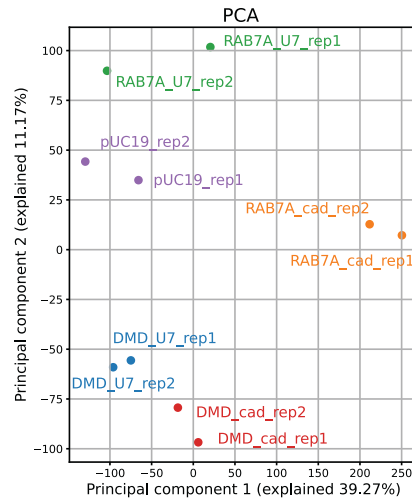


Enhancing RNA base editing on mammalian transcripts with small nuclear RNAs

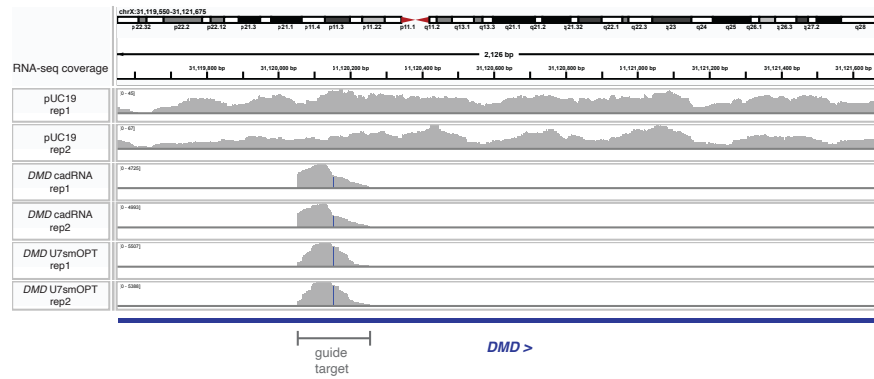
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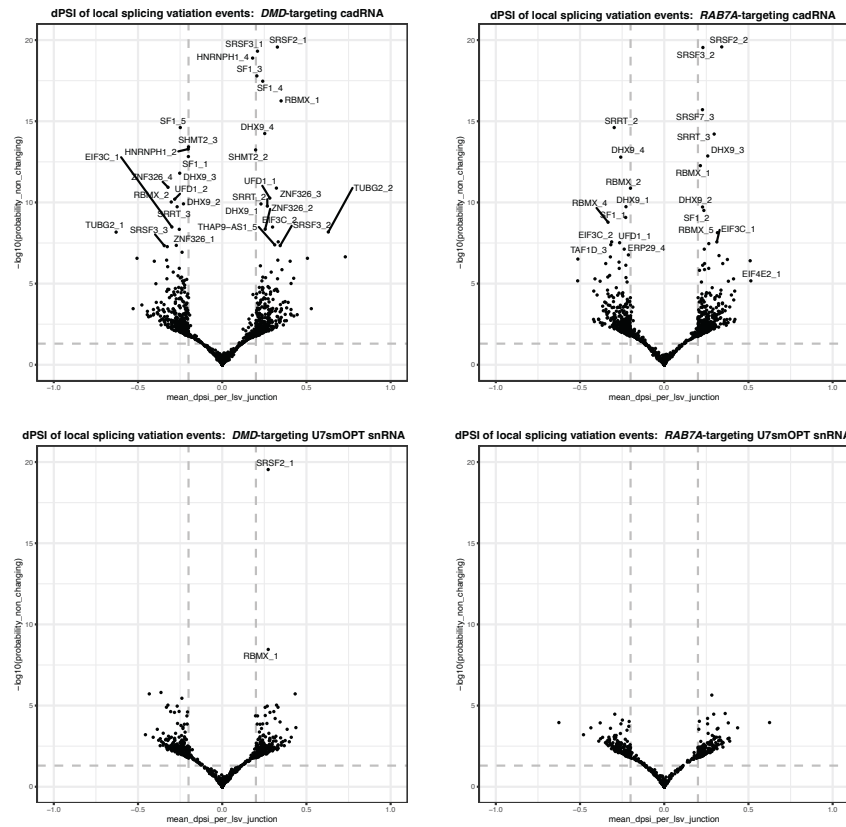
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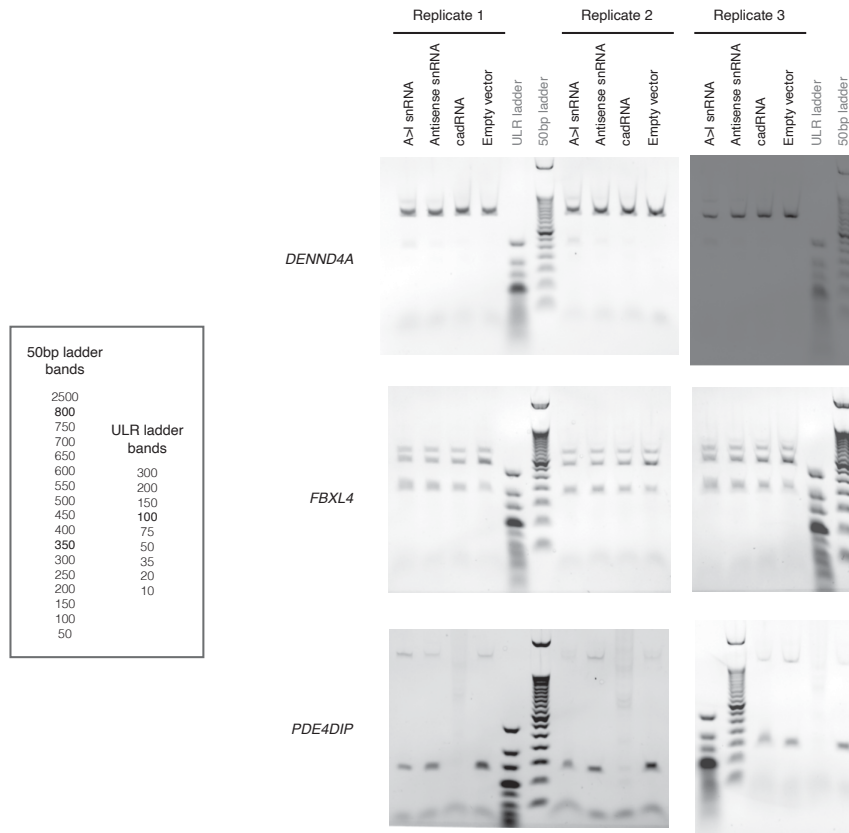
Supplementary Figure 1: Principal Component Analysis (PCA) plot of RNA-guided A>I base editor RNA sequencing data. PCA plot of RNA sequencing samples analyzed in Fig. 2, with each of two replicates colored by sample (pUC19, *RAB7A*-targeting cadRNA, *RAB7A*-targeting U7smOPT snRNA, *DMD*-targeting cadRNA, and *DMD*-targeting U7smOPT snRNA).



