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Reporting Summary

Nature Research wishes to improve the reproducibility of the work that we publish. This form provides structure for consistency and transparency in reporting. For further information on Nature Research policies, see Authors & Referees and the Editorial Policy Checklist.

Statistical parameters

		Methods section).			
n/a	Cor	Confirmed			
	\boxtimes	The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement			
	\boxtimes	An indication of whether measurements were taken from distinct samples or whether the same sample was measured repeatedly			
	\boxtimes	The statistical test(s) used AND whether they are one- or two-sided Only common tests should be described solely by name; describe more complex techniques in the Methods section.			
X		A description of all covariates tested			
\times		A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons			
	\boxtimes	A full description of the statistics including <u>central tendency</u> (e.g. means) or other basic estimates (e.g. regression coefficient) AND <u>variation</u> (e.g. standard deviation) or associated <u>estimates of uncertainty</u> (e.g. confidence intervals)			
	\boxtimes	For null hypothesis testing, the test statistic (e.g. <i>F</i> , <i>t</i> , <i>r</i>) with confidence intervals, effect sizes, degrees of freedom and <i>P</i> value noted <i>Give P values as exact values whenever suitable.</i>			
\times		For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings			
\times		For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes			
\times		Estimates of effect sizes (e.g. Cohen's d, Pearson's r), indicating how they were calculated			
	\boxtimes	Clearly defined error bars State explicitly what error bars represent (e.g. SD, SE, CI)			
. Our web collection on statistics for biologists may be useful,					

Software and code

Policy information about availability of computer code

Data collection None used Graphpad Prism 7, Crispresso2 Data analysis

For manuscripts utilizing custom algorithms or software that are central to the research but not yet described in published literature, software must be made available to editors/reviewers upon request. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Research guidelines for submitting code & software for further information.

Data

Policy information about availability of data

All manuscripts must include a data availability statement. This statement should provide the following information, where applicable:

- Accession codes, unique identifiers, or web links for publicly available datasets
- A list of figures that have associated raw data
- A description of any restrictions on data availability

All editor plasmids have been submitted to Addgene for public release; HTS data for base editing and indel quantification will be available from the NCBI sequence read archive (PRJNA498804)

Field-spe	cific re	porting		
Please select the be	est fit for your r	research. If you are not sure, read the appropriate sections before making your selection.		
∑ Life sciences ☐ Behavioural & social sciences ☐ Ecological, evolutionary & environmental sciences				
For a reference copy of t	he document with	all sections, see <u>nature.com/authors/policies/ReportingSummary-flat.pdf</u>		
Life scier	nces stu	udy design		
All studies must disclose on these points even when the disclosure is negative.				
Sample size	All samples were evaluated as biological triplicates			
Data exclusions	No data was excluded			
Replication	Biological triplicate experiments were done with fresh aliquots of cells at intervals ranging from weeks to months between experiments and carried out by up to three different researchers. All findings have been replicated.			
Randomization	Not relevant to	these experiments		
Blinding	Not relevant to these experiments			
Materials & experimental systems Methods				
Unique biolo	ogical mat	erials		
Policy information about <u>availability of materials</u>				
Obtaining unique materials All materials are readily available.				
Eukaryotic c	ell lines			
Policy information about <u>cell lines</u>				
Cell line source(s	ell line source(s) ATCC HEK293T (CRL-3216)			
Authentication		Cells were authenticated by the supplier		

No mycoplasma was detected in the HEK293T cells

No commonly misidentified lines were used

Mycoplasma contamination

Commonly misidentified lines (See <u>ICLAC</u> register)