUGCUCCCA	CCUCCACCUG	11500
CCCCACUCAC	CACCUCUGCU	AGUUCCAGAC	ACCUCCCAAG	CACGCAGCAA	11550
UGCAGCUCAA	AACGCUUAGC	CUAGCCACAC	CCCCACGGGA	AACAGCAGUG	11600
AUUAACCUUU	AGCAAUAAAC	GAAAGUUUAA	CUAAGCUAUA	CUAACCCAG	11650
GGUUGGUCAA	UUUCGUGCCA	GCCACACCGC	GGCCGCAUGA	AUACAGCAGC	11700
AAUUGGCAAG	CUGCUUACAU	AGAACUCGCG	GCGAUUGGCA	UGCCGCCUUA	11750
AAAUUUUUUAU	UUUAUUUUUU	CUUUUCUUUU	CCGAAUCGGA	UUUUGUUUUU	11800
AAUAUUUCAA	AAAAAAAAAA	AAAAAAAAAA	AAAAAAAAAGC	AUAUGACUAA	11850
AAAAAAAAAA	AAAAAAAAAA	AAAAAAAAAA	AAAAAAAAAA	AAAAAAAAAA	11900
AAAAAAAAAA	AAAAAAA				11918