

1 **SUPPLEMENTARY INFORMATION**

2

3 **Supplementary Figure S1. Effect of 1C8 on viral RNA levels in the 24NLESG cell** 4 **line.**

5 Schematic of HIV-1 provirus in the 24NLESG cell line is shown on top. HIV-1 was
6 induced in 24NLESG cells by addition of PMA in the presence of DMSO or 10 μ M 1C8.
7 After 24 h, cells were harvested and total RNA extracted. HIV-1 unspliced (US) RNA
8 levels were assayed using the Aurum Total RNA Lysis Kit (Bio-Rad), as per
9 manufacturer's instructions with the addition of Turbo DNase (Ambion). Purified RNA
10 (0.5-2 μ g) was reverse transcribed using M-MLV (Invitrogen). HIV-1 and actin mRNA
11 levels in DMSO- and compound-treated samples were quantified by qPCR using the
12 Mastercycler ep realplex (Eppendorf). Gene quantification was evaluated using the
13 absolute quantification method, normalized to β -actin expression, and expressed relative
14 to DMSO treatment.

15

16 **Supplementary Figure S2. 1C8 does not affect hTRA2 β and hnRNP F protein** 17 **expression.**

18 Western blot analysis of HeLa-HIV cells treated with different concentrations of 1C8.
19 Protein extracts were fractionated by SDS-PAGE and transferred to a nitrocellulose
20 membrane that was decorated with anti-hTRA2 β and hnRNP F antibodies.

21

22 **Supplementary Figure S3. RT-PCR analysis of splicing changes induced by 1C8.**

23 Scatter plot comparing the percent splicing index (PSI) for 96 alternative splicing events
24 in HeLa-HIV cells treated (*y*-axis) or not (*x*-axis) with 10 μ M of 1C8. Values were
25 obtained by quantitating amplification products obtained endpoint RT-PCR and
26 fractionated on a Caliper LC-90 automated microfluidic station. Each point represents an
27 average of triplicates with standard variations shown. Diagonals that run parallel to the
28 central diagonal depict a \pm 10% PSI window. *q*-values for the 1C8-induced difference
29 are shown in Supplementary Table S1. The red dot represents the only splicing event
30 (*INF2*) that is shifted with a *q*-value \leq 0.01. White dots represent events with *q*-values \leq
31 0.05 as listed in Supplementary Table S1.

32

33 **Supplementary Figure S4. RNA-Seq analysis of splicing changes induced by 1C8.**

34 Scatter plot comparing the percent splicing index (PSI) of more than 62 000 alternative
35 splicing events in HeLa-HIV cells not treated (*x*-axis) or treated (*y*-axis) with 1, 5, 10 and
36 20 μ M of 1C8. Diagonals that run parallel to the central diagonal depict a \pm 30% PSI
37 window. All replicates of a given treatment were merged into a single file before MISO
38 analysis of RNA-Seq data. The analysis was performed for all concentrations of 1C8,
39 revealing a total of 86 unique events significantly affected (red dots indicating Δ PSI
40 larger than 30 percentage points and with Bayesian coefficient superior to 20 ; see
41 Supplementary Table S2 for a list of significant events).

42

43 **Supplementary Figure S5. Validation of five cellular splicing events affected by** 44 **1C8.**

45 **A.** Graphic presentation of the splicing unit (bottom of the left panel) and the results of
46 the RNA-Seq and MISO analysis for *ITGB3BP* unit from samples of HeLa-HIV cells
47 treated with 1C8 (top of the left panel). The percent splicing index (PSI) of the *ITGB3BP*
48 unit was determined by endpoint RT-PCR on samples of HeLa-HIV cells treated with the
49 indicated concentrations of 1C8 and these values are shown in the histograms of the
50 right panel. **B-E.** Similar descriptions for the alternative splicing units of *DOM3Z*, *ESYT2*,
51 *RBM41* and *SPTAN1*, respectively.

52

53 **Supplementary Figure S6. Impact of siRNA-mediated knockdown of SRSF10 on**
54 **five cellular splicing events affected by 1C8.**

55 **A.** Immunoblot showing the siRNA-mediated depletion of SRSF10 in 293 cells. **(B-F)**
56 Histograms with percent splicing index (PSI) values for splicing units in *ITGB3BP*,
57 *DOM3Z*, *ESYT2*, *RBM41* and *SPTAN1*.

58

59 **Supplementary Figure S7. MA plots depicting expression changes (log₂FC) of**
60 **human genes caused by different concentrations of 1C8 relative to their relative**
61 **expression levels (logCPM) in untreated samples.**

62 The plots contain all expressed genes (15 867) based on a cpm cutoff of >1 in at least 4
63 experimental samples. All the MA plots are on the same scale (fold change from +4 to -
64 4). Genes whose expression increases and decreases by more than 2-fold are in red
65 and blue, respectively.

66

67 **Supplementary Figure S8. Bar plots of expression hits at 1 μm of 1C8.**

68 The distribution of genes whose expression changes. Approximately 4000 expressed
69 genes were binned in each category according to their global expression level in
70 untreated samples (low, medium, high and super high). The number of genes whose
71 expression was affected (increased in red and repressed in blue) at least two-fold is
72 represented by bar graphs.

73

74 **Supplementary Table S1. Alternative splicing of cellular transcripts affected by**
75 **1C8 as measured by endpoint RT-PCR.**

76 A set of 92 events was analyzed for alternative splicing using endpoint RT-PCR. Genes
77 were selected for their ability to yield two amplicons of sizes that provide adequate
78 resolution by microfluidic analysis using a RT-PCR screening platform (sequences of
79 forward and reverse primers are indicated). HeLa-HIV cells were treated with 1C8 (1 and
80 10 μ M) or DMSO as a control (in triplicates). Total RNA was extracted and RT-PCR
81 analysis was carried out. For each sample, percent splicing index (PSI) values are given
82 and changes relative to the control (Δ PSI) were calculated. We used a statistical
83 approach tailored to the analysis of genomic data based on the false discovery rate
84 (FDR) expressed as a q -value. For example, a 5% FDR, q value ≤ 0.05 indicates that
85 5% of the results judged to be significant could represent false positive signals. The
86 section in green indicates q values that are smaller than 0.05, while only one value (in
87 red) is inferior to 0.01.

88

89 **Supplementary Table S2. Cellular transcripts whose alternative splicing are**
90 **affected by 1C8 as revealed by MISO analysis of RNA-Seq data.**

91

92 MISO analysis of RNA-Seq data revealed a set of 86 alternative splicing units affected
93 by more than 30 percentage points in samples of HeLa-HIV cells treated with 1C8
94 (Bayesian coefficient score greater than 20) in at least one level of drug treatment.
95 Essentially, 106 significant shifting events were discovered respectively distributing as 9,
96 16, 25 and 56 in the 1, 5, 10 and 20 μM treatment with 1C8 (20 shifting events are
97 occurring at more than one concentration). Event types are alternative 3' splice site
98 (A3SS) alternative 5' splice site (A5SS) and skipped exon (SE), retained intron (RI) and
99 mutually exclusive exon (MXE).

100

101 **Supplementary Table S3. Cellular transcripts with expression affected by 1 μM of**
102 **1C8 based on RNA-Seq data.**

103

104 List of transcripts whose expression was affected more than 2-fold (\log_2 greater than 1
105 or smaller than -1) with FDR values inferior to 0.05.

106

107 **Supplementary Table S4. Primers used in the study.**

108

Gene	q-values of Δ PSI	
	CTRL – 1 μ M 1C8	CTRL – 10 μ M 1C8
<i>INF2</i>	6.470E-1	4.171E-3
<i>CLK1</i>	7.744E-1	1.283E-2
<i>AP2B1</i>	5.116E-1	1.719E-2
<i>ABI1</i>	5.116E-1	3.226E-2
<i>ADARB1</i>	6.102E-1	3.226E-2
<i>EPB41L1</i>	8.901E-1	3.228E-2
<i>GRHL1</i>	6.470E-1	3.228E-2
<i>ADARB1</i>	6.102E-1	3.298E-2
<i>AMACR</i>	6.102E-1	3.298E-2
<i>ATG16L1</i>	9.847E-1	3.298E-2
<i>C3orf17</i>	9.847E-1	3.298E-2
<i>ECT2</i>	9.991E-1	3.298E-2
<i>KCNAB2</i>	8.901E-1	3.298E-2
<i>MAPKAP1</i>	8.901E-1	3.298E-2
<i>SMN2</i>	6.102E-1	3.298E-2
<i>HNRNPAB</i>	8.040E-1	4.118E-2
<i>ASB3</i>	5.116E-1	4.425E-2
<i>PALM</i>	7.711E-1	4.425E-2
<i>CASC4</i>	6.102E-1	4.747E-2
<i>ZMIZ2</i>	9.991E-1	4.747E-2
<i>APC</i>	9.991E-1	4.763E-2
<i>BRD8</i>	9.991E-1	4.763E-2
<i>DCUN1D4</i>	8.040E-1	4.763E-2
<i>NKTR</i>	9.991E-1	4.763E-2
<i>BRD8</i>	9.314E-1	4.980E-2
<i>LRP8</i>	7.744E-1	4.980E-2
<i>PARL</i>	6.470E-1	5.107E-2
<i>LLGL2</i>	7.744E-1	5.219E-2
<i>LONRF3</i>	8.040E-1	5.267E-2
<i>TBC1D25</i>	7.711E-1	5.349E-2
<i>PLOD2</i>	8.794E-1	5.471E-2
<i>GEMIN7</i>	9.847E-1	5.650E-2
<i>MKNK1</i>	9.847E-1	6.202E-2
<i>OSBPL3</i>	8.040E-1	6.202E-2
<i>NFAT5</i>	6.470E-1	6.303E-2
<i>PPIP5K1</i>	7.744E-1	6.951E-2
<i>AGAP1</i>	9.991E-1	7.385E-2
<i>MTMR3</i>	8.040E-1	7.385E-2
<i>BOLA3</i>	6.040E-1	7.435E-2
<i>MBD1</i>	6.470E-1	7.488E-2
<i>CLCN6</i>	6.470E-1	9.172E-2
<i>FBF1</i>	8.794E-1	9.466E-2
<i>POLB</i>	9.702E-1	9.466E-2
<i>PML</i>	6.244E-1	9.800E-2
<i>CAMKK2</i>	7.744E-1	9.829E-2
<i>ODF2L</i>	9.991E-1	9.829E-2

