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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	il statistical analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section	٦.
n/a	Confirmed	
X	\Box The exact sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement	
x	A statement on whether measurements were taken from distinct samples or whether the same sample was measured repeat	edly
×	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	
X	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 AND variation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)	ı coefficient)
×	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 <i>Give P values as exact values whenever suitable.</i>	noted
×	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	
X	\square 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

All data used in this study are publicly available and were downloaded from the websites of the original studies / resources, see Data section below for a detailed list.

Data analysis

Source code and pre-compiled executables for the VRNetzer platform are available on https://github.com/menchelab/VRNetzer. We used open source softwares and programming environments Python (version 3.6), JavaScript (version ES6), MySQL (version 8.0), as well as the EPIC Unreal engine (version 4). Detailed software and package dependencies can be found in the dependencies section on https://github.com/menchelab/VRNetzer.

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 <u>availability of data</u>

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

All data used in this study are publicly available and were downloaded from the websites of the referenced original studies.

We used the following publicly available data sources:

Gene ontology (GO) taxonomy, from http://geneontology.org/docs/download-ontology/

GO annotations, from http://current.geneontology.org/products/pages/downloads.html

Disease ontology (DO) taxonomy, from https://github.com/DiseaseOntology/HumanDiseaseOntology/tree/main/src/ontology/releases						
DisGeNET, from https://www.disgenet.org/downloads Online Mendelian Inheritance in Man (OMIM), from https://www.omim.org/						
	Ontology (HPO) annotation, from https://hpo.jax.org/app/download/annotation					
	HPO taxonomy, from https://hpo.jax.org/app/download/ontology					
	Kyoto Encyclopedia of Genes and Genomes (KeGG), from https://www.genome.jp/kegg/genes.html					
	epository for Interaction Datasets (BioGRID), from https://downloads.thebiogrid.org/					
	tps://reactome.org/download-data					
	pression (GTEx) data, from https://gtexportal.org/home/datasets					
	Pubmed articles from INDRA, from https://github.com/sorgerlab/indra Human Integrated Protein-Protein Interaction Reference (HIPPIE) PPIs, from http://cbdm-01.zdv.uni-mainz.de/~mschaefer/hippie/download.php					
Weblinks, version numbers and release dates can also be found in the data availability section on https://github.com/menchelab/VRNetzer.						
Eiold spe	ocific reporting					
	ecific reporting					
	ne 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>					
Lite scier	nces study design					
All studies must dis	sclose on these points even when the disclosure is negative.					
Sample size	Study does not involve any analyses to which sample size calculations are applicable.					
Data exclusions	No data were excluded in the study.					
Replication	Study does not involve experimental results.					
Randomization	Study does not involve any samples / organisms / participants.					
Blinding	Study does not involve any samples / organisms / participants.					
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Reportin	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, ted 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 & ex	perimental systems Methods					
n/a Involved in th	ne study n/a Involved in the study					
✗ ☐ Antibodies	ChIP-seq					
Eukaryotic	cell lines Flow cytometry					
	Palaeontology and archaeology MRI-based neuroimaging					
Animals and other organisms						
Human research participants						
X Clinical data						
Dual use research of concern						