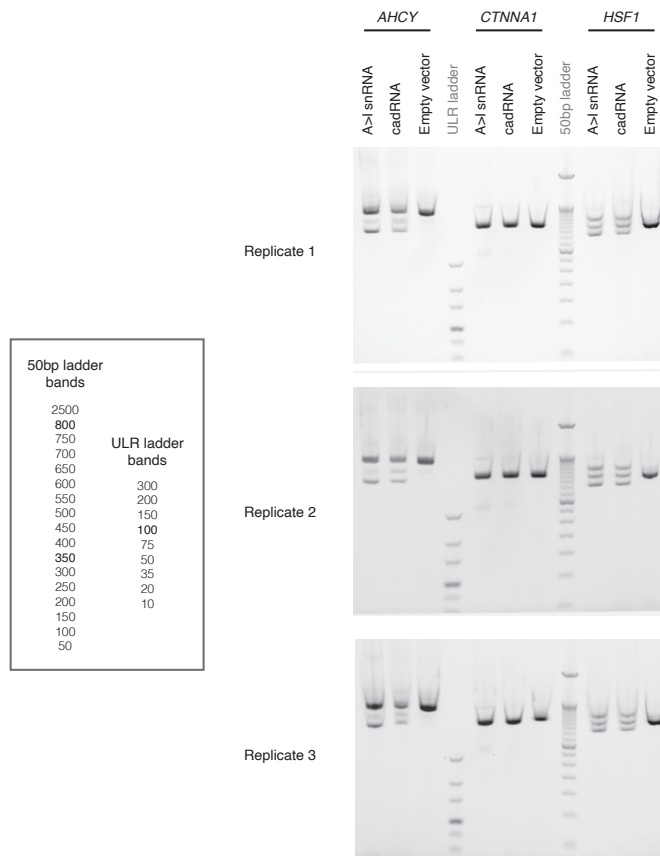
Supplementary Figure 2: Alignment of RNA-guided A>I base editor RNA sequencing reads to *DMD* gene. Alignment showing RNA sequencing read pile-up across *DMD*-targeting cadRNA and U7smOPT snRNA replicates at the guide target on *DMD*.



Supplementary Figure 3: Differential splicing analysis of RNA-guided A>I base editor RNA sequencing data. Scatterplots of local splicing variations (LSVs) against empty control (pUC19) of cadRNA backbone vs. U7smOPT snRNA backbone for *RAB7A*- and *DMD*-targeting guides. Cutoffs for significance are p -value < 0.05 and various dPSI (differential Percent Spliced In) values used in Fig. 2.



Supplementary Figure 4: Splicing gels for A>I editing-targeted pre-mRNA of *DENND4A*, *FBXL4*, and *PDE4DIP*. Full RT-PCR gels for all replicates of experiments in Fig. 4c. n = 3 biological replicates per condition.



Supplementary Figure 5: Splicing gels for A>I editing-targeted pre-mRNA of *AHCY*, *CTNNA1*, and *HSF1*. Full RT-PCR gels for all replicates of experiments in Figure 4d. n = 3 biological replicates per condition.

Supplementary Dataset 1: DESeq2 output for Fig. 2.

Spreadsheet containing standard DESeq2 output for conditions tested in Fig. 2 (RAB7A-targeting cadRNA, RAB7A-targeting U7smOPT snRNA, DMD-targeting cadRNA, and DMD-targeting U7smOPT snRNA) relative to pUC19 control.

Supplementary Table 1: Plasmid sequences.

