

SUPPLEMENTARY INFORMATION

Auto-Regulatory RNA Editing Fine-Tunes mRNA Re-Coding and Complex Behavior in *Drosophila*

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Supplementary Figure S1 | Conserved RNA editing sites. Shown below are the editing sites in the 23 mRNAs analyzed in this study, alongside the surrounding local sequence. Sensitivity to dADAR auto-editing was defined based on data from male heads.

*= intron location

A= editing site

A= non-sensitive site

A= G-sensitive site

A= S- and G-sensitive site

syt (CG3139)

TTGAAGAAGATGGACGTGGGCGGATTGTCTG*ATCCATATGTGAAA¹TTGCAATCA
TGCAAAATGGCAAACGTTTGAAAAAGA²GAAGACAAGT³TCAA⁵A⁶AAATGCACCC
TCAACCCTTACTATAATGAGTCGTTCTCATTGGAAGTACCATTGGAACAAAT⁴CAA*
AAAATCTGTCTCGTTGTGACCGTCGTGGACTACGATCGTATTGGCACCTCCGAACCC
ATCGGCCGCTG

cap (CG33653)

GACACAAGGTCTAAGCGGTAAACTT¹TGTCTGTGCTAGAGTCGACTTTGTC

cpx (CG32490)

AACTGAAAAATCAAAT¹GAAACGCAAGTAA²A³TGAGCTAAAAACTCAAATAGAGG
GAAAATGTGTCATGCAGTGA

unc (CG2999)

ACGCAATTGACGCAGTGTTTCGATGTTGTT¹GCAAACCTCGAGTGTCTGATCCAGA
AATTGGA

stn (CG40306)

GTTTACTATGCCCCGCCACGCAGGTCTCCCAT¹CCACCGTGCGCTCGGTGAGTGTCCA
GGATTCCGGATGGCGATGAACCGC

lap (CG2520)

ACTAGCAGCGGTGCTGCTGGCGCTAGCGCTGCACTA¹CAAATCCATTTCTATCGTC
GCCGCCAGCCGCGCAGGCTGGCCAGCCGAT²GTTGATCTGTTCCGGTGCCGCGTCCG

dsc (CG34405)

TATGAAGGAGGAGTGTTGGAA^A¹TGTTTCTCACCGAATCTCAAAAACACTA

sbd (CG6798)

GCCCAGCGATTTCGGGCGAAAAG^G¹GTT^A¹CGTT^A²TGTATTTCGATACTCGTGTC

stj (CG12295)

TTCAATGTTGGGACAGAGGCTAGAT^A¹TCGCAAAGCATGTTGTCAATACGATATTAG
GTACAAATGACTTTGTGAAC^A²TCTTCACCTTTGATAAGGAAGTGAG

ACTGAAAGAAGGGATTGAACTGTTT^A³GACCCAAATCGATCGCCAATTATAC

cat (CG15899)

GGTGGCCATTTTGGTTGAGGGATTC^A¹G TTCAGAG^G¹CGAAATGAACGTCGCGA

cad (CG4894)

CAGCGGAATGTGTTATGAAAATTTT^A¹G CATATGGTTTTGTGTTACAT^A²ATGGTGCA
TATCT^A³^A⁴GAAATGGATGGAATTTATTAGATTTTATTAGATTTTACAATTGTAGTTAT
^A⁵GG^G¹GGCGATAAGTACTGCACTCTCCC

shb (CG1066)

GGTGGTGCAGGTCTTTCGCATCATGCGCATCCTGCG^A¹^A²TCCTTAAGCTGG
CCCGTCACTCAACGGGCCTG

TCGTTTCAATACCGGAA^A³CATTTTGGTGGGCGGGTATTACAATGACAACCT
GTTGGCTACGGGGACATCT^A⁴TCCCACTGCACTGGGAAAGGTTATTGGT^A⁵C

CTGTGTGTTGCATATGCGGTGTTCTGGT^A⁶^A⁷TCGCTTTCCTATTCCCA
TCATCGTTAACAATTTTGCTGAATTTTATAAGAATCAG

eag (CG10952)

TCGCCGCGGAGACAGACAACGAGA^A¹GGTGTTCACCATCTGCATGATGATC
CTGGATACCGAGAAG*^GTA^A²CTAACT^A²TTGTCCGAAAGATATGAAGGCTGAC
TATGTGTTTCATCTAAATCGCAAAGT^A⁵TTT^A⁴ACGAGCATCCGGC^A⁵TTTCGTCTGGCC
TCGGATGGTTGTC
CTCAGACATCGACTGATTTTTTCGCA^A⁶GGTGGCCGATGTGAAGCGCGAAAAA
GTGATTTGCATGCCATCAAACGTGATA^A⁷AATTGCTCGAAGTCCTCGATTTCT

slo (CG10693)

GA^A¹ACTCTGGCGATCCGCTGGATTTT^A¹ATAATGCTCATCGTTTATCGTATTG
CATGATAACAG*^AAACTGGTCAATGATA^A²GTAACGTGCAGTTTCTCGATCAAGA

rdl (CG10537)

GTCAATTTTGGACCGATCCTCGTTT^A¹GCGTAT^A²GAAAACGACCTGGTGTAGAAACA
CT
CTACATACCCTCTGGACTGATCGTT^A³TTATATCATGGGTATCATTTTGGCTATGGGT
ATCATTTTGGCTCAATCGC^A⁴ATGCAACGCCGGCGCGTGTGGCGCT
GTCGGCTACATGGCAAACGAATTCA^A⁵^A⁶TGCGAAAACAAAGATTTATGGCGAT

daf (CG32975)

CTACTTCCGATGCGGTGCCATTGCTGG*^GTA^A⁹CATATTTCAATTGCATAATGTTTATG
GTAGCTTCATCCGTTGTGTCAACGATTTT^A¹^A²TATTAAATTATCATCATCGAAATGC
TGACAGACTGTGCCTTATCATATTCACAATGTTCA^A³CAA^A⁴TATT^A⁵GCCACAAT^A⁶GCT
GTACTACT^A⁷TC^A⁸GCACCACATATTATTGTCTCGT

das (CG4128)

