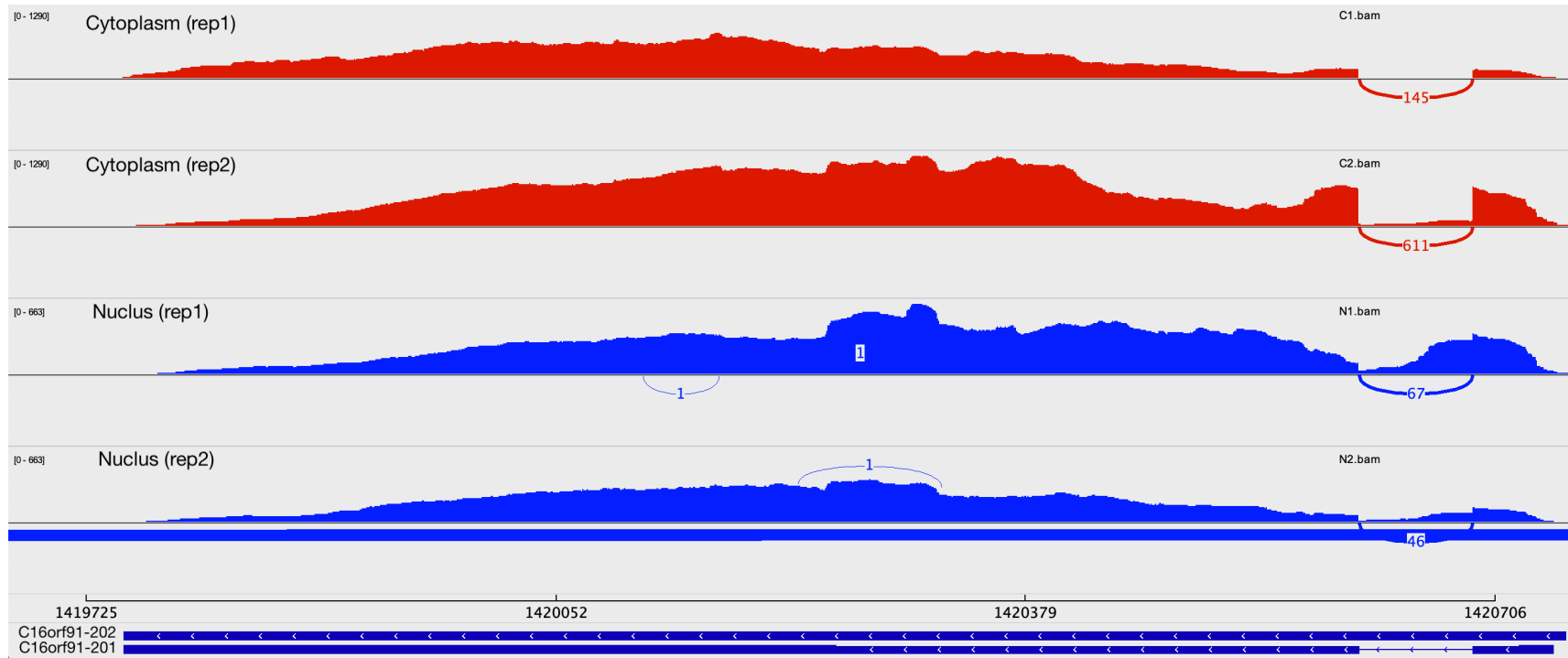


Supplementary Data 1. RNA-seq data used in this study

| Cell line | Tissue | Cytosolic_rep1 (C1) | Cytosolic_rep2 (C2) | Nuclear_rep1 (N1) | Nuclear_rep2 (N2) |
|-----------|-------------------|---------------------|---------------------|-------------------|-------------------|
| A549 | Lung | ENCFF000EKW | ENCFF000ELC | ENCFF000EMA | ENCFF000ELQ |
| | | ENCFF000EKX | ENCFF000ELF | ENCFF000EMD | ENCFF000EMB |
| GM12878 | Blood | ENCFF000EZH | ENCFF000EZF | ENCFF000FCG | ENCFF000FCH |
| | | ENCFF000EZR | ENCFF000EZT | ENCFF000FCI | ENCFF000FCU |
| HeLa-S3 | Cervix | ENCFF000FQK | ENCFF000FQW | ENCFF000FSU | ENCFF000FTB |
| | | ENCFF000FQX | ENCFF000FQV | ENCFF000FTA | ENCFF000FTJ |
| HepG2 | Liver | ENCFF000FXU | ENCFF000FYG | ENCFF000GAF | ENCFF000GAH |
| | | ENCFF000FYF | ENCFF000FYH | ENCFF000GAT | ENCFF000GAS |
| HT1080 | Connective tissue | ENCFF447UEE | ENCFF331FNA | ENCFF028JQU | ENCFF365ZYO |
| | | ENCFF058MHC | ENCFF503XDE | ENCFF381YLG | ENCFF087HWA |
| HUVEC | Umbilical vein | ENCFF000GTX | ENCFF000GUJ | ENCFF000GWF | ENCFF000GWS |
| | | ENCFF000GUH | ENCFF000GUI | ENCFF000GWT | ENCFF000GWW |
| IMR-90 | Lung | ENCFF000HCB | ENCFF000HCI | ENCFF000HDD | ENCFF000HDE |
| | | ENCFF000HCC | ENCFF000HCJ | ENCFF000HDF | ENCFF000HDH |
| MCF-7 | Breast | ENCFF000HSB | ENCFF000HSC | ENCFF000HTE | ENCFF000HTD |
| | | ENCFF000HSI | ENCFF000HSK | ENCFF000HVN | ENCFF000HVM |
| NHEK | Skin | ENCFF000IDC | ENCFF000IDF | ENCFF000IFF | ENCFF000IFI |
| | | ENCFF000IDI | ENCFF000IDH | ENCFF000IFH | ENCFF000IFW |
| SK-MEL-5 | Skin | ENCFF470ZTS | ENCFF687XKK | ENCFF005NLJ | ENCFF394ODW |
| | | ENCFF299CXA | ENCFF570YTU | ENCFF635CQM | ENCFF365MLV |
| SK-N-DZ | Brain | ENCFF960DLP | ENCFF138HQU | ENCFF482SFO | ENCFF993CAL |
| | | ENCFF288WWD | ENCFF822LCB | ENCFF691TRA | ENCFF689LXP |
| SK-N-SH | Brain | ENCFF000INM | ENCFF000INN | ENCFF000IOD | ENCFF000IOP |
| | | ENCFF000INT | ENCFF000INV | ENCFF000IOQ | ENCFF000IPB |
| K562 | Bone marrow | ENCFF000HJP | ENCFF000HJF | ENCFF000HOC | ENCFF000HOD |
| | | ENCFF000HJW | ENCFF000HJX | ENCFF000HOE | ENCFF000HOQ |

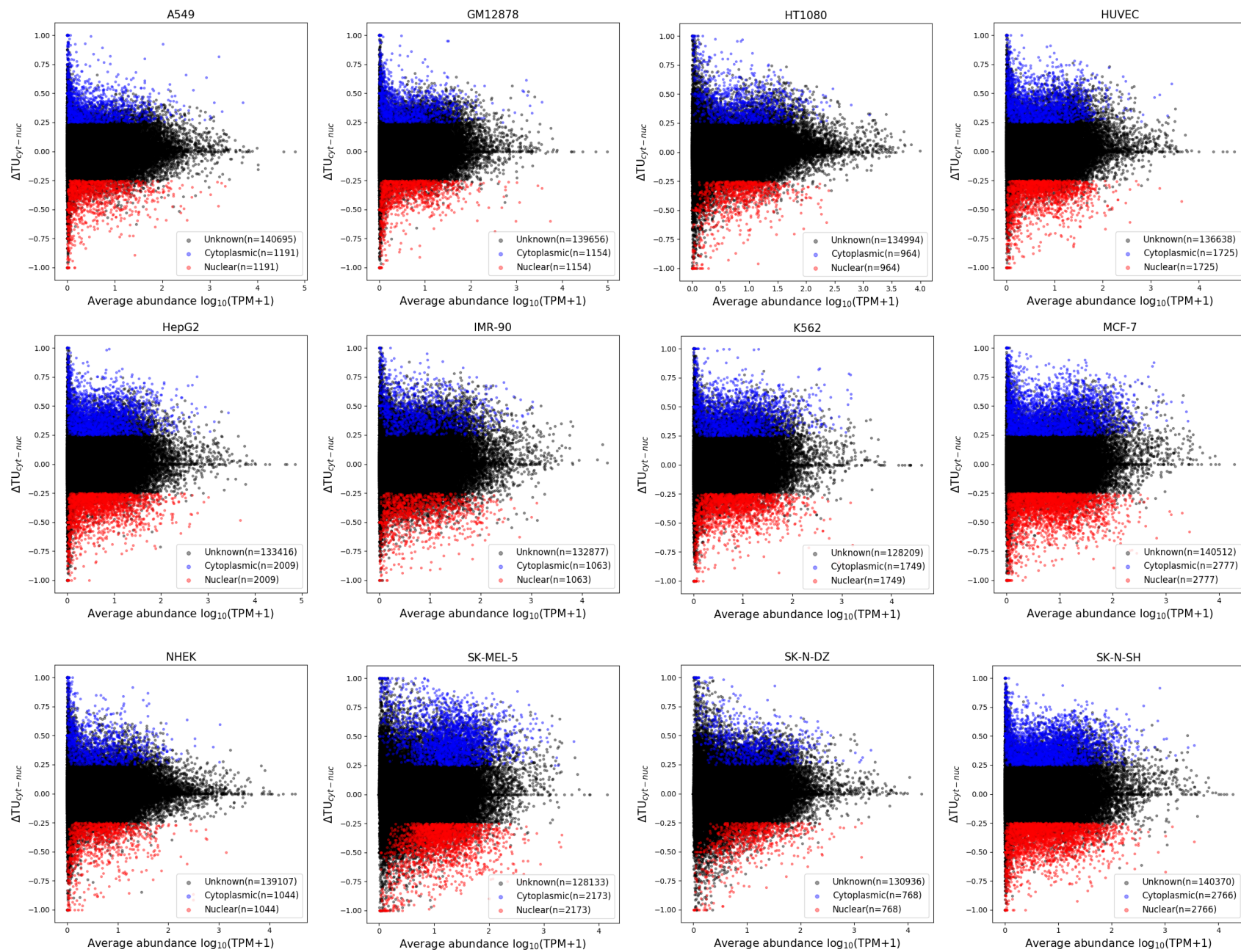
- The libraries were constructed from rRNA-depleted Poly-A+ RNA (> 200bp in size)
- Strand-specific, PE101nt or PE76nt