Name	Sequence
U1 promoter-terminator-U1_snRNA backbone	taacacaggctaaggaccagcttcttgggagagaacagacgcaggggcgggagggaaaaagggagaggcagacgtcacttccccttggcggctctggcagcagattggtcggttgagtggcagaaaaggcagacggggactgggcaaggcactgtcgggtgacatcacggacagggcgacttctatgtagatgaggcagcgcagaggctgctgcttcgcacttgcgtcttcaccacgaaggagtccccgtgccctgggagcgggttcaggaccgctgatcggaaagtgagaatcccagctgtgtgtcagggctggaaagggtcgggagtgcgcggggcaagtgaccgtgtgtgtaaagagttaggcgtatgaggctgtgtcggggcagaggcccaagatctca<guide>gcaggggagataccatgatcacgaaggtggtttccaggcgaggcttatccattgcactccggatgtgctgacccctgcgatttcccaaatgtgggaaactcgactgcataatttggtgtagtgggggactgcgttcgcgctttccctgactttctggagtttcaaaagtagactgtacgctaa
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Lentiviral empty vector containing Ef1a_core- eGFP- PuroR cassette	gaagatcctttgatctttctacggggtctgacgctcagtggaaacgaaaactcacgttaagggattttggatcagaga ttataaaaaaggatcttcacctagatccttttaataaaaaatgaagtttaaatcaatctaaagtatatatgagtaaactt ggtctgacagttaccaatgcttaacagtgaggcacctatctcagcgatctgtctatttcgttcacatagttgcctg actccccgtcgtgtagataactacgatacgggagggcttaccatctggccccagtgtgcaatgataaccgcgaga cccacgctcaccggctccagatttatcgcaataaaccagccagccggaaggggccgagcgagaaagtgtcct gcaactttatccgctccatccagcttattaattgttgcgggaagctagagtaagttagtcgccagttaatagttgc gcaacgttgttgccattgctacaggcatcgtggtgtcacgctcgtcgtttggtatggcttcattcagctccggtccc aacgatcaaggcgagttacatgatccccatgttgtgcaaaaaagcggttagctccttcggtctccgatcgttgtc agaagtaagttggccgagtggtatcactcatggttatggcagcactgcataattctcttactgtcatgccatccgtaa gatgcttttctgtgactggtgagtactcaaccaagtcattctgagaatagtgtatgcggcgaccgagttgctcttgc cggcgtcaatacgggataataaccgcgccacatagcagaactttaaaagtgtcatcattggaaaacgttcttcggg gcgaaaactctcaaggatcttaccgctgttgatccagttcgtatgtaaccactcgtgcacccaactgatcttcag catcttttactttcaccagcggttctgggtgagcaaaaacaggaaggcaaaatgccgcaaaaaagggaataaggg cgacacggaaatgtgaatactcatacttctcttttcaatattattgaagcatttatcaggggtattgtctcatgagcg gatacataattgaatgtatttagaaaaataaacaataaggggtccgcgcacattccccgaaaagtgccacctgac gtcgacggatcgggagatctccgatccctatggtgcactctcagtacaatctgctctgatccgcatagttaag ccagtatctgctccctgcttgtgtgttgaggtcgtgagtgtgcgcgagcaaaatttaagctacaacaaggcaa ggcttgaccgacaattgcatgaagaatctgcttaggggttaggcgttttgcgtgcttcgcgatgtacgggccagata tacgcgttgacattgattattgactagtatttaataagtaataacacggggtcattagttcatagcccatataggagtt ccgcgttacataactacggtaaatggcccgcctggctgaccgccaacgacccccgccattgacgtcaataat gacgtatgttccatagtaacgccaatagggactttccattgacgtcaatgggtggagttattacggtaaacgtccc acttggcagtagcatcaagtgtatcatatgccaaagtacgccccctattgacgtcaatgacggtaaatggcccgcctg gcattatgccagtagcatgacctatgggacttctacttggcagtagcatctacgtattagtcacgtattaccatgg