GGATTCGATGGCACGTATCACACCAA^A¹CA^A²TTGTGGTCAAACAT^A³^A⁴CGGCAG
TTGTCTGTACGTGCCCCCT

GTATCTTCAAGAGCACATGCAAGATA⁵GACATCACGTGGTTCCCATTTGATG
GGAAATCAG*TTGGATTTGGTTTTGA⁶TTCCGAAGATGGAGGGGATCTTTC

ard (CG11348)

AGAATATGACACAAAAAGTTGGAGT¹²GATTTGGTTTGGCGTTCGTACAGCT
TAATCAATGTC*AATGAGAAAAATCA³⁴TTATGAAATCAAACGTTTGGTTACG

sha (CG12348)

ATGTCCCTTTAGACGTATTTAGTGAAGAAATA¹²ATTTTATGAATTAGGTGATCAA
GCA
CGAATCAGGCTATGTCCTTGGCAAT³TTACGAGTGATACGATTAGTTCGAG
AAAATTGTCGGCTCTTTGTGCGTG⁴TCGCTGGTGTGCTGACAATCGCACT
TTTCAATTACTTCTATCACCGCGAA⁵CGGATC⁶GGAGGAGATGCAGAGCCA

adr (CG12598)

CAGCACCTTGTGGGGATGCACGGATATTT¹GTCCTCACGAAAACGACACTGGTGT
GATAAAC

cac (CG1522)

TTCAACCGATTTGATTGCGTTGTCATT¹GTGGTTCGATATTCGAAGTGATCTGGTCC
G
ATGCCCAAGAACTAACAGCAGCCGAAGAGG²ACAA¹³GTCGAA³GAGGATAAAGAG
AAACAACCTGCAGGAGC
ACCCTGGCAGCTATTTAAGAGAATTCTGGA⁴TATTATGGATGCTGTGGTCGTTATAT
GCGC
GCGTCGTGAACTCATTGAAAAATGTTGTAA⁵CATTCTAATCGTGTACATATTGTTTC
AATT
AATGGAAAATTTTTTTTATTGTACGGACGAA⁶GTAACATACTTCCGCAGAGTGCCA*
GGGCT

CATGATATCGGGTTCTCCGATACCGTATCTA¹¹ ATGTTGTAGAGATGGTCAAGGAGAC
TCGTC

AGGCATGGCAACAGTCATCCGCGGTATCCA¹² GAG*⁶ GTTCATGGTCAGCATCGACA
AGTCCGG

par (CG9907)

CAATTTACGACTGAAAAC³TTTC⁴ TTAATTGA⁵⁶ATAAATATTTTGAAACAGCT
GTTATCACTATG

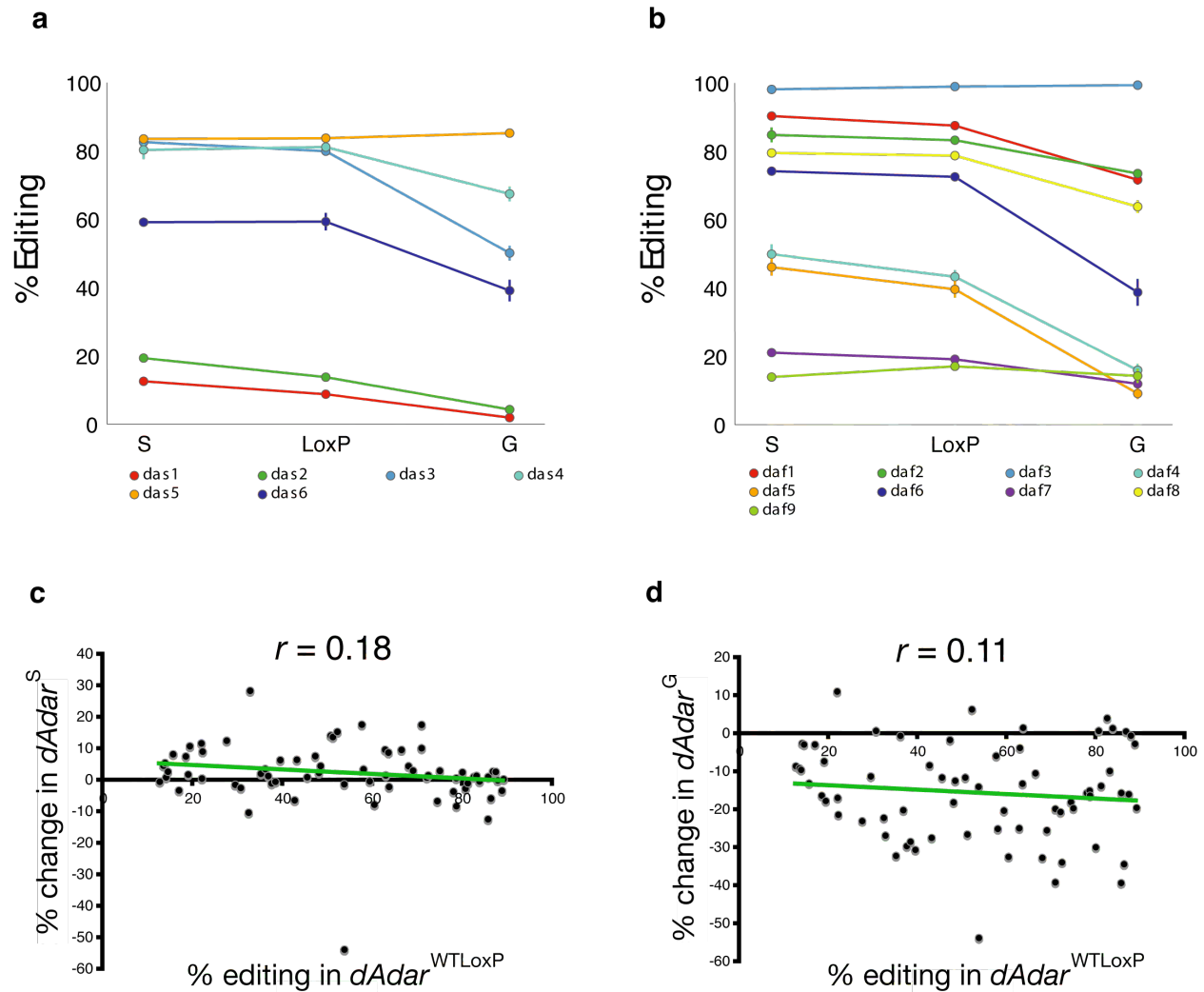
glu (CG7535)

