

Supplemental information

**Splice site m⁶A methylation prevents binding
of U2AF35 to inhibit RNA splicing**

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Table S1. List of all deep-sequencing libraries created in this study. Related to STAR Methods and Figures 1-4.

Data is available from Gene Expression Omnibus (GEO: GSE146873).

sample	experiment	description	reads	sequencing type
RR723	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	INPUT-Total RNA-1	78748292	single-end
RR724	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	INPUT-Total RNA-2	109227705	single-end
RR725	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	INPUT-Total RNA-3	98509826	single-end
RR726	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	INPUT-PolyA+ RNA-1	111319107	single-end
RR727	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	INPUT-PolyA+ RNA-2	111369816	single-end
RR728	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	INPUT-PolyA+ RNA-3	104925272	single-end
RR729	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	m6A IP-Total RNA-1	97238279	single-end
RR730	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	m6A IP-Total RNA-2	116365247	single-end
RR731	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	m6A IP-Total RNA-3	103682866	single-end
RR732	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	m6A IP-PolyA+ RNA-1	99423158	single-end
RR733	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	m6A IP-PolyA+ RNA-2	90657183	single-end
RR734	m6A profiling on RNA mix from mouse, worm and BmN4-SID1	m6A IP-PolyA+ RNA-3	105299920	single-end
RR618	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_totalRNA_WT-1	107952378	pair-end
RR619	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_totalRNA_WT-2	104276535	pair-end
RR620	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_totalRNA_WT-3	99043875	pair-end
RR621	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_totalRNA_WT-1	88177640	pair-end
RR622	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_totalRNA_WT-2	92603501	pair-end
RR623	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_totalRNA_WT-3	80301018	pair-end
RR624	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_polyA_2_WT-1	51187060	pair-end
RR625	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_polyA_2_WT-2	52291052	pair-end
RR626	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_polyA_2_WT-3	33117898	pair-end
RR627	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_polyA_2_KO-1	39691391	pair-end
RR628	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_polyA_2_KO-2	35240657	pair-end
RR629	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_polyA_2_KO-3	39991008	pair-end
RR630	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_polyA_2_WT-1	12022240	pair-end
RR631	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_polyA_2_WT-2	36551759	pair-end
RR632	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_polyA_2_WT-3	25683360	pair-end
RR633	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_polyA_2_KO-1	54841480	pair-end
RR634	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_polyA_2_KO-2	44893258	pair-end
RR635	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_polyA_2_KO-3	43057847	pair-end
RR600	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_polyA_1_WT-1	22391643	pair-end
RR601	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_polyA_1_WT-2	23096755	pair-end
RR602	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	input_polyA_1_WT-3	19042425	pair-end
RR603	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_polyA_1_WT-1	18362889	pair-end
RR604	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_polyA_1_WT-2	18425770	pair-end
RR605	m6A profiling of the worm Mett-10 KO and WT on NA22 plates	m6A_IP_polyA_1_WT-3	18620691	pair-end
RR892	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	input_polyA_WT-1	44602975	single-end
RR893	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	input_polyA_WT-2	57428958	single-end
RR894	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	input_polyA_WT-3	63145019	single-end
RR895	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	input_polyA_KO-1	56220445	single-end
RR896	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	input_polyA_KO-2	46443279	single-end
RR897	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	input_polyA_KO-3	42188533	single-end
RR898	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	m6A_IP_polyA_WT-1	11536036	single-end
RR899	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	m6A_IP_polyA_WT-2	11339449	single-end
RR900	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	m6A_IP_polyA_WT-3	11065165	single-end
RR901	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	m6A_IP_polyA_KO-1	14331371	single-end
RR902	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	m6A_IP_polyA_KO-2	13228517	single-end
RR903	m6A profiling of the worm Mett-10 KO and WT on OP50 plates	m6A_IP_polyA_KO-3	12318564	single-end

Table S3. DNA primers and RNA oligonucleotides used in this study. Related to STAR Methods and Figures 1-6.