Gene	q-values of Δ PSI	
	CTRL – 1 μ M 1C8	CTRL – 10 μ M 1C8
<i>ARFIP1</i>	7.076E-1	1.028E-1
<i>ERBB2IP</i>	9.991E-1	1.028E-1
<i>GTF2H2C</i>	8.794E-1	1.028E-1
<i>POGZ</i>	9.847E-1	1.028E-1
<i>ATG5</i>	5.116E-1	1.046E-1
<i>CDK5RAP2</i>	6.102E-1	1.072E-1
<i>NDEL1</i>	7.744E-1	1.072E-1
<i>NFAT5</i>	5.116E-1	1.072E-1
<i>DPP8</i>	7.744E-1	1.131E-1
<i>GAB1</i>	9.314E-1	1.201E-1
<i>ARHGEF11</i>	7.076E-1	1.416E-1
<i>FAM86A</i>	7.711E-1	1.494E-1
<i>AKIP1</i>	7.556E-1	1.586E-1
<i>GIT2</i>	9.991E-1	1.586E-1
<i>INSR</i>	6.102E-1	1.586E-1
<i>TMEM218</i>	9.991E-1	1.586E-1
<i>EIF6</i>	5.116E-1	1.834E-1
<i>PAOX</i>		1.843E-1
<i>MRPL33</i>	8.040E-1	1.922E-1
<i>PITPNC1</i>	5.116E-1	1.922E-1
<i>ATP2A1</i>	7.744E-1	1.993E-1
<i>GTF2I</i>	9.012E-1	2.018E-1
<i>SYNE2</i>	6.040E-1	2.018E-1
<i>AP3M1</i>	7.711E-1	2.226E-1
<i>N6AMT1</i>	9.991E-1	2.226E-1
<i>BTRC</i>	8.673E-1	2.231E-1
<i>ATP5S</i>	7.744E-1	2.334E-1
<i>ADNP</i>	9.991E-1	2.361E-1
<i>CD151</i>	5.116E-1	2.467E-1
<i>CTBP1</i>	8.927E-1	2.467E-1
<i>LRRC23</i>	8.040E-1	2.467E-1
<i>ABCD4</i>	7.711E-1	2.617E-1
<i>MEIS2</i>	5.116E-1	2.668E-1
<i>LHX6</i>	6.470E-1	2.874E-1
<i>HPS1</i>	9.770E-1	3.180E-1
<i>OSBPL9</i>	7.744E-1	3.312E-1
<i>MCAT</i>	9.277E-1	3.323E-1
<i>DTNB</i>	9.991E-1	3.662E-1
<i>MTMR2</i>	9.314E-1	3.662E-1
<i>NFATC2</i>		3.710E-1
<i>ANAPC11</i>	5.116E-1	4.092E-1
<i>MARK2</i>	9.036E-1	4.101E-1
<i>ARHGAP17</i>	8.040E-1	4.104E-1
<i>MAPKAP1</i>	9.991E-1	4.104E-1
<i>PDE9A</i>	8.040E-1	4.145E-1
<i>BTC</i>	5.116E-1	4.310E-1