Supplementary Data 2. Read coverage of gene *C16orf91* in HeLa-S3. IGV (Robinson et al 2011) was used to visualize the read coverage.

Reference:

Robinson, J.T., Thorvaldsdóttir, H., Winckler, W., Guttman, M., Lander, E.S., Getz, G. and Mesirov, J.P., 2011. Integrative genomics viewer. *Nature biotechnology*, 29(1), pp.24-26.



Supplementary Data 3. Transcriptome-wide identification of transcript variant switches in 12 human cell lines. We applied $|\Delta TU| > 0.25$ and $p < 0.05$ to filter out cytoplasmic ($|\Delta TU| > 0$, blue) and nuclear ($|\Delta TU| < 0$, red) transcript variants. Unknown (black) are transcripts filtered as no significant.

Supplementary Data 4. List of switching genes identified across 13 cell lines in this study.

Online resource: <https://doi.org/10.6084/m9.figshare.11999658>

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (sheared, continued)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|---|------------|------------------|------------------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 1.55E-16 | |
| GO:MF | molecular transducer activity | GO:0060089 | 5.59E-16 | |
| GO:MF | signaling receptor activity | GO:0038023 | 5.68E-16 | |
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 6.56E-12 | |
| GO:MF | passive transmembrane transporter activity | GO:0022803 | 0.000493696 | |
| GO:MF | channel activity | GO:0015267 | 0.000493696 | |
| GO:BP | multicellular organismal process | GO:0032501 | 3.31E-12 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 2.10E-11 | |
| GO:BP | system process | GO:0003008 | 3.32E-09 | |
| GO:BP | sensory perception of chemical stimulus | GO:0007606 | 1.85E-07 | |
| GO:BP | nervous system process | GO:0050877 | 2.73E-07 | |
| GO:BP | detection of stimulus involved in sensory perception | GO:0050906 | 1.36162E-06 | |
| GO:BP | detection of chemical stimulus involved in sensory perception | GO:0050907 | 1.61549E-06 | |
| GO:BP | signaling | GO:0023052 | 2.52304E-06 | |
| GO:BP | signal transduction | GO:0007165 | 1.66204E-05 | |
| GO:BP | detection of chemical stimulus | GO:0009593 | 2.20905E-05 | |
| GO:BP | sensory perception | GO:0007600 | 2.65966E-05 | |
| GO:BP | cell communication | GO:0007154 | 2.81869E-05 | |
| GO:BP | defense response | GO:0006952 | 6.05683E-05 | |
| GO:BP | detection of stimulus | GO:0051606 | 6.68613E-05 | |
| GO:BP | response to external stimulus | GO:0009605 | 0.000100343 | |
| GO:BP | multicellular organism development | GO:0007275 | 0.000100343 | |
| GO:BP | adaptive immune response | GO:0002250 | 0.000171355 | |
| GO:BP | cell differentiation | GO:0030154 | 0.000179869 | |
| GO:BP | system development | GO:0048731 | 0.000207557 | |
| GO:BP | ion transport | GO:0006811 | 0.000405413 | |
| GO:BP | positive regulation of immune system process | GO:0002684 | 0.000439648 | |
| GO:BP | anatomical structure development | GO:0048856 | 0.000518076 | |
| GO:BP | regulation of multicellular organismal process | GO:0051239 | 0.000537973 | |
| GO:BP | defense response to bacterium | GO:0042742 | 0.000820237 | |
| GO:CC | plasma membrane | GO:0005886 | 3.50E-13 | |
| GO:CC | cell periphery | GO:0071944 | 4.22E-13 | |
| GO:CC | extracellular region | GO:0005576 | 1.17977E-05 | |
| GO:CC | plasma membrane region | GO:0098590 | 0.000637715 | |
| GO:CC | intrinsic component of membrane | GO:0031224 | 0.000637715 | |

GO(Gene Ontology) enrichment analysis for 1219 sheared switching genes (detected as switch gene ≥ 5 cell lines in this study) was conducted by g:Profiler(FMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

* MF, BP, CC represent Molecular Function, Biological Process, Cellular Component, respectively.

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (A549, continued)




| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|---|------------|------------------|------------------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 2.58E-09 | |
| GO:MF | signaling receptor activity | GO:0038023 | 9.26E-08 | |
| GO:MF | molecular transducer activity | GO:0060089 | 9.26E-08 | |
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 1.27132E-05 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 1.535E-05 | |
| GO:BP | sensory perception of chemical stimulus | GO:0007606 | 2.3181E-05 | |
| GO:BP | detection of chemical stimulus involved in sensory perception | GO:0050907 | 2.3181E-05 | |
| GO:BP | sensory perception of smell | GO:0007608 | 3.06052E-05 | |
| GO:BP | detection of chemical stimulus | GO:0009593 | 3.06052E-05 | |
| GO:BP | detection of stimulus involved in sensory perception | GO:0050906 | 8.11614E-05 | |
| GO:CC | no significant results | | | |

GO(Gene Ontology) enrichment analysis for 1191 switching genes in A549 was conducted by g:Profiler(PMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

* MF, BP, CC represent Molecular Function, Biological Process, Cellular Component, respectively.

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (GM12878, continued)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|--|------------|------------------|---|
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 4.27792E-05 |  |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 3.80158E-05 |  |
| GO:BP | system process | GO:0003008 | 0.000132801 |  |
| GO:CC | no significant results | | | |

GO(Gene Ontology) enrichment analysis for 1154 switching genes in GM12878 was conducted by g:Profiler(PMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (HT1080, continued)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|--|------------|------------------|------------------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 6.49781E-05 | |
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 0.000178858 | |
| GO:MF | signaling receptor activity | GO:0038023 | 0.000211021 | |
| GO:MF | molecular transducer activity | GO:0060089 | 0.000211021 | |
| GO:BP | signal transduction | GO:0007165 | 4.50381E-06 | |
| GO:BP | multicellular organismal process | GO:0032501 | 4.50381E-06 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 5.06491E-06 | |
| GO:BP | signaling | GO:0023052 | 5.06491E-06 | |
| GO:BP | cell communication | GO:0007154 | 2.71825E-05 | |
| GO:CC | extracellular region | GO:0005576 | 4.51E-12 | |
| GO:CC | extracellular space | GO:0005615 | 5.10E-10 | |
| GO:CC | cell periphery | GO:0071944 | 1.51052E-06 | |
| GO:CC | plasma membrane | GO:0005886 | 2.08264E-06 | |
| GO:CC | extracellular organelle | GO:0043230 | 6.65978E-05 | |
| GO:CC | extracellular vesicle | GO:1903561 | 6.65978E-05 | |
| GO:CC | extracellular exosome | GO:0070062 | 9.1387E-05 | |

GO(Gene Ontology) enrichment analysis for 964 switching genes in HT1080 was conducted by g:Profiler(PMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (HUVEC, continued)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|---|------------|------------------|------------------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 1.04E-08 | |
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 1.04E-08 | |
| GO:MF | signaling receptor activity | GO:0038023 | 1.73E-07 | |
| GO:MF | molecular transducer activity | GO:0060089 | 1.96E-07 | |
| GO:BP | detection of stimulus involved in sensory perception | GO:0050906 | 5.37E-10 | |
| GO:BP | detection of chemical stimulus involved in sensory perception | GO:0050907 | 9.09E-10 | |
| GO:BP | detection of stimulus | GO:0051606 | 5.62E-09 | |
| GO:BP | sensory perception of chemical stimulus | GO:0007606 | 2.26E-08 | |
| GO:BP | sensory perception of smell | GO:0007608 | 2.26E-08 | |
| GO:BP | multicellular organismal process | GO:0032501 | 1.40E-07 | |
| GO:BP | system process | GO:0003008 | 2.72E-07 | |
| GO:BP | detection of chemical stimulus | GO:0009593 | 2.90E-07 | |
| GO:BP | sensory perception | GO:0007600 | 4.30E-07 | |
| GO:BP | nervous system process | GO:0050877 | 1.37468E-05 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 7.79881E-05 | |
| GO:BP | cell differentiation | GO:0030154 | 0.000238532 | |
| GO:CC | extracellular region | GO:0005576 | 2.7509E-06 | |
| GO:CC | extracellular space | GO:0005615 | 0.000209584 | |

GO(Gene Ontology) enrichment analysis for 1725 switching genes in HUVEC was conducted by g:Profiler(PMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini–Hochberg correction
- FDR(False Discovery Rate) < 0.001

* MF, BP, CC represent Molecular Function, Biological Process, Cellular Component, respectively.