Name	Sequence	Name in the study/Comments
Worm mutants		
oKDM42	GGTGATTGTTGTCAGTGC	Reverse primer for detecting and sequencing of intron 2 deletion in <i>sams-3</i> . If edit is present, it creates 473 bp with oKDM43
oKDM43	GTCACAGGTAAGTAACCAACG	Forward primer for detecting and sequencing of intron 2 deletion in <i>sams-3</i> . If edit is present, it creates 473 bp with oKDM42
oKDM41	GACTCCAGCAGCGATTCAG	Forward primer for detecting and sequencing of intron 2 deletion in <i>sams-4</i> . If edit is present, it creates 433 bp with oKDM44
oKDM44	ACGTTATCTCCGAAGCGTAG	Reverse primer for detecting and sequencing of intron 2 deletion in <i>sams-4</i> . If edit is present, it creates 433 bp with oKDM41
oKDM12	GCAACGTGTCGCTAAAGAGG	Forward primer for detecting and sequencing of intron 2 deletion in <i>sams-5</i> . If edit is present, it creates 526 bp with oKDM39
oKDM39	TCCGTCTGCTTACCTTCGAG	Reverse primer for detecting and sequencing of intron 2 deletion in <i>sams-5</i> . If edit is present, it creates 526 bp with oKDM12
oKDM17	CCGTCATTCCCAAACCTCCATC	Forward primer to detect and sequence C-terminal HA tag in <i>sams-3</i> . If edit is present, it creates 554 bp band with oKDM17
oKDM18	ATGAACCAACCGCTGAGAAC	Reverse primer to detect and sequence C-terminal HA tag in <i>sams-3</i> . If edit is present, it creates 554 bp band oKDM17 to be sequenced
oFSa0131	TGTGGTGTGACAGGTAG	Forward primer to detect and sequence deletion of <i>mett-10</i> . If edit is present, it creates around 260 bp band with oFSa0195
oFSa0195	ATGTCCATGCCAACATCCAGC	Reverse primer to detect and sequence deletion of <i>mett-10</i> . If edit is present, it creates around 260 bp band with oFSa0131
oKDM5	AAACTTGATTGACAATGTCGAG	sgRNAs for C-terminal HA tagging of <i>sams-3</i>
oKDM6	TCTTGTGACAGCATGTACAATCAA	
oKDM7	CGATCTAACCGGAATGGAATTGCTCATGGAGGAGGATCAATGGAGGATAACCCATATGATGTTCCAGATTACGGCAGTACCCATATGATGTTGACATGTCTGATTCTGGTT	Repair template for the C-terminal tagging of <i>sams-3</i>
oKDM19	AAACTTCTGTAATCAAAGTAACC	
oKDM20	TCTTGGTTACTTTGATTACAGAAA	sgRNA for removing intron 2 of <i>sams-3</i> , <i>sams-4</i> and <i>sams-5</i>
oKDM21	AAACTTGTAAACAGGTAAGAAC	sgRNA for removing intron 2 of <i>sams-3</i> , <i>sams-4</i> and <i>sams-5</i>
oKDM22	TCTTGGTTCTTACCTGTTACAA	
oKDM23	AAACCTGAATCACTCACCAACAGAC	sgRNA for removing intron 2 of <i>sams-4</i>
oKDM24	TCTTGTGCTTGTGGTAGTTGAGG	
oKDM25	AAACCAAGCAACCTTAGCGTGAGC	
oKDM26	TCTTGTCACTGAAAGTTGCTG	