Gene Symbol	Event type	Treatment	Psi. Diff.Score	Bayes Factor	Event name
AC005077.12	A3SS	10uM	0.40	660327.00	chr7:76090972:76091153:+@chr7:76092862:76093026:+@chr7:76109738:76110094:+@chr7:76111825:76112464:+
AC068533.7	A5SS	5uM	0.30	638.00	chr7:65592691:65592727:+@chr7:65595114:65595235:+@chr7:65595731:65595829:+
ALG13	A5SS	1uM	0.34	67080.00	chrX:110928193:110928331:+@chrX:110932315:110932425:+@chrX:110951255:110951621:+@chrX:110952193:110952276:-
ARFGAP2	MXE	10uM	-0.31	21.00	chr11:47198067:47198185:-@chr11:47197402:47197474:+@chr11:47196733:47196864:-@chr11:47193832:47193884:-
ARFGAP2	MXE	20uM	-0.30	425.00	chr11:47198067:47198185:-@chr11:47197402:47197474:+@chr11:47196733:47196864:-@chr11:47193832:47193884:-
ARL13B	A3SS	10uM	0.36	20.00	chr3:93698983:93699326:+@chr3:93714722:93714788:+@chr3:93754175:93754285:+@chr3:93755396:93755598:+
ASAH2B	A3SS	5uM	-0.32	29.00	chr10:52502675:52502770:+@chr10:52504962:52505034:+@chr10:52509103:52509162:+
ATP5G2	SE	20uM	-0.35	54.00	chr12:54069960:54070512:-@chr12:54067480:54067596:-@chr12:54066360:54066429:-
ATPAF1	A5SS	1uM	0.36	158.00	chr1:46655029:46654852 46654914:-@chr1:46654353:46654652:-
ATPAF1	A5SS	20uM	0.30	29.00	chr1:46655029:46654852 46654914:-@chr1:46654353:46654652:-
BTD	MXE	5uM	-0.39	39.00	chr3:15643255:15643401:+@chr3:15643644:15643817:+@chr3:15676931:15677195:+
C19orf47	SE	20uM	0.41	248.00	chr19:40839758:40839836:-@chr19:40838429:40838550:-@chr19:40834320:40834457:-
C19orf47	SE	5uM	0.42	299.00	chr19:40839758:40839836:-@chr19:40838429:40838550:-@chr19:40834320:40834457:-
CAMK2D	A5SS	10uM	-0.51	1.00E+12	chr4:114378719:114378447 114378491:-@chr4:114372188:114375671:-
CAMK2D	A5SS	20uM	-0.43	8.75E+06	chr4:114378719:114378447 114378491:-@chr4:114372188:114375671:-
CAMK2D	A5SS	5uM	-0.44	406.00	chr4:114378719:114378447 114378491:-@chr4:114372188:114375671:-
CC2D1B	RI	1uM	-0.31	12805.00	chr1:52824133:52824003:-@chr1:52823570:52823453:-
CNOT1	MXE	20uM	0.31	49.00	chr16:58663632:58663790:-@chr16:58658577:58658698:-@chr16:58657186:58657313:-@chr16:58633140:58633415:-
DCHS1	A3SS	20uM	0.32	31.00	chr11:6624634:6624880:-@chr11:6623445 6623494:6623082:-
DDX41	RI	20uM	0.33	1.00E+12	chr5:176942822:176942686:-@chr5:176942340:176942187:-
DHX34	SE	1uM	0.39	81.00	chr19:47879700:47879817:+@chr19:47880172:47880246:+@chr19:47880357:47880463:+
DOM3Z	A3SS	20uM	0.40	76764.00	chr6:31938689:31938924:-@chr6:31938529 31938602:31938383:-
EDC3	SE	20uM	0.34	530.00	chr15:74988221:74988386:-@chr15:74979432:74979520:-@chr15:74967302:74967483:-
ELP5	SE	10uM	-0.35	1.00E+12	chr17:7157902:7158122:+@chr17:7160176:7160357:+@chr17:7161907:7162002:+
ENGASE4	A5SS	20uM	-0.31	1.00E+12	chr17:77077980:77078145 77078959:+@chr17:77079102:77079205:+
ERBB2	SE	10uM	0.33	32.00	chr17:37868575:37868701:+@chr17:37869406:37869522:+@chr17:37871539:37871612:+
ESYT2	SE	10uM	-0.31	1.00E+12	chr7:158552177:158552272:-@chr7:158545472:158545534:-@chr7:158542340:158542414:-
ESYT2	SE	20uM	-0.31	1.00E+12	chr7:158552177:158552272:-@chr7:158545472:158545534:-@chr7:158542340:158542414:-
FGFR1	SE	20uM	0.38	91.00	chr8:38314874:38315052:-@chr8:38287200:38287466:-@chr8:38285864:38285953:-
FN1	SE	20uM	0.36	1.00E+12	chr2:216259251:216259442:-@chr2:216257654:216257926:-@chr2:216256355:216256537:-
GALT	A5SS	10uM	0.41	1.00E+12	chr9:34646674:34646783:+@chr9:34647086:34647255:+
GALT	A5SS	20uM	0.33	2003.00	chr9:34646642:34646783 34646916:+@chr9:34647086:34647255:+
GALT	A5SS	20uM	0.35	2.28E+06	chr9:34646674:34646783:+@chr9:34647086:34647255:+
GALT	A5SS	5uM	0.36	2.32E+07	chr9:34646642:34646783 34646916:+@chr9:34647086:34647255:+
GALT	A5SS	5uM	0.31	2216.00	chr9:34646674:34646783:+@chr9:34647086:34647255:+
GUSB	SE	5uM	0.34	41.00	chr7:65445211:65445396:-@chr7:65444821:65444898:-@chr7:65444386:65444528:-
HDAC6	RI	10uM	0.31	778342.00	chrX:48664038:48664078:+@chrX:48664775:48664871:+
HDLBP	SE	20uM	-0.31	76.00	chr2:242254985:242255117:-@chr2:242208244:242208710:-@chr2:242207892:242207956:-
IL17RC	RI	20uM	0.45	340.00	chr3:9965565:9965704:+@chr3:9965916:9965975:+
IRF9,RNF31	SE	10uM	0.35	24.00	chr14:24633823:24634164:+@chr14:24634266:24634392:+@chr14:24635056:24635171:+
IRF9,RNF31	SE	20uM	0.31	20.00	chr14:24633823:24634164:+@chr14:24634266:24634392:+@chr14:24635056:24635171:+
ITGB3BP	SE	10uM	-0.34	1.00E+12	chr1:63919589:63919645:-@chr1:63913236:63913285:-@chr1:63912477:63912527:-
ITGB3BP	SE	20uM	-0.42	1.00E+12	chr1:63919589:63919645:-@chr1:63913236:63913285:-@chr1:63912477:63912527:-
KIAA0528	SE	20uM	0.32	52.00	chr12:22612426:22612476:-@chr12:22611418:22611519:-@chr12:22609905:22610095:-
KIF21A	SE	20uM	-0.31	101.00	chr12:39713708:39713815:-@chr12:39711875:39712003:-@chr12:39705219:39705355:-
LAS1L	SE	10uM	0.35	1.00E+12	chrX:64744845:64744930:-@chrX:64744444:64744494:-@chrX:64743936:64744142:-
LAS1L	SE	20uM	0.35	1.00E+12	chrX:64744845:64744930:-@chrX:64744444:64744494:-@chrX:64743936:64744142:-
LINC00665	MXE	20uM	0.31	68.00	chr19:36822349:36822620:-@chr19:36821155:36821394:-@chr19:36813032:36813286:-@chr19:36812895:36812932:-
LMBR1L	MXE	1uM	0.44	93973.00	chr12:49500744:49500828:-@chr12:49499707:49499740:-@chr12:49498529:49498668:-@chr12:49497470:49497596:-
LTBP1	SE	20uM	0.31	72.00	chr2:33540211:33540336:+@chr2:33567905:33568030:+@chr2:33572434:33572577:+
MEGF6	SE	5uM	0.40	3020.00	chr1:3413797:3413925:-@chr1:3413552:3413683:-@chr1:3413219:3413347:-
MLL3	A5SS	1uM	-0.36	30.00	chr7:151842380:151842238 151842250:-@chr7:151841798:151841966:-
MTMR14	SE	1uM	-0.32	84.00	chr3:9730628:9730766:+@chr3:9739395:9739550:+@chr3:9743474:9744078:+
MTMR2	SE	10uM	0.41	142.00	chr11:95657039:95657371:-@chr11:95647406:95647476:-@chr11:95621320:95621425:-
MTMR2	SE	20uM	0.33	36.00	chr11:95657039:95657371:-@chr11:95647406:95647476:-@chr11:95621320:95621425:-
MTMR2	SE	20uM	0.31	131.00	chr11:95621320:95621425:-@chr11:95620776:95620848:-@chr11:95598765:95598840:-
MUC1	MXE	5uM	-0.32	50.00	chr1:155160484:155160539:-@chr1:155160198:155160329:-@chr1:155159931:155160052:-@chr1:155159701:155159850:-
NAA60	MXE	10uM	-0.37	138.00	chr16:34993668:34993837:+@chr16:3497482:3497620:+@chr16:3498471:3498540:+
NAA60	MXE	10uM	-0.36	1.09E+10	chr16:34993649:34993827:+@chr16:3497482:3497620:+@chr16:3498471:3498540:+
NAA60	MXE	20uM	-0.34	16042.00	chr16:34993668:34993837:+@chr16:3497482:3497620:+@chr16:3498471:3498540:+
NAA60	MXE	5uM	-0.37	1205.00	chr16:34993668:34993837:+@chr16:3497482:3497620:+@chr16:3498471:3498540:+
NAA60	MXE	5uM	-0.37	1.00E+12	chr16:34993649:34993827:+@chr16:3497482:3497620:+@chr16:3498471:3498540:+
NAP1L4	SE	20uM	-0.35	1.00E+12	chr11:2972489:2972545:-@chr11:2970457:2970494:-@chr11:2965660:2966876:-
NDUFV3	MXE	10uM	-0.30	1.00E+12	chr21:44317037:44317157:+@chr21:44323292:44324386:+@chr21:44328974:44329773:+
NIP2A	MXE	20uM	0.36	992.00	chr15:23034147:23034427:-@chr15:23033894:23034006:-@chr15:23027801:23027922:-@chr15:23021198:23021429:-
NIP2A	MXE	20uM	0.37	2.20E+08	chr15:23034147:23034427:-@chr15:23033894:23034006:-@chr15:23033278:23033413:-@chr15:23021198:23021429:-
NIP2A	MXE	20uM	-0.41	27.00	chr15:23034147:23034427:-@chr15:23033894:23034006:-@chr15:23033278:23033413:-

