

Supplementary Figure 1 (a) Profiling of TRF2-S-associated transcriptomes in cortical neurons. Ribonucleoprotein (RNP) immunoprecipitation (RIP)-assay and subsequent microarray analysis identified 140 putative TRF2-S bound transcripts in cultured rat cerebral cortical neurons. A bar graph shows the top gene ontology (GO) terms that are enriched in TRF2-S targets. The GO terms are ranked by -log *P* values. (b) A list of putative TRF2-S target axonal mRNAs.



Supplementary Figure 2 (a) CoIP of TRF2-S and FMRP with brain cytoplasmic extracts using 450 mM KCl IP buffer. The immunoblots demonstrate that endogenous TRF2-S and FMRP interact with each other under the high salt IP condition. (b) WT and *Fmr1* KO brain lysates were subjected to immunoblotting using FMRP and TRF2-S antibodies. Relative protein loading was assessed by bActin. Bottom, densitometry analysis of TRF2-S normalized to bActin. n=3 per group. All values are mean<u>+</u>SD. P values are based on Student's *t*-test. (c) RNA mobility shift assay showing that the binding of FMRP with TRF2-S inhibited the formation of TRF2-S-mRNA complex. The lysate from HEK293 cells expressing HA-TRF2-S was mixed either with or without recombinant GST-FMRP protein prior to HA-CoIP. The purified protein complexes were than incubated with TRF2-S target RNA fragments *Trf2*-S-CR1 (upper), *Rab3*a-CR2 (middle) and a non-target RNA fragment *Trf2*-S-3'UTR (lower) for RNA-EMSA. Lane 1, GST-FMRP only; Lane 2, GST-FMRP plus HA-TRF2-S; lane 3, HA-TRF2-S only. Arrow and arrowhead point to bound and unbound mRNA fragments, respectively.



Supplementary Figure 3 (a) Top, biotin pulldown and immunoblotting showing that neither recombinant GST empty vector (lane 1-3) nor GST-FMRPct (lane 4-6) that contains a truncated mouse FMRP with RNA binding RGG box were capable of binding to TRF2-S target mRNAs. GST-TRF2-S (lane7) served as positive control. Bottom, input of GST tagged recombinant proteins. (b) Biotin pulldown and immunoblotting showing that recombinant GST-FMRP was capable of binding to biotinylated *App1* and *App2*, two *App* fragments known as FMRP targets^{41,42}.



Supplementary Figure 4 (a-b) On culture day 6 cortical neurons were infected for 4 days with adenovirus bearing β Gal and TRF2-S (a), and lentivirus bearing shRNA#1 (865) for TRF2-S silencing and non-target control (b). The whole cell extracts were subjected for RT-qPCR analysis of TRF2-S target mRNAs. The values are the fold change normalized to *18S rRNA*. *Gapdh* is a non-target mRNA of TRF2-S. n=3. All values are mean<u>+</u>SD. P values are based on Student's *t*-test.



Supplementary Figure 5 | TRF2-S exhibits a monosomal sedimentation profile, and has little or no global effects on translations of its target mRNA. (a) On culture day 6 cortical neurons were infected for 4 days with lentivirus carrying non-target (NT) scramble shRNA and TRF2s targeting shRNA#1. Whole cell lysates were fractionated through linear sucrose gradients (15-60%). The arrow indicates the direction of sedimentation from monosomes to polysomes. (b) The relative distributions (percentages) of *Aplp1, Rab3a* and *Actb* mRNAs were quantified by RT-qPCR analysis of RNA in each of 12 gradient fractions. (c) TRF2-S, FMRP and eEF2 protein distributions in each of 12 gradient fractions from the cortical neuron lysate (9 d in culture) were analyzed by immunoblotting. (d) On culture day 6 cortical neurons were subjected to lentiviral mediated TRF2-S silencing as described in a. Four days later neurons were treated with actinomycin D. The RNAs were collected at the indicated times. The half-life of *Aplp1m*RNA was analyzed by RT-qPCR upon normalization to *Gapdh*, defined as the time needed to reach 50% of its original abundance at time 0 h. The linear-fit lines are for Lv. NT-control (black) and Lv. TRF2-S/sh#1(red), respectively. The graph represents two independent experiments.



