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Reporting Summary

Nature Portfolio 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 Portfolio policies, see our <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

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For	all statistical and	alyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.		
n/a	Confirmed			
	The exact :	sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement		
	A stateme	nt on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly		
	The statist Only commo	ical test(s) used AND whether they are one- or two-sided on tests should be described solely by name; describe more complex techniques in the Methods section.		
\boxtimes	A descripti	on of all covariates tested		
	🔀 A descripti	on of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons		
\boxtimes	A full desc AND variat	A full description of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) AND variation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)		
	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>			
\boxtimes	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings			
\boxtimes	For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes			
\boxtimes	Estimates of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated			
Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.				
Software and code				
Policy information about <u>availability of computer code</u>				
Da	ata collection	StereoInvestigator software from Microbrightfield LLC was used for cell counts.		
Da	ata analysis	GraphPad Prism software was used for statistical anlayses.		

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 and reviewers. We strongly encourage code deposition in a community repository (e.g. GitHub). See the Nature Portfolio guidelines for submitting code & software for further information.

Data

Policy information about <u>availability of data</u>

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

- Accession codes, unique identifiers, or web links for publicly available datasets
- A description of any restrictions on data availability
- For clinical datasets or third party data, please ensure that the statement adheres to our policy

All data needed to evaluate the results of this article are present in the paper.

Field-spe	ecific reporting		
	ne below that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.		
Life sciences	Behavioural & social sciences		
	the document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>		
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Life scier	nces study design		
All studies must dis	close on these points even when the disclosure is negative.		
Sample size	The sample size was determined based on our previous studies for similar experiments. The sample sizes chosen in the study were sufficient based on the demonstrated statistical significance between groups.		
Data exclusions	No data were excluded		
Replication	he reproducibility of the experimental design is apparent from our findings on seizures. In the study, grafting of human interneuron rogenitors into the brain reduced seizures in the temporal lobe epilepsy model but blocking the activity of graft-derived interneurons using a esigner drug resulted in increased seizures. Furthermore, following the washout of the designer drug, the seizures were reduced again. Also, ne ability of human interneuron progenitors to reduce seizures in the temporal lobe epilepsy model is consistent with our previous study in the status epilepticus model (Upadhya et al., PNAS, 2019).		
Randomization	Chronically epileptic rats having a similar range of seizures were randomly assigned to different groups; one group received grafts expressing designer receptors and designer drug treatment, the second group received no grafts, and the third group received only the designer drug.		
Blinding	Investigators who quantified EEG traces were blinded to group allocations.		
We require informati system or method list Materials & ex n/a Involved in th Antibodies Eukaryotic Palaeontol Animals an Human res Clinical dat Dual use re	cell lines cell lines mathematicipants ChIP-seq Flow cytometry MRI-based neuroimaging dother organisms search participants		
Antibodies			
Antibodies used	Mouse anti-human nuclear antigen (Millipore, MAB1281), mouse anti-NeuN (Millipore, MAB377), rabbit anti-GABA (Sigma, A2052), mouse anti-Parvalbumin (Sigma, P3088), Rabbit anti-Neuropeptide Y (Peninsula Laboratories, T-4070). Goat anti-PSD95 (Abcam, ab12093); Mouse anti-human Synaptophysin (ThermoFisher, 14-6525-80), Rabbit anti-MAP2 (Millipore, AB5622).		
Validation	All antibodies were validated using positive control and negative control brain tissue sections before employing them in the study.		
Animals and	other organisms		
Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research			
Laboratory anima	mals Male Fischer 344 rats		
Wild animals	N/A		

All experiments were performed as described in the animal protocol, approved by the institutional animal care and use committee of

the Texas A&M Health Sciences Center and Central Texas Veterans Health Care System.

N/A

Field-collected samples

Ethics oversight

Note that full information on the approval of the study protocol must also be provided in the manuscript.