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (HeLa-S3, continued)

| source | term_name | term_id | adjusted_p_value | -log10 (adjusted_p_value) |
|--------|---|------------|------------------|---------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 6.36E-20 | |
| GO:MF | signaling receptor activity | GO:0038023 | 3.32E-18 | |
| GO:MF | molecular transducer activity | GO:0060089 | 4.04E-18 | |
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 1.43E-16 | |
| GO:MF | gated channel activity | GO:0022836 | 0.00014682 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 4.22E-14 | |
| GO:BP | detection of chemical stimulus involved in sensory perception | GO:0050907 | 4.22E-13 | |
| GO:BP | detection of stimulus involved in sensory perception | GO:0050906 | 5.95E-13 | |
| GO:BP | sensory perception of chemical stimulus | GO:0007606 | 6.90E-13 | |
| GO:BP | multicellular organismal process | GO:0032501 | 7.57E-13 | |
| GO:BP | nervous system process | GO:0050877 | 1.73E-12 | |
| GO:BP | sensory perception | GO:0007600 | 4.13E-12 | |
| GO:BP | sensory perception of smell | GO:0007608 | 1.13E-11 | |
| GO:BP | detection of chemical stimulus | GO:0009593 | 1.59E-11 | |
| GO:BP | system process | GO:0003008 | 4.10E-11 | |
| GO:BP | detection of stimulus | GO:0051606 | 2.85E-07 | |
| GO:BP | adaptive immune response | GO:0002250 | 4.41341E-05 | |
| GO:BP | signaling | GO:0023052 | 0.000978609 | |
| GO:CC | plasma membrane | GO:0005886 | 3.22E-20 | |
| GO:CC | cell periphery | GO:0071944 | 6.49E-20 | |
| GO:CC | integral component of plasma membrane | GO:0005887 | 1.44E-09 | |
| GO:CC | integral component of membrane | GO:0016021 | 1.84E-09 | |
| GO:CC | intrinsic component of membrane | GO:0031224 | 3.40E-09 | |
| GO:CC | intrinsic component of plasma membrane | GO:0031226 | 6.90E-09 | |
| GO:CC | receptor complex | GO:0043235 | 3.00457E-06 | |
| GO:CC | plasma membrane region | GO:0098590 | 2.56347E-05 | |
| GO:CC | extracellular matrix | GO:0031012 | 3.55608E-05 | |
| GO:CC | plasma membrane protein complex | GO:0098797 | 0.000121469 | |
| GO:CC | plasma membrane signaling receptor complex | GO:0098802 | 0.000238214 | |
| GO:CC | collagen-containing extracellular matrix | GO:0062023 | 0.000679446 | |
| GO:CC | extracellular region | GO:0005576 | 0.000679446 | |
| GO:CC | synaptic membrane | GO:0097060 | 0.000679446 | |

GO(Gene Ontology) enrichment analysis for 2073 switching genes in HeLa-S3 was conducted by g:Profiler(FMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

* MF, BP, CC represent Molecular Function, Biological Process, Cellular Component, respectively.

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (HepG2, continued)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|--|------------|------------------|------------------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 3.54E-15 | |
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 9.45E-13 | |
| GO:MF | olfactory receptor activity | GO:0004984 | 1.15E-12 | |
| GO:MF | signaling receptor activity | GO:0038023 | 1.15E-12 | |
| GO:MF | molecular transducer activity | GO:0060089 | 1.15E-12 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 5.33E-14 | |
| GO:BP | sensory perception of chemical stimulus | GO:0007606 | 9.52E-14 | |
| GO:BP | sensory perception of smell | GO:0007608 | 1.60E-12 | |
| GO:BP | detection of chemical stimulus involved in sensory perception | GO:0050907 | 5.18E-12 | |
| GO:BP | detection of chemical stimulus involved in sensory perception of smell | GO:0050911 | 1.43E-11 | |
| GO:BP | detection of chemical stimulus | GO:0009593 | 2.53E-10 | |
| GO:BP | system process | GO:0003008 | 4.56E-09 | |
| GO:BP | multicellular organismal process | GO:0032501 | 7.88E-09 | |
| GO:BP | detection of stimulus involved in sensory perception | GO:0050906 | 9.69E-09 | |
| GO:BP | sensory perception | GO:0007600 | 1.23E-08 | |
| GO:BP | nervous system process | GO:0050877 | 2.19E-08 | |
| GO:BP | detection of stimulus | GO:0051606 | 8.61E-08 | |
| GO:BP | signaling | GO:0023052 | 6.64E-07 | |
| GO:BP | signal transduction | GO:0007165 | 9.84594E-06 | |
| GO:BP | cell communication | GO:0007154 | 1.09238E-05 | |
| GO:BP | keratinization | GO:0031424 | 0.000190632 | |
| GO:BP | regulation of response to stimulus | GO:0048583 | 0.000408547 | |
| GO:BP | humoral immune response | GO:0006959 | 0.000504249 | |
| GO:BP | keratinocyte differentiation | GO:0030216 | 0.000527629 | |
| GO:BP | gene silencing by miRNA | GO:0035195 | 0.000527629 | |
| GO:CC | cell periphery | GO:0071944 | 4.18E-14 | |
| GO:CC | plasma membrane | GO:0005886 | 1.57E-13 | |
| GO:CC | integral component of membrane | GO:0016021 | 1.44303E-06 | |
| GO:CC | intrinsic component of membrane | GO:0031224 | 2.02693E-06 | |
| GO:CC | integral component of plasma membrane | GO:0005887 | 0.000194967 | |
| GO:CC | intrinsic component of plasma membrane | GO:0031226 | 0.00034656 | |
| GO:CC | plasma membrane protein complex | GO:0098797 | 0.000698663 | |

GO(Gene Ontology) enrichment analysis for 2009 switching genes in HepG2 was conducted by g:Profiler(PMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

* MF, BP, CC represent Molecular Function, Biological Process, Cellular Component, respectively.

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (IMR-90, continued)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|--|------------|------------------|------------------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 5.05E-11 | |
| GO:MF | signaling receptor activity | GO:0038023 | 7.05E-11 | |
| GO:MF | molecular transducer activity | GO:0060089 | 1.38E-10 | |
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 3.27E-09 | |
| GO:MF | olfactory receptor activity | GO:0004984 | 5.7643E-05 | |
| GO:BP | multicellular organismal process | GO:0032501 | 3.2905E-06 | |
| GO:BP | sensory perception | GO:0007600 | 4.58447E-05 | |
| GO:BP | system process | GO:0003008 | 6.72981E-05 | |
| GO:BP | nervous system process | GO:0050877 | 0.000186145 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 0.000239019 | |
| GO:BP | detection of stimulus involved in sensory perception | GO:0050906 | 0.000612089 | |
| GO:BP | detection of chemical stimulus involved in sensory perception of smell | GO:0050911 | 0.000612089 | |
| GO:BP | sensory perception of chemical stimulus | GO:0007606 | 0.000618172 | |
| GO:BP | detection of chemical stimulus involved in sensory perception | GO:0050907 | 0.000618172 | |
| GO:CC | plasma membrane | GO:0005886 | 1.24E-07 | |
| GO:CC | cell periphery | GO:0071944 | 1.24E-07 | |

GO(Gene Ontology) enrichment analysis for 1063 switching genes in IMR-90 was conducted by g:Profiler(PMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

* MF, BP, CC represent Molecular Function, Biological Process, Cellular Component, respectively.

