

Supplementary Information for

Reverse-transcribed SARS-CoV-2 RNA can integrate into the genome of cultured human cells and can be expressed in patient-derived tissues

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Datasets S1 to S4

Supplementary Figures

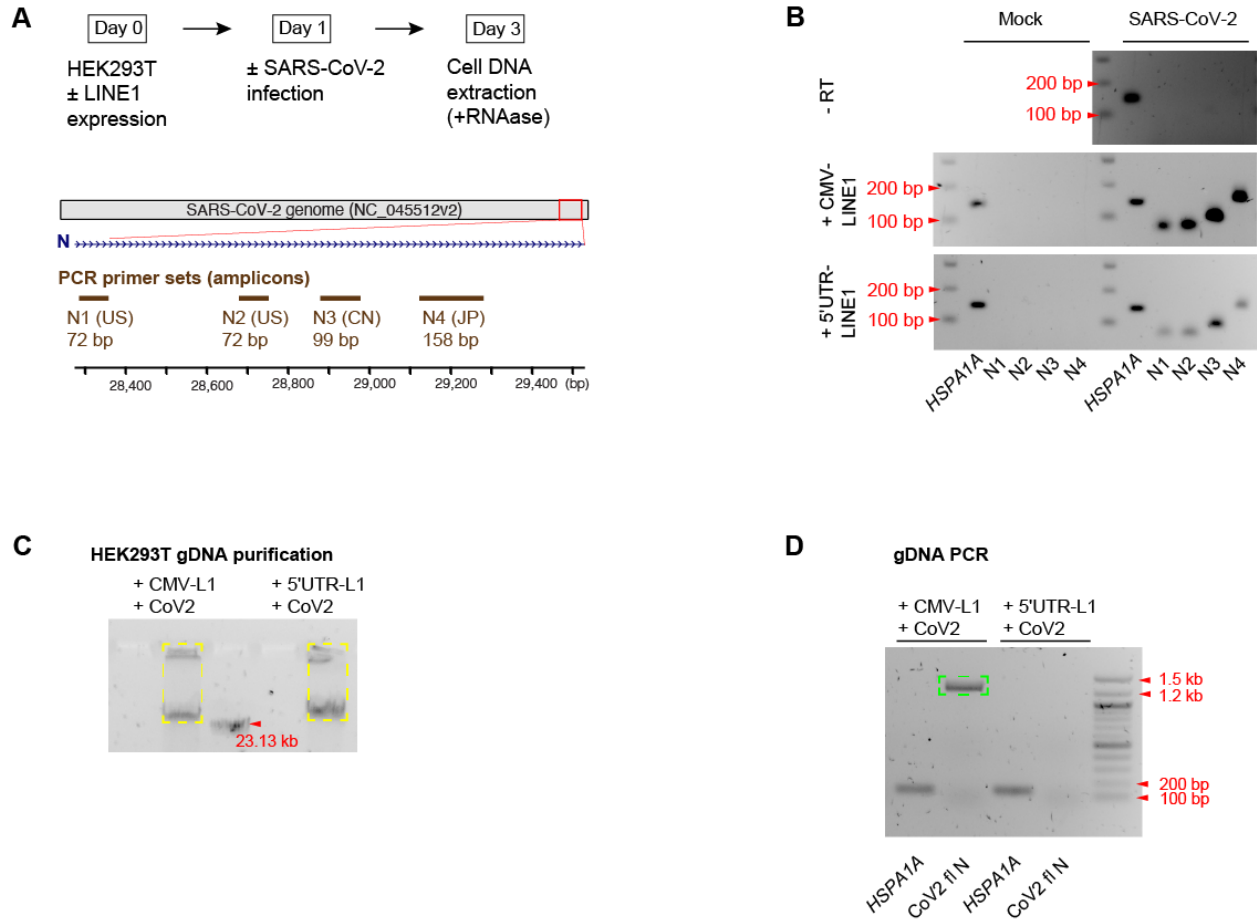


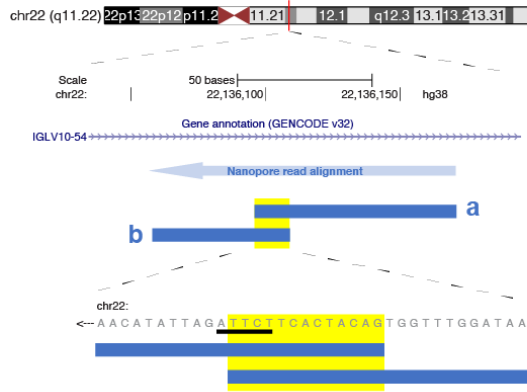
Fig. S1. Detection of DNA copies of SARS-CoV-2 RNA in infected LINE1-overexpressing cells. **A)** Experimental workflow (top) and PCR primer sets (bottom) used to detect reverse-transcription and integration of SARS-CoV-2 RNA. **B)** PCR detection of SARS-CoV-2 NC sequences in DNA purified from mock (left) or SARS-CoV2 (right) infected HEK293T cells without or with transfection of human LINE1 (CMV-LINE1 or 5'UTR-LINE1) plasmids. *HSPA1A*: human *HSPA1A* gene as control; N1 – N4: SARS-CoV-2 NC amplicons as shown in **A**). N1 – N4 PCR products were loaded on gel three times the amount of *HSPA1A* PCR product. Note that we didn't detect DNA copies of SARS-CoV-2 sequences in cells without LINE1 overexpression by this low-sensitive PCR assay. **C)** Gel purification of large fragments of genomic DNA (yellow boxes) from SARS-CoV-2 infected HEK293T cells that were transfected

with CMV-LINE1 or 5'UTR-LINE1. **D)** Cloning of a DNA copy of a complete SARS-CoV-2 NC gene sequence (CoV2 fl N, green box) from gel-purified HEK293T genomic DNA.

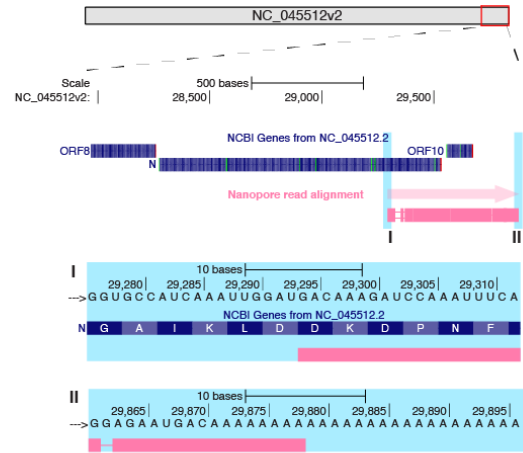
A “Human-CoV2-human” chimeric read (Nanopore)



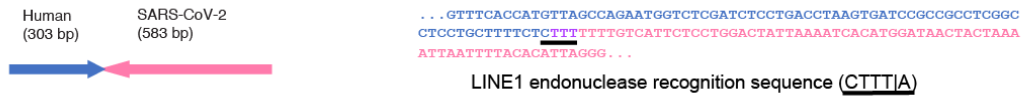
B “Human-CoV2-human” chimeric read (Nanopore) alignment on Human Chr22



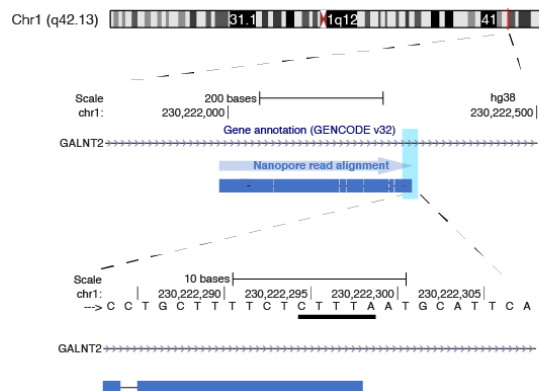
C “Human-CoV2-human” chimeric read (Nanopore) alignment on the SARS-CoV-2 genome