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	cgggtcaccgagctgcaagaactcttctcacgcgcgtcgggctcgacatcggaaggtgtgggtcgcggacg acggcgccgcggtggcggtctggaccacgccggagagcgtcgaagcggggcggtgttcgccgagatcggc ccgcgcacatggccgagttgagcgggtcccggtggccgcgcagcaacagatggaaggcctctggcgccgcac cggcccaaggagcccgctgggttctggccaccgtcggagttctcgccgaccaccaggggaagggtctgggc agcgccgtcgtgctccccggagtgaggcgccgagcgcgcgggggtccccgccttctggagacctccgc gccccgaacctcccccttctacgagcggctcggcttcaccgtcaccgccgacgtcgaggtgccgaaggaccg cgcacctggtgcatgaccgcaagcccggtgcgggttcggcgcaacaaacttctctctgctgaacaagccgg agatgtcgaagagaatcctggaccggtgagcaaggcgagggagctgttcaccgggggtggtgccatctgtgtc gagctggacggcgacgtaaacggccacaagttcagcgtgtccggcgaggcgagggcgatgccacctacgg caagctgacctgaagttcatctgcaccaccggcaagctgcccgtgccctggcccacctcgtgaccacctga ctacggcgtgcatgtcttcagccgtaccccgaccacatgaagcagcacgacttctcaagtccgcatgccc gaaggctacgtccaggagcgcaccatcttctcaaggacgacggcaactacaagaccgcgccgaggtgaagt tcgagggcgacacctgtgaaccgcatcgagctgaaggcgatcgacttcaaggaggacggcaacatctgg ggcacaagctggagtacaactacaacagccacaacgtctatatcatggccgacaagcagaagaacggcatcaa ggtgaactcaagatccgccacaacatcgaggacggcagcgtgcagctcgccgaccactaccagcagaacac ccccatcggcgacggccccgtgctgctgcccgaaccactacctgagcaccagtcgccctgagcaagac cccaacgagaagcgcgatcacatggctctgctggagttcgtgaccgccggggtacactctcgcatggacg agctgtacaagtaactcgagcgcgttaagtcgacaatcaacctctggattacaaaatttgtaaagattgactggta ttcttaactatgttgctcctttacgctatgtggatacgtgctttaatgcccttgatcatgtattgcttcccgatggctt tcattttctcctctgtataaatcctggtgctgctcttcttatgaggagttgtggccggtgtcaggcaacgtggcggtg gtgtgactgtgttgcgtgacgcaacccccactggttggggcattgccaccacctgtcagctccttccgggacttt cgcttccccctccctattgccacggcggaactcatcgccgctgccttggccgctgctggacaggggctcggt gttgggcactgacaattccgtggtgtgtcggggaaatcatcgtccttcttggctgctcgctgtgttgcacctg gattctgcgcgggacgtccttctgtacgtcccttcggccctcaatccagcggaccttcttccgcggcctgtg ccggctctgcggcctcttccgcgtcttcgccttcgccctcagacgagtcggatctcccttggggccgctccccgc gtcactttaagaccaatgacttacaaggcagctgtagatcttagccactttttaaagaaaaggggggactggaa gggctaattcaactccaacgaagacaagatctgcttttgcctgtactgggtctctctggttagaccagatctgagcc tgggagctctctggctaactaggggaacccactgcttaagcctcaataaagcttgccttgagtgttcaagtagtgtg tgcccgctgtgtgtgactctggttaactagatccctcagaccctttagtcagtgtgaaaatctctagcagggc ccgtttaaacccgctgatcagcctcactgtgccttctagtggcagccatctgtgttggccctccccgtgccttc cttgacctggaaggtgccactcccactgtccttctcctaataaaatgaggaaattgcacgcattgtctgagtaggtg tcattctattctgggggggtgggggtggggcaggacagcaaggggaggattgggaagacaatagcaggcatgct ggggatgcgggtgggctctatggcttctgaggcggaaagaaccagctggggctctagggggtatccccacgcgc cctgtagcggcgcatlaagcgcggcggtgtgtgtgttacgcgcagcgtgaccgctacacttgcagcgcctta gcgcccgtccttctgcttcttcccttcttctcgcacgttcgcgggttccccgtcaagctctaaatcgggggc tcccttaggggtccgatttagtgctttacggcacctcgacccaaaaaacttgattaggggtgatggtcacgtagtg ggccatcgccctgatagacggttttcgcctttgacgttggagtcacggttcttaatagtggactcttgttcaaaat ggaacaacactcaacctatctcggtctattctttgattataagggattttgccgatttcggcctattgttaaaaat gagctgatttaacaaaaattaacgcgaattaattctgtggaatgtgtgcagttagggtgtggaaagtccccaggct ccccagcaggcagaagtatgcaaagcatgcattcaattagtcagcaaccaggtgtggaaagtccccaggctcc ccagcaggcagaagtatgcaaagcatgcattcaattagtcagcaaccatagtcggcccttaactccgcccac ccgcccctaactccgcccagttccgcccattctccgcccattggctgactaatttttttatatgcagaggccgag gccgctctgcctctgagctattccagaagtagtgaggaggctttttggaggcctaggcttttgcaaaaagctccc gggagcttgtatataccatttccggtctgatcagcacgtgttgacaattaatcatcgccatagtatatcgccatagtat aatacacaaggtgaggaactaaacatggccaagttagaccagtgcggttccggtgctaccgcgcgcgacgtc gccggagcgggtcagttctggaccgaccggctcgggttctccgggacttcgtggaggacgacttcgccggtgt
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	gggccgggacgacgtgacctgttcacagcgcggtccaggaccaggtgggtgccggacaacacctggcctgg gtgtgggtgcgcggcctggacgagctgtacgccgagtggtcggaggtcgtgtccacgaactccgggacgcct ccgggcccggccatgaccgagatcggcgagcagccgtggggggcgggagttcgcctgcgcgacccggccgg caactgcgtgcaacttcgtggccgaggagcaggactgacacgtgctacgagatttcgattccaccgccgccttctat gaaaggttgggcttcggaatcgtttccgggacgcggtggtgatcctccagcgcggggatctcatgtggag ttctcgcaccccaactgtttattgcagcttataatggttacaataaagcaatagcatcacaatttcacaaataa agcattttttcactgcattctagtgtgtgttgcctcaaacatcaatgtatcttatcatgtctgtataccgtcgacctta gctagagcttggcgtaatcatggtcatagctgttccctgtgtgaaattgttatccgtcacaattccacacaacatac agccggaaagcataaagtgtaaagcctggggtgcctaatagtgagtaactcacattaattgcgttgcgtcactg cccgtttccagtcgggaaacctgtcgtgccagctgcattaatgaatcgccaacgcgcggggagagggcggtt gcgtattgggcgtcttccgttcctcgtcactgactcgtcgcgtcgggtcgttcgggtcgggcgagcgggtatca gctcactcaaaggcggtataacggttatccacagaatcaggggataacgcaggaaagaacatgtgagcaaaag gccagcaaaaggccaggaaccgtaaaaaggccgcgttgcgtggcgtttttccataggctccgccccctgacgag catcacaataatcgacgtcaagtcagaggtggcgaaaccgcagaggactataaagataaccaggcgtttcccc tggaagctccctcgtgcgtctcctgttccgacctgccgcttaccggatacctgtccgcctttctccctcgggaa gcgtggcgctttctcatagctcacgtgtaggtatctcagttcgggtgtaggtcgttcgctccaagctgggctgtgtgc acgaacccccgttcagcccagcgtgcgccttatccggttaactatcgtcttgagtccaacccggtaagacacg acttatcgccactggcagcagccactggtaacaggattagcagagcgaggtatgtaggcggtgctacagagttct tgaagtggcctaactacggctacactagaagaacagtatttggtatctgcgctctgctgaagccagttaccttc ggaaaaagagttggtagctcttgatccggcaaaacaaccaccgctggtagcgggtggtttttgtttgcaagcagc agattacgcgcagaaaaaaaggatctcaa
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Supplementary Table 2: Guide and snoRNA sequences.

