Reporting Checklist for Nature Neuroscience

This checklist is used to ensure good reporting standards and to improve the reproducibility of published results. For more information, please read Reporting Life Sciences Research.

Please note that in the event of publication, it is mandatory that authors include all relevant methodological and statistical information in the manuscript.

Statistics reporting, by figure

- Please specify the following information for each panel reporting quantitative data, and where each item is reported (section, e.g. Results, & paragraph number).
- Each figure legend should ideally contain an exact sample size (n) for each experimental group/condition, where n is an exact number and not a range, a clear definition of how n is defined (for example x cells from x slices from x animals from x litters, collected over x days), a description of the statistical test used, the results of the tests, any descriptive statistics and clearly defined error bars if applicable.
- For any experiments using custom statistics, please indicate the test used and stats obtained for each experiment.
- Each figure legend should include a statement of how many times the experiment shown was replicated in the lab; the details of sample collection should be sufficiently clear so that the replicability of the experiment is obvious to the reader.
- For experiments reported in the text but not in the figures, please use the paragraph number instead of the figure number.

Note: Mean and standard deviation are not appropriate on small samples, and plotting independent data points is usually more informative. When technical replicates are reported, error and significance measures reflect the experimental variability and not the variability of the biological process; it is misleading not to state this clearly.

<table>
<thead>
<tr>
<th>TEST USED</th>
<th>n</th>
<th>DESCRIPTIVE STATS (AVERAGE, VARIANCE)</th>
<th>P VALUE</th>
<th>DEGREES OF FREEDOM &amp; F/T/Z/R/ETC VALUE</th>
</tr>
</thead>
<tbody>
<tr>
<td>Fig. 1a</td>
<td>one-way ANOVA</td>
<td>9, 9, 10, 15, mice from at least 3 litters/group</td>
<td>Methods para 8</td>
<td>error bars are mean +/- SEM</td>
</tr>
<tr>
<td>Results para 6</td>
<td>unpaired t-test</td>
<td>15, slices from 10 mice</td>
<td>Results para 6</td>
<td>error bars are mean +/- SEM</td>
</tr>
<tr>
<td>Supp Fig 1c</td>
<td>Kolmogorov-Smirnov test</td>
<td>p. 2, 510 (WT) vs. 548 (PDAPP)</td>
<td>layer 2/3 neurons</td>
<td>fig. legend</td>
</tr>
<tr>
<td>FIGURE NUMBER</td>
<td>WHICH TEST?</td>
<td>SECTION &amp; PARAGRAPH</td>
<td>EXACT VALUE</td>
<td>n</td>
</tr>
<tr>
<td>---------------</td>
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</tr>
<tr>
<td>Supp Fig 1e</td>
<td>2-sample t-test</td>
<td>p. 3</td>
<td>4 WT vs. 4 PDAPP mice</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Supp Fig 2e</td>
<td>2-sample t-test</td>
<td>p. 3</td>
<td>4 WT vs. 4 PDAPP mice</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Fig 1a</td>
<td>2-sample t-test</td>
<td>p. 3</td>
<td>10 controls vs. 9 treated mice</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Supp Fig 2a</td>
<td>2-sample t-test</td>
<td>p. 3</td>
<td>8 controls vs. 8 treated mice</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Fig 1b</td>
<td>2-sample t-test</td>
<td>p. 3</td>
<td>9 controls vs. 7 treated mice</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Supp Fig 2b</td>
<td>2-sample t-test</td>
<td>p. 3</td>
<td>7 controls vs. 7 treated mice</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Fig 1c</td>
<td>2-sample t-test</td>
<td>p. 3</td>
<td>9 controls vs. 7 treated mice</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Supp Fig 2c</td>
<td>2-sample t-test</td>
<td>p. 3</td>
<td>7 controls vs. 7 treated mice</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Fig 1h</td>
<td>Kolmogorov-Smirnov test</td>
<td>p. 4</td>
<td>1414 controls vs. 1393 treated layer 2/3 neurons</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Fig 1i</td>
<td>2-sample t-test</td>
<td>p. 4</td>
<td>13 controls vs. 10 treated mice</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Fig 1j</td>
<td>2-sample t-test</td>
<td>p. 4</td>
<td>13 controls vs. 10 treated mice</td>
<td>fig. legend</td>
</tr>
<tr>
<td>Fig 1k</td>
<td>2-sample t-test</td>
<td>p. 4</td>
<td>3 controls vs. 5 treated mice</td>
<td>fig. legend</td>
</tr>
</tbody>
</table>
### Representative figures