D “Human-CoV2” chimeric read (Nanopore)



E “Human-CoV2” chimeric read (Nanopore) alignment on Human Chr1



F “Human-CoV2” chimeric read (Nanopore) alignment on the SARS-CoV-2 genome

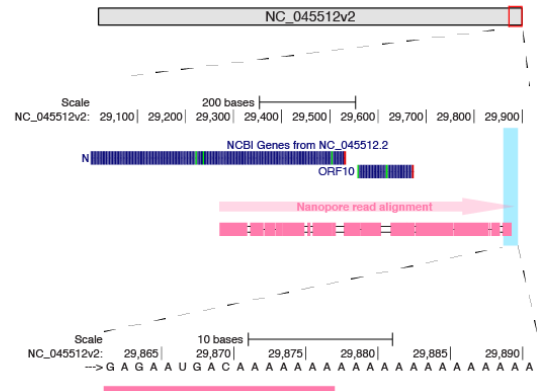


Fig. S2. Nanopore sequencing reads provide evidence for integration of SARS-CoV-2 sequences. **A)** A Nanopore sequencing read showing integration of a SARS-CoV-2 NC sub-genomic RNA sequence (magenta) and human genomic sequences (blue) flanking both sides of the integrated viral sequence. Features indicative of LINE1 mediated “target-primed reverse transcription” include: the target site duplication (yellow highlight) and the LINE1 endonuclease recognition sequence (underlined). Sequences that could be aligned to both genomes are shown in purple and sequences that cannot be aligned are shown in black. Arrows indicate sequence orientations with regard to the human and SARS-CoV-2 genomes as shown in **B, C**), with the two sides labeled as “a” and “b” in blue. **B)** Alignment of the Nanopore read in **A)** with the human genome (chromosome 22) showing the integration site. The human sequences at the junction region show the target site which was duplicated when the SARS-CoV-2 cDNA was integrated (yellow highlight) and the LINE1 endonuclease recognition sequence (underlined). **C)** Alignment of the Nanopore read in **A)** with the SARS-CoV-2 genome showing the integrated viral DNA represents a DNA copy of a portion of the NC sub-genomic RNA. Light blue highlighted regions are enlarged to show the two ends of the read. **D)** A Nanopore sequencing read showing the integrated portion of a SARS-CoV-2 RNA sequence (magenta) and human genomic sequences (blue) from one side of the junction with a LINE1 endonuclease recognition sequence (underlined). Sequences that could be mapped to both genomes are shown in purple. Arrows indicate sequence orientations with regard to the human and SARS-CoV-2 genomes. **E)** Alignment of the Nanopore read in **D)** with the human genome (chromosome 1) showing the integration site. The light blue highlighted region is enlarged to show a LINE1 endonuclease recognition sequence (underlined). **F)** Alignment of the Nanopore read **D)** with the SARS-CoV-2 genome showing the integrated viral sequence. The light blue highlighted region is enlarged to show the 3’ end of the viral sequence at the junction with human sequence.

**SARS-CoV-2 DNA read coverage
(Illumina paired-end whole genome sequencing)**

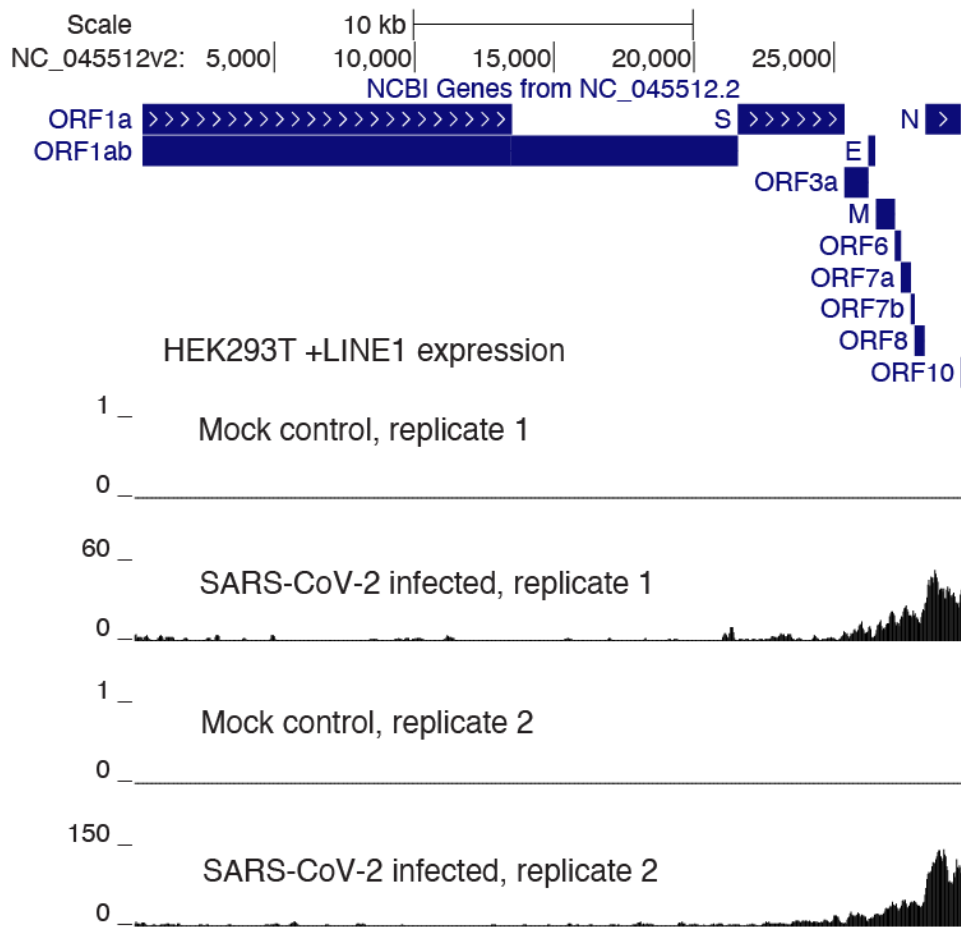


Fig. S3. Reverse transcribed viral sequences are predominant from the 3' end of SARS-CoV-2 genome. Viral reads were obtained using Illumina whole-genome paired-end sequencing of DNA from HEK293T cells that overexpressed LINE1. Genomic tracks showing the number of viral reads binned at 10 bp.