Supplementary Figure 6 | (a) RT-qPCR analysis of axonal mRNAs harvested from the high-throughput microfluidic system after 9 d cortical neuron cultures. Upon the normalization with *Gapdh*, the axonal mRNA *synaptophysin* (*Syp*) was readily detectable, whereas the levels of dendritic mRNA *calcium/calmodulin-dependent protein kinase II* (*CamK2a*) and nuclear mRNA *Histone1(H1*) were barely measurable (average Ct values are larger than 33.6 and 32.3, respectively). n=3. All values are mean<u>+</u>SD. P values are based on Student's *t*-test. (b) Top, schematic of RNA trap assay and necessities of TRF2-S GAR domain and MS2-GST for precipitation of a (*ms2*)24 fused TRF2-S target RNA chimera. Bottom, HEK293 cells were co-transfected with the plasmid expressing (*ms2*)24 tagged *Aplp1*-5'CR1 RNA chimera, accompanied with either MS2-GST or HA-TRF2-S variants. The existence of the exogenous RNA chimera in each GST- or HA-precipitates was assessed by RT-PCR using a primer set (T7 and Aplp1). An arrow points to PCR products; an arrowhead points to primer dimmers. (**c**-**d**) DIV7 cortical neurons were co-transfected with MS2-YFP-NLS and (*ms2*)24-*Rab3a*-CR2 without (**c**) or with mCherry-FMRP (**d**). Overexpression of mCherry-FMRP failed to translocate YFP labeled RNP from the nucleus. The images are representative from 50 transfected neurons per group. Scale bar=20µm.





Supplementary Figure 7 (a) Immunoblots showing the expression patterns of TRF2-S and FMRP during the course of neuronal maturation. Synaptophysin (Synap) and PSD95 are pre-and post-synaptic markers, respectively. (**b**-**c**) Neurons (6 days in culture) were infected for 4 days with lentivirus carrying shRNAs for the silencing of rat TRF2-S (shRNA#1, 865 and shRNA#2, 774) in **b**; for the silencing of rat FMRP (shRNA#1, 2624 and shRNA#2, 2623) in **c**. A non-target (NT) scramble shRNA was used as controls. Cell lysates were then subjected to immunoblot analysis of endogenous levels of Rab3a, along with TRF2-S (**b**) and FMRP (**c**). Densitometry analysis showed as bar graphs at bottom of b and c. n=3; *p<0.05 and **p<0.005. All values are mean<u>+</u>SD. P values are based on Student's *t*-test.

Full scan for western blot in Figure 1 c





Full scan for blots in Figure 1 e



Supplementary Figure 8. Full blots for Figures 1c, d and e.

Supplementary Figure 9



Full scan for western blots in Fig. 2b



* Endogenous TRF2 in 293 cells

TRF2-S/FMRP IP 16 THEY FINGE THEY FINGE 188-IB: FMRP TRF2S NR 6 98 -IB: eEF2 62 -62 -49 -IB: TRF2-S Input 38 -28 FMRP TRF2-S eEF2 16 -* lgG

Full scan for western blots in Fig. 2c





Supplementary Figure 10. Full blots for Figures 3 a, b c, d and e.



Full scan for western blots in Supplementary Fig. 2a

62 49 38 28 17 10 WT1 WT2 K01 K02 WT1 WT2 K01 K02 WT1 WT2 K01 K02 WT1 WT2 K01 K02

Full scan for western blots in Supplementary Fig. 2b



Kb

12

Full scan for western blots in Supplementary Fig. 2c

RNA EMSA

2

3



Supplementary Figure 12. Full blots for Supplementary Figures 3 a and b.



Supplementary Figure 13. Full blot for Supplementary Figure 5c.