1. Are any representative images shown (including Western blots and immunohistochemistry/staining) in the paper?
   
   *If so, what figure(s)?*

2. For each representative image, is there a clear statement of how many times this experiment was successfully repeated and a discussion of any limitations in repeatability?
   
   *If so, where is this reported (section, paragraph #)?*

### Statistics and general methods

1. Is there a justification of the sample size?
   
   *If so, how was it justified?*

   Where (section, paragraph #)?

   Even if no sample size calculation was performed, authors should report why the sample size is adequate to measure their effect size.

   *yes,*  
   
   the major aim of the study was to obtain high quality imaging recordings from a commonly used and statistically sufficient sample size. The sample size was chosen on the basis of 1) pilot experiments, 2.) previous experiences with similar types of experiments, and 3.) are similar and, in some cases, even larger to those reported in previous publications (see e.g. refs. 6-8, 27,28).
2. Are statistical tests justified as appropriate for every figure?
   Where (section, paragraph #)?
   yes, we used the statistical tests only under standard conditions and for standard purposes.

   a. If there is a section summarizing the statistical methods in the methods, is the statistical test for each experiment clearly defined?

   b. Do the data meet the assumptions of the specific statistical test you chose (e.g. normality for a parametric test)?
       Where is this described (section, paragraph #)?
       All data meet the assumptions of the chosen statistical tests (e.g. normality and equal variances).

   c. Is there any estimate of variance within each group of data? Is the variance similar between groups that are being statistically compared?
       Where is this described (section, paragraph #)?
       yes, summary plots contain error bars that illustrate SEM.

   d. Are tests specified as one- or two-sided?

   e. Are there adjustments for multiple comparisons?
       t-tests were two-sided
       not applicable

3. Are criteria for excluding data points reported?
   Was this criterion established prior to data collection?
   Where is this described (section, paragraph #)?
   No data were excluded from the analysis.

4. Define the method of randomization used to assign subjects (or samples) to the experimental groups and to collect and process data. If no randomization was used, state so.
   Where does this appear (section, paragraph #)?
   Mice were randomly assigned to receive either anti-Aβ or isotype-matched control antibody.

5. Is a statement of the extent to which investigator knew the group allocation during the experiment and in assessing outcome included? If no blinding was done, state so.
   Where (section, paragraph #)?
   Investigators who performed and analyzed the experiments did not know which mice received anti-Aβ antibody or isotype-matched control antibody. Additionally, all data analysis was performed using semi-automated standard methods, developed prior to data acquisition.

6. For experiments in live vertebrates, is a statement of compliance with ethical guidelines/regulations included?
   Where (section, paragraph #)?
   yes, Online methods section, first paragraph

7. Is the species of the animals used reported?
   Where (section, paragraph #)?
   yes, see Main text and Online methods section, "Immunization design"

8. Is the strain of the animals (including background strains of KO/transgenic animals used) reported?
   Where (section, paragraph #)?
   yes, see Main text and Online methods section, "Immunization design"
9. Is the sex of the animals/subjects used reported?
   Where (section, paragraph #)?
   yes, Online methods section, ”Immunization design”

10. Is the age of the animals/subjects reported?
    Where (section, paragraph #)?
    yes, see Main text and Online methods section, ”Immunization design”

11. For animals housed in a vivarium, is the light/dark cycle reported?
    Where (section, paragraph #)?
    Mice were reared in 12/12 h light/dark cycles.