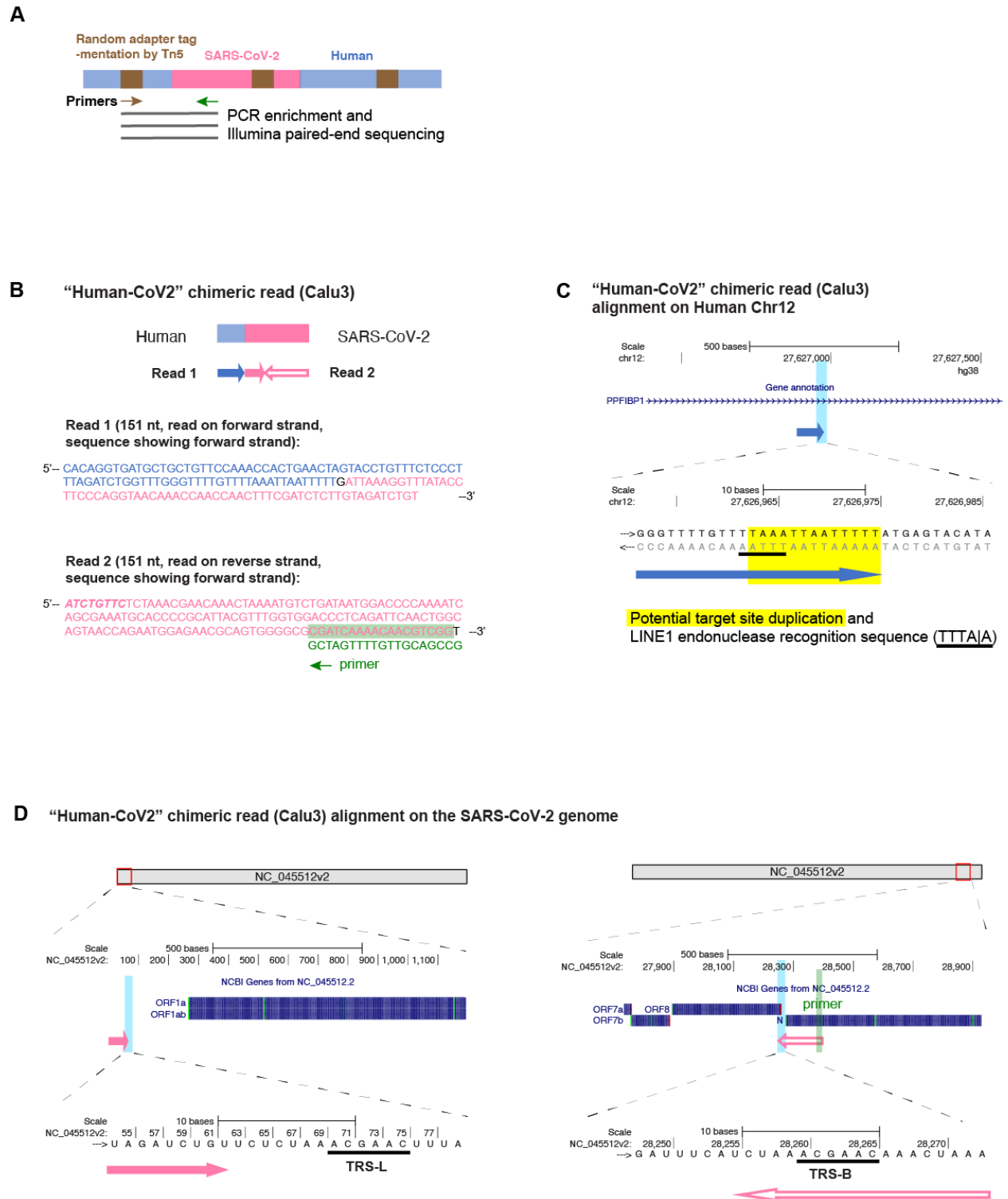


Fig. S4. Evidence for integration of SARS-CoV-2 cDNA in Calu3 cells. A) Experimental design for the Tn5 tagmentation mediated enrichment sequencing method used to map

integration sites in the host cell genome. The viral primer (reverse) was designed to target near-5' end of the viral NC gene (green arrow). **B)** A human-viral chimeric read pair supporting viral integration. The reads are aligned with the human (blue) and SARS-CoV-2 (magenta) genomic sequences. Arrows indicate read orientations relative to the human and SARS-CoV-2 genomes as shown in **C, D)**. Closed arrows show read 1 in the pair that was mapped to both human (blue) and SARS-CoV-2 (magenta) sequences. The open arrow (magenta) shows read 2 in the pair that was mapped to the SARS-CoV-2 genome. The sequence corresponding to the viral primer in read 2 is shown with green highlight (corresponding to the green arrow illustrated in **A)**. **C)** Alignment of the read pair in **B)** with the human genome (chromosome 12, blue arrow). The highlighted (light blue) region of the human sequence is enlarged to show the LINE1 recognition sequence (underlined) and the potential target site duplication (highlighted in yellow) that would be generated by LINE1 mediated retroposition. **D)** Alignment of the read pair in **B)** with the SARS-CoV-2 genome (magenta arrows). The closed arrow (left) corresponds to read 1 in **B)**, aligned to the viral leader sequence. The open arrow (right) corresponds to read 2 in **B)**, aligned to the NC gene body (the beginning 8 bases in this read is aligned to the viral leader sequence, shown in italics in **B)**). The highlighted (light blue) regions of the SARS-CoV-2 sequences are enlarged to show the TRS-L (left) and TRS-B (right) sequences (underlined, these are the sequences where the viral polymerase jumps to generate the sub-genomic RNA). The viral primer sequence is shown with green highlight.