Full scan for western blots in Supplementary Fig. 7a



NT sh#2 sh#1 NT sh#2 sh#1 NT sh#2 sh#1 49 49 IB:Rab3a 62 IB:TRF2-S B:BActin -38 38 49 -+ 28 -28 = 38 - 17 -17 -28

Full scan for western blots in Supplementary Fig. 7b

Full scan for western blots in Supplementary Fig. 7c



Supplemental table1. Global analysis of TRF2-S bound neural mRNAs Gene Symb

	group 1 Z ratio	group 2 Z ratio	RefSeq ID	GI	Description
MLPH	14.07	8.5	NM_001012135.1	58865839	melanophilin (Mlph)
Akt1	9.67	5.85	NM_033230	402692490	thymoma viral proto-oncogene 1 (Akt1)
NNAT	9.12	9.85	NM_181687	399498546	neuronatin (Nnat)
Gcfc2	8.79	5.37	NM_001134554	197386558	GC-rich sequence DNA-binding factor 2 (Gcfc2) (RGD1304792_predicted)
LOC501280	8.59	7.88	XR_007805	109486608	similar to myosin regulatory light chain-like (LOC501280)
EIF4EBP2	8.41	6.8	NM_001033069.1	74271858	eukaryotic translation initiation factor 4E binding protein 2 (Eif4ebp2)
MLLT11	8.4	7.71	NM_001013912.1	62078554	myeloid/lymphoid or mixed-lineage leukemia (trithorax homolog
LOC500095	8.33	4.95	XM_575446	62647113	similar to T-cell receptor beta chain V region 86T1 precursor (LOC500095)
TEKT1	7.96	3.54	NM_053508.2	51036667	tektin 1 (Tekt1)
EPN1	7.88	8.21	NM_057136.1	16923989	Epsin 1 (Epn1)
MAPK3	7.41	1.95	NM_017347	68537200	mitogen activated protein kinase 3 (Mapk3)
MLF2	7.03	9.95	NM_001107889	157823872	myeloid leukemia factor 2 (predicted) (Mlf2_predicted)
SLC39A2	6.94	4.33	NM_001107260	157822388	solute carrier family 39 (zinc transporter)
R3hdm2	6.54	3.1	NM_001130557	194474025	R3H domain containing 2
PFN1	6.306	7.966	NM_022511.2	42476143	profilin 1
RGD1564087	6.1	2.47	XR_009360.1	109467794	similar to 14-3-3 protein tau (14-3-3 protein theta) (predicted)
RGD1559566	5.99	4.83	XM_001056263.1	293350680	similar to 60S ribosomal protein L9 (predicted) (RGD1559566_predicted)
ATP5G3	5.997	7.804	NM_053756	16758591	ATP synthase, H+ transporting, mitochondrial Fo complex, subunit C3 (subunit 9)
CIC	5.98	5.89	NM_001107490.3	215272395	capicua homolog (Drosophila) (predicted) (Cic_predicted)
Brsk1	5.93	7.05	NM_001127337	187960159	BR serine/threonine kinase 1
CAMK2N2	5.63	4.92	NM_021678.1	11067400	calcium/calmodulin-dependent protein kinase II inhibitor 2 (Camk2n2)
RGMA	5.51	4.83	XM_218791.3	157819830	RGM domain family
Ccdc125	5.25	3.49	NM_001134761	197927213	CCDC125 coiled-coil domain containing 125
GNG10	5.21	7.74	NM_053660.1	399154111	guanine nucleotide binding protein (G protein)
P11 protein	5.05	2.51	XM_579635.1	62659845	similar to P11 protein (LOC497693)
Tmem189	5.02	2.58	NM_001113752	166091516	transmembrane protein 189
LOC365025	5.15	6.63	XM_577671.1	62665500	similar to alpha tubulin subunit (LOC365025)
SNURF	5.12	10.02	NM_130738.1	397787561	SNRPN upstream reading frame (Snurf)
LOC300760	5.03	3.57	XM_236308.3	62653977	similar to H3 histone
Yaf2	5.063	5.468	XM_001073895.1	199559858	similar to YY1-associated factor 2 (LOC690262)
LOC500344	4.9	2.49	XM_575696.1	62648245	LOC500344 (LOC500344)
MRPS2	4.88	3.8	NM_001108576	157819458	mitochondrial ribosomal protein S2