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GCCTCAAAGTCGATCTACTATTCA³ GCGAGAATTCTCATATTACTTAATACAAATT

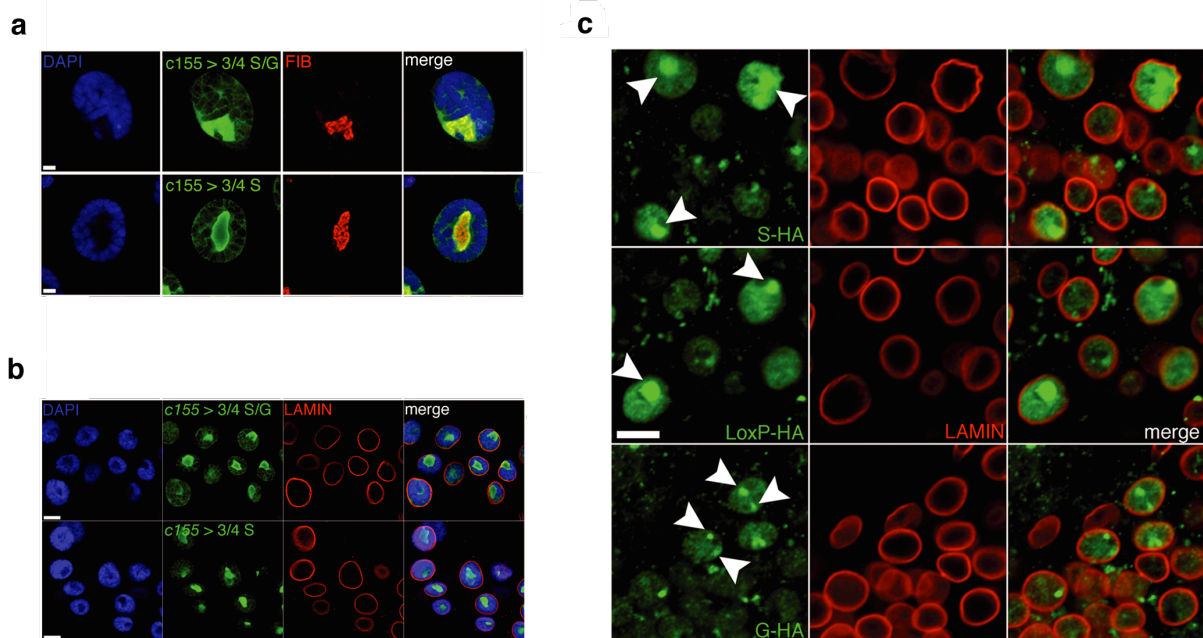
dop (CG18314)

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GTCTTTGCATTCTGGGTGTCTGGCTGCCATGGATTCTG

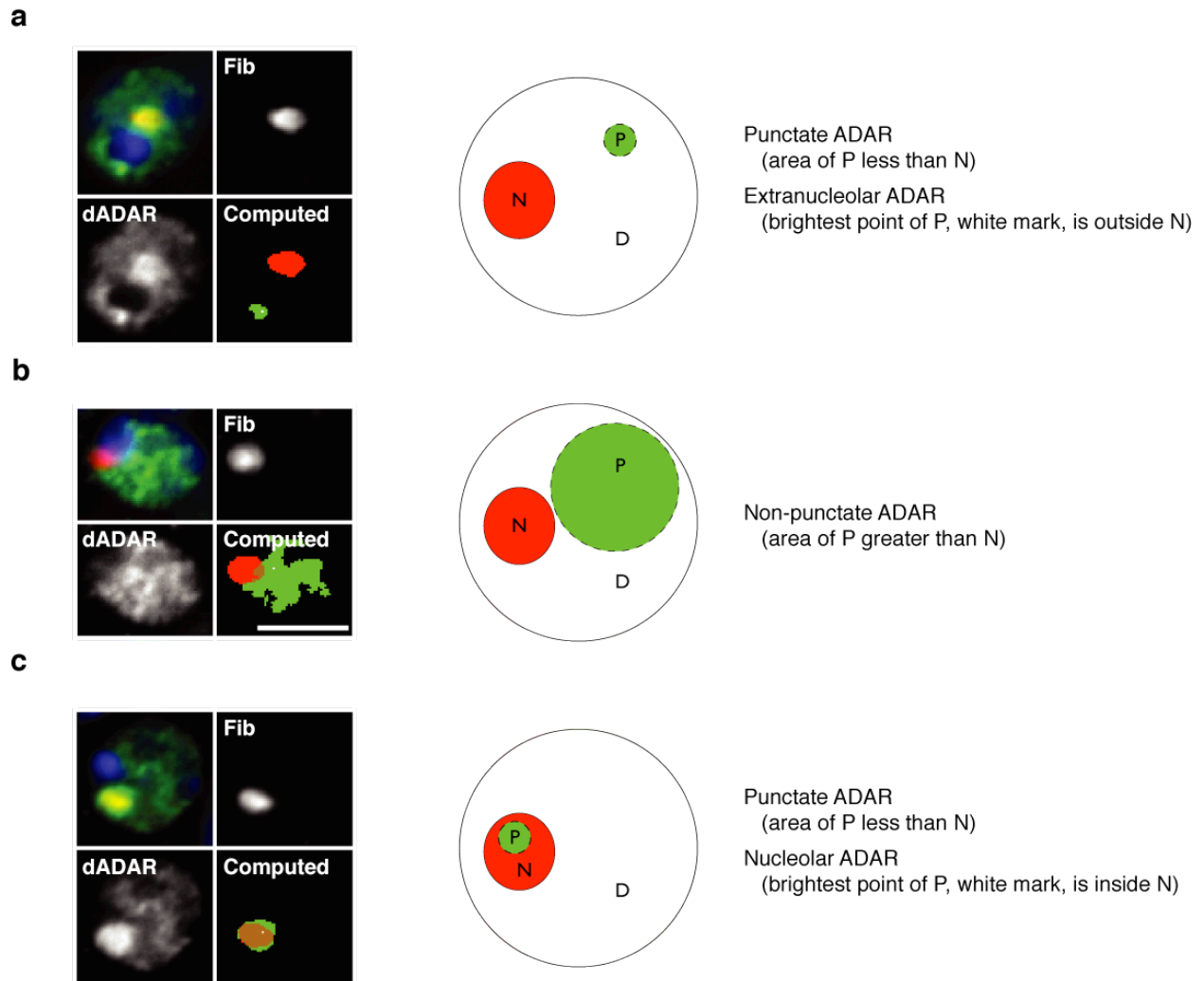
TGCTTAACCATTTGTTGTAAGACTA⁵⁶ GGGCCGTTTGCAAGCAGAGCTA⁷⁸ TCGGGT
TGGACCCAGATGACTA⁹ GAGTTAGATTTAGATTTTCATTCTCC