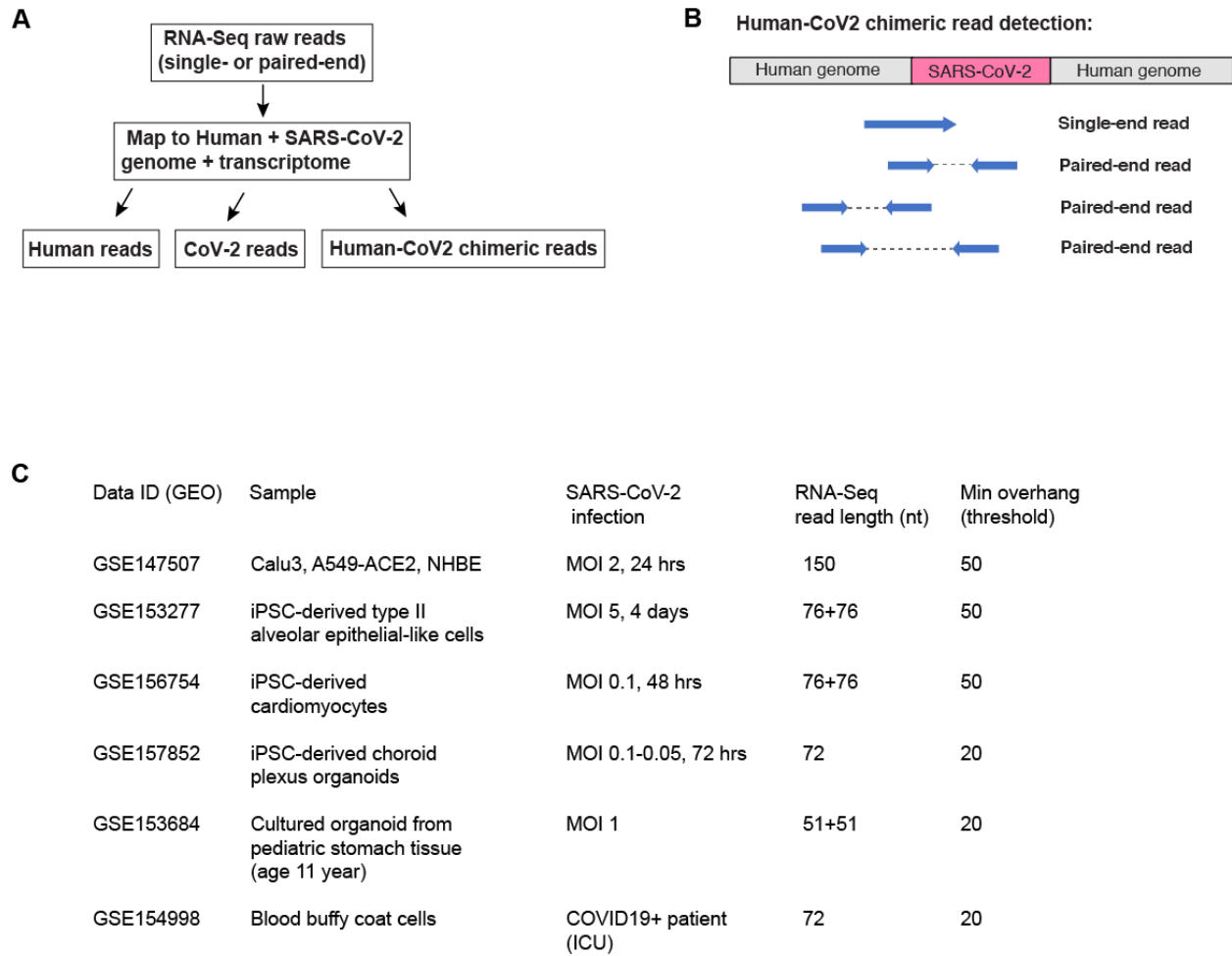


Fig. S5. Analysis of published data for human-viral chimeric transcripts. **A)** The pipeline used to identify human-CoV2 chimeric RNA-Seq reads. **B)** Schema showing human- SARS-CoV2 chimeric RNA-seq reads mapped to potential SARS-CoV-2 integration sites. **C)** Published data used to identify human-viral chimeric reads: Data ID (GEO accession number), sample type, infection method/type (MOI: Multiplicity of Infection), RNA-Seq format (single or paired-end with read length), and threshold to call chimeric reads (Min overhang: minimum number of bases mapped to either human or SARS-CoV-2 genome/transcriptome to call a chimeric reads).

A Human - CoV2 chimeric read from Calu3 (infected) RNA-Seq:

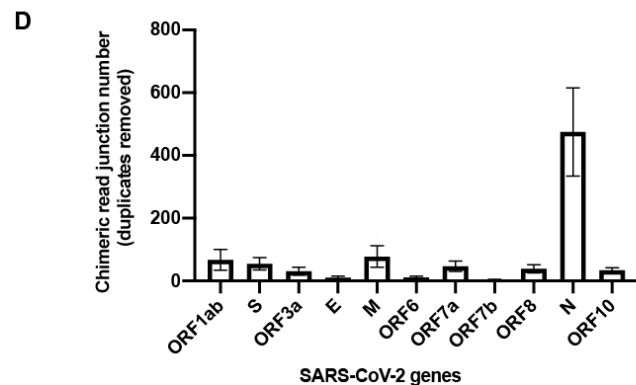
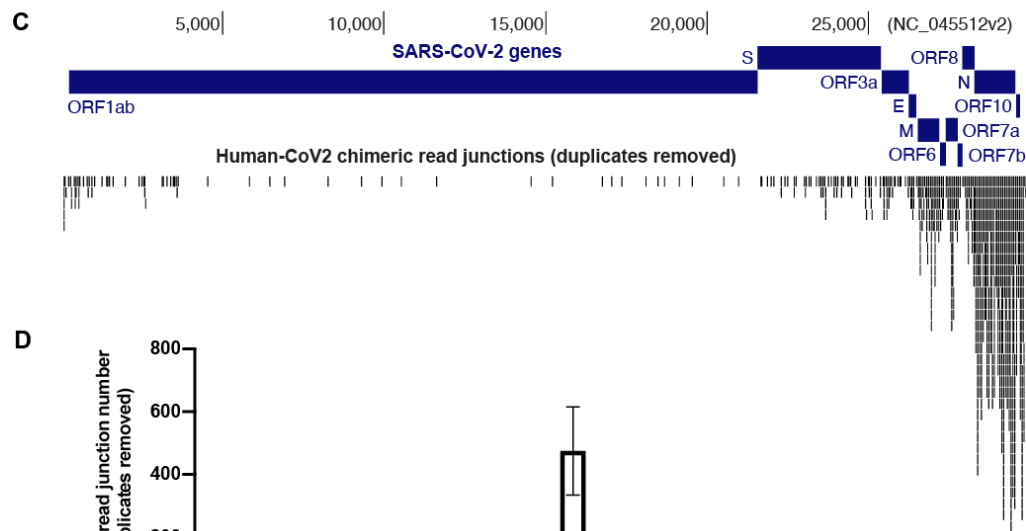
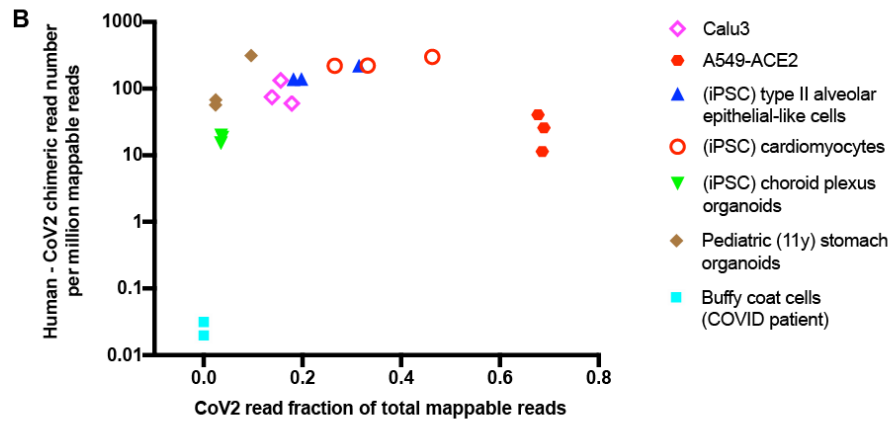
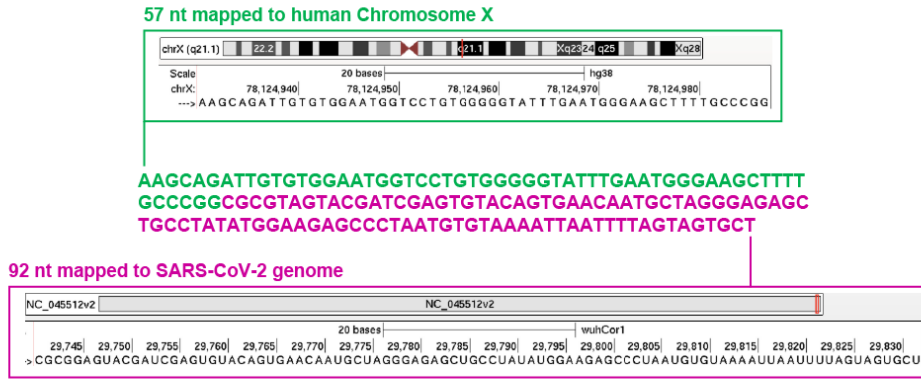


Fig. S6. Human-viral chimeric reads from published RNA-seq data. **A)** A chimeric RNA read (149 nt) from (SARS-CoV-2) infected Calu3 RNA-Seq with 57 nt mapped to human Chromosome X (green) and 92 nt (magenta) mapped to the SARS-CoV-2 genome. **B)** Scatter plot showing the number of human-CoV2 chimeric reads (per million total mappable reads, y-axis) versus the fraction of SARS-CoV-2 reads in total mappable reads (x-axis) in published RNA-Seq datasets from different SARS-CoV-2 infected samples. **C-D)** Human-CoV2 chimeric read junctions (duplicates removed) mapped to the SARS-CoV-2 genome (**C**) and distribution among SARS-CoV-2 genes (**D**, three biological replicates; mean \pm s.e.m.). RNA-Seq data is from SARS-CoV-2 infected Calu3 cells (GSE147507). A chimeric read junction is defined by the “break point” of the sequences mapped to human or SARS-CoV-2 genome/transcriptome in a given RNA-Seq read.