oKDM27	TCTTGTCAACGCAAAGGTGCTG	
oKDM28	AAACCATGCAACCTTGGCTGAG	
oKDM29	TCTTGAAGGTTGCTGTTGGTAAG	
oKDM30	AAACCTTACACATGCAACCTTC	
oKDM31	AAACACAGTTCTGAAACCTTAC	
oKDM32	TCTTGTAAAGTTTACAGAACTGT	
oKDM33	AAACCAAGCAACCTTGGCTGAGC	
oKDM34	TCTTGTCAACGCCAAGGTGCTG	
oKDM35	CGACAGCTTTCGAGGTGATCTCACCGCACACATGATCATACCGTTAGTCACTGTTCA	Repair template for removing intron 2 in <i>sams-4</i>
oKDM36	CAAGAACCTTACGGTCTGGCTGAGCATG	Repair template for removing intron 2 in <i>sams-3</i>
oKDM37	CAGCTTTGAGGTGATCTCACCGCACACATGATCATACCGTTGGTCACTGTTCACAA	Repair template for rem+A22:C38oving intron 2 in <i>sams-5</i>
sams-3 artificial constructs		
DH298	ACGAAGTGTTCACCTT	Forward primer for <i>sams-3</i> FL amplification
DH299	CTCAATAATGATTTTATAAG	Reverse primer for <i>sams-3</i> FL amplification
DH300	CCCTAGAACACGGCTGTCACATCAAAGAACAGCGTCCGAGAT	Primer to introduce specific sequences into exon2
DH301	TGAACGACCGTGTCTAGGGACGAAAGGTTGATGT	Primer to introduce specific sequences into exon2
DH302	TACGGCACTAAAGTAGGTGTCGGACAGAACCTGGTAGTC	Primer to introduce specific sequences into exon3
DH303	ACAGCCTACTTGTAGTGGCTAATACGATGATTCTCGAAAG	Primer to introduce specific sequences into exon3
DH304	CTTATAAAAATCTTATTGAGCTCTCAGGACGCTGGCATGC	Forward primer for pUC19 fragment amplification
DH305	AAGGTGAAACACTCGTTGTTCTGGAAATTGAGA	Reverse primer for pUC19 fragment amplification
MM320	GATTACAGACTGGTACTAAACGGGTATG	Forward primer for METT-10 consensus motif mutation
MM321	TTAGTCACCAAGTGTAACTAAAGAACCG	Reverse primer for METT-10 consensus motif mutation
HeLa splicing constructs		
MM415	ATGGCTAGCCACCAAGTGTTCACCTTACAAC	Forward primer for <i>sams-3</i> FL clonning into phRL-TK vector
MM416	CATCGGGCGCTCTAGAACTCAATAATGATTTTTATAAGGTC	Forward primer for <i>sams-3</i> FL clonning into phRL-TK vector
Splicing		
MM378	GGTGAACGTGGATGAGTTGGTGGTGGAGGCCCTGGGAGGTTGGTATCAAGGTTACAAGACA GGTTAACGGAGACCAATAGAAACTGGGCATGTGGAGACAGAGAACACTTGGCCTCGAGGT TTCTGATAGGCACGTACTCTCTGGCTATTGGCTATTTCACCCCTAGGCTGCTGGTG GTCTACCCCTGGGCCAGAGGGTCTTGAGGATATCCGTACACCATCAGGGTACGAGCTAGC CCATGGCGTACACCATCAGGGTACGACTAGTAGATCTCGTACACCATCAGGGTACGAGAATTC	Human B-globin with T7 promoter
MM379	TAATACGACTCACTATAGGGTAACCGTGGATGAAGTTG	Forward primer to amplify 3' fragment of B-globin
MM460	/52MOeRg//i2MOeRg/GAAAATAGACCAATAGGCAG	Reverse primer to amplify 3' fragment of B-globin
MM_RNA_59	/5phos/ACCTTAGGCTGCTGGTACCCCTGGACCCAGAGGTCTTGAG	Unmethylated oligo for B-globin pre-mRNA ligation