Gene Symbol	Event type	Treatment	Psi. Diff. Score	Bayes Factor	Event name
PCMC1	SE	10uM	0.36	172.00	chr8:17797200:17797370:+@chr8:17797551:17797667:+@chr8:17804695:17804872:+
PDCD10	MXE	10uM	-0.41	6.81E+06	chr3:167452594:167452651:-@chr3:167452002:167452034:-@chr3:167443189:167443261:-@chr3:167437850:167438061:-
PDCD10	MXE	10uM	-0.36	41052.00	chr3:167452002:167452138:-@chr3:167443189:167443261:-@chr3:167437850:167438061:-
PDCD10	MXE	20uM	-0.36	243.00	chr3:167452594:167452651:-@chr3:167452002:167452034:-@chr3:167443189:167443261:-@chr3:167437850:167438061:-
PLSCR3	SE	20uM	0.35	249.00	chr17:7296786:7296828:-@chr17:7296463:7296683:-@chr17:7296110:7296271:-
PIIP5K2	SE	20uM	-0.30	13821.00	chr5:102515722:102515889:+@chr5:102518935:102519108:+@chr5:102520373:102520445:+
PREPL	SE	20uM	0.36	1372.00	chr2:44588663:44589001:-@chr2:44586636:44586889:-@chr2:44573407:44573529:-
RBM23	SE	10uM	0.34	19811.00	chr14:23388208:23388358:-@chr14:23380995:23381112:-@chr14:23380537:23380612:-
RBM41	MXE	20uM	0.31	1.00E+12	chrX:106358582:106358786:-@chrX:106356627:106356698:-@chrX:106331666:106332069:-
RCOR3	SE	20uM	0.32	249.00	chr1:211477425:211477482:+@chr1:211486062:211486177:+@chr1:211486766:211489725:+
RHBD2	SE	1uM	-0.35	37.00	chr7:75508317:75508578:+@chr7:75510683:75510804:+@chr7:75511147:75511554:+
RHBD2	SE	20uM	-0.30	64.00	chr7:75508317:75508578:+@chr7:75510683:75510804:+@chr7:75511147:75511554:+
RP4-564F22.2	A5SS	20uM	0.32	1.00E+12	chr20:37063976:37063670 37063840:-@chr20:37063147:37063269:-
SCNN1D	SE	20uM	-0.51	35428.00	chr1:1219358:1219470:+@chr1:1220951:1221044:+@chr1:1221306:1221658:+
SCNN1D	SE	5uM	-0.38	40.00	chr1:1219358:1219470:+@chr1:1220951:1221044:+@chr1:1221306:1221658:+
SDR39U1	A5SS	20uM	-0.30	76.00	chr14:24911466:24911304 24911384:-@chr14:24910880:24911001:-
SDR39U1	A5SS	20uM	-0.30	115.00	chr14:24911466:24911315 24911384:-@chr14:24910880:24911001:-
SDR39U1	A5SS	20uM	-0.40	4.54E+09	chr14:24911466:24911315 24911384:-@chr14:24909989:24910132:-
SDR39U1	A5SS	20uM	-0.36	1.50E+10	chr14:24911466:24911304 24911384:-@chr14:24909989:24910132:-
SEC22C	MXE	10uM	0.37	27.00	chr3:42610357:42610565:-@chr3:42608236:42608328:-@chr3:42605257:42605472:-@chr3:42605015:42605178:-
SEPT7P2	SE	5uM	0.31	27.00	chr7:45786814:45786910:-@chr7:45786426:45786547:-@chr7:45784596:45784647:-
SGSH	MXE	1uM	-0.33	22.00	chr17:78190831:78190991:-@chr17:78188832:78188937:-@chr17:78188414:78188564:-@chr17:78187971:78188127:-
SGSM2	SE	10uM	-0.38	1440.00	chr17:2268509:2268635:+@chr17:2270565:2270699:+@chr17:2274556:2274709:+
SPTAN1	SE	10uM	-0.40	6076.00	chr9:131353756:131353904:+@chr9:131355262:131355321:+@chr9:131356454:131356652:+
ST3GAL6	MXE	5uM	-0.30	28.00	chr3:98491657:98491760:+@chr3:98492764:98492827:+@chr3:98503789:98503884:+@chr3:98506880:98507066:+
TAPBP	MXE	20uM	-0.31	3.73E+10	chr6:33281782:33282164:-@chr6:33281471:33281641:-@chr6:33280994:33281254:-@chr6:33272766:33273164:-
TOR1AIIP1	A3SS	20uM	0.34	87.00	chr1:179853827:179853904:+@chr1:179858445 179858448:179858504:+
TPM1	A3SS	10uM	0.31	1444.00	chr15:63334838:63335142:+@chr15:63335668 63335905:63336030:+
TPM1	A3SS	20uM	0.43	1.00E+12	chr15:63334838:63335142:+@chr15:63335668 63335905:63336030:+
TPM1	A3SS	5uM	0.34	269.00	chr15:63334838:63335142:+@chr15:63335668 63335905:63336030:+
TRIM5	SE	20uM	0.34	115.00	chr11:5706098:5706339:-@chr11:5705216:5705369:-@chr11:5700991:5701468:-
UEVLD	MXE	10uM	0.36	27.00	chr11:18610160:18610293:-@chr11:18600271:18600355:-@chr11:18596894:18596959:-@chr11:18591761:18591924:-
WASF2	SE	20uM	-0.49	1.00E+12	chr1:27739066:27739221:-@chr1:27736186:27736700:-@chr1:27730734:27734840:-
WDR45	A3SS	20uM	-0.33	125.00	chrX:48935302:48935406:-@chrX:48934409 48934484:48934304:-
ZFYVE26	RI	20uM	0.35	1.17E+09	chr14:68229129:68228920:-@chr14:68228301:68228083:-
ZNF195	MXE	20uM	0.33	375.00	chr11:3392205:3392300:-@chr11:3383027:3383119:-@chr11:3381950:3382018:-@chr11:3379157:3381795:-
ZNF266	SE	20uM	0.37	28.00	chr19:9545784:9545869:-@chr19:9545474:9545535:-@chr19:9530741:9530894:-
ZNF330	A5SS	20uM	0.31	55.00	chr4:142143520:142143631 142143645:+@chr4:142145641:142145660:+
ZNF562	MXE	20uM	0.36	46.00	chr19:9771396:9771550:-@chr19:9770055:9770143:-@chr19:9768685:9768811:-@chr19:9767223:9767329:-

Gene	1 μ M 1C8	
	Fold change (Log ₂)	FDR value
<i>GPR183</i>	2.650443	2.48E-08
<i>SPINK1</i>	2.574378	7.71E-14
<i>AC068057.1</i>	2.149233	0.001587
<i>INPP5D</i>	2.075333	8.42E-09
<i>KLHDC1</i>	1.983792	0.003443
<i>LPL</i>	1.969363	4.94E-60
<i>APOBEC3H</i>	1.905028	1.13E-05
<i>RP11-611O2.5</i>	1.891939	0.000476
<i>CHL1</i>	1.840547	0.000369
<i>EDN1</i>	1.808004	5.82E-22
<i>KCNN2</i>	1.764675	2.13E-08
<i>HSF2BP</i>	1.724695	1.50E-47
<i>FGF</i>	1.674479	3.50E-06
<i>RP11-809N8.5</i>	1.658597	0.016428
<i>TM4SF1</i>	1.627575	2.11E-49
<i>LIX1</i>	1.626405	0.003474
<i>HSPA9P1</i>	1.603818	0.032998
<i>RP5-1198O20.4</i>	1.591723	2.08E-05
<i>PDE2A</i>	1.586868	4.06E-56
<i>INSL4</i>	1.579286	1.25E-21
<i>ANO2</i>	1.573054	1.57E-20
<i>SCG2</i>	1.557212	1.53E-06
<i>PTGER3</i>	1.527466	8.70E-08
<i>RP11-807H7.2</i>	1.483031	0.03631
<i>CUL4A</i>	1.468718	1.01E-43
<i>C5orf46</i>	1.415557	6.41E-09
<i>CRISPLD2</i>	1.406103	2.21E-11
<i>RP11-115D19.1</i>	1.403545	0.034854
<i>GRHL3</i>	1.400786	0.018813
<i>TNFAIP3</i>	1.398859	2.60E-39
<i>RP11-78A19.4</i>	1.394101	1.87E-05
<i>ITGA11</i>	1.377958	0.004201
<i>FGG</i>	1.37735	3.92E-09
<i>FGFBP1</i>	1.377072	0.00014
<i>PSG5</i>	1.342883	3.54E-05
<i>HRCT1</i>	1.341015	3.32E-11
<i>CCDC68</i>	1.303014	6.09E-20
<i>SCG5</i>	1.294259	5.36E-05
<i>FTH1P11</i>	1.292303	0.010688
<i>MYT1L</i>	1.290632	2.88E-11
<i>AC023137.2</i>	1.282045	0.001057
<i>SPINK6</i>	1.262633	0.0005
<i>DNMT3L</i>	1.247603	2.46E-08
<i>PLEKHG4B</i>	1.246211	0.018468
<i>ALPK2</i>	1.238146	2.20E-05
<i>RGS2</i>	1.231964	1.03E-23
<i>SLC16A6</i>	1.231223	1.23E-29
<i>RP11-421L10.1</i>	1.226661	0.008277
<i>GLIPR1L2</i>	1.218035	0.024921
<i>RP11-144N1.1</i>	1.21039	0.005083
<i>RXFP1</i>	1.208839	2.40E-05
<i>VWA5A</i>	1.207525	4.28E-33
<i>ACTA2</i>	1.202727	1.24E-05
<i>RP1-140K8.5</i>	1.188101	7.77E-09
<i>RP11-399H11.2</i>	1.185094	0.00139
<i>PSG9</i>	1.174724	0.009592
<i>RP11-347C18.3</i>	1.171276	0.040768
<i>OLFM1</i>	1.154358	3.80E-19
<i>PSG1</i>	1.151686	0.009213
<i>CTC-295J13.3</i>	1.150792	0.005046
<i>CRB1</i>	1.14419	1.19E-13
<i>GAGE2A</i>	1.142999	0.043287
<i>FOXQ1</i>	1.135102	1.02E-18
<i>CYR61</i>	1.12529	3.35E-24
<i>EID3</i>	1.124967	0.010763