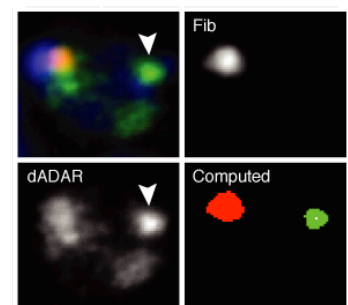
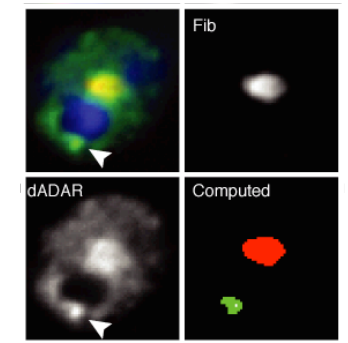
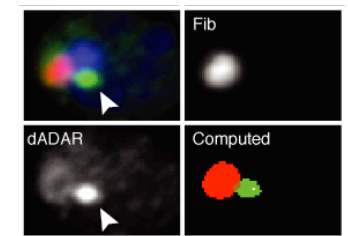
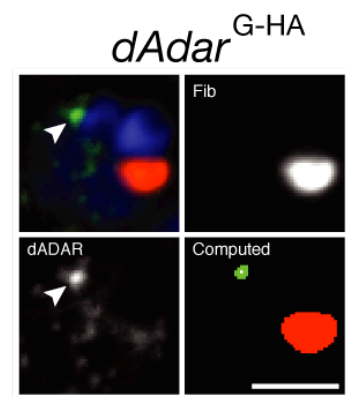
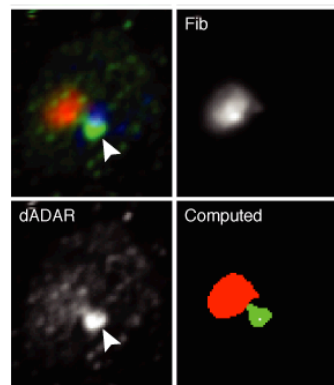
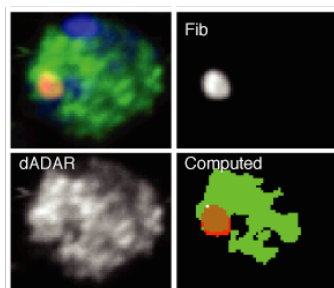
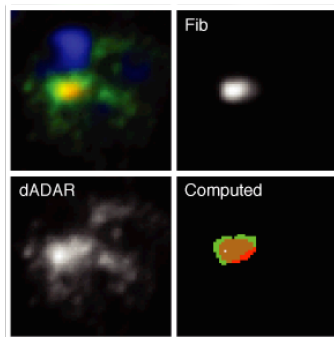
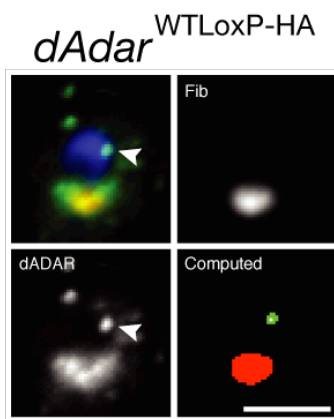
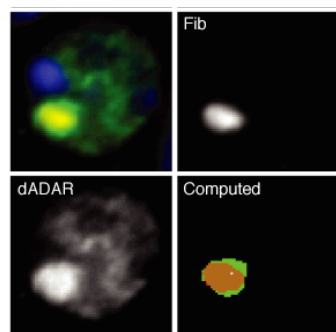
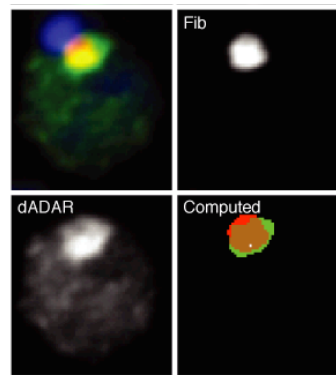
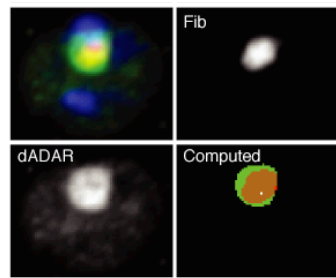
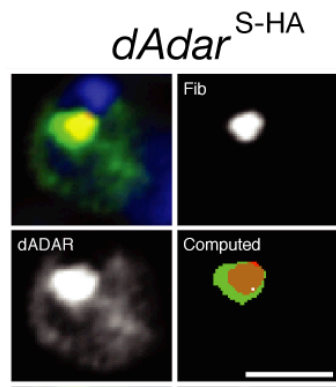
Supplementary Figure S2 | Effects of altering dADAR auto-editing are not biased to sites with high or low editing levels. (a) Six editing sites in the *Da6* acetylcholine receptor transcript show differential regulation in *dAdar*^S and *dAdar*^G when compared to *dAdar*^{WTLoxP}. One site in *Da6* shows no alteration, three sites show a mono-directional alteration, and two sites show a bi-directional alteration in response to abolishing or hard-wiring *dAdar* auto-editing. **(b)** Nine editing sites in the *Da5* acetylcholine receptor transcript show differential regulation in *dAdar*^S and *dAdar*^G when compared to *dAdar*^{WTLoxP}. Two sites in *Da5* show no alteration, five sites show a mono-directional, and two sites a bi-directional alteration upon changes in *dAdar* auto-editing. mRNAs were amplified from male head cDNA. Mean values for each site were defined as significantly different ($P < 0.05$) from *dAdar*^{WTLoxP} using one-way ANOVA with Dunnett post-hoc test ($n = 3-8$ PCRs per site). Error bars, s.e.m. **(c-d)** For all sites analyzed with editing levels between 10-90%, increased editing in *dAdar*^S males heads **(c)** and reduced editing in *dAdar*^G male heads **(d)** are not correlated with the endogenous level of editing for each site in *dAdar*^{WTLoxP} male heads.