MM_RNA_60	/5phos/ACCCTTm6AGGCTGCTGGTGGTCAACCTTGAGCCAGAGGTCTTGAG	Splice site methylated oligo for B-globin pre-mRNA ligation
MM_RNA_61	/5phos/ACCCTTAGGCTGCTGGTGGTCTm6ACCCTTGAGCCAGAGGTCTTGAG	Exon methylated oligo for B-globin pre-mRNA ligation
MM453	CTGGGTCAGGGTAGACCACCAGCACGCTAACGGGGAAATAGACCAATAGGCAGA	Splint oligo for B-globin ligation
MINX_FL	CACTCTGGATCGGAAACCGCTCGGCCGAACGGTAAGAGCCTAGCATGTAGAACTGGTTACCTGCAGCCAAAGCTGCTGCACGCTAGGGCAGTAGTCAGGGTTCTGATGATGTCATACCTTATCCCTGCTCCAGACAGCTGGTGGAGGACGACAACACTTCGGCGTCTGGGG	MINX pre-mRNA sequence
MM_RNA_85	/5phos/cacagctcgccgttggaggacaacttcgcggctttccagtgg	Unmethylated oligo for MINX pre-mRNA ligation
MM_RNA_86	/5phos/cacm6Agctcgccgttggaggacaacttcgcggctttccagtgg	Splice site methylated oligo for MINX pre-mRNA ligation
MM_RNA_87	/5phos/cacagctcgccgttggagg/iN6Me-rA/caaaacttcgcggctttccagtgg	Exon methylated oligo for MINX pre-mRNA ligation
MM578	gagtttgtctcaaccgcgagctgtgtgtgttaaagg	Splint oligo for MINX ligation
MM617	TAATACGACTCACTATAGGCCACTTGGATCGGAAACCC	Forward primer to amplify 3' fragment of MINX
MM582	/52MOErG//i2MOErA/aaaaaaaaaaaaaaaaggacaggaaTaagtatgc	Reverse primer to amplify 3' fragment of MINX
MM583	/52MOErG//i2MOErT/gtgtaaaaaggacaggaaTaagtatgc	Reverse primer to amplify 3' fragment of the weak polypyrimidine tract MINX
MM585	gagtttgtctcaaccgcgagctgtgtgtgttaaagg	Splint oligo for the weak polypyrimidine tract MINX ligation
sams-3 RT-PCR		
MM395	CTCGGACGCTGTTCTGATG	PCR for intron retention, endogenous sams-3
MM396	AACCTGGTAGTCGACGACAG	PCR for intron retention, endogenous sams-3
MM363	TGTGAACGACCCGTGTTCTAGG	PCR for intron retention in mini-gene constructs
MM364	CGTATTACGCACCTCAAAGTAGGC	PCR for intron retention in mini-gene constructs
Mettl16 knock-in mice		
tracerRNA		tracerRNA, 67 bases, cat no 1072533, IDT
CD.Cas9.PGDV2419.AB , RNA	CTCAAAGGAACCACTCAGCG GUUUUAGAGCUAUGCU	gRNA for cat-dead and RNA-binding mutant Mettl16 mouse
MM233	TCATCTCCCTAGTGGTAAAGTGGCAGAAGACACTCTGATGGATGGCTTAAGAAGAGTCTGAGATAGTCTATGACTCTGTGAAACCCCTCCGGCTTCGCCAACCAATTGGAAAGCCAAGTACAGTACCTTAAAGTTACCAATGCTTAGCTAGCTTACATCATTGAGAGATA TCAACAAAAGGCAG	ssDNA template for homology recombination for F187G modification
MM232	TCATCTCCCTAGTGGTAAAGTGGCAGAAGACACTCTGATGGATGGCTTAAGAAGAGTCTGAGATAGTCTATGATTCTGTATGTCGAAACCCCTCCGGCTTCGCCAACCAATTGGAAAGCAAGGTACAGTACCTTAAAGTTACCAATGCTTAGCTAGCTTACATCATTGAGAGATA AGAACAAAAGGCAG	ssDNA template for homology recombination for 185AA-PP186 modification
MM342	TCTCCATTGCTGCAGGTCTG	Forward primer for F187G genotyping
MM343	CTTCAATTGGTGGCGAACG	Reverse primer for F187G genotyping
MM348	CGTCTCCATTGCTGCAGGTCTG	Forward primer for PP185-186AA genotyping
MM349	ATTTGGTGGCAAAAATGGCGC	Reverse primer for PP185-186AA genotyping
MM340	AACTTGATGCGCTCCATTG	Forward primer to amplify the whole catalytic site region for sequencing