12. For animals housed in a vivarium, is the housing group (i.e. number of animals per cage) reported?
    Where (section, paragraph #)?
    Mice were housed in groups of 2 - 4 mice per cage.

13. For behavioral experiments, is the time of day reported (e.g. light or dark cycle)?
    Where (section, paragraph #)?
    not applicable

14. Is the previous history of the animals/subjects (e.g. prior drug administration, surgery, behavioral testing) reported?
    Where (section, paragraph #)?
    no previous history

   a. If multiple behavioral tests were conducted in the same group of animals, is this reported?
      Where (section, paragraph #)?
      not applicable

15. If any animals/subjects were excluded from analysis, is this reported?
    Where (section, paragraph #)?
    No animals were excluded from the analysis unless imaging data could not be obtained from them (e.g. bad quality of neuronal staining)

   a. How were the criteria for exclusion defined?
      Where is this described (section, paragraph #)?
      not applicable

   b. Specify reasons for any discrepancy between the number of animals at the beginning and end of the study.
      Where is this described (section, paragraph #)?
      not applicable

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Reagents

1. Have antibodies been validated for use in the system under study (assay and species)?
   yes, we used antibodies that are widely used and have been well-validated for the applications used here.

   a. Is antibody catalog number given?
      Where does this appear (section, paragraph #)?
      We received 3D6 antibodies from Janssen Alzheimer Immunotherapy (see Acknowledgments) and β1 antibodies from Novartis
b. Where were the validation data reported (citation, supplementary information, Antibodypedia)?

Where does this appear (section, paragraph #)?

We choose the antibodies according to previous publications, citations are provided in the text.

2. If cell lines were used to reflect the properties of a particular tissue or disease state, is their source identified?

Where (section, paragraph #)?

a. Were they recently authenticated?

Where is this information reported (section, paragraph #)?

not applicable

Data deposition

Data deposition in a public repository is mandatory for:

a. Protein, DNA and RNA sequences
b. Macromolecular structures
c. Crystallographic data for small molecules
d. Microarray data

Deposition is strongly recommended for many other datasets for which structured public repositories exist; more details on our data policy are available here. We encourage the provision of other source data in supplementary information or in unstructured repositories such as Figshare and Dryad.

We encourage publication of Data Descriptors (see Scientific Data) to maximize data reuse.

1. Are accession codes for deposit dates provided?

Where (section, paragraph #)?

not applicable

Computer code/software

Any custom algorithm/software that is central to the methods must be supplied by the authors in a usable and readable form for readers at the time of publication. However, referees may ask for this information at any time during the review process.

1. Identify all custom software or scripts that were required to conduct the study and where in the procedures each was used.

We used custom-written software based on LabView for imaging and custom-made routines of the Igor Pro software for analysis of the data. Similar routines are used by many other labs in our field.

2. If computer code was used to generate results that are central to the paper's conclusions, include a statement in the Methods section under "Code availability" to indicate whether and how the code can be accessed. Include version information as necessary and any restrictions on availability.

no, but we would be happy to share. researchers can directly contact the corresponding authors.

Human subjects

1. Which IRB approved the protocol?

Where is this stated (section, paragraph #)?

not applicable
2. Is demographic information on all subjects provided?
   Where (section, paragraph #)? not applicable

3. Is the number of human subjects, their age and sex clearly defined?
   Where (section, paragraph #)? not applicable

4. Are the inclusion and exclusion criteria (if any) clearly specified?
   Where (section, paragraph #)? not applicable

5. How well were the groups matched?
   Where is this information described (section, paragraph #)? not applicable

6. Is a statement included confirming that informed consent was obtained from all subjects?
   Where (section, paragraph #)? not applicable

7. For publication of patient photos, is a statement included confirming that consent to publish was obtained?
   Where (section, paragraph #)? not applicable

fMRI studies

For papers reporting functional imaging (fMRI) results please ensure that these minimal reporting guidelines are met and that all this information is clearly provided in the methods:

1. Were any subjects scanned but then rejected for the analysis after the data was collected?
   not applicable

   a. If yes, is the number rejected and reasons for rejection described?
      Where (section, paragraph #)?