Supplementary Figure S3 | dADAR localization in 3rd instar larval salivary glands and adult neurons. (a) Two separate transgenes representing the predominant adult-stage isoforms of dADAR, which lacks the alternatively spliced 3a exon²³ (termed ‘3/4’) were expressed in larval salivary glands using the *elav^{c155}*-Gal4 driver. At the auto-editing site, the serine codon was either wild-type (AGT) (‘S/G’) and is capable of being edited by the dADAR transgene, or was mutated to an un-editable synonymous TCT codon (‘S’). When expressed in the *dAdar* null adult nervous system, the wild-type dADAR transgene is robustly auto-edited²³. Both the 3/4 S/G (upper panels) and the 3/4 S transgene (lower panels) primarily co-localized with the nucleolus, labeled with an anti-fibrillar (FIB) antibody. Scale bar, 5 μ m. (b) Both the 3/4 S/G (upper panel) and the 3/4 S (lower panel) transgenes localize internally to the nuclear envelope (labeled with an anti-Lamin antibody) in 3rd instar larval salivary glands. Scale bars, 20 μ m. (c) In adult neuronal nuclei, WTLoxP-HA, S-HA and G-HA dADARs also localize to within the nuclear envelope. Arrows indicate the primary concentrations of nuclear dADAR. Note the presence of multiple punctae within the nucleus in neurons expressing G-HA dADAR, but not S-HA dADAR. Scale bar, 5 μ m.

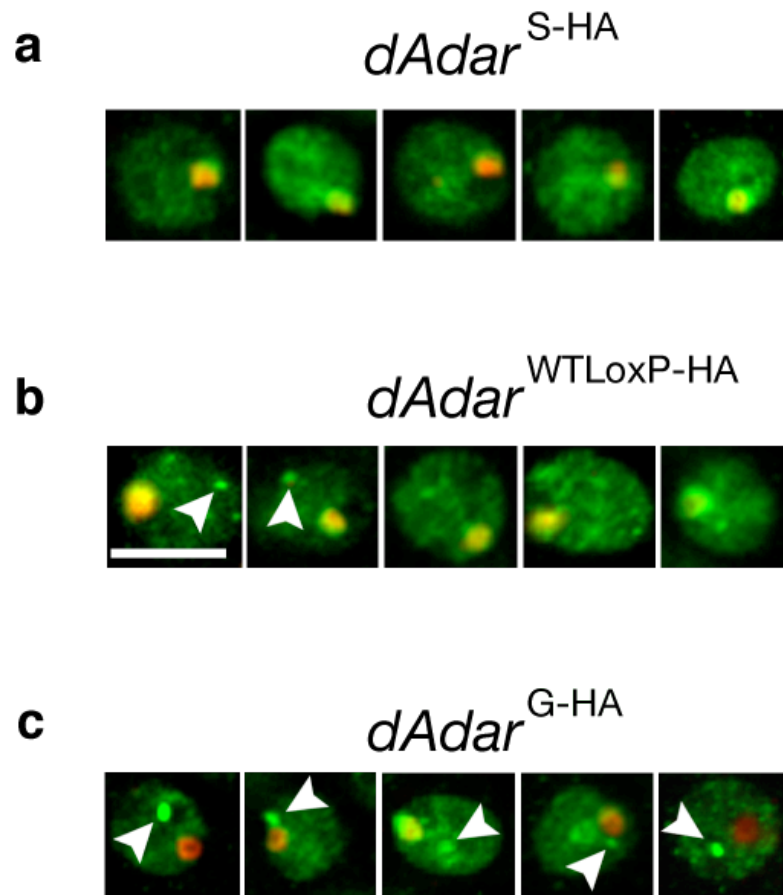


Supplementary Figure S4 | Computational analysis of dADAR localization in adult neurons. Automated categorization of extranucleolar punctate dADAR staining. Each category contains an example image, a model of its category, and the definition of the category. The tetrad of images show a merged image with DAPI (blue), dADAR (green), and fibrillar (red), the individual dADAR and fibrillar channels, and the computer's identification of the nucleolus and puncta. The computer's identification shows measures A (white), P (green), and N (red) as described in the methods. Next is a model of the computer's categorization of the cell followed by the definitions of that category. **(a)** Extranucleolar punctate dADAR staining. **(b)** Non-punctate dADAR staining. **(c)** Punctate non-extranucleolar dADAR staining. Scale bar, 5 μm .