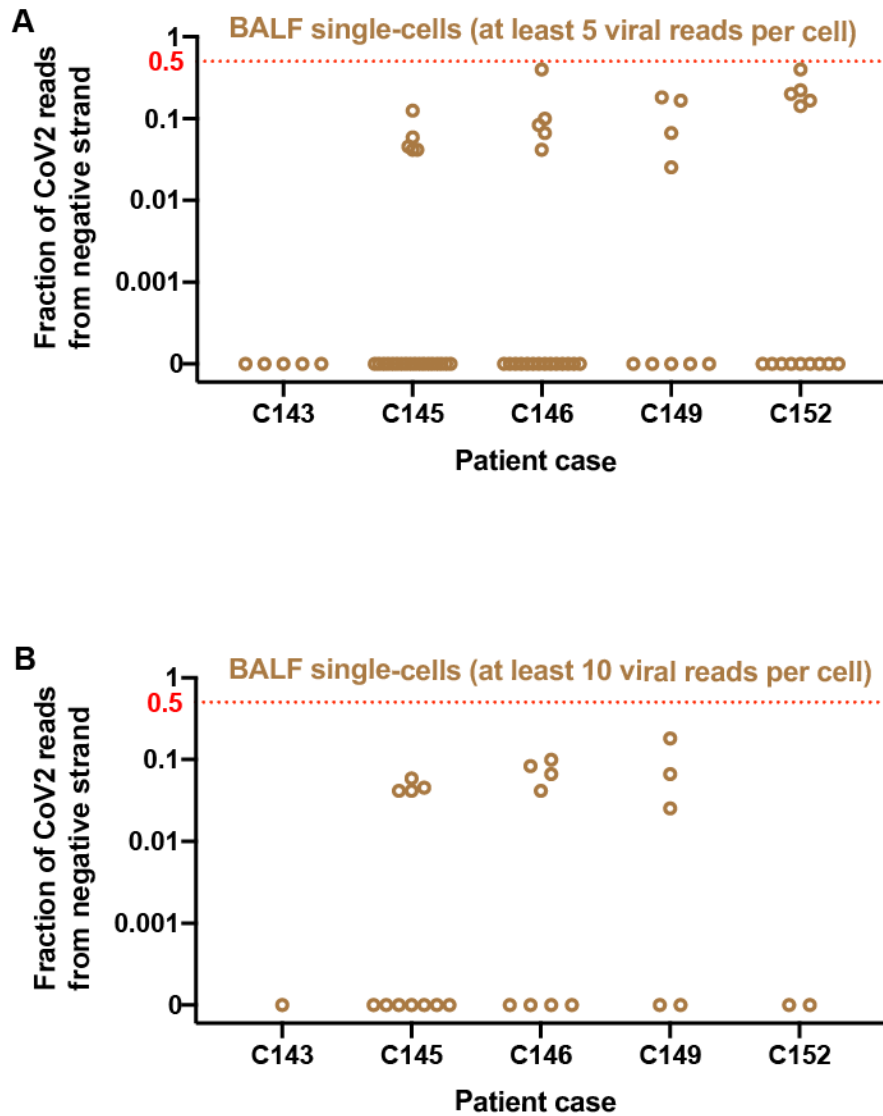


Fig. S7. Negative strand viral RNA-seq reads from patient single-cells suggest that integrated SARS-CoV-2 sequences are expressed. **A)** Fraction of viral reads that are derived from negative strand viral RNA in single BALF cells from patients with at least 5 viral reads per cell (published RNA-seq data, GSE145926). **B)** Fraction of viral reads that are derived from negative strand viral RNA in patient single BALF cells with at least 10 viral reads per cell (published RNA-seq data, GSE145926). Red dashed lines indicate the level at which 50% of all viral reads were from negative strand viral RNAs, a level expected if all the viral sequences were derived from integrated sequences.

Supplementary Tables

Table S1. Summary of negative strand viral and human-viral chimeric RNA-seq reads from acutely infected lung cells or organoids

Sample	CoV2 reads	Negative strand CoV2 read fraction	Human-CoV2 chimeric reads	Negative strand CoV2 (in chimeric RNAs) read fraction
Calu3, rep1	52,962,587	0.08%	8,170	0.81%
Calu3, rep2	61,256,542	0.10%	6,218	0.76%
Calu3, rep1 (Blanco-Melo et al.)	32,10,542	0.01%	4,430	0
Calu3, rep2 (Blanco-Melo et al.)	2,378,641	0.01%	1,859	0
Calu3, rep3 (Blanco-Melo et al.)	4,320,681	0.01%	9,702	0.01%
Lung organoid, rep1 (Han et al.)	615	0	1	0
Lung organoid, rep2 (Han et al.)	12,752	0.04%	15	0
Lung organoid, rep3 (Han et al.)	1,320	0.08%	2	0

Table S2. Summary of negative strand viral and human-viral chimeric RNA-seq reads from tissues of deceased COVID-19 patients (published RNA-seq data, Desai et al., GSE150316)

Sample	GEO accession number	CoV2 reads	Negative strand CoV2 read fraction	Human-CoV2 chimeric reads	Negative strand CoV2 (in chimeric RNAs) read fraction
Case1-lung1 LUL	GSM4546576	51,4418	6.7%	108	1.9%
Case1-lung2 RML	GSM4546577	54,598	8.4%	9	0.0%
Case1-lung3 RUL	GSM4546578	20,746	13.0%	6	0.0%
Case1-lung4 LLL	GSM4546579	37,232	2.4%	4	0.0%
Case2-lung1 RLL	GSM4546581	483	12.4%	0	
Case2-lung2 LUL	GSM4546582	42	0.0%	0	
Case2-jejunum1	GSM4546583	10	0.0%	0	
Case2-lung3 RUL	GSM4546584	10	0.0%	0	
Case3-lung1 LUL	GSM4546586	16	25.0%	0	
Case3-lung2 RLL	GSM4546588	4	0.0%	0	
Case5-lung1 LLL	GSM4546596	220	0.0%	0	
Case5-lung2 RML	GSM4546597	38	47.4%	0	
Case5-lung3 LUL	GSM4546598	648	0.9%	0	
Case5-lung4 RML	GSM4546599	514	0.4%	0	
Case5-lung5 RUL	GSM4546601	722	4.4%	0	
Case5-liver1	GSM4546604	6	0.0%	0	
Case6-lung1 LUL	GSM4698531	14	0.0%	0	
Case7-lung5 LUL	GSM4698540	1,284	0.6%	0	
Case8- bowel1	GSM4698541	24	0.0%	0	
Case8- heart1	GSM4698542	78	0.0%	0	
Case8-lung1 RLL	GSM4698544	3,150	5.0%	0	
Case8-lung2 RUL	GSM4698545	200	51.0%	0	
Case8-lung3 LLL	GSM4698546	24,527	1.8%	46	0.0%
Case8-lung4 RML	GSM4698547	5,820	2.1%	0	
Case8-lung5 LUL	GSM4698548	102	0.0%	0	
Case9- lung1RLL	GSM4698549	45,539	6.4%	56	5.4%
Case9- lung2 RML	GSM4698550	154,157	9.6%	405	42.5%
Case9- lung3RUL	GSM4698551	138,578	8.1%	173	3.5%
Case9-lung4 LUL	GSM4698552	361,535	4.8%	652	5.8%
Case9- lung5LLL	GSM4698553	179,729	14.5%	145	35.2%
Case10- lung2 LLL	GSM4698522	112	0.0%	0	
Case10- lung3 RLL	GSM4698523	22	0.0%	0	

Case11 -bowel1	GSM4698524	92	0.0%	0	
Case11-lung1RML	GSM4698526	72,029	3.2%	141	2.8%
Case11-lung32RUL	GSM4698527	17,256	12.7%	13	0.0%
Case11-lung3RLL	GSM4698528	1,328	3.5%	0	
CaseA-lung	GSM4698554	1,202	4.0%	0	
CaseB-lung	GSM4698555	6	33.3%	0	
CaseC-lung	GSM4698556	168,935	1.5%	47	0.0%
CaseD-lung	GSM4698557	43,750	6.8%	9	0.0%