Name	Sequence
RAB7A A>I guide	AGACAGTTGTCCCCCTGGAGAGATGAAATCGATGTTGGCTCTT AATGGAAAGATAAAAGGCGTACATTCAAGATGTGTCTACTGTA CAGAATACTGCCGCCAGCTGCTAATCCCAATTCTGAGTATGTGT CTGCAATCCAAACACCCATCAACCCTCCACCTTTGTGCGCCTGCA TTACAGGAGAATAACACATAATCCAA
DAXX A>I guide	GCTCCTGTAACCTGATGCCACATCTCGGAAGGCATCCTGAGC CATGAGCTGGAGCTGCTGTCTGGGGGAGGCCAAGGCTGTGTCTG GGCAGCTGCCTTCTCCACAGCCCGAAGCACATCCCCATAGTCA GGGAAGGTATCAGGCCCTGGCTTGTTGATGAGCCGCTCAATGC GCCTGTTAACCTCTGGGTAGCGGGTGCCA
GAPDH A>I guide	GGCCATCCACAGTCTTCTGGGTGGCAGTCTACGCATGGACTGT GGTCTACTGTCCTTCCACGATACGTTCTGTTGTCATGGATGACCT TGGCCAGGGGTGCCAAGCACAACTGGTGCAGGAGGCAAACG TGATGATCTTGAGGCACAAGTCATACTTCTCATGCAAGACACCC ATGACGAACATGGGGGCATCAGCAGAG
TARDBP A>I guide	TGTGTTTCATATTCCGTAAAACGAACAATCGGAAACCCCTTTGA ATGTGGTGTCTTAAGATCTTTCAACTCCTGCACCATAAGAACTT CTCCAAAGGTACCAAAATTGAGTTTCAGGTCCTGTTCCCAAGT GTTCCATGGGAGAGGGTACACTATTAAATCGGTACAGTTCTGGA CTGCTCTTGTCACCTTTCCTGCTGA
ALDOA A>I guide	GTCCGTCCTTCTTGTACTGGGCACAGCGCTCAGACAGCCCATC CAACCCTTGGGTGGTAGTCTCGCCATTTGTCCCTGCCAGGGGG ACCACGCCCTTGTCCACCTTGATGCCCAACACCGCCCTTGG ATTTGATAACTTGGGGGAAGGGACGCCCATCATCCGCCTTCTGG TAGAGTGTCTCATGGAAGAGGATGACAC
SMAD4 A>I guide	TTGTAGTCCACCATCCTGATAAGGTAAACCCGCCCAACGGTAAA AGAGGAGAGTCTAAAGGTTGTGCCAGTGCAATCGGCATGGTAT GAAGTACTTCGTCCAGGAGGACCAGGGCCCGGTGTAAGACGG TTTCAATCCAGCAAGCACATTCTTTGATGCTCTGAGAAGGGTAA TCCGGTCCCCAGCCTTTCACAAAACCTC
FANCC A>I guide	GAGTCTGGGCTGAGGGACCTGGCTCTGCTAAATGTAAAATAGA TACTTCGTGATTGTCCCAAGATGACATCAGCTCATTCTCACAGC CCAGCGAGGGCACCTACTCGTGTAATGCGTGGCCACAGCAGTT CACCTGTCCTGTGGGGGAGGCGAGCCTGATCCCAGTGGCCGGG CACCACACGGCCTGCGTGCCTTCTAG
GAA A>I guide	tggtgtccagggggccggcagcgtcaccactgccctcgctgtggatggctggctcacggggagc tgcaggtgggggtgggaggtgccaaagggcctcCactggcaccgtctgcaggtcgtaccatgtgcc aaggggaagtagccagtcacttcggccttccggcctggagcactgggggtgatgagcagggcctccc
BLM A>I guide	gtggtgtaacaaatgattttgaagtctcagaagtatcaaagtcacatcatcccaatcattgatgtact taaagaatctggtgaagaactaaattcCaatttcttgagagcagtatcccgggatttctttacagttggtgt gttttgggtagtgcatacaacttccttcggagtctgcaagaaatctggcaataatg

CHD7 A>I guide	gttcatactgattatagtgatcaaaatgtgtcagctttgtttgattttgattagttgaaggatgatgaagggat ggctgtaaagaggcaaagccttggtcCattggcatttgctgacctataggattactggattttccgggta accacattctccgaggccttcaagaccttcactgaaaatattcccatcctcgccaa
FANCD2 A>I guide	acttaatctcttcaccctgcaagteccgggttttttagattgccagccagaaagcctctctacaattgttgag agtgagcatagctttgactctgcaaacCaaaagttccagggtcttttgagcagaggcacatgttgggtg agtctcgtgtcctggatcttggatgccacacaggtgatgaagcagccttgtgt
COL4A5 A>I guide	actgcacctaggaattttatatatatatcacattttaaacaacaaaaggaattctcaaatgttatgtc ctctcatgcacacttgacatcggcCaattcgtgtcctcaagtcctgtcttcagcgtttctgactgaggtt tactgaacatgtctgacacatctacagttgccagccaaaagctgtaggagtgtg
DMD A>I guide	aaagctaattacacttgatgtcagcccactctccaaaagctaattacacttgatgtcagaggtaacagattt gcaaaattataggtcacacgggtgtatcCattgaatgaatgatttaaaatcaaaaagaaataaaatggc atgaaagagtaaagcttttctaccagtccttagcttttctcttgagcttttctct
MDN1 A>I guide	acaataaaattttgtagtgtccaaaaggagcacctgggtaagcaatgtgacctctgaccacagtta agtctcactttggactcttcttctgttcCatgggtggtcagaggctgtcaccaactcaaacctgtctga gggcatcgctgagtgtctcaggaagtgcgtttacatctcgaagaatgatatagtatgg
UBR4 A>I guide	acaactcatggctcctaggtatgtacaggccctttgatggcttgggttacagacaacctcatagctggtgc accacacacagagataaaacaggaagccCaaaaacccaagccacaccaagaaaaatgagagag gggagggcggggtaacaatgcagcatcccgaggagggaacttaatgcacaaggaggaggagaacaga
HOTAIR A>I guide	ggaagttcaggcattgggaatggtaatcctcatctgtgattttttctgttctctgtactctattattctgt cttttaatacccccttctgtgtcCacatgcacacttatttaagtgttctctatgtctattttctactgcaactt tgtccaagctggggctatatttagagtgc aaagtcctcggttgca
MALAT1 A>I guide	gaaaaatctaaaaaaaggcttagcgccacctcacccctccacccacgccaacacagtttgctcac atgccagtactccagcataaaagctgaaatcCattcaatactattgtccataactgatctgactttgtatgt aaatacagaaaaagctgttcacctgttttctcattttgtccactgggtgaattcaact
XIST A>I guide	tgaataaactgttaaatgacttttgggtcggatctcacacctataattactcatttcctctgtgagcactctat aatgataacatcattttatatccCagggcattgtagtccgagccccacagaaagtaatcaccattcagta agccaatagttcattcctatctgtatagaactgtaggctttgtaaatctacaca
DENND4A A>I guide	gatgagggtactcccactaatatttgcggggcgcccataggagtagtacttgaataatttcacaacctgttt caatcttttccagtcataaaaacccCaaaaataaatttttaaaaaataatacacaaagcactttaaatat gaaaagtaattctcaggcatcatcaactgatccttaaatgtagtagcatggttaa
FBXL4 A>I guide	aggaagactggagactcctatccagcttcaaagcttcttgaataccaagcgaatgaagcaaatggaga aggcaagggaaccaggccaggaagaaaaatCaaaagcagagcaaaaatctcaaaaaccagaggt aaacagtaccactctcttcaggaatcaaggtagatctcaaaaatgcaaggaaataatccaaaaagt
PDE4DIP A>I guide	gagcctgaagtcttcgattgtcttcctgaggactctgttctcattgcagagctgaggtatggactccagg cctgactgtagaagttagaagtggatccCgtcccagaaagtagacatttgggtaacagaaactttcaa aagtttccattcggtatctctcaagatactgccccaaaccagcagaaaacccaaggac
DENND4A antisense guide	gatgagggtactcccactaatatttgcggggcgcccataggagtagtacttgaataatttcacaacctgttt caatcttttccagtcataaaaacccataaaataaatttttaaaaaataatacacaaagcactttaaatat gaaaagtaattctcaggcatcatcaactgatccttaaatgtagtagcatggttaa
FBXL4 antisense guide	aggaagactggagactcctatccagcttcaaagcttcttgaataccaagcgaatgaagcaaatggaga aggcaagggaaccaggccaggaagaaaaatctaaaagcagagcaaaaatctcaaaaaccagaggt aacagtaccactctcttcaggaatcaaggtagatctcaaaaatgcaaggaaataatccaaaaagt
PDE4DIP antisense guide	gagcctgaagtcttcgattgtcttcctgaggactctgttctcattgcagagctgaggtatggactccagg cctgactgtagaagttagaagtggatcctgtcccagaaagtagacatttgggtaacagaaactttcaa agtttccattcggtatctctcaagatactgccccaaaccagcagaaaacccaaggac