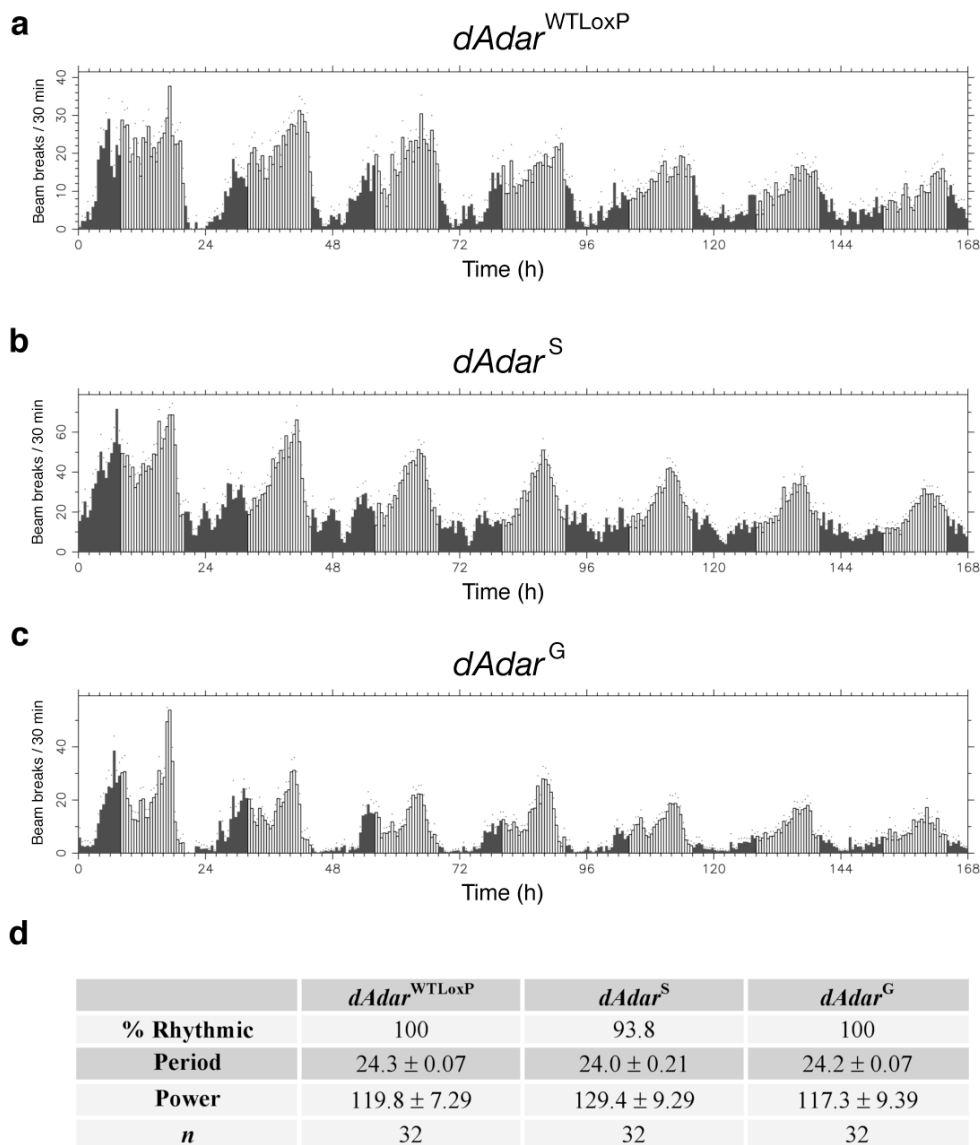


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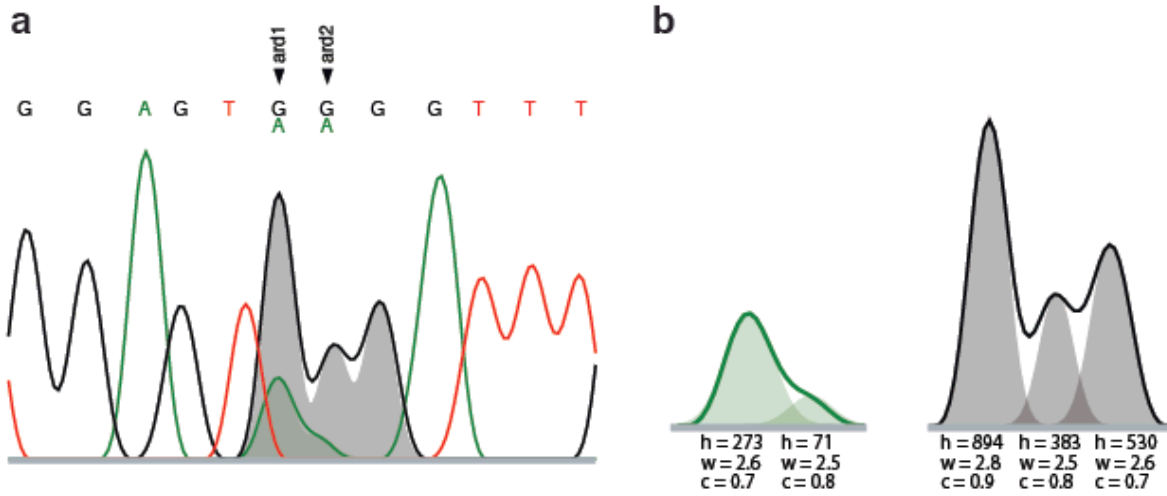
Supplementary Figure S5 | Examples of dADAR localization in *dAdar*^{S-HA}, *dAdar*^{WTLoxP-HA} and *dAdar*^{G-HA} adult neurons. For each example, confocal slices showing individual dADAR (green) and Fibrillarin (red) signals, alongside merged image with DAPI (blue), are shown, as well as the computed signals for dADAR and Fibrillarin. dADAR in *dAdar*^{S-HA} neurons is predominantly localized to the nucleolus and extranucleolar puncta are rarely observed, while in *dAdar*^{G-HA} neurons, strong extranucleolar puncta are frequently observed (arrowheads). Wild-type *dAdar*^{WTLoxP-HA} controls exhibit an intermediate phenotype, although dADAR is mainly non-punctal. Scale bars, 5 μ m.