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (K562, continued)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|---|------------|------------------|------------------------------------|
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 6.83E-11 | |
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 3.65E-09 | |
| GO:MF | signaling receptor activity | GO:0038023 | 2.52E-08 | |
| GO:MF | molecular transducer activity | GO:0060089 | 2.89E-08 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 1.94E-15 | |
| GO:BP | system process | GO:0003008 | 4.80E-12 | |
| GO:BP | nervous system process | GO:0050877 | 1.01E-10 | |
| GO:BP | detection of chemical stimulus | GO:0009593 | 1.18E-09 | |
| GO:BP | detection of chemical stimulus involved in sensory perception | GO:0050907 | 1.18E-09 | |
| GO:BP | multicellular organismal process | GO:0032501 | 9.58E-09 | |
| GO:BP | detection of stimulus involved in sensory perception | GO:0050906 | 1.36E-08 | |
| GO:BP | sensory perception of smell | GO:0007608 | 3.31E-08 | |
| GO:BP | sensory perception | GO:0007600 | 2.87E-07 | |
| GO:BP | sensory perception of chemical stimulus | GO:0007606 | 5.46E-07 | |
| GO:BP | detection of stimulus | GO:0051606 | 1.31782E-06 | |
| GO:BP | adaptive immune response | GO:0002250 | 0.000259623 | |
| GO:CC | cell periphery | GO:0071944 | 3.90E-09 | |
| GO:CC | plasma membrane | GO:0005886 | 4.12E-09 | |
| GO:CC | extracellular region | GO:0005576 | 0.000163539 | |
| GO:CC | intrinsic component of membrane | GO:0031224 | 0.000205003 | |
| GO:CC | extracellular matrix | GO:0031012 | 0.000216974 | |
| GO:CC | integral component of membrane | GO:0016021 | 0.000295277 | |

GO(Gene Ontology) enrichment analysis for 1749 switching genes in K562 was conducted by g:Profiler(PMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

* MF, BP, CC represent Molecular Function, Biological Process, Cellular Component, respectively.

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (MCF-7, continued)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|---|------------|------------------|------------------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 3.29E-22 | |
| GO:MF | signaling receptor activity | GO:0038023 | 9.46E-20 | |
| GO:MF | molecular transducer activity | GO:0060089 | 9.46E-20 | |
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 1.26E-17 | |
| GO:MF | receptor regulator activity | GO:0030545 | 0.000673771 | |
| GO:BP | detection of chemical stimulus involved in sensory perception | GO:0050907 | 9.20E-19 | |
| GO:BP | detection of stimulus involved in sensory perception | GO:0050906 | 4.21E-17 | |
| GO:BP | detection of chemical stimulus | GO:0009593 | 4.21E-17 | |
| GO:BP | multicellular organismal process | GO:0032501 | 1.97E-15 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 8.82E-15 | |
| GO:BP | sensory perception of chemical stimulus | GO:0007606 | 3.28E-14 | |
| GO:BP | sensory perception of smell | GO:0007608 | 1.02E-12 | |
| GO:BP | detection of stimulus | GO:0051606 | 5.08E-12 | |
| GO:BP | sensory perception | GO:0007600 | 5.29E-12 | |
| GO:BP | system process | GO:0003008 | 1.22E-11 | |
| GO:BP | nervous system process | GO:0050877 | 2.33E-11 | |
| GO:BP | gene silencing by miRNA | GO:0035195 | 2.77583E-05 | |
| GO:BP | posttranscriptional gene silencing | GO:0016441 | 3.61883E-05 | |
| GO:BP | posttranscriptional gene silencing by RNA | GO:0035194 | 3.66099E-05 | |
| GO:BP | response to external stimulus | GO:0009605 | 7.15327E-05 | |
| GO:BP | signaling | GO:0023052 | 7.15327E-05 | |
| GO:BP | adaptive immune response | GO:0002250 | 9.03152E-05 | |
| GO:BP | regulation of multicellular organismal process | GO:0051239 | 0.000169194 | |
| GO:BP | cell communication | GO:0007154 | 0.000223032 | |
| GO:BP | humoral immune response | GO:0006959 | 0.000229014 | |
| GO:BP | gene silencing by RNA | GO:0031047 | 0.000233736 | |
| GO:BP | keratinocyte differentiation | GO:0030216 | 0.000373816 | |
| GO:BP | keratinization | GO:0031424 | 0.000471737 | |
| GO:BP | system development | GO:0048731 | 0.000493893 | |
| GO:BP | regulation of developmental process | GO:0050793 | 0.000493893 | |
| GO:BP | defense response | GO:0006952 | 0.000572092 | |
| GO:BP | multicellular organism development | GO:0007275 | 0.000586089 | |
| GO:BP | signal transduction | GO:0007165 | 0.000639678 | |
| GO:BP | positive regulation of developmental process | GO:0051094 | 0.000859708 | |
| GO:BP | defense response to bacterium | GO:0042742 | 0.000886349 | |
| GO:BP | G protein-coupled receptor signaling pathway, coupled to cyclic nucleotide second messenger | GO:0007187 | 0.000886349 | |
| GO:BP | anatomical structure development | GO:0048856 | 0.000946523 | |
| GO:CC | plasma membrane | GO:0005886 | 2.37E-12 | |
| GO:CC | cell periphery | GO:0071944 | 2.37E-12 | |
| GO:CC | receptor complex | GO:0043235 | 1.57E-07 | |
| GO:CC | intrinsic component of membrane | GO:0031224 | 1.81786E-05 | |
| GO:CC | integral component of plasma membrane | GO:0005887 | 2.17165E-05 | |
| GO:CC | extracellular region | GO:0005576 | 2.17165E-05 | |
| GO:CC | integral component of membrane | GO:0016021 | 2.30812E-05 | |
| GO:CC | intrinsic component of plasma membrane | GO:0031226 | 2.30812E-05 | |
| GO:CC | intermediate filament | GO:0005882 | 6.0376E-05 | |
| GO:CC | plasma membrane protein complex | GO:0098797 | 0.000166003 | |
| GO:CC | plasma membrane region | GO:0098590 | 0.000387874 | |

GO(Gene Ontology) enrichment analysis for 2777 switching genes in MCF-7 was conducted by g:Profiler(FMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

* MF, BP, CC represent Molecular Function, Biological Process, Cellular Component, respectively.

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (NHEK, continued)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|---|------------|------------------|------------------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 0.000223642 | |
| GO:MF | signaling receptor activity | GO:0038023 | 0.000223642 | |
| GO:MF | molecular transducer activity | GO:0060089 | 0.000223642 | |
| GO:MF | olfactory receptor activity | GO:0004984 | 0.000669489 | |
| GO:BP | multicellular organismal process | GO:0032501 | 0.000809884 | |
| GO:CC | extracellular region | GO:0005576 | 0.000461703 | |
| GO:CC | plasma membrane | GO:0005886 | 0.00049703 | |
| GO:CC | cell periphery | GO:0071944 | 0.000666079 | |

GO(Gene Ontology) enrichment analysis for 2777 switching genes in NHEK was conducted by g:Profiler(PMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (SK-MEL-5, continued)


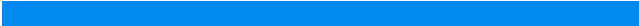
| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|---|------------|------------------|------------------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 2.50E-20 | |
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 2.50E-20 | |
| GO:MF | signaling receptor activity | GO:0038023 | 2.50E-20 | |
| GO:MF | molecular transducer activity | GO:0060089 | 2.50E-20 | |
| GO:MF | receptor regulator activity | GO:0030545 | 4.34223E-05 | |
| GO:MF | passive transmembrane transporter activity | GO:0022803 | 0.000322256 | |
| GO:MF | channel activity | GO:0015267 | 0.000322256 | |
| GO:MF | signaling receptor activator activity | GO:0030546 | 0.000481617 | |
| GO:MF | receptor ligand activity | GO:0048018 | 0.000631058 | |
| GO:MF | DNA-binding transcription factor activity | GO:0003700 | 0.000739906 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 3.38E-17 | |
| GO:BP | detection of stimulus involved in sensory perception | GO:0050906 | 3.39E-17 | |
| GO:BP | detection of chemical stimulus involved in sensory perception | GO:0050907 | 3.73E-16 | |
| GO:BP | multicellular organismal process | GO:0032501 | 1.60E-15 | |
| GO:BP | system process | GO:0003008 | 1.60E-15 | |
| GO:BP | detection of chemical stimulus | GO:0009593 | 2.10E-15 | |
| GO:BP | sensory perception | GO:0007600 | 5.11E-15 | |
| GO:BP | sensory perception of chemical stimulus | GO:0007606 | 4.57E-14 | |
| GO:BP | nervous system process | GO:0050877 | 1.96E-13 | |
| GO:BP | sensory perception of smell | GO:0007608 | 3.80E-12 | |
| GO:BP | detection of stimulus | GO:0051606 | 4.66E-11 | |
| GO:BP | adaptive immune response | GO:0002250 | 1.89E-08 | |
| GO:BP | signaling | GO:0023052 | 1.21971E-06 | |
| GO:BP | humoral immune response | GO:0006959 | 9.91038E-06 | |
| GO:BP | defense response to bacterium | GO:0042742 | 1.30477E-05 | |
| GO:BP | cell communication | GO:0007154 | 1.33747E-05 | |
| GO:BP | signal transduction | GO:0007165 | 4.62854E-05 | |
| GO:BP | response to bacterium | GO:0009617 | 8.22478E-05 | |
| GO:BP | cell differentiation | GO:0030154 | 0.000132096 | |
| GO:BP | cell adhesion | GO:0007155 | 0.000194183 | |
| GO:BP | biological adhesion | GO:0022610 | 0.000230205 | |
| GO:BP | cellular developmental process | GO:0048869 | 0.000259699 | |
| GO:BP | keratinization | GO:0031424 | 0.000331074 | |
| GO:BP | epidermal cell differentiation | GO:0009913 | 0.000362886 | |
| GO:BP | G protein-coupled receptor signaling pathway, coupled to cyclic nucleotide second messenger | GO:0007187 | 0.000362886 | |
| GO:BP | regulation of transmembrane transport | GO:0034762 | 0.000362886 | |
| GO:BP | biological regulation | GO:0065007 | 0.000400538 | |
| GO:BP | inflammatory response | GO:0006954 | 0.000400538 | |
| GO:BP | humoral immune response mediated by circulating immunoglobulin | GO:0002455 | 0.000400538 | |
| GO:BP | regulation of ion transport | GO:0043269 | 0.000451217 | |
| GO:BP | keratinocyte differentiation | GO:0030216 | 0.000664627 | |
| GO:BP | regulation of ion transmembrane transport | GO:0034765 | 0.000944702 | |
| GO:BP | regulation of cellular process | GO:0050794 | 0.000998044 | |
| GO:BP | response to stimulus | GO:0050896 | 0.000998044 | |
| GO:CC | plasma membrane | GO:0005886 | 2.21E-24 | |
| GO:CC | cell periphery | GO:0071944 | 3.52E-24 | |
| GO:CC | intrinsic component of plasma membrane | GO:0031226 | 6.58E-09 | |
| GO:CC | integral component of plasma membrane | GO:0005887 | 3.39E-08 | |
| GO:CC | intrinsic component of membrane | GO:0031224 | 1.36001E-06 | |
| GO:CC | plasma membrane protein complex | GO:0098797 | 1.73042E-06 | |
| GO:CC | integral component of membrane | GO:0016021 | 8.63394E-06 | |
| GO:CC | extracellular region | GO:0005576 | 0.00012106 | |
| GO:CC | receptor complex | GO:0043235 | 0.000138826 | |
| GO:CC | intermediate filament | GO:0005882 | 0.000227515 | |
| GO:CC | plasma membrane region | GO:0098590 | 0.000400338 | |