AHCY A>I guide	cacgaagctggggtggcccatggcacaaccaggtgaccagccgaccctcggccagcaggatgat gcggcgcccattcttcaaccgataccgggtccaccCacacgcaggcagggcaacagtgaaggcagggc agggccccgctccacagcccaccctctgatctggcaggcctctctgcctccaacagtgcccatgagg a
CTNNA1 A>I guide	catcaaggcagccccactaatgatgctttccagacgctcctccagggaaggcctaaagcgctcctcgct gaagctcaaggggtccacaatgatttgtttcCgcagagaaagtattaccgaatgttagtactcatctctgtt cccttctttattcaagggtgagaaaagtgttattagtaatatcaagatttaacacaaaaat
HSF1 A>I guide (~150 nt guide)	tcgtccccctcacccccgcctacgcacactggccaggctgctgtcaaggtcaggcaggctca tgtcgggcacggtcaccgaggggctgaacagcCgcggaggaggagggaagtcagaacagcacc cgggggcggggcggggagg
IDUA H/ACA box snoRNA	GTGCACATcttcgccGACCTGCTTTCTTCTATGTGAGTAGTGTgactgc ATGTGCTATACAAATAATTGAAGGctttggtcGCAGTATAACTATAAA TAGTAATGCTGCGagttgcCCTTCAGACAAAA
CFTR H/ACA box snoRNA	GTGCACATCTTTCCTTGACCTGCTTTCTTCTATGTGAGTAGTGTGTC CTGTTGCATGTGCTATACAAATAATTGAAGGCTTTCCTTGCAGT ATAACTATAAATAGTAATGCTGCGCTGTTCCCTTCAGACAAAA
ACTB-U1226 H/ACA box snoRNA	GTGCACATAACGCAACGACCTGCTTTCTTCTATGTGAGTAGTGT CAGTCATAATGTGCTATACAAATAATTGAAGGACGCAACGCAGT ATAACTATAAATAGTAATGCTGCAGTCAACCTTCAGACAAAA
EEF2-U2881 H/ACA box snoRNA	GTGCACATTAAGTCCCGACCTGCTTTCTTCTATGTGAGTAGTGT CCTAAGAGATGTGCTATACAAATAATTGAAGGAAGTCCCGCAG TATAACTATAAATAGTAATGCTGCGCTAAGACCTTCAGACAAAA
RPS6-U514 H/ACA box snoRNA	GTGCACATGGCTTTCTGACCTGCTTTCTTCTATGTGAGTAGTGT TCAACATAATGTGCTATACAAATAATTGAAGGCCTTTCTGCAGT ATAACTATAAATAGTAATGCTGCCAACATCCTTCAGACAAAA

Supplementary Table 3: Oligonucleotide sequences for PCR, qPCR, Sanger sequencing, NGS, pseudouridylation standardization, and RCA FISH.

Name	Sequence
RAB7A A>I FOR	CCTCCCTCCTTGAAGGCTACC
RAB7A A>I REV	AAGCTCCGCTAACCTAAGAATACC
GAPDH A>I FOR	GATGCTGGCGCTGAGTACGT
GAPDH A>I REV	CACCACTGACACGTTGGCAGT
TARDBP A>I FOR	CCATCGGAAGACGATGGGACG
TARDBP A>I REV	CCTGAATGGCTTGGGGATGAAG
FANCC A>I FOR	CCTGCACAACAGCTGATCAGGC
FANCC A>I REV	TTCTTTAATGGTTCATGACCAAATTCTTGG
ALDOA A>I FOR	GACAGCTGACGACCGCGTGAACC
ALDOA A>I REV	CCCCAATCTTCAGCACACAACGCCA
DAXX A>I FOR	CTGACCGGCCGTGTCATAGAGCAGC
DAXX A>I REV	GTGAGGTGGCAGCCAAAGTTGTAGATGA
SMAD4 A>I FOR	GCGTGCACCTGGAGATGCTG
SMAD4 A>I REV	ACAGGTGAAGAATTAATAAGAATGTGTTTCTCCT
GAA A>I FOR	ttctacccttcacatgcggaac
GAA A>I REV	ggtgaagtggagacagggga
BLM A>I FOR	cccaacaccacaaatcagcaa
BLM A>I REV	aggagggtggaggcaaatca
CHD7 A>I FOR	gtgaagtgaagcacaggcaag
CHD7 A>I REV	gagcagtagaggggtgtga
FANCD2 A>I FOR	tcttccgtgtgatgatggct
FANCD2 A>I REV	tctcatctgctgtgctctcc
COL4A5 A>I FOR	agtttcgttcagtccttca
COL4A5 A>I REV	ccgggatacagcaggattag
MDN1 A>I FOR	cagaggaagaccaagacccc
MDN1 A>I REV	AAAACCTCAGCCCAGGCAAA
UBR4 A>I FOR	GAGTACATCCGCCACAACGA
UBR4 A>I REV	agtatgaagctggccggga
DMD A>I FOR	GCGAGTAGTTCCACACAGGT
DMD A>I REV	TCAGGAACACCCCAAAACCA
NEAT1 qPCR FOR	GCTGGACCTTTCATGTAACGGG
NEAT1 qPCR REV	TGAACTCTGCCGGTACAGGGAA
GAPDH qPCR FOR	GTCTCCTCTGACTTCAACAGCG
GAPDH qPCR REV	ACCACCCTGTTGCTGTAGCCAA
HOTAIR A>I FOR	gcaccgcttttctaactggc
HOTAIR A>I REV	acacaaagtgcatacctacca
MALAT1 A>I FOR	tgtgtgggtttctctctccc
MALAT1 A>I REV	ccgcttgagatttgggcttt
XIST A>I FOR	acgcaagggtgtgtgtct
XIST A>I REV	ctcaatcaggccaggaagca