Supplementary Figure S6 | Additional images of differences in dADAR localization following manipulation of dADAR auto-editing. In each case, 5 representative adult neuronal nuclei are shown. dADAR (green) was visualized using a mouse anti-HA antibody, rather than the rabbit anti-HA antibody used for all other confocal experiments in the adult brain. **(a)** Consistent with the data obtained using the rabbit anti-HA antibody, $dADAR^{S-HA}$ stained with mouse anti-HA shows a generally diffuse pattern of expression within the nucleus, and co-localizes with fibrillar (red). **(b)** This pattern was also predominantly observed in $dADAR^{WTLoxP-HA}$ neurons, although small puncta were sometimes observed. **(c)** In $dADAR^{G-HA}$ neurons, we additionally observed strong extranucleolar puncta (arrowheads), as was similarly seen using the rabbit anti-HA antibody. Scale bar, 5 μ m.



Supplementary Figure S7 | Altering *dAdar* auto-editing does not result in arrhythmic behavior in constant-dark (DD) conditions. (a-c) For each genotype indicated, activity plots were averaged over seven days in DD. Males bearing all *dAdar* alleles exhibited rhythmic locomotor patterns. Light bars - subjective day, dark bars – subjective night. Dots represent s.e.m. **(d)** Table indicating % rhythmicity, period and the power of rhythmicity for all genotypes. Values are presented as mean \pm s.e.m. *n*-values are also indicated.



Supplementary Figure S8 | Computational methods to determine editing levels. (a) Example electropherogram showing the edited adenosines ard1 and ard2 (sites 1 and 2 of the *ard* acetylcholine receptor), in which the best fit Gaussian curves for the A and G residues are shown as filled green and black, respectively. (b) A and G chromatograms on the left and right in which each best fit Gaussian curve and its parameters are shown.

Supplementary Table S1 | Nomenclature for the 23 transcripts analyzed in various *dAdar* allelic backgrounds. Sites (1, 2...) are labeled in a 5' to 3' order within the transcript.

Abbreviation	Locus	Synonyms/ full name	Molecular function
adr	<i>dAdar</i>	<i>Drosophila adenosine deaminase acting on RNA</i>	RNA editing enzyme
ard	<i>Ard</i>	nAcR β -64B	Nicotinic acetylcholine receptor α -subunit
cac	<i>cacophony</i>	Dm α 1A	Voltage-gated calcium channel α -subunit
cad	<i>Ca-alpha1D</i>	Dm α 1D	Voltage-gated calcium channel α -subunit
cap	<i>caps</i>	Calcium activated protein for secretion	Regulator of exocytosis
cat	<i>Ca-alpha1T</i>	Dm α 1T	Voltage-gated calcium channel α -subunit
cpx	<i>complexin</i>		Regulator of exocytosis
daf	<i>Dα5</i>	nAcR α -34E	Nicotinic acetylcholine receptor α -subunit
das	<i>Dα6</i>	nAcR α -30D	Nicotinic acetylcholine receptor α -subunit
dop	<i>DopEcR</i>		Dopamine/Ecdysteroid receptor
dsc	<i>dsc1</i>	NaCP60E	Voltage-gated sodium channel
eag	<i>Eag</i>	<i>Ether-a-go-go</i>	Voltage-gated potassium channel α -subunit
glu	<i>GluCla</i>		Glutamate-gated chloride channel
lap	<i>Lap</i>	like-AP180	Regulator of endocytosis
par	<i>para</i>	<i>paralytic</i>	Voltage-gated sodium channel α -subunit
rdl	<i>Rdl</i>	<i>resistance to dieldrin</i>	GABA receptor α -subunit
sbd	<i>Sbd</i>	nAcR β -96A	Nicotinic acetylcholine receptor β -subunit
sha	<i>shaker</i>		Voltage-gated potassium channel α -subunit
shb	<i>shab</i>		Voltage-gated potassium channel α -subunit
slo	<i>slowpoke</i>		Ca ²⁺ -activated potassium channel
stj	<i>straightjacket</i>	α 2 δ	Voltage-gated calcium channel
stn	<i>stoned-B</i>		Regulator of endocytosis
syt	<i>synaptotagmin-1</i>		Regulator of exocytosis
unc	<i>unc-13</i>		Regulator of exocytosis