2. Is the number of blocks, trials or experimental units per session and/or subjects specified?
   Where (section, paragraph #)? not applicable

3. Is the length of each trial and interval between trials specified?
   not applicable

4. Is a blocked, event-related, or mixed design being used? If applicable, please specify the block length or how the event-related or mixed design was optimized.
   not applicable

5. Is the task design clearly described?
   Where (section, paragraph #)? not applicable
<table>
<thead>
<tr>
<th>Question</th>
<th>Answer</th>
</tr>
</thead>
<tbody>
<tr>
<td>6. How was behavioral performance measured?</td>
<td>not applicable</td>
</tr>
<tr>
<td>7. Is an ANOVA or factorial design being used?</td>
<td>not applicable</td>
</tr>
<tr>
<td>8. For data acquisition, is a whole brain scan used?</td>
<td>not applicable</td>
</tr>
<tr>
<td>If not, state area of acquisition.</td>
<td></td>
</tr>
<tr>
<td>a. How was this region determined?</td>
<td></td>
</tr>
<tr>
<td>9. Is the field strength (in Tesla) of the MRI system stated?</td>
<td>not applicable</td>
</tr>
<tr>
<td>a. Is the pulse sequence type (gradient/spin echo, EPI/spiral) stated?</td>
<td></td>
</tr>
<tr>
<td>b. Are the field-of-view, matrix size, slice thickness, and TE/TR/flip angle clearly stated?</td>
<td></td>
</tr>
<tr>
<td>10. Are the software and specific parameters (model/functions, smoothing kernel size if applicable, etc.) used for data processing and pre-processing clearly stated?</td>
<td>not applicable</td>
</tr>
<tr>
<td>11. Is the coordinate space for the anatomical/functional imaging data clearly defined as subject/native space or standardized stereotaxic space, e.g., original Talairach, MNI305, ICBM152, etc? Where (section, paragraph #)?</td>
<td>not applicable</td>
</tr>
<tr>
<td>12. If there was data normalization/standardization to a specific space template, are the type of transformation (linear vs. nonlinear) used and image types being transformed clearly described? Where (section, paragraph #)?</td>
<td>not applicable</td>
</tr>
<tr>
<td>13. How were anatomical locations determined, e.g., via an automated labeling algorithm (AAL), standardized coordinate database (Talairach daemon), probabilistic atlases, etc.?</td>
<td>not applicable</td>
</tr>
<tr>
<td>14. Were any additional regressors (behavioral covariates, motion etc) used?</td>
<td>not applicable</td>
</tr>
<tr>
<td>15. Is the contrast construction clearly defined?</td>
<td>not applicable</td>
</tr>
<tr>
<td>16. Is a mixed/random effects or fixed inference used?</td>
<td>not applicable</td>
</tr>
<tr>
<td>a. If fixed effects inference used, is this justified?</td>
<td></td>
</tr>
<tr>
<td>17. Were repeated measures used (multiple measurements per subject)?</td>
<td>not applicable</td>
</tr>
<tr>
<td>a. If so, are the method to account for within subject correlation and the assumptions made about variance clearly stated?</td>
<td></td>
</tr>
</tbody>
</table>
18. If the threshold used for inference and visualization in figures varies, is this clearly stated?  
   not applicable

19. Are statistical inferences corrected for multiple comparisons?
   a. If not, is this labeled as uncorrected?  
      not applicable

20. Are the results based on an ROI (region of interest) analysis?
   a. If so, is the rationale clearly described?
   b. How were the ROI’s defined (functional vs anatomical localization)?  
      not applicable

21. Is there correction for multiple comparisons within each voxel?  
    not applicable

22. For cluster-wise significance, is the cluster-defining threshold and the corrected significance level defined?  
    not applicable

### Additional comments

Additional Comments

none