RAB7A guide qPCR FOR	AGTTGTCCCCCTGGAGAGATG
RAB7A guide qPCR REV	CCTGTAATGCAGGCGACAAA
GAPDH guide qPCR FOR	CGCATGGACTGTGGTCTACT
GAPDH guide qPCR REV	CCATGTTTCGTCATGGGTGTC
TARDBP guide qPCR FOR	GAGTTTCAGGTCCTGTTCCCA
TARDBP guide qPCR REV	AAGTGACAAGAGCAGTCCAGAA
DENND4A A>I FOR	GCCAAAATGCCCAGTTGCTT
DENND4A A>I REV	TACTCATTGCTTTTGTAGTGGGC
FBXL4 A>I FOR	TTCCTTCAGTTCCTTTTCTGCTT
FBXL4 A>I REV	AGTGTCTAGCATGGAACTACTCA
PDE4DIP A>I FOR	GTAAGGGTCCTCCCACCTCT
PDE4DIP A>I REV	GCTGTGAGTTTGGGTGTTG
DENND4A RT-PCR FOR	TTACCCCAACTGGATTGTCAG
DENND4A RT-PCR REV	TACAGTGTTTCGTCTTTGCCAC
FBXL4 RT-PCR FOR	TTGTGCCGCATCTAGAGAGTC
FBXL4 RT-PCR REV	CCAGGTACCCTGAACTTTGC
PDE4DIP RT-PCR FOR	GAACACCGGCTGACCTCTAC
PDE4DIP RT-PCR REV	CTTCGATTGTCTTCCCTGAGGAC
AHCY A>I FOR	GAGACGGGCTTTCAGTGTGTTG
AHCY A>I REV	AACGGGGTACTTGTCTGGATGG
CTNNA1 A>I FOR	GTAGGCCATCTTCTGTGGGACA
CTNNA1 A>I REV	TGTACTCCGAAAGCAGGTCCTG
HSF1 A>I FOR	CTGTTCTGACTTCCCTCCCTCC
HSF1 A>I REV	TGGGACTTGGCTCACCTGAATC
AHCY RT-PCR FOR	gtcaagtggctcaacgagaacg
AHCY RT-PCR REV	tccaagaccactgagctcatgg
CTNNA1 RT-PCR FOR	caccctgatgtcgcagcctata
CTNNA1 RT-PCR REV	ctgaaacgtggtccatgacagc
HSF1 RT-PCR FOR	tgcctggacaagaatgagctca
HSF1 RT-PCR REV	ctctaggagacagtggggctcct
CFTR snoRNA guide FOR	ACATCTTTCCTTGACCTGCTTTC
CFTR snoRNA guide REV and cDNA primer	AGGGAACAGGCAGCATTACTA
ACTB snoRNA guide FOR	CGCAACGACCTGCTTTCTTC
ACTB snoRNA guide REV and cDNA primer	ATACTGCGTTGCGTCCTTCA
EEF2 snoRNA guide FOR	TAAGTCCCGACCTGCTTTCTT
EEF2 snoRNA guide REV and cDNA primer	CTGAAGGTCTTAGGCAGCATT
RPS6 snoRNA guide FOR	CATGGCTTTCTGACCTGCTTT
RPS6 snoRNA guide REV and cDNA primer	TCTGAAGGATGTTGGCAGC
PuroR qPCR FOR	atgaccgagtacaagccac
PuroR qPCR REV	acaccttgccgatgtcgag

ANXA5 qPCR FOR	CAAGCCTTTTCATAGCCTTCC
ANXA5 qPCR REV	AGTCTGGTCCTGCTTCACCT
Targeted amplicon CMC sequencing ACTB-U1226-TAG-F	ACACTCTTTCCCTACACGACGCTCTTCCGATCTN NNNtccatcggtccaccgcaaatgct
Targeted amplicon CMC sequencing ACTB-U1226-TAG-R	TGGAGTTCAGACGTGTGCTCTTCCGATCTNNNN NNgccaatctcatcttgttttctgcgca
Targeted amplicon CMC sequencing EEF2-U2881-TAG-F	ACACTCTTTCCCTACACGACGCTCTTCCGATCTN NNNcagagtccggaggcagcag
Targeted amplicon CMC sequencing EEF2-U2881-TAG-R	TGGAGTTCAGACGTGTGCTCTTCCGATCTNNNN NNaaagtgttggtgtcccatccc
Targeted amplicon CMC sequencing RPS6-U514-TAA-F	ACACTCTTTCCCTACACGACGCTCTTCCGATCTN NNNcgcaaaccttttcaatctctcTAAa
Targeted amplicon CMC sequencing RPS6-U514-TAA-R	TGGAGTTCAGACGTGTGCTCTTCCGATCTNNNN NNaggacacgtggagtaacaag
Targeted amplicon CMC sequencing Universal NGS Primer 1	AATGATACGGCGACCACCGAGATCTACACTCTTT CCCTACACGACGCTCTTCCGATCT
Targeted amplicon CMC sequencing Index NGS Primer 2-1	CAAGCAGAAGACGGCATAACGAGATCATACCACG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-2	CAAGCAGAAGACGGCATAACGAGATGAAGTTGGG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-3	CAAGCAGAAGACGGCATAACGAGATATGACGTCG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-4	CAAGCAGAAGACGGCATAACGAGATTTGGACGTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-5	CAAGCAGAAGACGGCATAACGAGATAGTGGATCG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-6	CAAGCAGAAGACGGCATAACGAGATGATAGGCTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-7	CAAGCAGAAGACGGCATAACGAGATTGGTAGCTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-8	CAAGCAGAAGACGGCATAACGAGATCGCAATCTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-9	CAAGCAGAAGACGGCATAACGAGATGATGTGTGG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC

Targeted amplicon CMC sequencing Index NGS Primer 2-10	CAAGCAGAAGACGGCATAACGAGATGATTGCTCG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-11	CAAGCAGAAGACGGCATAACGAGATCGCTCTATG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-12	CAAGCAGAAGACGGCATAACGAGATTATCGGTGCG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-13	CAAGCAGAAGACGGCATAACGAGATAACGTCTGG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-14	CAAGCAGAAGACGGCATAACGAGATACGTTTCAGG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-15	CAAGCAGAAGACGGCATAACGAGATCAGTCCAAG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-16	CAAGCAGAAGACGGCATAACGAGATTTGCAGACG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-17	CAAGCAGAAGACGGCATAACGAGATCAATGTGGG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-18	CAAGCAGAAGACGGCATAACGAGATACTCCATCG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-19	CAAGCAGAAGACGGCATAACGAGATGTTGACCTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-20	CAAGCAGAAGACGGCATAACGAGATCGTGTGTAG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-21	CAAGCAGAAGACGGCATAACGAGATACGACTTGG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-22	CAAGCAGAAGACGGCATAACGAGATCACTAGCTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-23	CAAGCAGAAGACGGCATAACGAGATACTAGGAGG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-24	CAAGCAGAAGACGGCATAACGAGATGTAGGAGTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC

Targeted amplicon CMC sequencing Index NGS Primer 2-25	CAAGCAGAAGACGGCATAACGAGATCCTGATTGG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-26	CAAGCAGAAGACGGCATAACGAGATATGCACGAG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-27	CAAGCAGAAGACGGCATAACGAGATCGACGTTAG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-28	CAAGCAGAAGACGGCATAACGAGATTACGCCTTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-29	CAAGCAGAAGACGGCATAACGAGATCCGTAAGAG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-30	CAAGCAGAAGACGGCATAACGAGATATCACACGG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-31	CAAGCAGAAGACGGCATAACGAGATCACCTGTTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-32	CAAGCAGAAGACGGCATAACGAGATCTTCGACTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-33	CAAGCAGAAGACGGCATAACGAGATTGCTTCCAG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-34	CAAGCAGAAGACGGCATAACGAGATAGAACGAGG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-35	CAAGCAGAAGACGGCATAACGAGATGTTCTCGTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Targeted amplicon CMC sequencing Index NGS Primer 2-36	CAAGCAGAAGACGGCATAACGAGATTCAGGCTTG TGACTGGAGTTCAGACGTGTGCTCTTCCGATC
Pseudouridylation standardization EEF2 standard (Ψ site underlined)	CACAAAGAGCCGGACACAG <u>T</u> AGAGAcGcAGGA cAcGAAGGccacacgggcaccaaggac
Pseudouridylation standardization RPS6 standard (Ψ site underlined)	CACAAAGAGCCGGACACCG <u>T</u> AAAGAcGcAGGA cAcGAAGGccacacgggcaccaaggac
Pseudouridylation standardization ACTB standard (Ψ site underlined)	CACAAAGAGCCGGACACCT <u>T</u> AGAGAcGcAGGA cAcGAAGGccacacgggcaccaaggac

Pseudouridylation standardization CFTR standard (Ψ site underlined)	CACAAAGAGCCGGACCACCGTGAAGAcGcAGGA cAcGAAGGccacacgggcaccaaggac
Pseudouridylation standardization FOR primer for generating IVT template (PCR1)	TAATACGACTCACTATAGGGAGGGAACAGAAAC AGAGACTAGGAGGGGCGCGTAGGATAAATAATT CACACAAAGAGCCGGACCACA
Pseudouridylation standardization FOR primer for generating IVT template (PCR2)	TAATACGACTCACTATAGGGAGGGAACAGA
Pseudouridylation standardization REV primer for generating IVT template (PCR1, PCR2)	TGTGTGGTCCTTGGTGCCCGTGTG
BID-Seq FOR primer for Ψ standard	ACACTCTTTCCCTACACGACGCTCTTCCGATCTAT AAACGGGCGGAGGGAACAGAAACAGAGA
BID-Seq REV primer for Ψ standard	GACTGGAGTTCAGACGTGTGCTCTTCCGATCTTA TTAGTTATGTGTGGTCCTTGGTGCCCGTGTG
BID-Seq FOR primer for CFTR luciferase reporter	ACACTCTTTCCCTACACGACGCTCTTCCGATCTA AGTGAGACACCGGGTACGA
BID-Seq REV primer for CFTR luciferase reporter	GACTGGAGTTCAGACGTGTGCTCTTCCGATCTGC ATCGGCAGAACCTATCACT
RCA FISH GAPDH A>I RT primer	A+GA+AG+TA+TG+AC+TT+GTGCCTCAAGAT
RCA FISH GAPDH A>I padlock probe	/5phos/AGGAGGCAAACGTGATGATCTTGAGctcctgt tcgacacctaccacctcatccactcttcaGTGCCAAGCACAAAC GTGGTGC
RCA FISH GAPDH A>I hybridization probe	/5TYE665/cacctcatccactcttcaAGGAGGCAAACGTGA
RCA FISH EEF2 U>Ψ RT primer	A+CT+AT+TT+AT+AG+TT+ATACTGCGGGACTTC
RCA FISH EEF2 U>Ψ padlock probe	/5phos/AATAATTGAAGGAAGTCCCGCctcctgttcgacac ctaccacctcatccactcttcaGTCCTAAGAGATGTGCTATA CA
RCA FISH EEF2 U>Ψ hybridization probe	/5TYE665/cacctcatccactcttcaGTCCTAAGAGATGTG