Table S3. Summary of negative strand viral and human-viral chimeric RNA-seq reads from BALF cells of COVID-19 patients (bulk analysis of published single-cell RNA-seq data, Liao et al., GSE145926)

Patient	GEO accession number	CoV2 reads	Negative strand CoV2 read fraction	Human-CoV2 chimeric reads	Negative strand CoV2 (in chimeric RNAs) read fraction
C143 (severe)	GSM4339771	1,525	18.6%	1	0.0%
C145 (severe)	GSM4339773	34,439	12.7%	16	0.0%
C146 (severe)	GSM4339774	893,327	15.3%	223	1.4%
C148 (severe)	GSM4475051	1,251	22.0%	0	
C149 (severe)	GSM4475052	89,928	19.6%	23	0.0%
C152 (severe)	GSM4475053	32,665	18.5%	5	0.0%

Table S4. PCR primers used in this study

Name	Sequences
N1	Forward: GACCCCAAATCAGCGAAAT Reverse: TCTGGTACTGCCAGTTGAATCTG
N2	Forward: GGGAGCCTTGAATACACCAAAA Reverse: TGTAGCACGATTGCAGCATTG
N3	Forward: GGGGAACCTTCTCCTGCTAGAAT Reverse: CAGACATTTTGCTCTCAAGCTG
N4	Forward: AAATTTTGGGGACCAGGAAC Reverse: TGGCACCTGTGTAGGTCAAC
N (for cloning complete NC gene)	Forward: ATGTCTGATAATGGACCCCAAAT Reverse: TTAGGCCTGAGTTGAGTCAGC
<i>HSPA1A</i>	Forward: ATCTCCACCTTGCCGTGTT Reverse: ATCCAGTGTTCCGTTTCCAG

Dataset S1 (separate file). Sanger sequencing results of the complete SARS-CoV-2 NC gene cloned from large-fragment cell genomic DNA from LINE1-overexpressing, SARS-CoV-2-infected HEK293T cells.

Dataset S2 (separate file). Summary of chimeric read sequences from Nanopore sequencing of DNA from LINE1-overexpressing, SARS-CoV-2-infected HEK293T cells.

Dataset S3 (separate file). Summary of chimeric sequences from Illumina paired-end whole genome sequencing of DNA from LINE1-overexpressing, SARS-CoV-2-infected HEK293T cells.

Dataset S4 (separate file). Summary of chimeric sequences from Tn5 tagmentation-mediated DNA integration site enrichment sequencing of DNA from SARS-CoV-2 infected HEK293T or Calu3 cells.

Update for Zhang et al., Reverse-transcribed SARS-CoV-2 RNA can integrate into the genome of cultured human cells and can be expressed in patient-derived tissues.

In our publication¹, we reported a total of 63 integrations of SARS-CoV-2 sequence in the genome of HEK293T cells overexpressing LINE1, by Nanopore whole-genome sequencing (Table 1). We recovered 2 integrations with human sequences flanking a viral sequence on both sides (Fig. 1B-D, S2A-C), showing a 20-bp or 13-bp target-site duplication as strong evidence for LINE1 mediated integration^{2,3}. For the other 61 integrations we recovered only one-side flanking the human sequence. As we reported, ~67% of all 63 integrations included a LINE1 endonuclease recognition sequence⁴ at the flanking human sequence that were either directly linked to the 3' end (poly-A tail) of the viral sub-genomic RNA sequence, or within a potential target site duplication (8-27 bp) that were linked to the 5' end of viral sequences, as evidence for LINE1 mediated integration. Another prediction of LINE1-mediated retrointegration is the presence of a poly-A tract at the 3' end of integrant⁵⁻⁹. As summarized in the table we found “poly-A tracts” ranging from 2 – 65 bp for these integrations at the junctions of the LINE1 endonuclease recognition sequence (e.g. 5'-TTTT|A-3') and the shared 3' end of the integrated viral sub-genomic RNA sequence (lower strand: 5'-polyT-GTCATTCTC...-3'). This supports a LINE1 mediated integration mechanism. The figure details the sequences of all integrants.

Table S5. Summary of Nanopore chimeric sequences with 3'-end of SARS-CoV-2 sub-genomic RNA sequence directly linked to a LINE1 endonuclease recognition sequence in the human genome

	Chromosome	L1 endonuclease recognition sequence ⁴	"poly-A tract" length (bp)*	Genomic feature	Illustrated in figures ¹
1	chr1	CTTT A	7	intron (GALNT2)	Fig. S2D-F
2	chr1	CTTT C	31	intron (PEX14)	
3	chr1	TCTT A	21	intron (AL359924.1, non-coding RNA)	
4	chr1	TTTTTCTTT A	40	exon (ZNF644)	
5	chr1	TTTATTTT A	21	intron (KIAA0319L)	
6	chr2	TTTTATTT A	3	intron (EML4-AS1, non-coding RNA)	
7	chr3	TTTCTT C(?)	16	intron (AC012020.1, non-coding RNA)	
8	chr4	TTTTCT G	35	intron (AC097528.1, non-coding RNA)	
9	chr5	TTTT A	2	intron (CCDC127)	
10	chr6	TTCT A	4	intergenic	
11	chr6	TTTTC A	7	intergenic	
12	chr7	TTTTT C	8	intron (NRCAM)	
13	chr7	AGTT C	2	intergenic	
14	chr9	TTTT C	19	intergenic	
15	chr9	TTGTT A	4	exon (3'UTR) (SCAI)	
16	chr10	TTTTTTT C	16	intergenic	
17	chr11	TTTTC A	8	exon (NEAT1, non-coding RNA)	
18	chr11	TTTCTT A	51	intron (TRIM44)	
19	chr12	TTTTTC A	22	intron (AC048352.1, LINC02378, non-coding RNAs)	
20	chr12	TTTT C	65	intergenic	

21	chr12	TTTT A	3	intergenic	
22	chr12	TTTTCTT A	4	intron (POLR3B)	
23	chr13	TTCT A	13	intergenic	
24	chr13	TCTT A	5	intergenic	
25	chr14	TTTTTTTTTTTTTT G	18	intergenic	
26	chr16	TTTT A	10	intron (CDR2)	
27	chr18	TTTTCTT G	21	intron (LINC01915, non-coding RNA)	
28	chr22	TTCTT A	10	intron (IGLV10-54)	Fig. S2A-C
29	ChrX	TTTT A	33	intron (DMD)	
30	ChrX	TTTTTT C	36	intergenic	
31	ChrX	TTCT A	4	intergenic	Fig. 1B-D
32	ChrX	TTTTCTT C(?)	61	exon (SLITRK2)	

*The “poly-T” of the human sequence at the junction is counted. This is the site where the viral RNA “poly-A” sequence annealed for “target-primed reverse transcription”.

References

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Chr1:

Read: 07573090-2aaf-49c6-8a1d-d094d8a66847

Human: ...CTCGGCCTCTGCTTTTCT**CTTTA**...

Chi read: 5'...CTCGGCCTCTGCTTTTCTCTTTTTTTTGTCAATTCTCCT...3'

CoV2: **TTTTTTT**GTCAATTCTCCT...

LINE1 endonuclease recognition sequence: CTTT|A

"PolyA tract" length: 7 bp

Integration site: intron (GALNT2)

Read: e05cbd8d-36b5-4dc2-b54b-f5ef36ca35a0

Human: ...CAGTGGCGCTGGCTTCTGC**CTTTC**...

Chi read: 5'...CAGTGGCGCTGGCTTCTGCCTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTGTCAATTCTCCT...3'

CoV2: **TTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTT**GTCAATTCTCCT...