GO(Gene Ontology) enrichment analysis for 2173 switching genes in SK-MEL-5 was conducted by g:Profiler(FMD: 31066453) with following parameters:

- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

* MF, BP, CC represent Molecular Function, Biological Process, Cellular Component, respectively.

Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (SK-N-DZ, continued)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|---|------------|------------------|---|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 0.000446565 |  |
| GO:BP | multicellular organismal process | GO:0032501 | 2.27025E-05 |  |
| GO:CC | no significant results | | | |

GO(Gene Ontology) enrichment analysis for 768 switching genes in SK-N-DZ was conducted by g:Profiler(PMID: 31066453) with following parameters:

- Hypergeometric test
- Benjamini–Hochberg correction
- FDR(False Discovery Rate) < 0.001

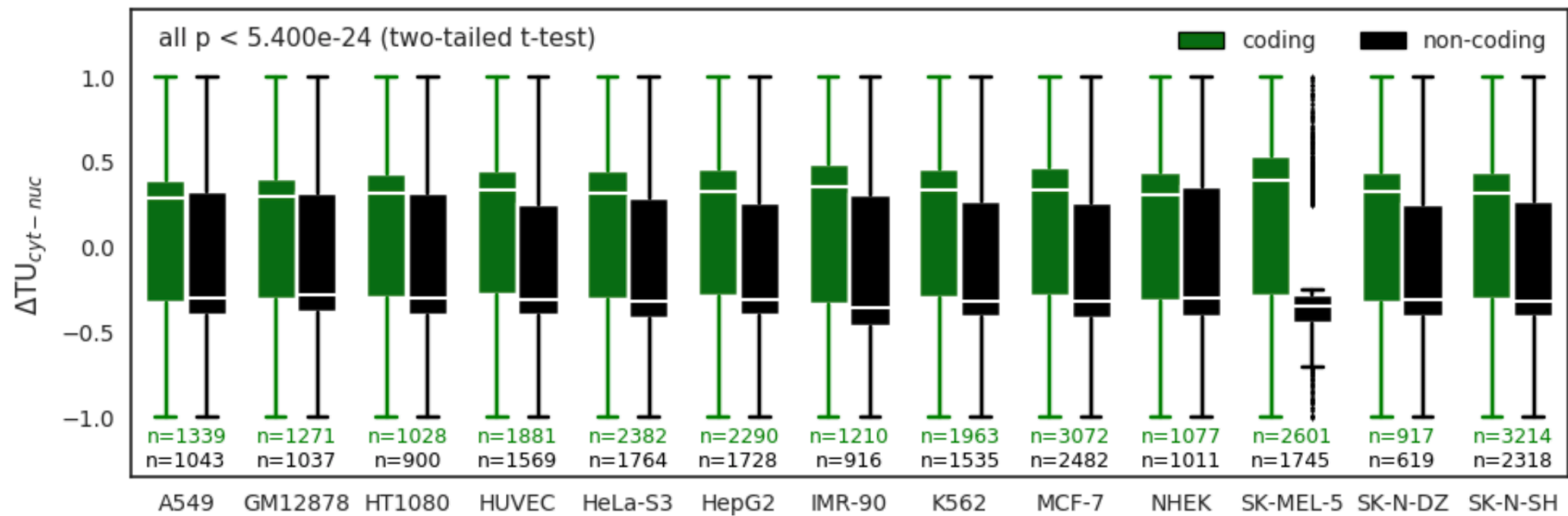
Supplementary Data 5. Gene Ontology enrichment analysis for switch genes (SK-N-SH)

| source | term_name | term_id | adjusted_p_value | negative_log10_of_adjusted_p_value |
|--------|--|------------|------------------|------------------------------------|
| GO:MF | transmembrane signaling receptor activity | GO:0004888 | 3.19E-16 | |
| GO:MF | signaling receptor activity | GO:0038023 | 3.19E-16 | |
| GO:MF | G protein-coupled receptor activity | GO:0004930 | 5.19E-16 | |
| GO:MF | molecular transducer activity | GO:0060089 | 5.19E-16 | |
| GO:MF | olfactory receptor activity | GO:0004984 | 2.89E-15 | |
| GO:MF | antigen binding | GO:0003823 | 0.000131859 | |
| GO:BP | detection of chemical stimulus | GO:0009593 | 3.47E-15 | |
| GO:BP | detection of chemical stimulus involved in sensory perception | GO:0050907 | 8.52E-15 | |
| GO:BP | G protein-coupled receptor signaling pathway | GO:0007186 | 8.77E-15 | |
| GO:BP | sensory perception of chemical stimulus | GO:0007606 | 1.30E-14 | |
| GO:BP | detection of chemical stimulus involved in sensory perception of smell | GO:0050911 | 6.08E-14 | |
| GO:BP | sensory perception of smell | GO:0007608 | 2.32E-12 | |
| GO:BP | nervous system process | GO:0050877 | 5.53E-12 | |
| GO:BP | detection of stimulus involved in sensory perception | GO:0050906 | 8.87E-12 | |
| GO:BP | system process | GO:0003008 | 1.37E-10 | |
| GO:BP | sensory perception | GO:0007600 | 2.82E-10 | |
| GO:BP | multicellular organismal process | GO:0032501 | 1.60E-09 | |
| GO:BP | detection of stimulus | GO:0051606 | 2.96E-07 | |
| GO:BP | keratinization | GO:0031424 | 5.33503E-06 | |
| GO:BP | defense response to bacterium | GO:0042742 | 2.54298E-05 | |
| GO:BP | adaptive immune response | GO:0002250 | 8.74689E-05 | |
| GO:BP | signaling | GO:0023052 | 0.000829655 | |
| GO:BP | humoral immune response mediated by circulating immunoglobulin | GO:0002455 | 0.000881868 | |
| GO:CC | plasma membrane | GO:0005886 | 2.24E-19 | |
| GO:CC | cell periphery | GO:0071944 | 2.11E-18 | |
| GO:CC | extracellular region | GO:0005576 | 1.40E-08 | |
| GO:CC | intrinsic component of membrane | GO:0031224 | 6.73E-08 | |
| GO:CC | integral component of membrane | GO:0016021 | 7.91E-08 | |
| GO:CC | membrane | GO:0016020 | 2.26958E-05 | |
| GO:CC | plasma membrane region | GO:0098590 | 8.71218E-05 | |
| GO:CC | receptor complex | GO:0043235 | 8.71218E-05 | |
| GO:CC | extracellular space | GO:0005615 | 8.71218E-05 | |
| GO:CC | cell surface | GO:0009986 | 8.72698E-05 | |
| GO:CC | intrinsic component of plasma membrane | GO:0031226 | 0.000264325 | |
| GO:CC | integral component of plasma membrane | GO:0005887 | 0.000455522 | |