TOMM22	4.63	4.9 NM_212514.2	71534292 translocase of outer mitochondrial membrane 22 homolog (yeast) (Tomm22)
Arglul	4.63	4.5 NM_001024972.1	68163363 arginine and glutamate rich 1 ()
TMEM158	4.61	6.33 NM_057212.1	17933495 transmembrane protein 158 (Tmem158)
ГЛЕН	4.44	8.82 NM_001134839.1	198386344 lymphocyte antigen 6 complex
LENG4	4.31	3.55 NM_001134978	201066394 leukocyte receptor cluster (LRC) member 4
AKT1S	4.25	3.54 NM_001106259	408772021 AKT1 substrate 1 (proline-rich)
Zgc:56193	4.23	5.55 XR_008163.1	109477420 similar to Zgc:56193 (predicted) (RGD1562162_predicted)
WBP2	4.21	3.32 NM_138975.1	20376817 WW domain binding protein 2 (Wbp2)
CYP46A1	4.11	3.77 NM_001108723.	157817886 cytochrome P450
Btbd2	4.11	5.43 XM_001076264.1	392341333 similar to BTB (PO)Z domain containing 2 (predicted) (RGD1566094_predicted)
CSDC2	4.08	3.7 XM_001077363.1	293348836 cold shock domain containing C2
RPL13A	4.05	8.07 NM_173340.1	27545437 ribosomal protein L13A (Rpl13a)
RGD1564980	4.02	4.16 XM_575373.2	293358409 similar to 60S ribosomal protein L29 (P23) (predicted) (RGD1564980_predicted)
Prrc2a	3.99	3.82 NM_212462.2	260166645 proline-rich coiled-coil 2A (Prrc2a)
PPP2R5B	3.93	7.09 NM_181379.2	60593006 protein phosphatase 2
LOC681314	3.91	5.43 XM_001061020.1	109489050 similar to BAX protein
TXN1	3.87	3.53 NM_053800.2	42476285 thioredoxin 1 (Txn1)
MFAP2	3.81	3.89 NM_001107989	157818962 microfibrillar-associated protein 2 (predicted) (Mfap2_predicted)
HMG-2	3.8	3.27 XM_573272.1	392351998 similar to High mobility group protein 2 (HMG-2) (LOC498072)
Hmgb2	3.8	2.76 NM_017187	158711705 high mobility group box 2 (Hmgb2)
LOC364253	3.76	4.27 XM_344296.1	34869079 similar to RIKEN cDNA 1110003E01 (LOC364253)
ADCK2	3.72	1.75 NM_001107855	157822890 aarF domain containing kinase 2 (predicted) (Adck2_predicted)
RPS7	3.71	2.15 XM_213053.2	62650073 ribosomal protein S7 (Rps7)
Ppp1r18	3.69	4.46 NM_001126287.2	261399901 PPP1R18 protein phosphatase 1, regulatory subunit 18
B3GAT1	3.66	4.35 NM_054003.1	16758917 beta-1,3-glucuronyltransferase 1
RGD1565149	3.66	4.35 XM_001077968.1	293343422 similar to chromosome 16 open reading frame 7 (predicted) (RGD1565149_predicted)
Fam57b	3.62	7.49 NM_001106296.2	182888556 family with sequence similarity 57, member B
LOC306766	3.61	3.23 NM_001014007.1	62078742 hypothetical LOC306766 (LOC306766)
RGD1311783	3.59	4.89 XM_001059424.1	197386190 similar to RIKEN cDNA 2010012005 (predicted) (RGD1311783_predicted)
Ptbp2	3.52	1.87 NM_001005555.1	53850637 polypyrimidine tract binding protein 2 (Ptbp2)
NME 2	3.51	4.75 NM_031833.2	55926144 NME/NM23 nucleoside diphosphate kinase 2
Agap3	3.47	5.59 NM_001108616	157824114 ArfGAP with GTPase domain, ankyrin repeat and PH domain 3
Rab2a	3.468	1.64 NM_031718.1	13929005 member RAS oncogene family (Rab2a)
MGST3	3.46	4.62 XM_001076104.1	109498992 microsomal glutathione S-transferase 3 (predicted) (Mgst3_predicted)
Gprin1	3.45	3.49 XM_001070221.1	293342617 G protein-regulated inducer of neurite outgrowth 1 (predicted) (Gprin1_predicted)
LOC500721	3.45	5.65 XM_580114.1	62651251 LOC500721 (LOC500721)
Gstp1	3.45	4.18 NM_012577.1	169646324 glutathione-S-transferase
Nckapl	3.41	2.03 XM_230038.4	158138548 NCK-associated protein 1 (Nckap1)
RRS1	3.394	6.446 _{NM} _001079699.1	119226199 RRS1 ribosome biogenesis regulator homolog (S.cerevisiae)
NREP	3.38	7.02 NM_178096.2	31342528 neuronal regeneration related protein (Nrep)
RGD1561956	3.37	3.52 XM_344287.1	34869055 similar to RIKEN cDNA 1110003E01 (predicted) (RGD1561956_predicted)