Gene	1 μ M 1C8	
	Fold change (Log ₂)	FDR value
<i>RP11-64D22.2</i>	1.115672	5.69E-06
<i>KB-1991G8.1</i>	1.11473	0.001487
<i>RP11-154J22.1</i>	1.110004	0.00724
<i>CD83</i>	1.101461	2.12E-32
<i>FMO5</i>	1.100955	0.032603
<i>SGK1</i>	1.09107	4.27E-37
<i>SPATA18</i>	1.089338	1.11E-07
<i>TM4SF1-AS1</i>	1.086954	3.94E-06
<i>CHKB-AS1</i>	1.083088	0.019742
<i>RP11-134G8.8</i>	1.06926	0.040298
<i>GRAMD3</i>	1.068721	8.32E-11
<i>MYPN</i>	1.067613	3.75E-19
<i>PCID2</i>	1.062882	3.26E-32
<i>CTD-2139B15.5</i>	1.062001	3.21E-05
<i>NUPR1</i>	1.061691	5.55E-15
<i>RP5-1028K7.2</i>	1.060282	0.019401
<i>FAM43A</i>	1.059855	8.74E-15
<i>CPM</i>	1.05938	9.93E-22
<i>CHMP1B</i>	1.050854	5.77E-18
<i>ARL4A</i>	1.047929	2.43E-18
<i>RP11-467D6.1</i>	1.035512	0.000982
<i>MFF</i>	1.034311	1.47E-29
<i>AC003104.1</i>	1.030898	4.19E-06
<i>SPINK13</i>	1.028151	1.55E-07
<i>ITPR1</i>	1.025354	1.04E-17
<i>RP11-356I2.4</i>	1.015603	1.19E-08
<i>LINC00471</i>	1.008444	0.012322
<i>SMC1B</i>	1.003701	0.034351
<i>FTH1P20</i>	1.003479	0.026094
<i>RP11-383H13.1</i>	1.002233	0.040768
<i>CFH</i>	-1.00233	1.09E-05
<i>RN7SK</i>	-1.0043	0.040578
<i>GPR50</i>	-1.02022	2.83E-09
<i>RP11-95O2.5</i>	-1.02056	0.03365
<i>EPGN</i>	-1.02859	0.022321
<i>AF003625.3</i>	-1.03325	1.92E-05
<i>KLRC2</i>	-1.04006	4.89E-05
<i>RP3-395M20.12</i>	-1.04413	3.59E-05
<i>HOXA-AS2</i>	-1.05449	2.55E-15
<i>RP11-715J22.2</i>	-1.07652	0.025005
<i>ITGB4</i>	-1.09704	0.048179
<i>PPL</i>	-1.10616	2.23E-05
<i>RP13-238F13.5</i>	-1.12623	0.000755
<i>ARC</i>	-1.13624	0.003111
<i>RAB6C-AS1</i>	-1.14205	0.02
<i>C20orf166-AS1</i>	-1.14246	0.006623
<i>LIPH</i>	-1.15069	0.016412
<i>RAPGEF3</i>	-1.15633	6.11E-06
<i>G0S2</i>	-1.16014	0.003552
<i>FAP</i>	-1.17566	1.26E-15
<i>GAS6-AS1</i>	-1.18169	0.018203
<i>ZCCHC12</i>	-1.18399	2.50E-16
<i>GRB7</i>	-1.241	0.013022
<i>ELF3</i>	-1.25264	4.71E-10
<i>RP11-284F21.9</i>	-1.26778	6.16E-05
<i>FOSB</i>	-1.27149	7.83E-06
<i>FAM231D</i>	-1.27846	0.009667
<i>H19</i>	-1.28402	1.98E-05
<i>CXCL3</i>	-1.29749	0.001547
<i>NMUR1</i>	-1.38076	0.024921
<i>SSC5D</i>	-1.47675	1.95E-07
<i>ANGPTL4</i>	-1.62486	0.000903
<i>RP11-474O21.5</i>	-1.64184	0.000216
<i>AL161626.1</i>	-1.64267	0.034338
<i>MIR663A</i>	-1.72027	0.007235
<i>LINC00319</i>	-2.71392	2.49E-71

1. Primers for HIV-1

For analysis of HIV-1 transcripts, primers were designed based on Human immunodeficiency virus 1, complete genome sequence: NCBI Reference Sequence: NC_001802.1

1.1. Primers for quantitative real-time PCR

HIV-US.1.FWD	5'- CTG AAG CGC GCA CGG CAA -3'
HIV-US.1.REV	5'- GAC GCT CTC GCA CCC ATC TC -3'
HIV-US.2.FWD	5'- GGA CAG CTA CAA CCA TCC CT -3'
HIV-US.2.REV	5'- TGA CCT GAT TGC TGT GTC CT -3'
HIV-US.4.FWD	5'- GGG AAA GCT AGG GGA TGG TT -3'
HIV-US.4.REV	5'- TGA CCC AAA TGC CAG TCT CT -3'
HIV-SS.1.FWD	5'- GGC GGC GAC TGG AAG AAG C -3'
HIV-SS.1.REV	5'- CTA TGA TTA CTA TGG ACC ACA C -3'
HIV-SS.1.2.FWD	5'- GGC GGC GAC TGG AAG AAG C -3'
HIV-SS.1.2.REV	5'- CCC CCA TCT CCA CAA GTG CTG A -3'
HIV-SS.2.FWD	5'- GGC GGC GAC TGA CTC TGC TAT AAG -3'
HIV-SS.2.REV	5'- TTG GTC TTC TGG GGC TTG TT -3'
HIV-SS.3.FWD	5'- GGC GGC GAC TG T TTG TTT CAT AAC -3'
HIV-SS.3.REV	5'- AGA GAA GCT TGA TGA GTC TGA CTG -3'
HIV-SS.4.FWD	5'- GCG GCG ACT G AA TTG GGT GTC -3'
HIV-SS.4.REV	5'- GGC TGA CTT CCT GGA TGC TTC C -3'

1.2. Primers for HIV-1 endpoint PCR

HIV-AS-2.FWD	5'- TCT CTC GAC GCA GGA CTC G -3'
HIV-AS-2.REV	5'- TTG GGA GGT GGG TTG CTT TGA TAG -3'

2. Primers for *SRSF10* quantitative RT-PCR

FUSP1_G_1_F	5' AGAAAGTGGATTTGTGGACGGC 3'
FUSP1_G_1_R	5'-ACATTCCTCCCTTCCTTGCTT-3'

3. Primers for quantitative RT-PCR of HIV fragments antibodies following RNA immunoprecipitation with anti-Flag

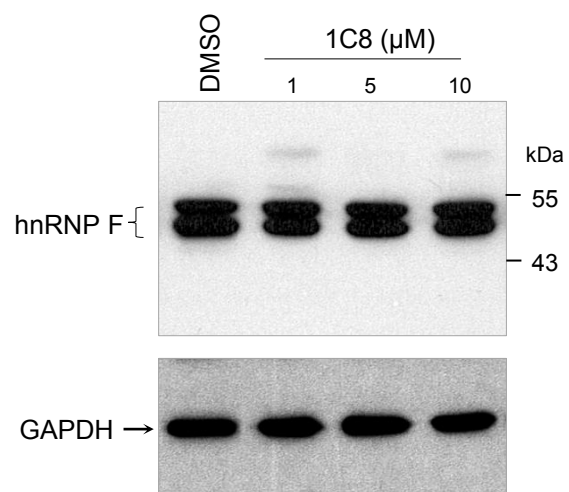
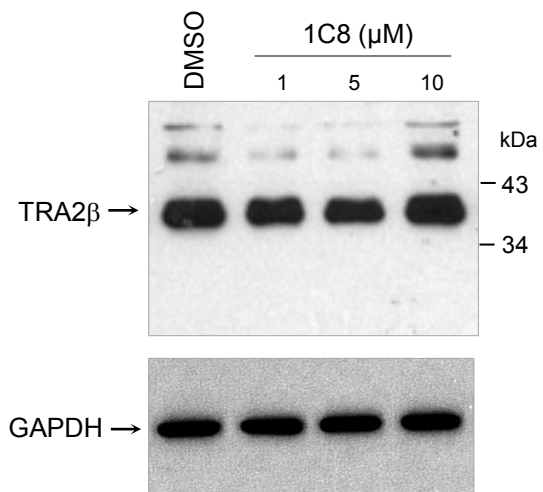
HIV_1-FWD	5'- GTGCAGAACATCCAGGGGCAAA-3'
HIV_1-REV	5'- TCTTGTGGGGTGGCTCCTTCT-3'
HIV_3-FWD	5'- TGTGAAAGGAAGGACACCAAATGA-3'
HIV_3-REV	5'- TCCTTGTCTATCGGCTCCTGCT -3'
HIV_5-FWD	5'- CCCC GGCTGGTTTTGCGATT-3'
HIV_5-REV	5'- ACCTCTTCTTCTGCTAGACTGCCAT-3'
HIV_8-FWD	5'- CCCGACAGGCCCGAAGGAATA-3'
HIV_8-REV	5'- CCCCTGCGTCCCAGAAGTT-3'

