nature portfolio

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Reporting Summary

Nature Portfolio 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 Portfolio policies, see our <u>Editorial Policies</u> and the <u>Editorial Policy Checklist</u>.

Please do not complete any field with "not applicable" or n/a. Refer to the help text for what text to use if an item is not relevant to your study. For final submission: please carefully check your responses for accuracy; you will not be able to make changes later.

Statistics

For	all sta	atistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.
n/a	Con	firmed
	\boxtimes	The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
	\boxtimes	A statement on 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
	\boxtimes	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>
	\boxtimes	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	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 points above.

Software and code

Policy information about <u>availability of computer code</u>
Data collection www.Gorilla.sc

Data analysis

malysis (Matlab (vR2022B); RStudio (v2023.06.1+524); Bayesian First Aid (v1.0; R); CBM Toolbox (Matlab)

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 Portfolio 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 description of any restrictions on data availability
- For clinical datasets or third party data, please ensure that the statement adheres to our policy

All data collected is freely available on GitHub: https://github.com/josephmbarnby/Barnby_etal_2023_D2D3Modelling/Data

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Research involving human participants, their data, or biological material

Policy information about studies with <u>human participants or human data</u>. See also policy information about <u>sex, gender (identity/presentation)</u>, <u>and sexual orientation</u> and <u>race, ethnicity and racism</u>.

Reporting on sex and gender	Only men were recruited for this study due to potential pharmacodynamic interactions between Haloperidol and birth control medication. This is discussed in the manuscript.
Reporting on race, ethnicity, or other socially relevant groupings	We don't report on race, ethnicity, or social relevant groupings for analysis.
Population characteristics	We do not report on population characteristics in this manuscript.
Recruitment	We recruited 86 participants by convenience sampling. 28 were included for analysis
	following screening.
Ethics oversight	The study was approved by KCL ethics board (HR-16/17-0603)

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

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.

🔀 Life sciences 🛛 🗌 Behavioural & social sciences 🔄 Ecological, evolutionary & environmental sciences

For a reference copy of the document with all sections, see <u>nature.com/documents/nr-reporting-summary-flat.pdf</u>

Life sciences study design

All studies must disclose on these points even when the disclosure is negative.

Sample size	n=35
Data exclusions	n=2 excluded for non completion of experimental days. n=5 excluded for failing medical screening.
Replication	Participants completed three experimenal days each
Randomization	Participants were randomised using a Williams Square design.
Blinding	The study used a double blind proceedure (experimenter & participant blinded to drug).

Behavioural & social sciences study design

All studies must disclose on these points even when the disclosure is negative.

Study description	
Research sample	
Sampling strategy	
Data collection	
Timing	
Data exclusions	
Non-participation	
Randomization	

Ecological, evolutionary & environmental sciences study design

All studies must disclose on these points even when the disclosure is negative.

Study description	
Research sample	
Sampling strategy	
Sumpling Strategy	
Data collection	
Timing and spatial scale	
Data exclusions	
Reproducibility	
Randomization	
Blinding	
Did the study involve field	work? Yes No

Field work, collection and transport

Field conditions	
Location	
Access & import/export	
Disturbance	

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 material, 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 response.

Materials & experimental systems

N	let	h	\cap	Ч	c
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n/a	Involved in the study	n/a	Involved in the study
\bowtie	Antibodies	X	ChIP-seq
\bowtie	Eukaryotic cell lines	\bowtie	Flow cytometry
\bowtie	Palaeontology and archaeology	\mathbf{X}	MRI-based neuroimaging
\bowtie	Animals and other organisms		
\bowtie	Clinical data		
\boxtimes	Dual use research of concern		
\boxtimes	Plants		

Antibodies

Antibodies used
Validation

Eukaryotic cell lines

Policy information about <u>cell lines</u>	plicy information about <u>cell lines and Sex and Gender in Research</u>				
Cell line source(s)					
Authentication					
Mycoplasma contamination					
Commonly misidentified lines (See <u>ICLAC</u> register)					

Palaeontology and Archaeology

Specimen provenance	
Specimen deposition	
Dating methods	
Tick this box to confirm	n that the raw and calibrated dates are available in the paper or in Supplementary Information.
Ethics oversight	
Note that full information on th	a approval of the study protocol must also be provided in the manuscript

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Animals and other research organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research, and <u>Sex and Gender in</u> <u>Research</u>

Laboratory animals	
Wild animals	
Reporting on sex	
Field-collected samples	
Ethics oversight	

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

Clinical data

Policy information about <u>clinical studies</u> All manuscripts should comply with the ICMJE <u>guidelines for publication of clinical research</u> and a completed <u>CONSORT checklist</u> must be included with all submissions.

Clinical trial registration	
Study protocol	
Data collection	
Outcomes	

Dual use research of concern

Policy information about dual use research of concern

Hazards

Could the accidental, deliberate or reckless misuse of agents or technologies generated in the work, or the application of information presented in the manuscript, pose a threat to:

No	Yes
	Public health
	National security
	Crops and/or livestock
	Ecosystems
	Any other significant area

Experiments of concern

Does the work involve any of these experiments of concern:

No	Yes
	Demonstrate how to render a vaccine ineffective
	Confer resistance to therapeutically useful antibiotics or antiviral agents
	Enhance the virulence of a pathogen or render a nonpathogen virulent
	Increase transmissibility of a pathogen
	Alter the host range of a pathogen
	Enable evasion of diagnostic/detection modalities
	Enable the weaponization of a biological agent or toxin
	Any other potentially harmful combination of experiments and agents

Plants

Seed stocks	
Novel plant genotypes	
Hover plant Schotypes	
Authentication	

ChIP-seq

Data deposition

	Confirm that both raw and final p	processed data have been	n deposited in a public database such a	as <u>GEC</u>
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Confirm that you have deposited or provided access to graph files (e.g. BED files) for the called peaks.

Data access links May remain private before publication.	
Files in database submission	
Genome browser session	
(e.g. UCSC)	

Methodology

Replicates	
Sequencing depth	
Antibodies	
Peak calling parameters	
Data quality	

Flow Cytometry

Plots

Confirm that:

The axis labels state the marker and fluorochrome used (e.g. CD4-FITC).

The axis scales are clearly visible. Include numbers along axes only for bottom left plot of group (a 'group' is an analysis of identical markers).

All plots are contour plots with outliers or pseudocolor plots.

A numerical value for number of cells or percentage (with statistics) is provided.

Methodology

Sample preparation	
Instrument	
Software	
Cell population abundance	
Gating strategy	

Tick this box to confirm that a figure exemplifying the gating strategy is provided in the Supplementary Information.

Magnetic resonance imaging

Experimental design		
Design type		
Design specifications		
Behavioral performance measures		
Imaging type(s)		
Field strength		
Sequence & imaging parameters		
Area of acquisition		
Diffusion MRI Used	Not used	
Proprocessing		

Preprocessing

Preprocessing software	
Normalization	
Normalization template	
Noise and artifact removal	
Volume censoring	

Statistical modeling & inference

Model type and settings	
Effect(s) tested	

Specify type of analysis: 🗌 Whole brain 🗌	ROI-based Both
Statistic type for inference	
(See <u>Eklund et al. 2016</u>)	
Correction	
Models & analysis	
n/a Involved in the study Involved in the study Image: State of the study Image: State	5
Functional and/or effective connectivity	
Graph analysis	
Multivariate modeling and predictive analysis	

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