GDI1	3.35	5.95 NM_017088.2	71534275 guanosine diphosphate dissociation inhibitor 1 (Gdil)
RGD1564450	3.35	3.26 XM_001078024.1	157822410 RGD1564450 (predicted) (RGD1564450_predicted)
RGD1560015	3.28	4.26 XM_574311.2	392343688 trans-2,3-enoyl-CoA reductase-like
Npm3	3.26	3.95 XM_001058548.1	392338228 nucleophosmin/nucleoplasmin, 3
Zbtb45	3.26	3.37 NM_001107478	157819040 zinc finger and BTB domain containing 45
SPC25	3.26	2.23 NM_001009654.1	57527006 SPC25, NDC80 kinetochore complex component
LOC311629	3.21	3.81 XM_230844.3	62646356 similar to sorting nexin 21 isoform a; sorting nexin L; sorting nexin 21 (predicted)
PPFIA3	3.21	3.91 NM_001270985	401709972 protein tyrosine phosphatase, receptor type
Cox7b	3.2	2.29 NM_182819.2	65301489 cytochrome c oxidase subunit VIIb
Gtf2h5	3.17	1.82 NM_001126088	186910270 general transcription factor IIH, polypeptide 5
Atxn7l3	3.11	3.88 XM_213493.3	392351635 ataxin 7-like 3, transcript variant 2
MAP1B	3.166	2.525 NM_019217	158749619 microtubule-associated protein 1B
Sypl1	3.106	1.868 NM_001014263	62079260 synaptophysin-like 1
RPL18A	3.09	5.68 NM_212510.1	47059005 Rattus norvegicus ribosomal protein L18A (Rpl18a)
RGD1565784	3.05	2.47 XM_001059479.1	157817212 PREDICTED: Rattus norvegicus RGD1565784 (predicted) (RGD1565784_predicted)
TMEFF1	3.03	5.89 NM_023020.2	121583853 Rattus norvegicus transmembrane protein with two follistatin-like domains 1 (Tmeff1)
Fabp7	3	3.62 NM_030832.1	402744885 fatty acid binding protein 7, brain
Map11c3a	3	1.68 NM_199500	377652334 microtubule-associated protein 1 light chain 3 alpha
LOC499793	2.97	5.51 XM_575129.1	62645054 similar to 60S ribosomal protein L12
Cbr1	2.94	2.98 NM_019170.2	46048534 carbonyl reductase 1 (Cbr1)
PEA15A	2.896	2.177 NM_001013231.1	61557369 phosphopro cytoch
SLC25A11	2.89	4.96 NM_022398.1	11693169 solute carrier family 25 (mitochondrial carrier; oxoglutarate carrier)
Aplp1	2.89	-2.08 XM_001076030.1	198041598 similar to Amyloid beta (A4) precursor-like protein 1 (predicted)
Cox7a2l	2.88	5.51 NM_001106704	157821820 cytochrome c oxidase subunit VIIa polypeptide 2 like
BAX	2.87	3.103 NM_017059	386869282 Bcl2-associated X protein
S100a1	2.85	2.51 NM_001007636.1	218749895 S100 calcium binding protein A1 (S100a1)
Rpl7	2.82	2.53 NM_001100534	198278504 ribosomal protein L7
RGD1564372	2.77	4.31 XM_344301.1	34869098 similar to RIKEN cDNA 1110003E01 (predicted) (RGD1564372_predicted)
Dpp9	2.77	2.28 XM_001061312.1	392342223 dipeptidyl peptidase 9
IL-17D	2.77	2.54 XM_577439.1	62661954 similar to IL-17D (LOC502011)
RGD1565358	2.76	2.29 XM_235062.2	109480494 genes associated with retinoid-IFN-induced mortality 19 (predicted)
C1QL1	2.76	2.8 NM_001108838	157819268 complement component 1,q subcomponent-like 1
Lrrc73	2.75	5.11 NM_001024360.1	119310198 eucine rich repeat containing 73
Actb	2.75	6.06 NM_031144.2	42475962 beta actin
Mapre1	2.742	1.579 NM_138509.3	78097099 microtubule-associate
RAB3A	2.72	3.33 NM_013018.2	61098194 RAB3A, member RAS oncogene family
LOC301444	2.72	3.83 NM_001047872	114145717 pseudogene for diazepam binding inhibitor 1 (LOC301444)
LYPLA2	2.71	4.7 NM_031342.1	13786177 lysophospholipase 2 (Lypla2)
NDUFB8	2.66	5.9 NM_001106360	187937126 NADH dehydrogenase (ubiquinone) 1 beta subcomplex 8
Hm13	2.65	3.74 NM_001107789	157823208 histocompatibility 13
Hdac3	2.623	2.343 NM_053448.1	16758191 histone deacetylase 3