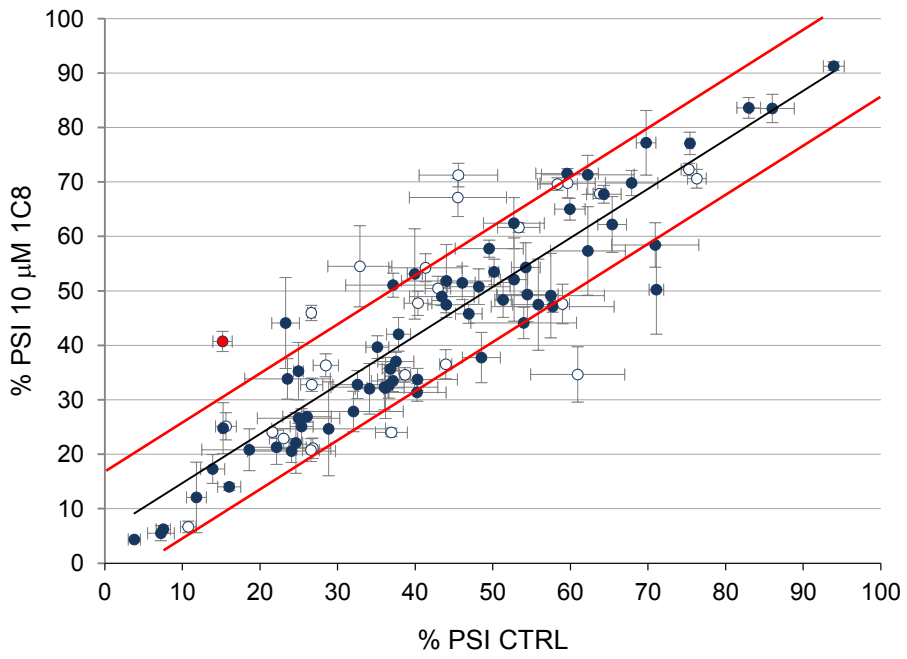
4. Primers for endpoint PCR of transcripts from *Bcl-x* minigenes

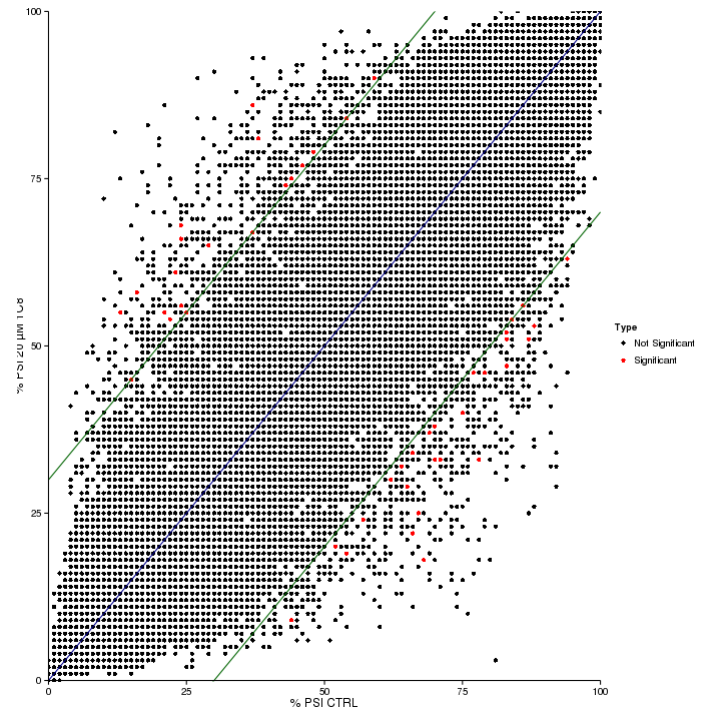
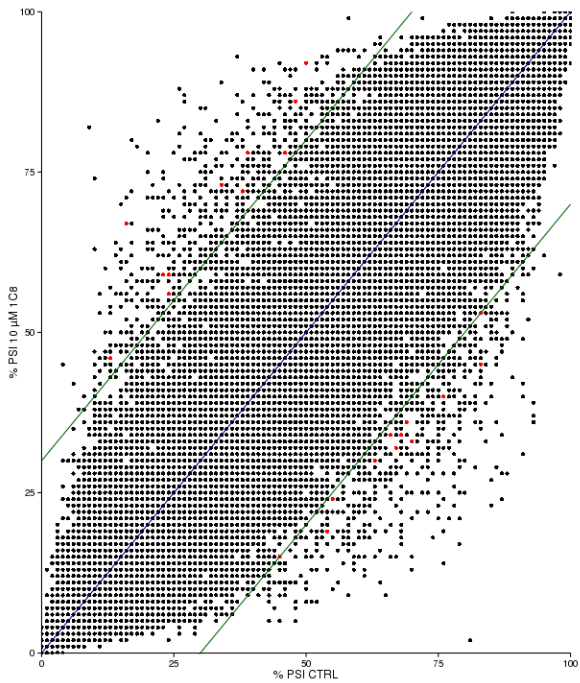
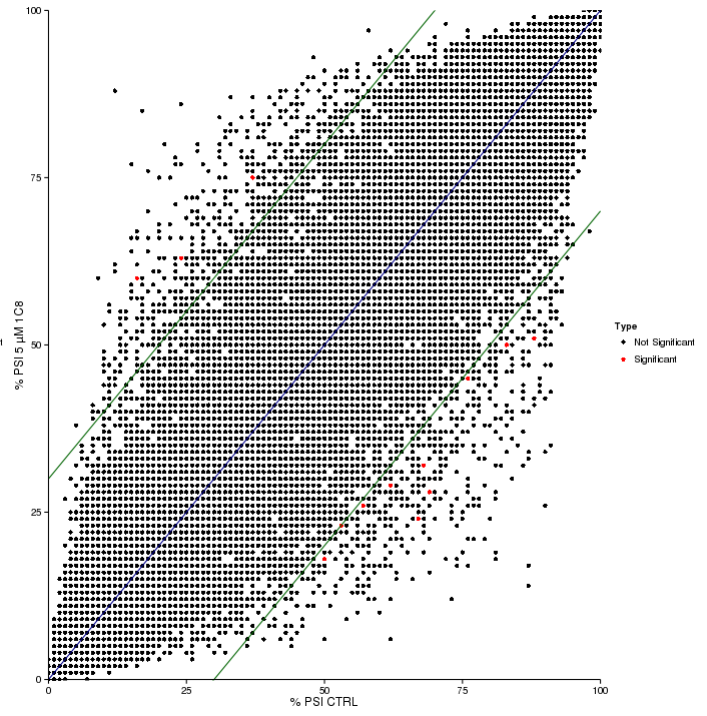
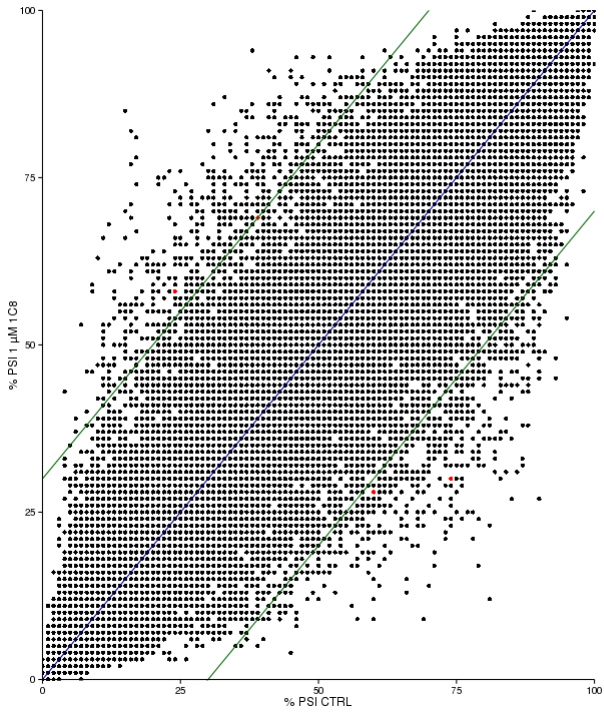
X34 5'-AGGGAGGCAGGCGACGGCGACGAGTTT-3'
X-Age-Rev 5'-CCGGTGGATCCCCCGGGCTGCAGGAATTCGAT-3'

