# nature portfolio

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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 Editorial Policies and the Editorial Policy Checklist.

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For	all statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.
n/a	Confirmed
	$oxed{oxed}$ The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
	🔀 A statement on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly
	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
	🔀 A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
	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>
$\times$	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings
	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.
So	ftware and code

Policy information about availability of computer code

PharmaEngine EDC software. Data collection

Data analysis All statistical analyses will be performed using SAS (SAS Institute, NC, USA) or R (R Foundation for Statistical Computing, Vienna, Austria).

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 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 description of any restrictions on data availability
- For clinical datasets or third party data, please ensure that the statement adheres to our policy

All requests for raw and analyzed data and materials are promptly reviewed by the Cedars-Sinai Board of Governors Institute of Regenerative Medicine to verify if the request is subject to any intellectual property or confidentiality obligations. Patient-related data not included in the paper were generated as part of clinical trials and may be subject to patient confidentiality. Any data and materials that can be shared will be released via a Material Transfer Agreement. All raw and analyzed sequencing data can be found at the NCBI Sequence Read Archive (accession number: pending).

Field-spe	ecific reporting
Please select the o	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 Ecological, evolutionary & environmental sciences
For a reference copy of t	the document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>
Life scier	nces study design
All studies must dis	sclose on these points even when the disclosure is negative.
Sample size	Detailed in methods section
Data exclusions	Detailed in methods section
Replication	Detailed in methods section
Randomization	Detailed in methods section
Blinding	Detailed in methods section
We require informati	g for specific materials, systems and methods  on from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each material, ted is relevant to your study. If you are not sure if a list item applies to your research, read the appropriate section before selecting a response.
	perimental systems  Methods
n/a Involved in th	· · · · · · · · · · · · · · · · · · ·
Antibodies	
Eukaryotic Palaeontol	cell lines Flow cytometry  logy and archaeology MRI-based neuroimaging
	nd other organisms
	search participants
Clinical dat	ta esearch of concern
Antibodies	
Antibodies used	Human nuclei (Stem101) 1:200 ab-101-u-050 Takara Bio Human cytoplasm (Stem121) 1:2000 ab-121-u-050 Takara Bio
	Human-specific Nestin 1:15,000 ABD69 EMD Millipore GFAP 1:500 Z0334 Dako
	ChAT 1:200 AB144P-1ML EMD Millipore GDNF 1:250 BAF212 R&D Systems
	AQP4 1:100 HPA014784 Sigma Glast 1:20 AF6048 R&D Systems
	Human-specific GDNF (biotinylated) 1:50 BAF212 R&D Systems Human-specific ChAT 1:200 AF3447 R&D Systems
	Iba-1 1:250 NB100-1028 Novus Bio Human-specific Nestin 1:2000 ABD69 EMD Millipore
	GFAP 1:500 Z0334 Dako
	Ki-67 1:200 RM-9106-S Thermo Scientific CD34 1:500 PAB18289 Abnova Collegen NV 1:150 CO 4:01 106 0:1 Registered
	Collagen IV 1:150 600-401-106-0.1 Rockland S100B 1:100 HPA015768 Sigma-Aldrich
	Human-specific NF-H 1:4000 AF3108 R&D Systems

Antibodies were validated and optimized based on the manufacturer's recommendations and using known positive controls.

Validation

#### Eukaryotic cell lines

Policy information about cell lines

Cell line source(s) Tissue Donation obtained by Dr. Guido Nikkah

Authentication HLA typing was performed for the line at multiple timepoints during manufacturing for authentication.

Cell lines are negative for mycoplasma and all adventitious testing Mycoplasma contamination

Commonly misidentified lines (See ICLAC register)

N/A

#### Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research

Laboratory animals Male and female SOD1G93A transgenic rats (NTac:SDTg (SOD1G93A) L26H) and wild-type littermates were acquired from Taconics (Hudson, NY, USA) and maintained as a colony by in-house breeding with Sprague-Dawley females (Taconics), Age ~100 days at study

Female and Male Yucatan Mini Pigs, 14-20Kg at study start.

N/A Wild animals

N/A Field-collected samples

Ethics oversight Detailed in methods section

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

#### Human research participants

Policy information about studies involving human research participants

Population characteristics Main table 1

Detailed in methods section Recruitment

Cedars-Sinai Office of Research Compliance and Quality Improvement - Study IRB# Pro00042350 Ethics oversight

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

#### Clinical data

Policy information about clinical studies

All manuscripts should comply with the ICMJE guidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.

Clinical trial registration NCT02943850

Requests can be made to, and promptly reviewed by, the Cedars-Sinai Board of Governors Institute of Regenerative Medicine Study protocol

Data collection Detailed in methods section and/or results section

Detailed in methods section and/or results section Outcomes

### Magnetic resonance imaging

#### Experimental design

Design type Longitudinal

Participants were imaged at different timepoints for safety evaluations Design specifications

Not Used Behavioral performance measures

Acquisition			
Imaging type(s)	Structural		
Field strength	3 Tesla		
Sequence & imaging parameters	Preoperative thoracolumbar MR in a Siemens Skyra 3T Magnet. The unenhanced spine MR included the following sequences: sagittal T1 turbo spin-echo(tse) 2mm, sagittal T2 tse 2mm, sagittal T2 space 1mm, coronal T2 space 1mm, axial T2 tse. For postoperative imaging, contrast enhanced MR with sagittal and axial T1 sequences following intravenous Gadovist (gadobutrol, 1.0mmol/mL, a nonionic macrocyclic agent; Bayer Shering Pharma) were added to the unenhanced protocol.		
Area of acquisition	Thoracolumbar		
Diffusion MRI Used	Not used		
Preprocessing			
Preprocessing software	Not used		
Normalization	Not used		
Normalization template	Not used		
Noise and artifact removal	Not used		
Volume censoring	Not used		
Statistical modeling & infere	nce		
Model type and settings	Not used		
Effect(s) tested	Not used		
Specify type of analysis: W	hole brain ROI-based Both		
Statistic type for inference (See <u>Eklund et al. 2016</u> )	Not used		
Correction	Not used		
Models & analysis			
n/a Involved in the study  Functional and/or effective	connectivity		

n/a	Involved in the study
$\boxtimes$	Functional and/or effective connectivity
$\boxtimes$	Graph analysis
$\boxtimes$	Multivariate modeling or predictive analysi