

Corresponding author(s): Marcus Fändrich and Matthias Schmidt

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 <u>Authors & Referees</u> and the <u>Editorial Policy Checklist</u>.

When statistical analyses are reported, confirm that the following items are present in the relevant location (e.g. figure legend, table legend, main

Statistical parameters

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	
	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.	
\boxtimes	A description of all covariates tested	
\boxtimes	A description of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons	
	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>	
X	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	\square 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 <u>statistics for biologists</u> may be useful.

Software and code

Data collection

Policy information about <u>availability of computer code</u>

Data analysis RELION v2.1, MotionCor2 v1, Gctf v1.06, IMOD v.4.9.4, COOT v0.8.9, PHENIX v1.12 and v1.14, Fiji (ImageJ v.1.51n)

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 <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 list of figures that have associated raw data
- A description of any restrictions on data availability

The reconstructed cryo-EM maps were deposited in the Electron Microscopy Data Bank with the accession codes EMD-8910 (murine) and EMD-9232 (human). The coordinates of the fitted atomic models were deposited in the Protein Data Bank under the accession codes 6DSO (murine) and 6MST (human).

Field-specific reporting			
Please select 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/authors/policies/ReportingSummary-flat.pdf</u>		
Life sciences study design			
All studies must disclose on these points even when the disclosure is negative.			
Sample size	Murine: Fibrils were extracted from a single NMRI mouse. Human: Fibrils were extracted from a renal tissue sample of a single patient.		
Data exclusions	No data were excluded from the analyses.		
Replication	Electron microscopy data is based on a single mouse or single human patient, respectively. SDS-PAGE and mass spectrometry data were replicated successfully in at least four independent experiments. Representative data is shown.		
Randomization	Not relevant to study. Single case Study.		
Blinding	Not relevant to study. Single case Study.		
Materials & experimental systems Methods n/a Involved in the study n/a Involved in the study ☑ Unique biological materials ☑ ChIP-seq ☑ Antibodies ☑ Flow cytometry ☑ Eukaryotic cell lines ☑ MRI-based neuroimaging ☑ Palaeontology ☑ Animals and other organisms ☑ Human research participants			
Unique biological materials			
Policy information about <u>availability of materials</u> Obtaining unique materials All unique materials used are readily available from the authors upon reasonable request or from standard commercial sources.			
Obtaining unique	Sources are presented in the methods section.		
Animals and	other organisms		
Policy information	Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research		
Laboratory anima	Female 6- to 8-week-old NMRI mice (Charles River Laboratories)		
Wild animals	The study did not involve wild animals.		

Human research participants

Policy information about studies involving human research participants

This study did not involve samples collected from the field.

Population characteristics

Field-collected samples

See Methods: Source of human AA amyloid fibrils; Age at death: 69; Gender: Female; Diagnosis: Chronic pulmonary obstructive disease, erosive seropositive rheumatoid arthritis, AA amyloidosis

Recruitment

Selected based on clinical findings at autopsy.