MM341	TGGTAACTTTAAAGGTACTGTACC	Reverse primer to amplify the whole catalytic site region for sequencing
Mettl16 conditional KO		
MM101	aaagatttgccttggcgtttcc	Forward primer to detect loxP insertion in the Mettl16 locus
MM102	cccaggaaatggcactatacaag	Reverese primer to detect loxP insertion in the Mettl16 locus
oIMR7643	CACGTGCAGCCGTTAACGGCGT	Forward primer for Ddx4-cre genotyping
oIMR7644	TTCCCATCTAACACACCCCTGAA	Reverse primer for Ddx4-cre genotyping
MM314	AAATTCAAATGGGATTGGG	Forward primer for Mettl16 deletion genotyping
MM315	CCGAATCCTCATCTATTACATTG	Reverse primer for Mettl16 deletion genotyping
In vitro methylation		
Name in the study/Comments		
MM RNA 5	CCCCCCCUGUAGGUACAGAGAACGGGGGG	Mouse 3'UTR ; 29nt
MM RNA 6	UGUUGGGCUAGGUACAGAGAACGGGUUCA	Mouse 3'UTR mut ; 29nt
MM RNA 31	ACAACGGUUACUUUAACAGAACCGGUGACUAAAACGG	RNA-0 ; 40nt
MM RNA 32	GGUUACUUUGAUUACAGAACCGGUGACUAA	RNA-1 ; 30nt
MM RNA 34	GGUUACUUUGAUUACAGAACCGGUGACUAA	RNA-2 ; 30nt
MM RNA 33	GGUUACUUUGGUUACAGAACCGGUGACUAA	RNA-3 ; 30nt
MM RNA 35	GGUUACUUUGAUUACAGACUUGGUGACUAA	RNA-4 ; 30nt
MM RNA 36	GGUUACUUUGAUUACAGUAACGGUGUGACUAA	RNA-5 ; 30nt
MM RNA 37	GGUUACUUUGAUUACAGAGACUUGGUGACUAA	RNA-6 ; 30nt
MM RNA 43	GGUUACUUUGAUUACAGAGACCGGUGACUAA	RNA-7 ; 30nt
MM RNA 38	CCCCCCCUACAGAACCCCCCCCC	RNA-8 ; 25nt
MM RNA 39	CCCCCCCUACAGAACCGGGGGGG	RNA-9 ; 25nt
MM RNA 100	GGUUACUUUGAUUACAGAACACUAGGAA	RNA-10 ; 30nt
MM RNA 101	UCAGUGGGUGAUUACAGAACACUAGUGAA	RNA-11 ; 30nt
MM RNA 120	GGUUACUUUGAUUACAGAACACGGUGUGUAA	RNA-12 ; 30nt
MM RNA 121	GCAUACUUUGAUUACAGAACACGGUGUGUAA	RNA-13 ; 30nt
MM RNA 122	GGUUACUUUGAUUACAGAACACGGACAGUAA	RNA-14 ; 30nt
MM RNA 123	GCAUUGGUUGAUUACAGAACACGUGUAA	RNA-15 ; 30nt
MM RNA 124	CCCCCCUUUGAUUACAGAACACGGGGGGAA	RNA-16 ; 30nt
MM RNA 44	AAACUUUGUUUUUUUAGAACUGUUGGUAAA	Mouse ; 30nt
MM RNA 45	AAACUUUGUUUUUUUAGAACUGUUGGUAAA	Mouse mut ; 30nt
MM RNA 46	UUUUUUUAACCUUACAGAACACCAUACUAA	Silkworm ; 30nt
MM RNA 47	UUUUUUUAACCUUACUGAACACCUUACUAA	Silkworm mut ; 30nt
MM RNA 48	CAUAAUAAAAAAACAGAACACGGUCCCAA	Fly ; 30nt
MM RNA 49	CAUAAUAAAAAAACAGAACACGGUCCCAA	Fly mut ; 30nt
MM RNA 67	UUUUUUUUUUUUUUUACAGAACAGAAAAGAGAAGAUGAA	RNA-67/Muc11 ; 40nt
MM RNA 68	AGAGUGGGUUUCUGUUUUUACAGAAAUGGAAACUGAAAUGJ	RNA-68/Spagl ; 40nt
MM RNA 69	AAUCUCUAAUGUUUUUACAGAAAUGCAUCAGAACGUG	RNA-69/Pla2r1 ; 40nt
MM RNA 70	UUUCUGGAACUUUUUUUACAGAAAAGUUUAGAGACUGUC	RNA-70/Gyg ; 40nt
MM RNA 71	GCAUGGGGGGGGUUACUUAACAGAACACGGCCGGAGAACAG	RNA-71/Des ; 40nt
MM RNA 72	AUGCCAAAACUUUCGUUUUACAGAACACCCGGUUUCUAA	RNA-72/Helt ; 40nt
MM RNA 73	ACUCUUGCUUCUGUUCACAGAACACGGGAAGGGUCCCUA	RNA-73/Pecam ; 40nt
MM RNA 74	AAUUCUUUUUUUUUCAUACAGAGAACAGAACAGCUAGAC	RNA-74/Gm4853 ; 40nt
MM RNA 75	CUUUUGGGCUUUUUUACAGAAAAGUUUAGGACACAUAG	RNA-75/Gm15810 ; 40nt
MM RNA 76	GAUGCUUUUUUGUUUACAGAGAAAUGACAGGGUGGUC	RNA-76/Tjp2 ; 40nt
MM RNA 80	UAGGAAUUUUUUUUUUCUUCUACAGAAAACGACUUGGGAGCCCCAGUA	Sorbs1 ; 45nt
MM RNA 81	UAAUUGGUUUUCAUUUACAGAGAACGACGUCUGAGCAGCC	Lrpprc ; 40nt