GO(Gene Ontology) enrichment analysis for 2766 switching genes in SK-N-SH was conducted by g:Profiler(PMID: 31066453) with following parameters:

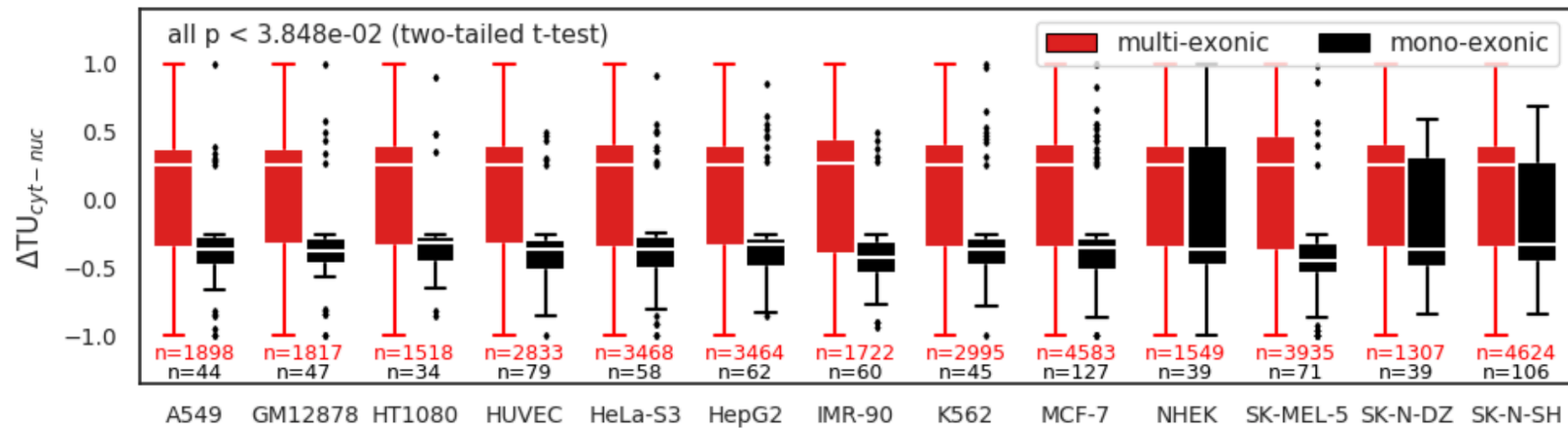
- Hypergeometric test
- Benjamini-Hochberg correction
- FDR(False Discovery Rate) < 0.001

* MF, BP, CC represent Molecular Function, Biological Process, Cellular Component, respectively.

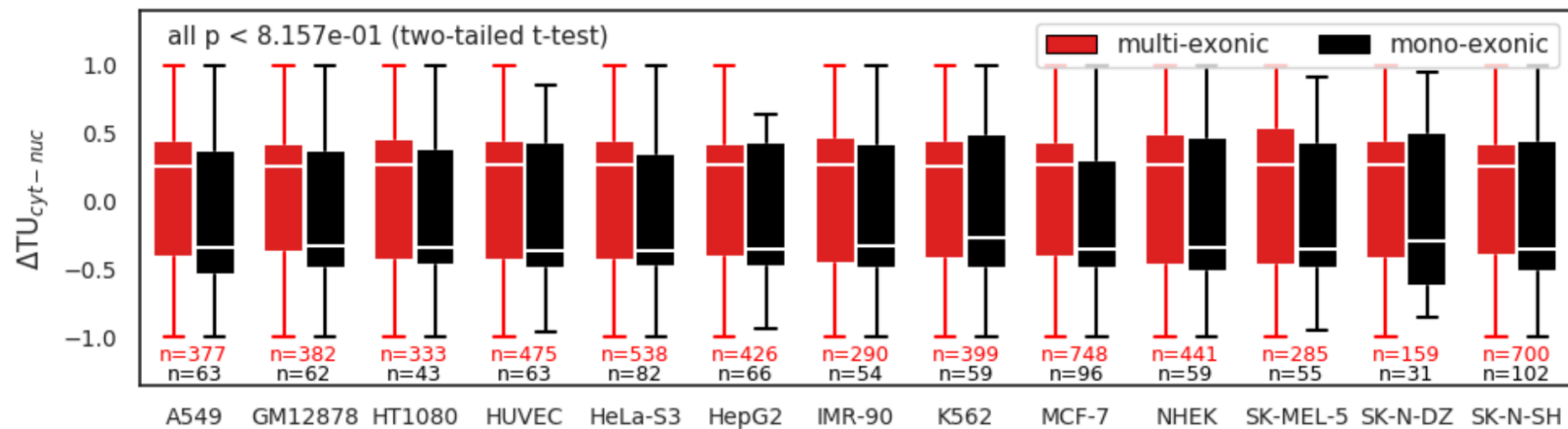


Supplementary Data 6. Comparison of ΔTU between protein-coding transcripts (green) and non-coding transcripts (black). Protein-coding transcripts possess higher ΔTU scores, which is consistent with our understanding that the transcript encoding protein is more prone to be located in the cytoplasm. Based on two-tailed t-test, we calculated the significant level of difference between ΔTU in protein-coding and non-coding transcripts in each cell line. The largest p value is 5.400e-24.