SNAP91	2.61	4.12 NM_031728.1	13994176 synaptosomal-associated protein 91 (Snap91)
TERF2	2.61	1.54 NM_001242355.1	334285045 telomeric repeat binding factor 2, transcript variant 2
Nras	2.6	4.8 NM_080766.1	18158430 neuroblastoma ras oncogene (Nras)
LOC691672	2.6	4.06 XM_001079224.1	109513656 similar to Discs large homolog 5 (Placenta and prostate DLG)
MLLT3	2.604	3.286 NM_053718.1	192447427 myeloid/lymphoid or mixed lineage-leukemia translocation to 3 homolog (Drosophila)
Zfp513	2.59	4.67 NM_001012110.1	58865789 zinc finger protein 513 (Zfp513)
Mgrnl	2.57	4.76 NM_001013964.1	62078656 mahogunin ring finger 1, E3 ubiquitin protein ligase
Ndufb2	2.56	2.54 XM_001070173.1	219277691 NADH dehydrogenase (ubiquinone) 1 beta subcomplex
Stk11	2.55	4.6 XM_234901.4	157823728 downstream of Stk11 (predicted) (Dos_predicted)
Hes5	2.55	4.53 NM_024383.1	13242286 hairy and enhancer of split 5 (Drosophila) (Hes5)
VAMP8	2.491	4.002 NM_031827	13929181 vesicle-associated membrane protein 8
Lmo3	2.43	3.91 NM_139112.1	20806164 LIM domain only 3 (Lmo3)
Mkrn3	2.41	3.8 XM_001057262.1	392337728 makorin, ring finger protein, 3
Gnal	2.357	2.641 NM_001191836	300797436 guanine nucleotide binding protein, alpha, stimulating, olfactory type
NDUFB3	2.259	3.999 NM_001106912	157824070 NADH dehydrogenase (ubiquinone) 1 beta subcomplex 3
Lemd2	2.18	4.78 NM_001039032.1	84781657 Rattus norvegicus LEM domain containing 2 (Lemd2)
Ube2c	2.18	1.86 NM_001106542	157818348 ubiquitin-conjugating enzyme E2C
Efna2	2.15	2.53 NM_001168670	274327666 ephrin A2 (Efna2)
LOC361128	2.08	4.17 XM_341415.2	62662451 similar to TR4 orphan receptor associated protein TRA16 (LOC361128)
Hnrpa2b1	1.97	4.15 XM_342684.3	157265558 heterogeneous nuclear ribonucleoprotein A2/B1 (predicted) (Hnrpa2b1_predicted)
MTAP2	1.939	1.862 NM_013066.1	6981181 microtubule-associated protein 2
Sdhb	1.82	2.12 NM_001100539	209915613 succinate dehydrogenase complex, subunit B, iron sulfur (Ip)

