# nature research

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

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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

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

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. F, t, r) with confidence intervals, effect sizes, degrees of freedom and P value noted Give P values as exact values whenever suitable.

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

Our web collection on <u>statistics for biologists</u> contains articles on many of the por

Software and code

Policy information about <u>availability of computer code</u> Data collection Spike 2.0 version 7; Olympus software; Dataquest A.R.T.; Cellvizio software 2.2

NCSS 2007; µCats Python software (http://doi.org/10.5281/zenodo.1242164; https://github.com/abrazhe/uCats); Dataquest Analysis

#### Data

on about availability of data

All manuscripts us tinduce a data valiability statement. This statement should provide the following information, where applicable:

- Accession codes, unique identifies, or web links for publicly available datasets

- A list of figures that have associated rare data

- A description of any restrictions on data availability

Not yet indicated in the text of the manuscript; The data that support the findings of this study are available from the corresponding author upon request.

### Field-specific reporting

Please select the one 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

Ethics	oversigh

All animal experimentations were performed in accordance with the European Commission Directive 86/609/EEC [Euro-Convention for the Protection of Vertebrate Animals used for Experimental and Other-Scientific Procedures] Act [1986] with project approval from the University College London Animal Care and Use Committee and the UK Home Office.

A flow sensitive alternating inversion recovery (FAIR) ASI, MRI sequence with a three shot segmented EPI readout was applied to measure the cerebrailbood flow. The following sequence parameters were used: TR = 5000 ms, TII=2000 m and trus issee-64-64, FOV-35 mm × 35 mm, TE=10 ms, single slice (slice thickness=2 mm), inversion pulse bandwidth=2 (shite. FAIR images vera exquired every 30 s. C.

Cerebral blood flow time-series data were extracted from manually drawn ROIs in the cortex and the striatum

perfusion 9.7 T

Not used

## Magnetic resonance imaging

Experimental design

Changes in cerebral blood flow in response to increased intracranial pressure were recorded in urethane-anesthetized Design type

In naive rats the CBF was measured using ASL MRI. The experimental setup was first pre-calibrated outside of the MRI scanner to deliver increases in ICP by 10-15 mmHg for 30-00 s. FAIR images were acquired every 30 seconds for "30 minutes with asseline period of 17 minutes, followed by 30-05 or for faised ICP and 10 minutes of recovery." Design specifications Behavioral performance measures N/A

Acquisition

Imaging type(s) Field strength

Sequence & imaging parameters

Area of acquisition Diffusion MRI Used

Preprocessing

Preprocessing software

Normalization template

Statistical modeling & inference

Model type and settings Effect(s) tested Specify type of analysis: Whole brain ROI-based Both

Correction

ICE	al location(s) Cortex and Striatum
	Specify voxel-wise or cluster-wise and report all relevant parameters for cluster-wise methods.
	Describe the transfer and form it is a beginning for a state of the second of the seco

ipe (mass univariate, multivariate, RSA, predictive, etc.) and describe essential details of the model at the first nd levels (e.g. fixed, random or mixed effects; drift or auto-correlation).

## Life sciences study design

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The paper describes the results of the experiments performed using in vivo animal preparations. Power calculations for the animal studies are described below. The significance was set at 0.05 and the beta was set at 0.20. From years of relevant research experience, we expect of detect a significant difference with a minimum of 5 animals per group, if the treatments cause difference between means that are as large as 2.25 standard deviations (SD), likely to be a physiologically significant difference. If the difference is as small as 1.75 SD, we increase sample sizes to 9. Although, increasing the sample sizes per group to 15 may enable to detect a difference between means of 1.25 SD, it is doubtful these differences will have a biological significance. Sample size

In the experiment illustrated on Fig 1f, the data obtained in one control animal was excluded from the analysis since the cannula used to deliver the experimental stimulas was blocked. In the experiment illustrated on Fig 4d, the data obtained in 1 control animals, 2 FeLC and 2 deliver the experimental stimulas was excluded from the analysis because either the cannula used to deliver the experimental stimulus was blocked or no transgene expression was detected in the targeted area of the braintening poposi-hoc histological examination. In the experiment summarized is Deplementary Table 1, the data were not obtained in 3 animals transduced to express TeLC because the batteries in the biotelemetry transducers run out of power during the course of the experiment.

in our strong opinion, replication of the whole group experiments is not essential for this study. It involved one individual experiment conduced per day and the effect of experimental treatment (ganglionic blockade [Fig II], TeLC or dinSNARE expression in the brainstem [Fig 4] completely abolished the physiological responses in all the individual animals. Assessment of satisfular lesponses in all the original responses in all one individual animals. Assessment of satisfular lesponses in one (Fig 2 and 3) demonstrated robust activations in all the experiments (In-7 and 8, respectively) conduced, therefore we felt that no further tests were required in this study.

Randomization is not relevant to this study, since experimental subjects (outbred Sprague-Dawley rats) are obtained from a genetically homogeneous colony and then assigned to different experimental groups according to the treatment.

All the experimenters were blinded to the condition or treatment of experimental subjects and to the identity of the experimental groups

## Reporting for specific materials, systems and methods

We require information from authors about some types of materials, experimental systems and methods used in many studies. Here, indicate whether each r system or method listed 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 resp

Materials & experimental systems		Methods		
n/a	Involved in the study	n/a	Involved in the study	
	Antibodies	$\boxtimes$	ChIP-seq	
$\boxtimes$	Eukaryotic cell lines	$\boxtimes$	Flow cytometry	
$\bowtie$	Palaeontology		MRI-based neuroimaging	
	Animals and other organisms			
$\boxtimes$	Human research participants			
V	Clinical data			

#### Antibodies

Antibodies used Chicken anti-GFP (1:500; Aves Labs), rabbit anti-tyrosine hydroxylase (1:500; Santa Cruz), rabbit anti-glial fibrillary acidic protein (1:500; Dako) and mouse anti-microtubule-associated protein 2 (1:500; Sigma) antibodies. The antibodies were validated using negative control protocols. We employ these antibodies routinely and we have published numerous research articles demonstrating their specificity. In the present study immunohatochemistry was used to identify anatomical landmarks and the spread of transgene expression, resulting in characteristic stating patterns. Validation

#### Animals and other organisms

Laboratory animals	Young adult Sprague-Dawley rats (200-300 g)
Wild animals	Provide details on animals observed in ar captured in the field; report species, sex and age where possible. Describe how animals were cought and transported and what happened to captive animals after the study (if silled, explain why and describe method, if reclased, say where and when (i) if state that the study did not involve wild animals.
Field-collected samples	For laboratory work with field-collected samples, describe all relevant parameters such as housing, maintenance, temperature,

#### Models & analysis

n/a	Involved in the study	
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Functional and/or effective connectivity

Graph analysis
Multivariate modeling or predictive analysis

Functional and/or effective connectivity

Graph analysis

Report the measures of dependence used and the model details (e.g. Pearson correlation, partial correlation, mutual information).