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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

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

n/a	Cor	nfirmed
	X	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
	x	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.
×		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 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)
	×	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>
×		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
×		Estimates of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated
	x	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 Microsoft Excel for database management.

Crystallographic software is described in the methods section

Graphpad Prism 7.0 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

The data that support the findings of this study are available from the corresponding author upon reasonable request.

The coordinates and structure factors for the ConM SOSIP.v7 structure in complex with PGT124 and 35022 have been deposited in the Protein Data Bank (PDB code

(6IEQ).						
Field spe	cific r	aparting				
Field-spe		. 9				
Please select the be		research. If you are not sure, read the appropriate sections before making your selection. Behavioural & social sciences				
For a reference copy of the	he document witl	h all sections, see <u>nature.com/authors/policies/ReportingSummary-flat.pdf</u>				
Life scien	ices st	udy design				
All studies must disc	close on these	e points even when the disclosure is negative.				
Sample size	No sample-siz	e calculations were performed.				
Data exclusions	No data was e	excluded from analyses				
Replication	Experiments v	were repeated and/or performed in duplicate.				
Randomization	on Allocation was random					
Blinding	Blinding was r	Blinding was not relevant for our studies				
Poporting	σ for c	pecific materials, systems and methods				
Neporting	g 101 3	pecific materials, systems and methods				
Materials & expe						
n/a Involved in the Unique biol	e study logical material	n/a Involved in the study				
X Antibodies	S	Flow cytometry				
x Eukaryotic	cell lines	MRI-based neuroimaging				
Palaeontolo	0,					
Animals and other organisms						
Unique biolo	gical ma	terials				
Policy information a	about <u>availabi</u>	lity of materials				
Obtaining unique	materials (Unique materials are available upon request.				
Antibodies						
Antibodies used		All antibodies used were provided by Michel Nussenzweig, Hermann Katinger, Mark Connors, James Robinson, Dennis Burton,				
Validation		ohn Mascola, Peter Kwong and William Olson directly or through the NIH AIDS Research and Reference Reagent Program.				
		N.A.				
Eukaryotic ce	ell lines					
Policy information a	about <u>cell line</u>	<u>s</u>				
Cell line source(s)		HEK293 cell lines were obtained from Thermo Fisher or ATCC as described in the methods. B cell lines were provided by Takayuki Ota and David Nemazee				
Authentication		None of the cell lines were validated				
Mycoplasma cont	tamination	All cell lines were tested negative for mycoplasma contamination				

Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for re	porting animal re	esearch
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Laboratory animals New Zealand White rabbits, female, 2.5-3kg Macaques, Macaca mulatta, female, 5-6 kg

Wild animals N.A.

Field-collected samples N.A.