Supplemental Table 2: TRF2-S targets related to axonogenesis mRNA identified in DIV 13 cortical axons⁸ RPL18A Ribosomal protein L18A RPL13A Ribosomal protein L13A LOC499793 60S ribosomal protein L12 Rpl7 Ribosomal protein L7 Hnrpa2b1 Heterogeneous nuclear ribonucleoprotein A2/B1 NNAT Neuronatin MAP1B Microtubule-associated protein 1B Map1lc3a Microtubule-associated protein 1 light chain 3 alpha ATP5G3 ATP synthase, H+ transporting, mitochondrial Fo complex, subunit 9 Cox7b Cytochrome c oxidase subunit VIIb COX7A2L Cytochrome c oxidase subunit VIIa polypeptide 2 CAMK2N2 Calcium/calmodulin-dependent protein kinase II inhibitor 2 GNG10 Guanine nucleotide binding protein (G protein) TXN1 Thioredoxin 1 NME2 Non-metastatic cells 2 NREP Neuronal regeneration related protein LOC691672 Discs large homolog 5 mRNA indentified in embryonic axons¹¹ RPL18A Ribosomal protein L18A RPL13A Ribosomal protein L13A LOC499793 60S ribosomal protein L12 Rpl7 Ribosomal protein L7 NNAT Neuronatin MAP1B Microtubule-associated protein 1B Map1lc3a Microtubule-associated protein 1 light chain 3 alpha MAPRE1 Microtubule-associated protein MTAP2 Microtubule-associated protein 2 NCK-associated protein Nckap1 ATP5G3 ATP synthase, H+ transporting, mitochondrial Fo complex, subunit 9 SLC25A11 Solute carrier family 25 (mitochondrial carrier; oxoglutarate carrier) COX7A2L Cytochrome c oxidase subunit VIIa polypeptide 2 BAX Bcl2-associated X protein Ndufb2 NADH dehydrogenase (ubiquinone) 1 beta subcomplex NDUFB3 NADH dehydrogenase (ubiquinone) 1 beta subcomplex 3 Cox7b Cytochrome c oxidase subunit VIIb TOMM22 Outer mitochondrial membrane 22 homolog (yeast) RAB3A RAB3A APLP1 Amyloid beta (A4) precursor-like protein 1 RAB2 RAB2 SYPL Synaptophysin-like 1 Guanosine diphosphate dissociation inhibitor 1 GDI1 GNG10 Guanine nucleotide binding protein (G protein) NREP Neuronal regeneration related protein Akt1 Thymoma viral proto-oncogene 1 Brsk1 BR serine/threonine kinase 1

PPP2R5B	Protein phosphatase 2
YY1	YY1-associated factor 2
HMGB2	High mobility group box 2
TXN1	Thioredoxin 1
CIC	Capicua homolog (Drosophila)