5. Primers for endpoint PCR of various splicing units

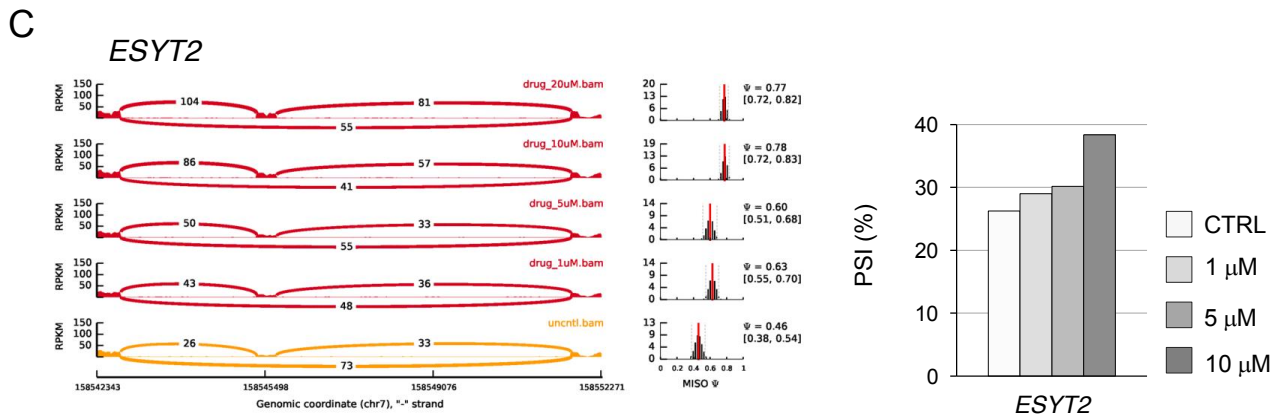
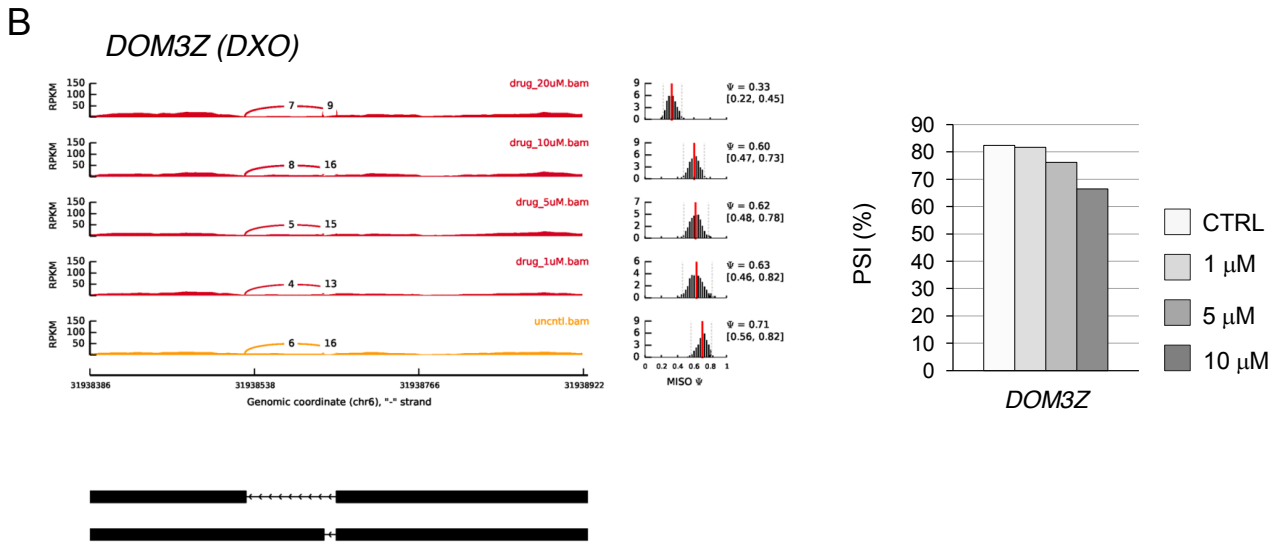
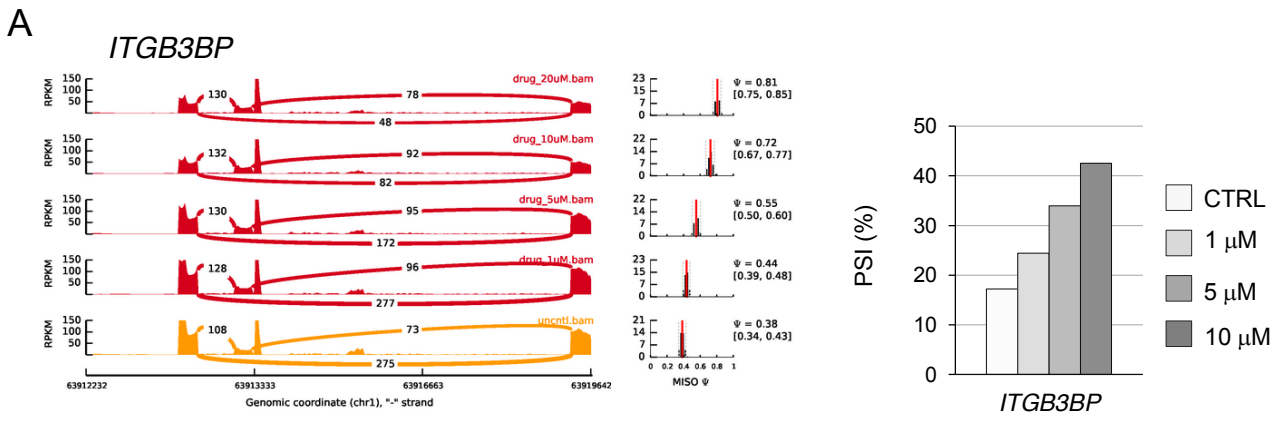
BCLAF1-FWD 5'-GTCTGGTTCTGTTGGAAATA-3'
BCLAF1-REV 5'-AGCAAGCAGCCTGTCTTTAGTC-3'
GLYR1-FWD 5'-GCCTGGATCAAAGTGGAAACA-3'
GLYR1-REV 5'-CTAGACTCCGGGATGGTGAG-3'
CHEK1-FWD 5'-GACTGGGACTTGGTGCAAAC-3'
CHEK1-REV 5'-TGCCATGAGTTGATGGAAGA-3'
SMN2-FWD 5'-CCTCCCATATGTCCAGATTCTC-3'
SMN2-REV 5'-GTGGTGTCAATTTAGTGCTGCTC-3'
DOM3Z-FWD 5'-ATAGTGACGTGGCGGGGGCA-3'
DOM3Z-REV 5'-GAGAAGAGCAGAGGGTGGCTTC-3'
ESYT2-FWD 5'-TCAAAGCTGACAAAGACCAAGC-3'
ESYT2-REV 5'-TGGACAACAGGATTTGGGTTGC-3'
ITGB3BP-FWD 5'-AACAAAACTGTTTGAAAAGAGTACAGG-3'
ITGB3BP-REV 5'-AAAATGGCTTTAAGGAATTCATAGCTGT-3'
NAP1L4-FWD 5'-CGAGGAGGGAGAAGACGAGGAT-3'
NAP1L4-REV 5'-GAAGTCCAGAGCTACAGGCAC-3'
RBM41-FWD 5'-TTTCAGGGAGCTGATCGTCACT-3'
RBM41-REV 5'-AGATCTCTTTCAGTCACCCGAGG-3'
SPTAN1-FWD 5'-GGTGGAAAGTGGAAAGTGAACGA-3'
SPTAN1-REV 5'-AACCTGCTCCAAGTCTGCTCC-3'