RP RNA 19	GGGCUUUACGGGCUUAAAACAAAACAAAACAAAACAAA	40 nt RNA marker
RP RNA 1	TGACATGAACACAGGTGCTCAGATAAGCTTT	30 nt RNA marker
RP RNA 3	UGACAUGAACACAGGUGUCAGAUAGCU	28 nt RNA marker
RP RNA 18	AGCACCGUAAGACGCC	16 nt RNA marker
ITC		
KM RNA 10	CUAGG	ITC experiment; 5 nt
KM RNA 11	CUm6AGG	ITC experiment; 5 nt
SCARLET		
MM437	GGAGAGACAACTTAAAGAGACTTAAAGATTAAATTAAATTTATCAAAAAGACTTATTGACT TAAAGTCTAACCTATAGGATACTTACAGCCATCGCCGGCTGTGAGTTAATAG	Universal DNA oligo for splint ligation
MM438	TGATCATACCGTTTAGTCACCGTTCTATTAACTCACAGGACCGGCATGGCTG	sams-3 splint oligo
MM698	mCmGmUmUmCmU GTAA mUmCmAmAmGmU	sams-3 chimera
MM614	CCTTGCGCAGGGGCATGCTAATCTCTCTATTAACTCACAGGACCGGCATGGCTG	U6snRNA splint oligo
MM702	mCmUmUmCmUmCmU GTAT mUmGmUmUmCmC	U6 snRNA chimera
MM616	CCACTTTTATATCTATAGCGCGCGTTCTATTAACTCACAGGACCGGCATGGCTG	Positive control splint oligo
MM615	mGmCmGmUmUmGmU TCAG mAmGmUmCmUmAmA	Positive control chimera
MM RNA 98	UUAGACUCUGAACACGCGCGCUAUAGAUUAAGGUU	Positive control oligo - unmethylated
MM RNA 99	UUAGACUCUGAm6ACAACCGCGCUAUAGAUUAAGGUU	Positive control oligo - methylated
MM RNA 96	GTTACACGGTTACTTTGATTACAGAAACCGGTGACTAAACCGGGT	sams-3 control oligo - unmethylated
MM RNA 97	GTTACACGGTTACTTTGATTACm6AGAAACGGTGACTAAACCGGGT	sams-3 control oligo - methylated