Supplemental Table 3 Oligomer pairs for biotin pull-down assays Forward

Trf2s: 5'UTR CCAAGCTTCTAATACGACTCACTATAGGGAGAAACAGCGGGCCCTGCTTC CGTGTTCGACCGCCGAG CR1 CCAAGCTTCTAATACGACTCACTATAGGGAGAATGGCGGGAGGAGGGG CR2 CCAAGCTTCTAATACGACTCACTATAGGGAGATTTGTATCAAAAACAAAGAATTTGA TTAGGGCTGTTCCACTTG 3'UTR CCAAGCTTCTAATACGACTCACTATAGGGAGACGGGCTTGAAGAAAAGGAAG CCCGCACTTCAAACAGCT Clip-WT ${\tt CCAAGCTTCTAATACGACTCACTATAGGGAGAGGGTCTGCGGGGCCTTTCGGAGTAGCCGGGACTTCAGGCAGATCCG}$ Clip-MT1 CCAAGCTTCTAATACGACTCACTATAGGGAGAGGT CTGCGGGCCTAAGCCTCATCCCGGTACCGGGACTTCAGGCAGATCCG

- Clip-MT2 CCAAGCTTCTAATACGACTCACTATAGGGAGAGGT CTGCGGGCCTTCGGAGTAGCCGGTACCGGGACTTCACCGTCTAGCG
- ClipMT1+2 CCAAGCTTCTAATACGACTCACTATAGGGAGAGGT CTGCGGGCCTAAGCCTCATCCCGGTACCGGGACTTCACCGTCTAGGG

Aplp1

Rab3a	
3'UTR	CCAAGCTTCTAATACGACTCACTATAGGGAGAGAAGAACGACCTTGACCCCAACCCT
CR5	CCAAGCTTCTAATACGACTCACTATAGGGAGAAGATGCCCCAGTGACCCTTCC
CR4	CCAAGCTTCTAATACGACTCACTATAGGGAGAGAGAGAGTTCTGATGGCCCTGAG
CR3	CCAAGCTTCTAATACGACTCACTATAGGGAGAGGTCAGCAGAGTCACTCCCAC
CR2	CCAAGCTTCTAATACGACTCACTATAGGGAGAGGTTGTGCCCTTCCATTGCCT
5'CR1	CCAAGCTTCTAATACGACTCACTATAGGGAGAGCAGGGCCGGGGGGGG

5'CR1	CCAAGCTTCTAATACGACTCACTATAGGGAGAGAGCCTCCACCGCCGCCTGCA
CR2	CCAAGCTTCTAATACGACTCACTATAGGGAGACCATGGGCTTCATTCTAATGT
3'UTR	CCAAGCTTCTAATACGACTCACTATAGGGAGAACCACACTTCCCTTCCCCACC

App

887-1150	CCAAGCTTCTAATACGACTCACTATAGGGAGAtgcagatgggagtgaagacaaag
1944-2150	CCAAGCTTCTAATACGACTCACTATAGGGAGAccgccctgctgccgaccgagga

Reverse

ATTTTTGAAGCCTTTTCAAATTCTT

CGGATCTGCCTGAAGTCCCGGTACC CGGATCTGCCTGAAGTCCCGGTACC CGCTAGACGGTGAAGTCCCGGTACC CGCTAGACGGTGAAGTCCCCGGTACC

AGGCAATGGAAGGGCACAACC GTGGGAGTGACTCTGCTGACC CTCAGGGCCATCAGAACTCTC GGAAGGGTCACTGGGGCATCT AGCTAGGGTTGGGGTCAAGGT GAAGAGGACTTTATTGGGATGTTAG

ACATTAGAATGAAGCCCATGG GGTGGGGAAGGGAAGTGTGGT TTCTCTCTTTAATGTCAAGGACTGC

tggaaatgggcatgttcattctc cttcttcagcatcaccaaggtg