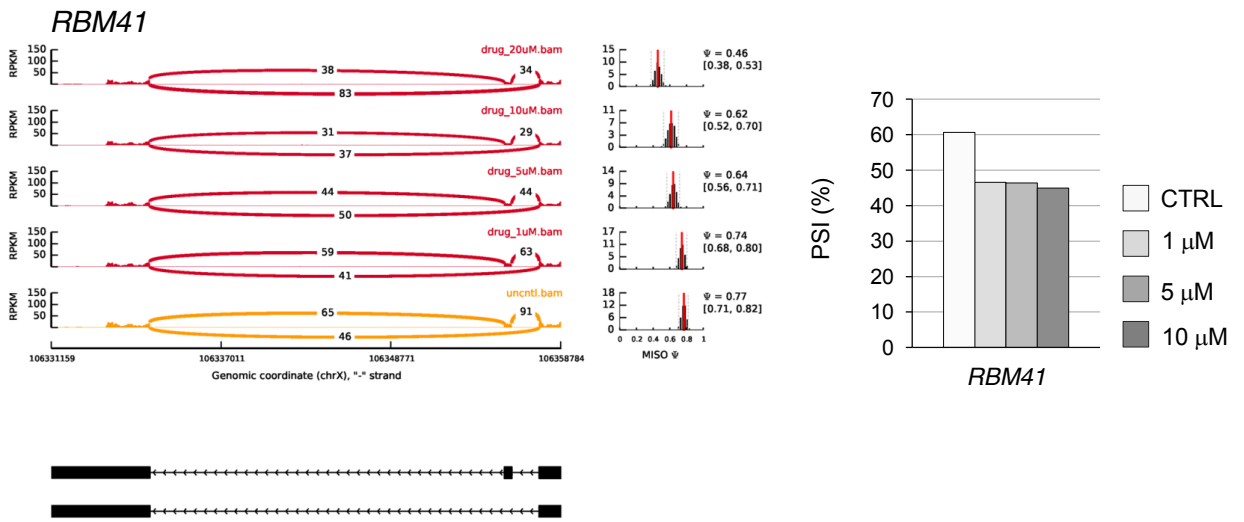




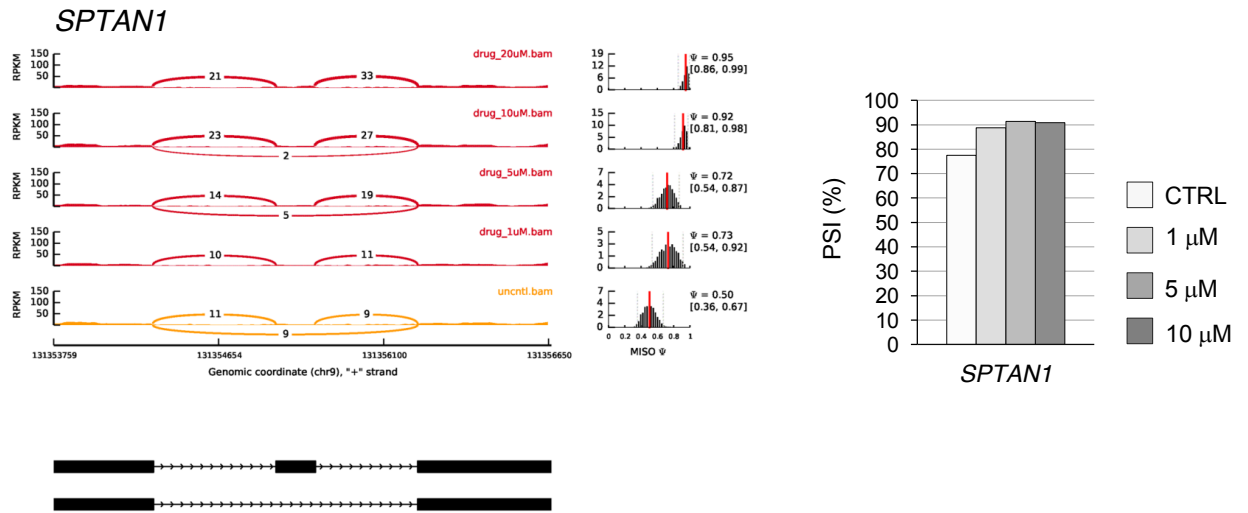
Supplementary Figure S4

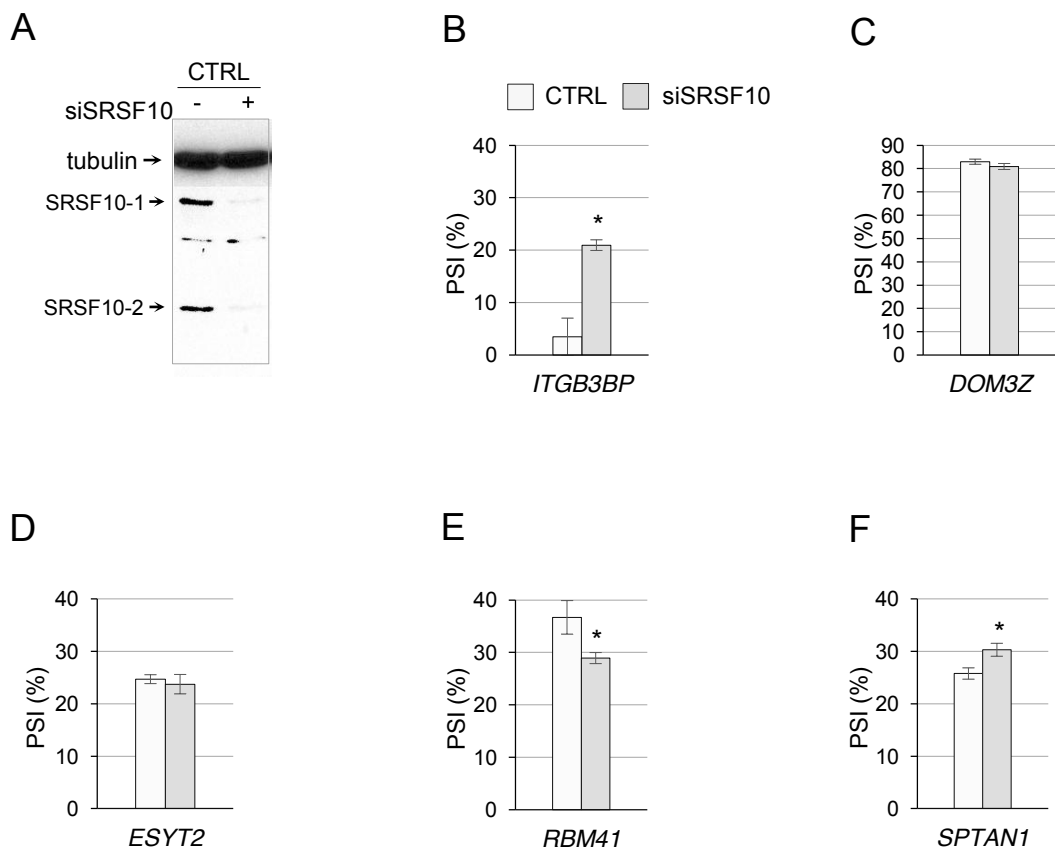


D



E

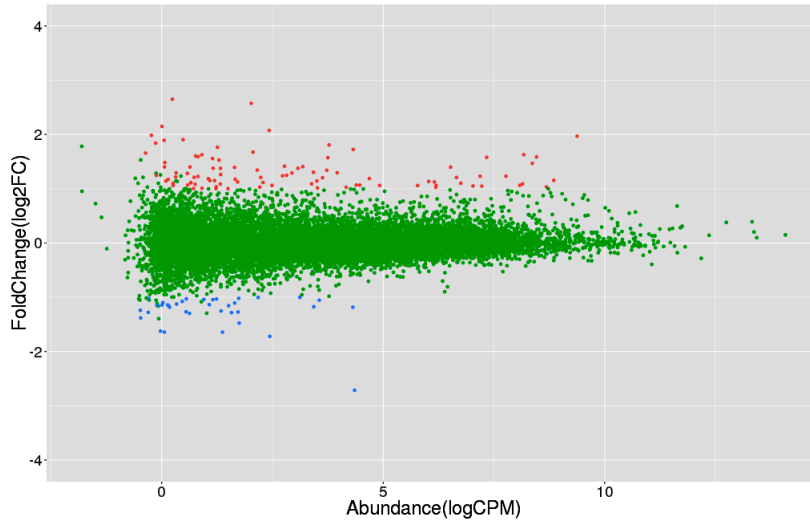




A

Differential Expression of mRNA in drug_1uM

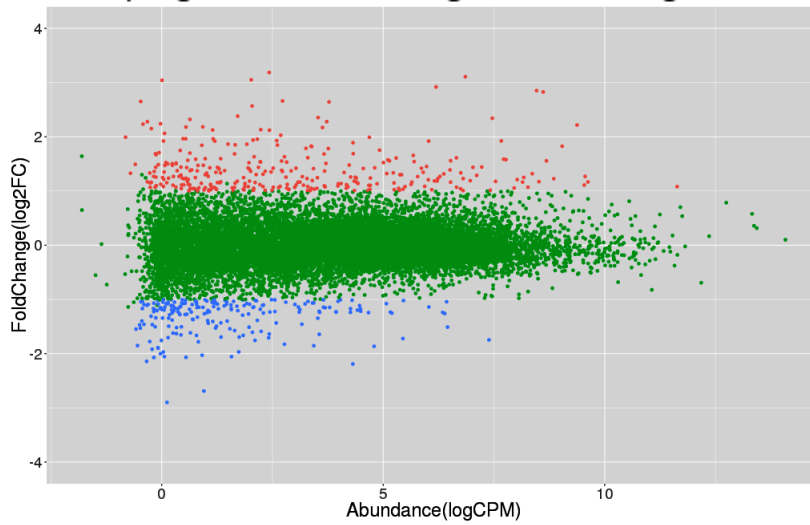
• Upregulated • Unchanged • Downregulated



B

Differential Expression of mRNA in drug_5uM

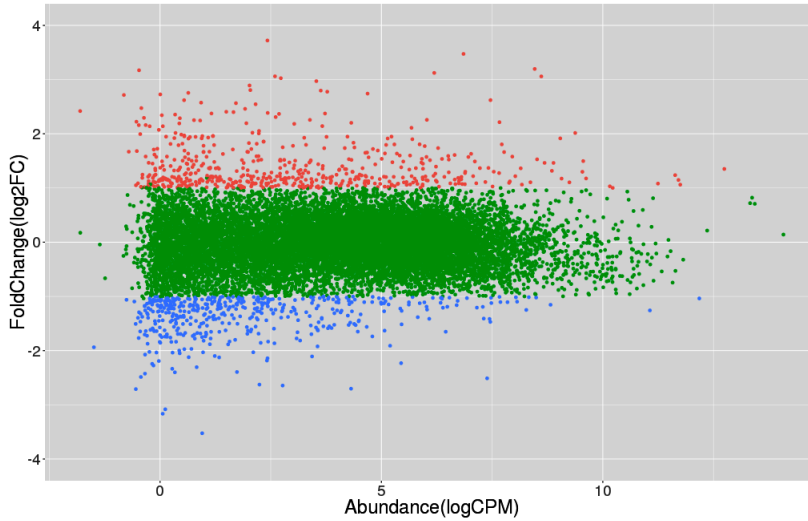
• Upregulated • Unchanged • Downregulated



C

Differential Expression of mRNA in drug_10uM

• Upregulated • Unchanged • Downregulated



D

Differential Expression of mRNA in drug_20uM

• Upregulated • Unchanged • Downregulated