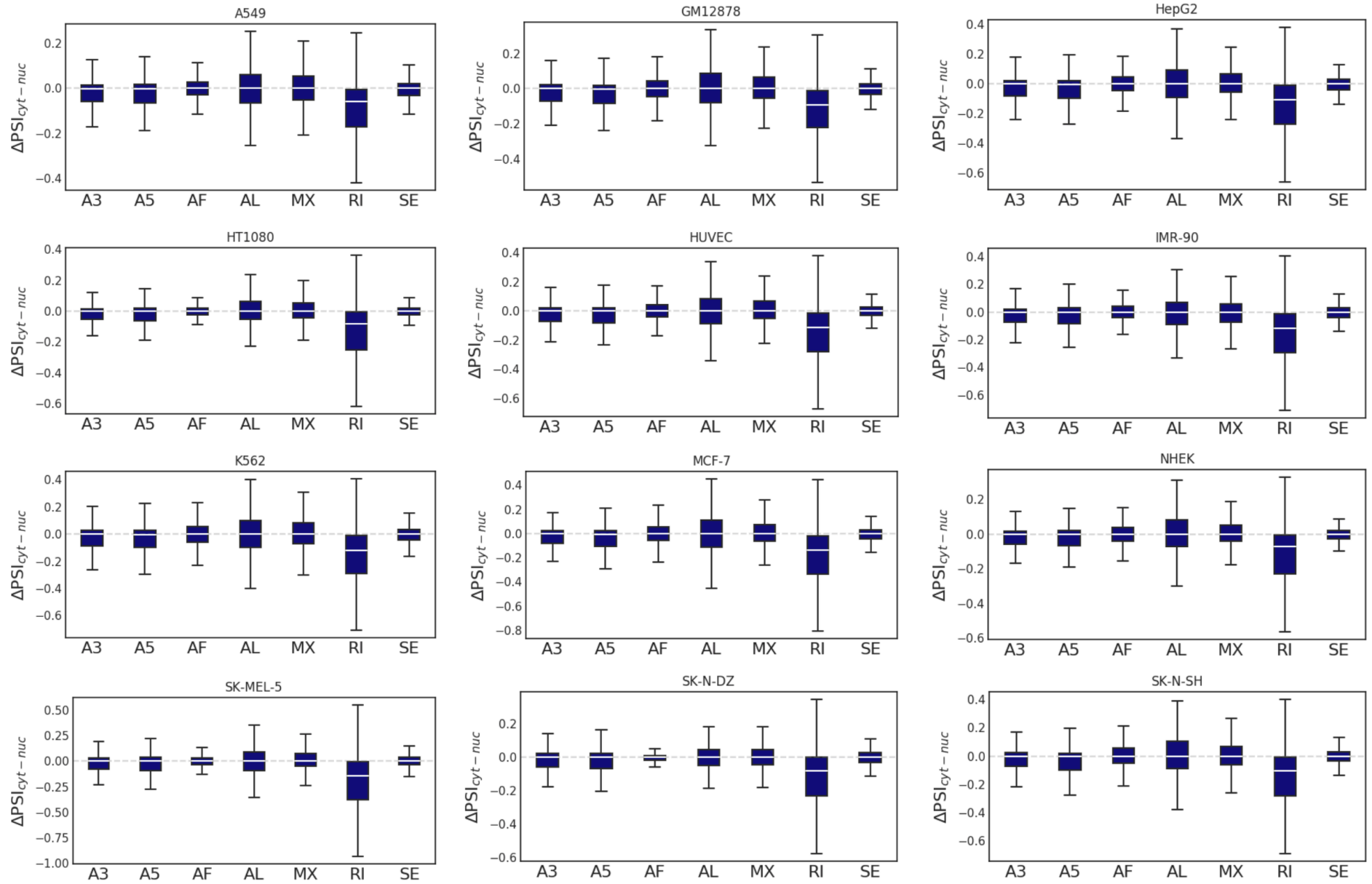
Protein coding genes



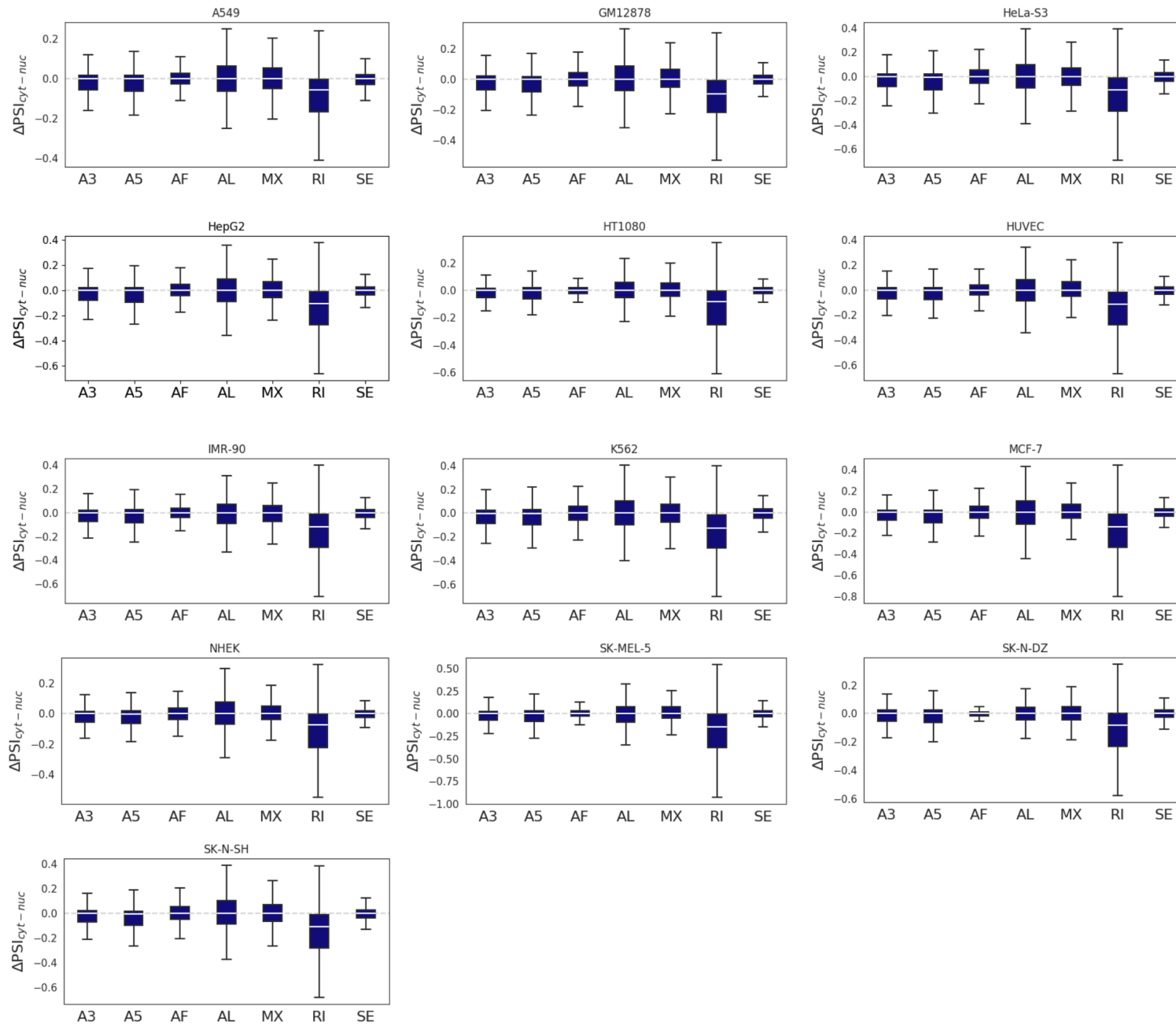
Noncoding genes



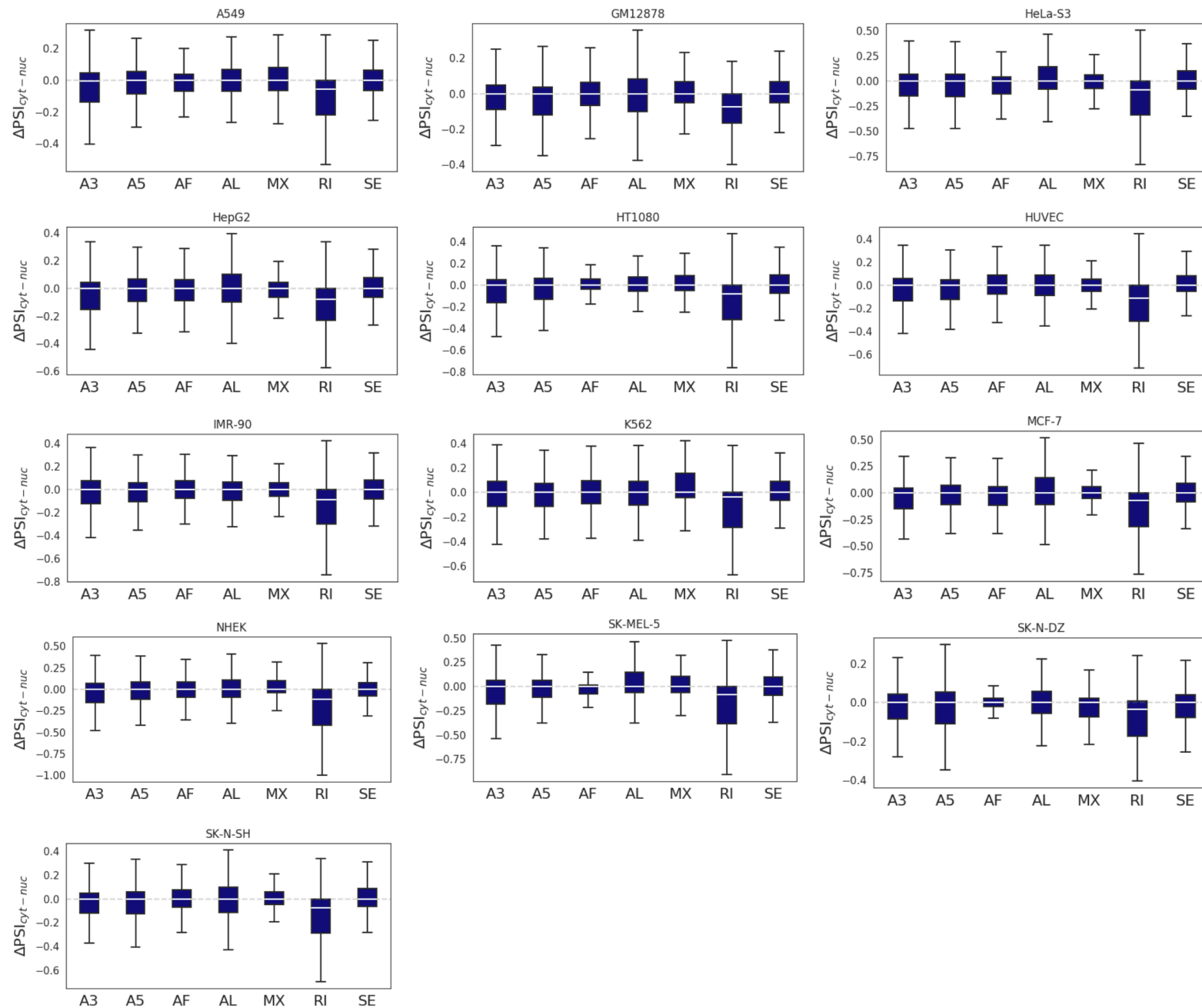
Supplementary Data 7. Comparison of ΔTU between mono-exonic (black) and multi-exonic (red) transcripts across 13 cell lines. Top and bottom panels for protein-coding and noncoding genes, respectively. ΔTU shows a significant positive correlation with splicing, indicating that splicing appears to be a dominant factor for RNA export from the nucleus. Based on two-tailed t-test, we calculated the significant level of difference between ΔTU in multi-exonic and mono-exonic transcripts in each cell line. The largest p value is 3.848e-02 and 8.157e-01 for protein-coding and non-coding genes, respectively.