LINE1 endonuclease recognition sequence: CTTT|C

"PolyA tract" length: 31 bp

Integration site: intron (PEX14)

Read: be93c20e-0591-4e2b-a633-df92b6a36e04

Human: ...CAAATGAACAAATACAT**TCTTA**...

Chi read: 5'... CAAATGAACAAATACATCTTGATTTTTTTTTTTTTTTTTTTTTTTTGTCAATTCTC...3'

CoV2: **T TT TTTTTTTTTTTTTTTTTTTTTTTTTTT**GTCAATTCTC...

LINE1 endonuclease recognition sequence: TCTT|A

"PolyA tract" length: 21 bp

Integration site: intron (AL359924.1, non-coding RNA)

Read: 80097bd4-2c11-40f8-9238-59826c54606b

Human: ...AAGGATGTCAATTTTT**CTTTA**...

Chi read: 5'...AAGGATGTCAATTTTTCTTGTCAATTCTCCT...3'

CoV2: **TTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTT**GTCAATTCTCCT...

LINE1 endonuclease recognition sequence: TTTTTCTTT|A

"PolyA tract" length: 40 bp

Integration site: exon (ZNF644)

Read: f40cad59-4beb-49a0-87c5-7642024de8e7

Human: ...AACAAAGACCAATTTATTTTTA...

Chi read: 5'...AACAAAGACCAATTTATTTTTTTTTTTGTTTTTTTTTTTGTTCATTCTCCTA...

CoV2: TTTTTTTT TTTTTTTTTTTTGTTCATTCTCCTA...

LINE1 endonuclease recognition sequence: TTTATTTTT|A

"PolyA tract" length: 21 bp

Integration site: intron (KIAA0319L)

Chr2:

Read: 40d42d5b-7125-4200-8d88-0ea079267ab0

Human: ...TCATGTCAGAAGAACTGCTTTTATTTA...

Chi read: 5'...TCATGTCAGAAGAACTGC TTTATTTGTCATTCTCCTAAGA...3'

CoV2: TTTGTCATTCTCCTAAGA...

LINE1 endonuclease recognition sequence: TTTTATTT|A

"PolyA tract" length: 3 bp

Integration site: intron (EML4-AS1, non-coding RNA)

Chr3:

Read: d9ff2a50-f4fb-482c-ba21-6a54d47c7bf1

Human: ...TTCAGGATCACCTTCTTC...

Chi read: 5'...TTCAGGATCACCTTCTTTTTTTTTTTTTTTGTCATTCTCCT...3'

CoV2: TTT TTTTTTTTTTTTTTGTTCATTCTCCT...

LINE1 endonuclease recognition sequence: TTTCTT|C

"PolyA tract" length: 16 bp

Integration site: intron (AC012020.1, non-coding RNA)

Chr4:

Read: 28ea733a-fb5f-484a-8f69-c60135693069

Human: ...AGAGACTTTCTG...

Chi read: 5'...AGAGACTTTCTGTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTGTTCATTCTC...3'

CoV2: TTTT T TTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTTGTTCATTCTC...

LINE1 endonuclease recognition sequence: TTTTCT|G

"PolyA tract" length: 35 bp

Integration site: intron (AC097528.1, non-coding RNA)

Chr5:

Read: da820d14-8fdd-400b-b319-751055955437

Human: ...TAGTCAGAGTAA**TTTTA**...

Chi read: 5'...TAGTCAGAGTAA TTGTCATTCT CTAAGAAGCTATT...3'

CoV2: **TT**GTCATTCTCCTAAGAAGCTATT...

LINE1 endonuclease recognition sequence: TTTT|A

"PolyA tract" length: 2 bp

Integration site: intron (CCDC127)

Chr6:

Read: 5b2b7b8d-79ec-4378-a055-246618f76bb4

Human: ...TGGAAGTATTAT**TTCTA**...

Chi read: 5'...TGGAAGTATTAT**TTCT**TGTCATTCTCCTAAG...3'

CoV2: **TT TT**GTCATTCTCCTAAG...

LINE1 endonuclease recognition sequence: TTCT|A

"PolyA tract" length: 4 bp

Integration site: intergenic

Read: fb8bd42f-a1d6-4e52-99ff-f2c0d021a59f

Human: ...GGATGCTTCAGTTAT**TTTCA**...

Chi read: 5'...GGATGCTTCAGTTATTTTCTTTGTCATTCTCCTAAG...3'

CoV2: **TTTT TTT**GTCATTCTCCTAAG...

LINE1 endonuclease recognition sequence: TTTTC|A

"PolyA tract" length: 7 bp

Integration site: intergenic

Chr7:

Read: d296e986-48f2-4d72-93e6-b53456f8e1c4

Human: ...ATGCATTGATTTTTGAACAGT**TTTTTC**...
Chi read: 5'...ATGCATTGAT TTTTGAACAGTTTTTTTTTGTTCATTCTCCTAAGAAG...3'
CoV2: **TTTTTTTT**GTCATT CTCCTAAGAAG...

LINE1 endonuclease recognition sequence: TTTTT|C

"PolyA tract" length: 8 bp

Integration site: intron (NRCAM)

Read: 9bf30fba-3195-4241-b2e3-e5ab595e6e1d

Human: ...CACCAGTGG**AGTTC**...
Chi read: 5'...CACCAGTGGAGTTGTCATTTTCTCCTAAGAAGCTATTAA...3'
CoV2: **TT**GTCA TTCTCCTA GAAGCTATTAA...

LINE1 endonuclease recognition sequence: AGTT|C

"PolyA tract" length: 2 bp

Integration site: intergenic

Chr9:

Read: 7a893df8-8b8f-47fd-9286-b7d4e7ebee4e

Human: ...CATAGAACTGGGTGGCTTAA**TTTTTC**...
Chi read: 5'...CATAGAACTGGGTGGCTTAATTTTTTTTTTTTTTTTTTTTTGTTCATTCTC TAAGAAGCTATT...3'
CoV2: **TTTTTTTTTTTTTTTTTTTT**GTCATTCTCCTAAGAAGCTATT...

LINE1 endonuclease recognition sequence: TTTT|C

"PolyA tract" length: 19 bp

Integration site: intergenic

Read: 6a43b132-635c-4c9b-9e29-04790f6770b1

Human: ...CAGACAAGCACAAT**TGTTA**...
Chi read: 5'...CAGACAAGCACAATTGTTGTCATTCTCCTA...3'
CoV2: **TT TG**TTCATTCTCCTA...

LINE1 endonuclease recognition sequence: TTGTT|A

"PolyA tract" length: 4 bp

Integration site: exon (3'UTR) (SCAI)

Chr10:

Read: 8975c59e-bc5e-4d71-ae28-55d1819e8032

Human: ...TACTTTCTTGCATATGATTT**TTTT**C...

Chi read: 5'...TACTTTCT GCATATGATTTTTCTTTTTTTTTTTTTGTTCATTCTCC...3'

CoV2: **TTTT TTTTTTTTTT**GTCATTCTCC..

LINE1 endonuclease recognition sequence: TTTTTTT|C

“PolyA tract” length: 16 bp

Integration site: intergenic

Chr11:

Read: b365ab7a-6514-459b-bd3c-a4689f41d9ab

Human: ...CAATTTCACTTCGCTCATGAT**TTTCA**...

Chi read: 5'...CAATTTCACTTCGCTCATGATTTTCTTTGTTCATTCTCCTAAG...3'

CoV2: **TTTT TTTT**GTCATTCTCCTAAG...

LINE1 endonuclease recognition sequence: TTTTC|A

“PolyA tract” length: 8 bp

Integration site: exon (NEAT1, non-coding RNA)

Read: c90e0881-3da9-4730-be43-227d9a2cbb05

Human: ...AGAGTGTAGTGTT**CTTA**...