Table S4. Composition of nutrient-low and nutrient-high plates used for *C. elegans* culture. Related to Figure 1-4

Nutrient-low media

NGM (nematode growth medium)	2 litres
Agar	35 g
Peptone	5 g
Phosphate Buffer pH 6,0 + NaCl	50 ml phosphate buffer + NaCl
MgSO ₄ 1 M	2 ml
CaCl ₂ 1 M	2 ml
Cholesterol in ETOH (5 mg/ml)	2 ml
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Buffer NGM Phosphate + NaCl 40X	2 litres
1 M Phosphate Buffer pH:6.0	72 g K ₂ HPO ₄
	216 g KH ₂ PO ₄
2 M NaCl	240 g NaCl

Nutrient-high media

Pepton-rich plates	2 litres
Agar	50 g
Peptone	40 g
Phosphate Buffer pH: 6,0 + NaCl	50 ml phosphate buffer + NaCl
MgSO ₄ 1M	2 ml
Cholesterol in ETOH (5 mg/ml)	2 ml
<hr/>	
Buffer Pepton Rich Phosphate + NaCl 40X	2 litres
1 M Phosphate Buffer	40 g K ₂ HPO ₄
	240 g KH ₂ PO ₄
820 mM NaCl	96 g NaCl

Table S5: List of worm strains used in this study. Related to Figure 1-4.

Strain	Genotype	Source	Comments
N2	wild type	CGC	wild type strain
VC1743	<i>mett-10 (ok2204) III</i>	CGC	<i>Mett-10</i> deletion
FAS198	<i>sams-3 (uge125) IV</i>	This study	<i>sams-3</i> intron 2 deletion
FAS199	<i>sams-4 (uge126) IV</i>	This study	<i>sams-4</i> intron 2 deletion
FAS200	<i>sams-3 (uge127) IV; sams-4 (uge126) IV</i>	This study	<i>sams-3</i> intron 2 deletion and <i>sams-4</i> intron 2 deletion
FAS201	<i>sams-4 (uge126) IV; sams-5 (uge128) IV</i>	This study	<i>sams-4</i> intron 2 deletion and <i>sams-5</i> intron 2 deletion
FAS202	<i>sams-3 (uge129) IV; sams-4 (uge126) IV; sams-5 (uge128) IV</i>	This study	<i>sams-3</i> intron 2 deletion, <i>sams-4</i> intron 2 deletion and <i>sams-5</i> intron 2 deletion
FAS203	<i>sams-3 ((uge130) [sams-3::HA]) IV</i>	This study	HA-tagged <i>sams-3</i>
FAS204	<i>sams-3 (uge131) IV; sams-3 ((uge130) [sams-3::HA]) IV</i>	This study	HA-tagged <i>sams-3</i> with intron 2 deleted
FAS206	<i>sams-3 ((uge130) [sams-3::HA]) IV; sams-1 ((uge132) [OLLAS::sams-1] X; mett-10 (ok2204) III</i>	This study	HA-tagged <i>sams-3</i> with <i>mett-10</i> deletion NOTE: This strain has <i>sams-1</i> tagged with OLLAS in the background, which was not used in this study