Supplementary Table S2 | Primers used in this study

Gene	CG Number	Sites	Primer Type	Se quence (5' - 3')
syt	CG3139	1 to 6	Forward	GCTGCGCTACGTGCCGACCGCCGG
			Reverse Sequencing	GTAGTCCACGACGGTCACAACGAG GCTGCGCTACGTGCCGACCGCCGG
cap	CG33653	1	Forward	GATTTGAAAAGAACGATGGGAAAG
			Reverse Sequencing	CAGCATATTAATCTGCTGCGAATACC CGGGTGTGCCACTTCTGAAGATTTAT
cpx	CG32490	1 to 3	Forward	AGCTAAGCAGATGGTTGGAAA
			Reverse Sequencing	TGCATGACACATTTTCCCTCT CCCCAAGAAGAGCCCAAT
unc	CG2999	1	Forward	TGGACAGTTATCAGCATCTTCAA
			Reverse Sequencing	ATTCGTGGCTCCAAACTGAT GCTGTGGACATGAAGTACGC
stn	CG12473	1	Forward	TCAAGGGTATCGAGCGAATC
			Reverse Sequencing	GGCCAAGATGCCTTTGATAA TGCATACACCACACATCAGC
lap	CG2520	1 and 2	Forward	CGATGCGTTGGATCTTTACA
			Reverse Sequencing	GGACAGCCAAGTATGATGGG TTGTTAGATGCCTTGGAGCA
dsc	CG34405	1	Forward	GCAAGGAATGCGGATTGTAG
			Reverse Sequencing	GCGTTGCTCACTTCCAGAAT CGGATCGTTCTTCACACTGA
sbd	CG6798	1 and 2	Forward	GACCTACAATGGTGCCCAAG
			Reverse Sequencing	CACATCGATCTCGTTGGTGT CCAAGTGGATCTGAAGCAT
stj	CG12295	1 to 3	Forward	CGTCCGGAATTCACAATAC
			Reverse Sequencing	CCTCCTTGCCAATCAGGTAG CGACGAGTCCGAAGGATATT
cat	CG15899	1	Forward	GTTGCTGCGAATCCTCAAAT
			Reverse Sequencing	GTTGGTGGTCGAGGAGTCTG TGTGGCACTAATGACGTTCCG
cad	CG4894	1 to 5	Forward	GCATCGATTCTATGGGCATT

			Reverse Sequencing	CAGTGGACGTAGCACTCGAA TTGCCAACTGTATTGCCTTG
shb	CG1066	1 to 7	Forward	GAAGGTAAATGCGCCGAGTA
			Reverse Sequencing	GTCCGTTTGCGAGAGATTGT GGAAACGAATAAGAATGCAACG
eag	CG10952	1 to 5	Forward	CAATACAGCTGGCTGTGGAA
			Reverse Sequencing	TCACCCTTCTCGACATCACTT GACGGCCCTATATTTACCA
		6 to 7	Forward	CAATACAGCTGGCTGTGGAA
			Reverse Sequencing	TCACCCTTCTCGACATCACTT GCGACGAAATTTGGAGAAGA
slo	CG10693	1	Forward	CAGCATTGCATCCCTCATT
			Reverse Sequencing	TGGTTCCTTGGAAGAACTCC CGGTCTTCGATTTCTTCGAG
		2	Forward	CTACCACGAGCTGAAACACG
			Reverse Sequencing	CGGGTGGGTTGGTTATTACA GATGACCTTTGACGACACGA
rdl	CG10537	1 and 2	Forward	CATGCTGGGTGACGTAAACA
			Reverse Sequencing	CATACCGACGCCACATT CGGAGTCACCATGTATGTGC
		3 to 6	Forward	CATGCTGGGTGACGTAAACA
			Reverse Sequencing	CATACCGACGCCACATT TGCCCCAATTTAAGGTCTTG
daf	CG32975	1, 2 and 9	Forward	CACTGGGTGTTACCATCTTGC
			Reverse Sequencing	CTACGAGACAATAATATGTGGTG ACTGGGTGTTACCATCTTGC
		3 to 8	Forward	CACTGGGTGTTACCATCTTGC
			Reverse Sequencing	CTACGAGACAATAATATGTGGTG CGTGCATCAAATCATCAACT
das	CG4128	1 to 6	Forward	AATCTGCGCTGGAATGAAAC
			Reverse Sequencing	CAATGTGAAGCCCAGTAGGG TGGAATGAAACGGAATACGG
ard	CG11348	1 to	Forward	GACCTACAATGGTGCCCAAG

		4	Reverse Sequencing	CACATCGATCTCGTTGGTGT CCCAAGTGGATCTGAAGCAT
sha	CG12348	1 and 2	Forward	CTGGTCATGGCTTTGGTGGCGGACC
			Reverse Sequencing	CCGTGTGATCAGTCAGACCTGGCG GGATCTGTGATGTCAGGCACCTCG
		3 to 6	Forward	CTGGTCATGGCTTTGGTGGCGGACC
			Reverse Sequencing	CCGTGTGATCAGTCAGACCTGGCG CGAGGTGCCTGACATCACAGATCC
adr	CG12598	1	Forward	CCACAGCATATCAGTCGATTT
			Reverse Sequencing	TGGAATCGTCCCCTCACCGGAC CCACAGCATATCAGTCGATTT
cac	CG1522	1	Forward	GCGGAGAAAAGGTTTCGTTT
			Reverse Sequencing	GAGTGTTGCAAGACCTGTGG TCAACAGTGCTATCGGGAAA
		2 to 5	Forward	CCACAGCATATCAGTCGATTT
			Reverse Sequencing	TGGAATCGTCCCCTCACCGGAC ATGCATTTACCGGCGTATTC
		6	Forward	GTACGAGGAGGAGGACGAACTGC
			Reverse Sequencing	CTGAAGTCTAGCGGGACTCG GTACGAGGAGGAGGACGAACTGC
		11	Forward	GTACGAGGAGGAGGACGAACTGC
			Reverse Sequencing	CTGAAGTCTAGCGGGACTCG CACTGGCCTACGCCTACTTC
		12	Forward	GTACGAGGAGGAGGACGAACTGC
			Reverse Sequencing	CTGAAGTCTAGCGGGACTCG CTCCAGTGGCCAGATCTCC
par	CG9907	1 to 4	Forward	CATTGGTGCAAATCGAACAA
			Reverse Sequencing	GCTCCGAATGGACATCTTCT CATTATTCATGCACACGACGA
glu	CG7535	1 to 2	Forward	GGCAGCGGACACTATTTCTG
			Reverse Sequencing	GCATCTAAACTGGCCTGCTC CTGACTATGGCGGGACCA
		3	Forward	GGCAGCGGACACTATTTCTG
			Reverse Sequencing	GCATCTAAACTGGCCTGCTC CCTACCTCGCTTCACACTGG

dop	CG18314	$\frac{1}{9}$ to	Forward	GGTGCCCTTCTCCGTGTAT
			Reverse	TTGAGGAGAATGAAAATCTAAATCTAA
			Sequencing	GACCGGAGAATGGATGTACG