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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 our Editorial Policies and the Editorial Policy Checklist.

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For a	all statistical analyses, 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 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>
x	For Bayesian analysis, information on the choice of priors and Markov chain Monte Carlo settings
x	For hierarchical and complex designs, identification of the appropriate level for tests and full reporting of outcomes
	\blacksquare Estimates of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated
	Our web collection on statistics for biologists contains articles on many of the points above.

Software and code

Policy information about availability of computer code

Data collection Microsoft Excel v16.37 for Macintosh; GraphPad Prism 8 (GraphPad Software, La Jolla, CA); IBM SPSS Statistics v25.0 for Macintosh (IBM, USA)

Data analysis Microsoft Excel v16.37 for Macintosh; GraphPad Prism 8 (GraphPad Software, La Jolla, CA); IBM SPSS Statistics v25.0 for Macintosh (IBM, USA); SPM8 for MATLAB R2013a (Wellcome Trust Institute for Neurology, UCL, UK)

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 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

Source data are provided with this paper using Mendeley Data (doi: 10.17632/tvxf6f9gj6.1). Raw neuroimaging data are available from the corresponding author upon request. (NB/ this doi is currently reserved and not published yet).

Field-spe	cific reporting			
Please select the or	e below that is the best fit for your research. If you are not sure, read the appropriate sections before making your selection.			
x Life sciences	Behavioural & social sciences Ecological, evolutionary & environmental sciences			
For a reference copy of t	ne document with all sections, see nature.com/documents/nr-reporting-summary-flat.pdf			
Life scier	ces study design			
All studies must dis	close on these points even when the disclosure is negative.			
Sample size	nple size of SEVEN (7) marmosets. Power calculations were not done to determine this sample size. The sample size was chosen based on vious paper (Alexander et al., 2019, Neuron) which used between 4-6 marmosets for experiments, sufficient to demonstrate an effect of C/25 over-activation.			
Data exclusions	No data were excluded from the manuscript.			
Replication	epeated-measures primate study such as this one does not have in-built replication. However, the human intruder data are a replication of revious study published in 2019 and was extended here investigating the anxiolytic properties of ketamine. Two out of five/seven animals re run independently by a postdoctoral scientist, the results of which matched those already obtained by the PhD student who ran the maining animals.			
	The entire PET study was not replicated separately; however, the PET neuroimaging study was carried out both by a PhD student and a postdoctoral scientist independently (each working with two animals of the cohort of four). Data were analyzed both by a PhD student and by a blinded neuroimaging scientist not involved in data collection.			
	Where replication was performed, it was successful.			
Randomization	This is not relevant to the study - it is a within-subject design.			
Blinding	For cardiovascular analysis, the investigator was not blinded because the data are objective readouts from a telemetry probe. These were analyzed using the independently designed Spike2 software as described in the methods.			
	For behavioral analysis, scoring was done by a independent research assistant who was blinded to drug effects.			
	Neuroimaging data were independently analysed by a blinded neuroimaging scientist.			
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, ed 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	ChIP-seq			
Eukaryotic				
✗ ☐ Palaeontol	pgy and archaeology MRI-based neuroimaging			
Animals an	d other organisms			
Human research participants				
X Clinical data				
Dual use re	search of concern			
Animals and	other organisms			
Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research				
Laboratory animals Common marmoset, Callithrix jacchus, three females, four males. Mean age: 2 years at the start of the study and 3 years months at the end.				

Wild animals

Field-collected samples

The study did not involve wild animals.

The study did not involve samples collected from the field.

Ethics oversight

All procedures were carried out in accordance with the UK Animals (Scientific Procedures) Act 1986 and the University of Cambridge Animal Welfare and Ethical Review Body.

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

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