Supplementary Data 8. Comparison of alternative splicing patterns between cytoplasmic and nuclear transcripts. (all genes)

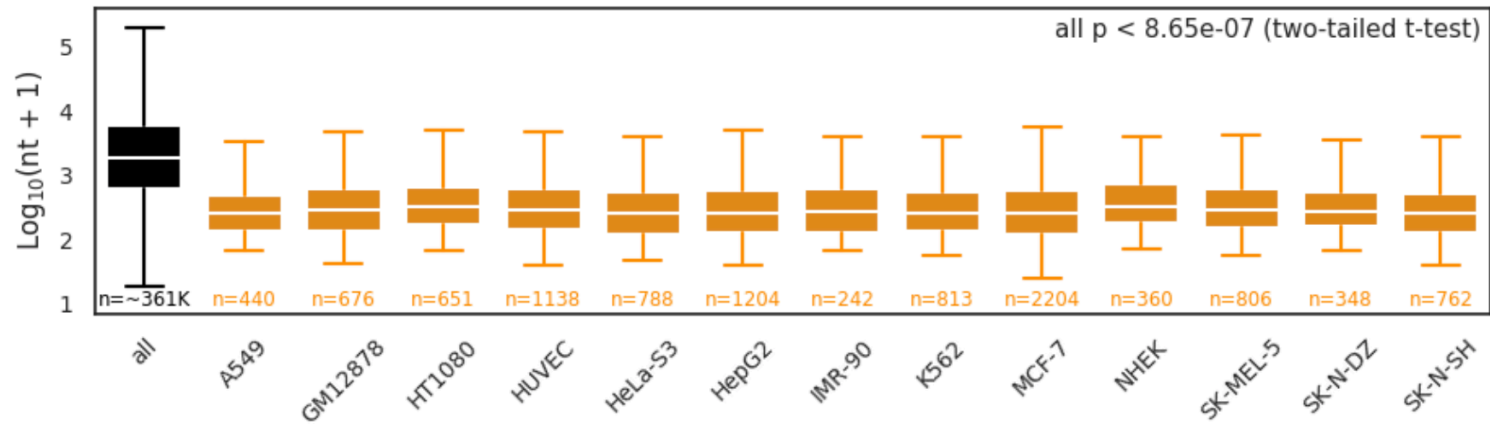


Supplementary Data 9. Comparison of alternative splicing patterns between cytoplasmic and nuclear transcripts. (protein-coding genes, continued)

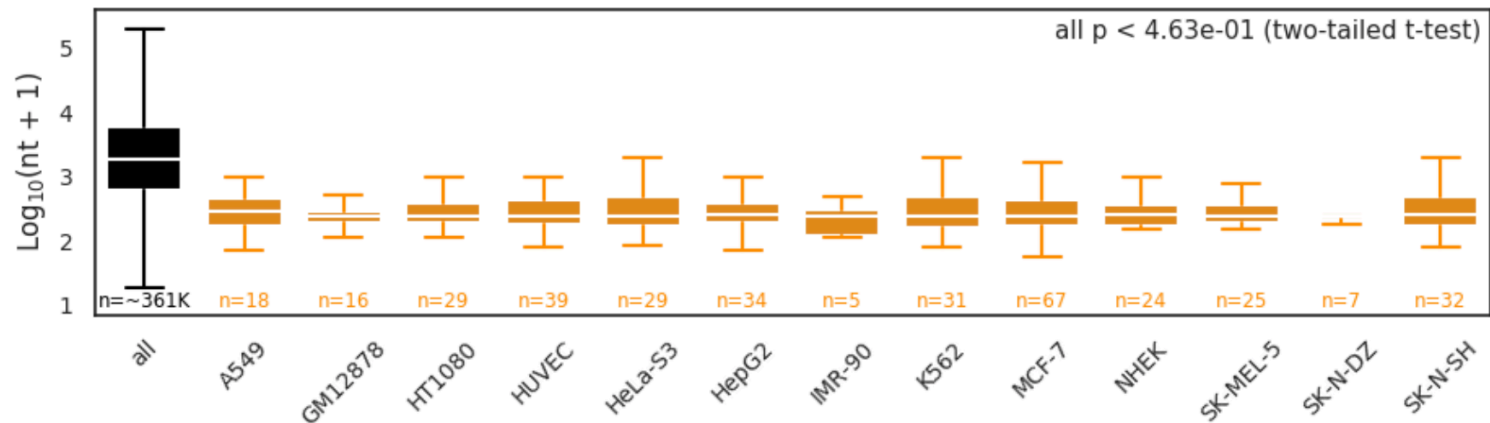


Supplementary Data 9. Comparison of alternative splicing patterns between cytoplasmic and nuclear transcripts. (noncoding genes)

Protein coding genes



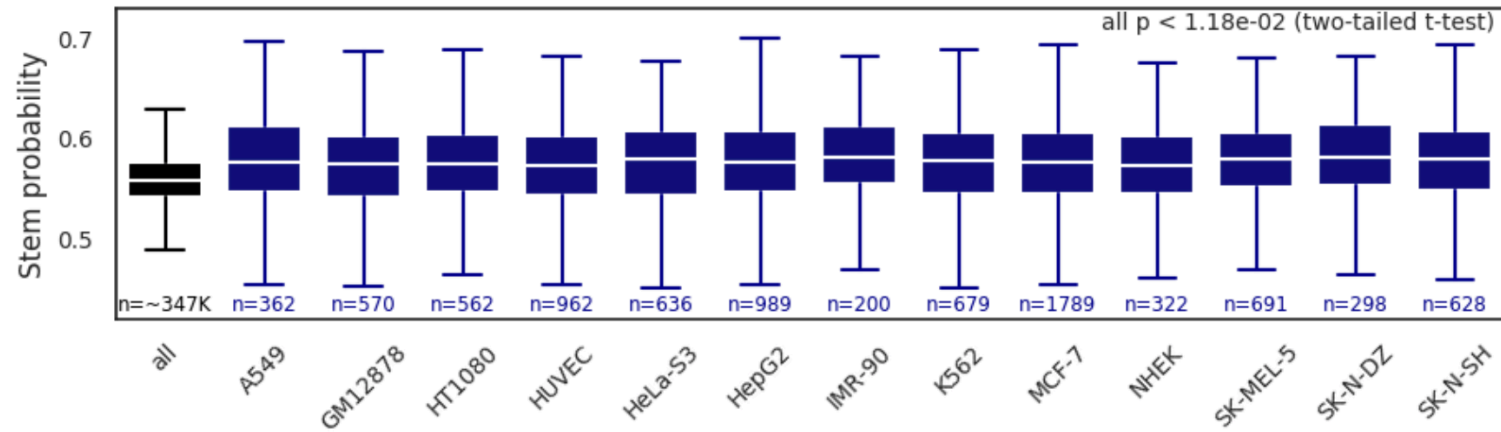
Noncoding genes



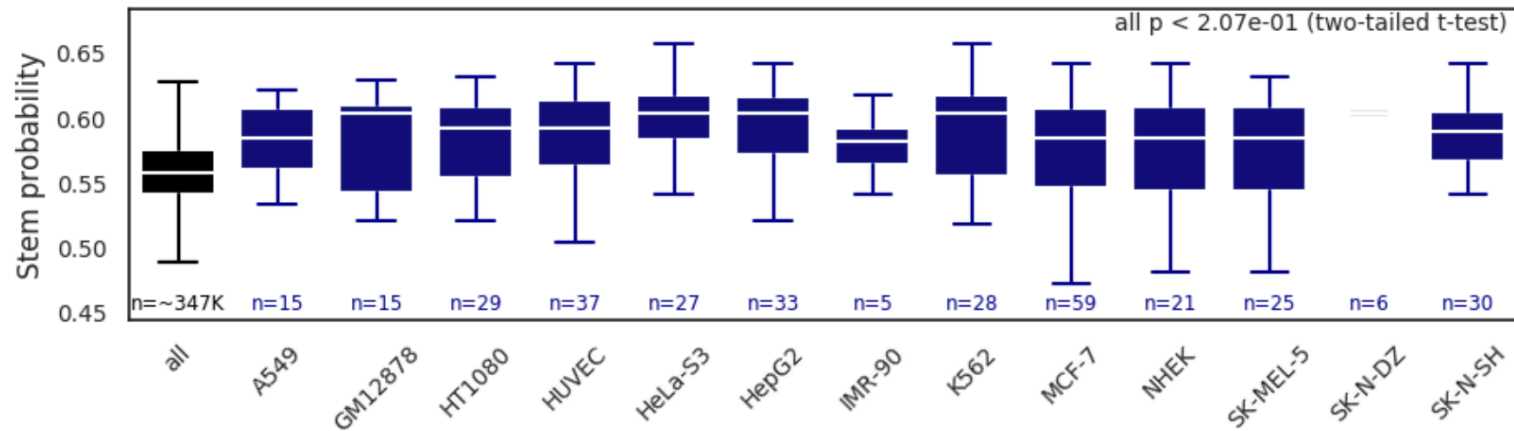
Supplementary Data 10. Comparison of length between all introns and nuclear RIs ($\Delta\Psi < 0$ and $p < 0.05$).

Top and bottom panels for protein-coding and noncoding genes, respectively. P values: two-tailed t-test between all introns and nuclear RIs. The largest p value is 8.65e-07 and 4.63e-01 for protein-coding and non-coding genes, respectively.

Protein coding genes



Noncoding genes



Supplementary Data 11. Comparison of RNA secondary structure (stem probability) between all introns and nuclear RIs ($\Delta\Psi < 0$ and $p < 0.05$). Top and bottom panels for protein-coding and noncoding genes, respectively. P values: two-tailed t-test between all introns and nuclear RIs. The largest p value is 1.18e-02 and 2.07e-01 for protein-coding and non-coding genes, respectively.