Chi read: 5'...AGAGTGTAGTGTTCTTTGTTCATTCTCCT...3'

CoV2: **TTT TT**GTCATTCTCCT...

LINE1 endonuclease recognition sequence: TTTCTT|A

“PolyA tract” length: 51 bp

Integration site: intron (TRIM44)

Chr12:

Read: 17c45e22-69ae-4f6e-93b4-31d3a81d6f9b

Human: ...GAATTTCCAGGCTGTT**TTTCA**...

Chi read: 5'...GAATTTCCAGGCTG TTTCTTTTTTTTTTTTTTTTTTTTTGTTCACATTCTCCTAAGAAGCTAT...3'

CoV2: **TTTT TTTTTTTTTTTTTTTTTT**GT CATTCTCCTAAGAAGCTAT...

LINE1 endonuclease recognition sequence: TTTTTC|A

"PolyA tract" length: 22 bp

Integration site: intron (AC048352.1, LINC02378, non-coding RNAs)

Read: 69963961-b72e-4193-8fd9-29350e837663

Human: ...ACCTCCATTGCTTTTTTC...

Chi read: 5'...ACCTTCTATTGTCTTTTTTCTTTTTTTTTT(Tx30)TTTTTTTTTTTTTTTTTTTGTTCATTCTCCTAAGAA...3'

CoV2: TTTTTT TTTTTTTTTT(Tx30)TTTTTTTTTTTTTTTTTTTGTTCATTCTCCTAAGAA...

LINE1 endonuclease recognition sequence: TTTT|C

"PolyA tract" length: 65 bp

Integration site: intergenic

Read: 0e6049f7-fdd2-49bd-b35b-131ed9791e88

Human: ...GCCTTTTCTGTTTCCTTTAAATTTTTA...

Chi read: 5'...GCCTTTTCTGTTTCCTTTAAA TTTGTTCATTCTCCTAAGAAGCT...3'

CoV2: TTTGTTCATTCTCCTAAGAAGCT...

LINE1 endonuclease recognition sequence: TTTTT|A

"PolyA tract" length: 3 bp

Integration site: intergenic

Read: c9e4ccb2-7edc-4535-af71-504f0670cf04

Human: ...TGTGCCACATTTTCTTA...

Chi read: 5'...TGTGCCACATCTTCTGTCAATTCTCCTAAGAAGCTATTA...3'

CoV2: TT TTGTCA TTCTCCTAAGAAGCTATTA...

LINE1 endonuclease recognition sequence: TTTTCTT|A

"PolyA tract" length: 4 bp

Integration site: intron (POLR3B)

Chr13:

Read: 94f90063-f0d8-4a56-9f78-cea4b239b5fa

Human: ...ACATTAGAGTTTTCTTCTA...

Chi read: 5'...ACATTAGAGTTTTCTTCTATTTAAGTTTTTTTGTTCATTCTCCTAAGAAGC...3'

CoV2: TT T TTT TTTTTTGTTCATTCTCCTAAGAAGC...

LINE1 endonuclease recognition sequence: TTCT|A

"PolyA tract" length: 13 bp

Integration site: intergenic

Read: 6e548530-47d4-4b2f-8665-b2107fc66d32

Human: ...GGTATTTCTAGGTTATCTTA...

Chi read: 5'...GGTATTTCTAGGTTATCTTATTGTCATTCTCCT...3'

CoV2: T TT TTGTCATTCTCCT...

LINE1 endonuclease recognition sequence: TCTT|A

"PolyA tract" length: 5 bp

Integration site: intergenic

Chr14:

Read: 50eef64-c7a5-4966-a519-3e5c611cc423

Human: ...CACATTGTTTGCTTTTTTTTTTTTTTTTGG...

Chi read: 5'...CACATTGTTTGCTTTTTTTTTTTTTTTTTTTGTC TACTCCTA GAAGC GTAAAAATCACATGGGGATAG...3'

CoV2: TTTTTTTTTTTTTTTTTTGTGTCATTCTCCTAAGAAGCTATTAATAATCACATGGGGATAG...

LINE1 endonuclease recognition sequence: TTTTTTTTTTTTTTTT|G

"PolyA tract" length: 18 bp

Integration site: intergenic

Chr16:

Read: 0c381657-f571-4cdc-b8de-17bc6a224ade

Human: ...AAAGCAACTTTTTA...

Chi read: 5'...AAAGCAACTTTTTTTTTTTGTCATTCTCCTA...3'

CoV2: TTTTTTTTTTGTGTCATTCTCCTA...

LINE1 endonuclease recognition sequence: TTTTT|A

"PolyA tract" length: 10 bp

Integration site: intron (CDR2)

Chr18:

Read: 028774db-8163-4dde-b7af-16c7d7b7cde2

Human: ...TCAACTTCAATTTATTTTTTAAAATTTCTTG ...
Chi read: 5'...TCAACTTCAATTTATTTTTTAAAATTTCTTCCTTTTTTTTTTTTTTTTGTTCATTCTCCTAAGAAGCTATT...3'
CoV2: TTTT TT TTTTTTTTTTTTTTTTGTTCATTCTCCTAAGAAGCTATT...

LINE1 endonuclease recognition sequence: TTTTCTT|G

"PolyA tract" length: 21 bp

Integration site: intron (LINC01915, non-coding RNA)

Chr22:

Read: f40cad59-4beb-49a0-87c5-7642024de8e7

Human: ...AGGTTTGGTGACATCACTTCTTA...
Chi read: 5'...AGGTTTGGTGACATCACTTCTTTTTTTTGTTCATTCTC TAAGAAGCTATT...3'
CoV2: TT TTTTTTTTGTTCATTCTCCTAAGAAGCTATT...

LINE1 endonuclease recognition sequence: TTCTT|A

"PolyA tract" length: 10 bp

Integration site: intron (IGLV10-54)

ChrX:

Read: 1233b662-c642-472b-9529-73286cc3f473

Human: ...TTAGGAATGCATATTCTGATTTTTA...
Chi read: 5'...TTAGGAATGCATATTCTGATTGTTCATTCTCCTAAGAAG...3'
CoV2: TTGTTCATTCTCCTAAGAAG...

LINE1 endonuclease recognition sequence: TTTTT|A

"PolyA tract" length: 33 bp

Integration site: intron (DMD)

Read: 502e6323-b284-442e-a039-c28e14a92c06

Human: ...GACTTTTTGCACCTTTTTTC...
Chi read: 5'...GACTTTTTGCACCTTGTTCATTCTCCTAAGAAG...3'
CoV2: TTGTTCATTCTCCTAAGAAG...

LINE1 endonuclease recognition sequence: TTTTTT|C

"PolyA tract" length: 36 bp

Integration site: intergenic

Read: b1c4ffb8-480b-4a10-9461-6f1393119003

Human: ...TCAATTGCTATTGCTTCTA...

Chi read: 5'...TCAATTGCTATTGCTTCTTGTTCATTCTC TAAGAA...3'

CoV2: TT TTGTCATTCTCCTAAGAA...

LINE1 endonuclease recognition sequence: TTCT|A

"PolyA tract" length: 4 bp

Integration site: intergenic

Read: 4757692e-e0cd-46b5-b3b2-163dc1d6a2ad

Human: ...TATTCAGTACGTTTTCTTC...

Chi read: 5'...TATTCAGTACGTTTTCTTC(Tx25)TTTTTTTTTTTTTTTTTGTTCATTCTCCTAA...3'

CoV2: TTTT TTTTTTTTTTTTTT(Tx25)TTTTTTTTTTTTTTTTTGTTCATTCTCCTAA...

LINE1 endonuclease recognition sequence: TTTTCTT|C

"PolyA tract" length: 61 bp

Integration site: exon (SLITRK2)