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 statistical analyses, confirm that the following items are present in the 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	
X		A statement on 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.	
	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	
	X	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)	
	x	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 Give <i>P</i> values as exact values whenever suitable.	
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	
	x	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	No software used	
Data analysis	All statistical analyses were performed with SAS software, version 9.4 TS1M5 (SAS Institute)	
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		

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

Data will be shared with bona fide researchers who submit a research proposal approved by the independent review board. Individual patient data will be shared in datasets in a de-identified and anonymised format. Data will be made available after research completion and approval of the product, and product use in the EU and the USA. Information about data access request proposals can be found at novonordisk-trials.com.

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	Biologic sex was reported for trial participants; information on gender was not collected
Reporting on race, ethnicity, or other socially relevant groupings	Race and ethnicity were reported for trial participants
Population characteristics	Reported in Supplementary Table 1
Recruitment	Site selection is handled by NN, however the actual recruitment process is handled independently by the sites/PI's/study coordinators
Ethics oversight	National and institutional regulatory and ethical authorities approved the protocol, and all patients provided written and informed consent

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.

X 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	17,604. Sample size determination was provided in the primary publication (Lincoff AM et al. NEJM 389; 2221-2232 (2023).
Data exclusions	No data were excluded
Replication	This is an international, randomized, double-blind trial; thus replication is not possible
Randomization	Participants were randomized in a 1:1 ratio
Blinding	Participants and investigators were blinded to treatment allocation. Investigators were blinded to group allocation during data collection and analysis.

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		
Sampling strategy		
Data collection		
Timing and spatial scale		
Data exclusions		
Reproducibility		
Randomization		
Blinding		
Did the study involve field work?		

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

Methods

n/a	Involved in the study	n/a	Involved in the study
x	Antibodies	X	ChIP-seq
X	Eukaryotic cell lines	x	Flow cytometry
X	Palaeontology and archaeology	x	MRI-based neuroimaging
X	Animals and other organisms		
	X Clinical data		
X	Dual use research of concern		
x	Plants		

Antibodies

Antibodies used
Validation

Eukaryotic cell lines

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

Palaeontology and Archaeology

Specimen provenance		
Specimen deposition		
Dating methods		
Tick this box to confirm 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	

rotocol must als

Animals and other research organisms

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

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 c	inical studies
All manuscripts should comply	with the ICMJE guidelines for publication of clinical research and a completed CONSORT checklist must be included with all submissions.
Clinical trial registration	NCT03574597
Study protocol	Provided in the primary publication (Lincoff AM et al. NEJM 389; 2221-2232 (2023)
Data collection	The SELECT study enrolled participants from 41 countries between October 2018 through March 2021.All potentially eligible patients were eligible to take part, thereby minimizing any potential self-selection bias
Outcomes	Study outcomes focused on several key areas related to kidney health, including changes in estimated glomerular filtration rate (eGFR), and modifications in urinary albumin-to-creatinine ratio (UACR). All of the primary and secondary outcomes, and the statistical methodologies used to analyze them, were prespecified in the Statistical Analysis Plan. All statistical

Dual use research of score formed for all outcome measures were thus predefined.

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:

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
x	Demonstrate how to render a vaccine ineffective
x	Confer resistance to therapeutically useful antibiotics or antiviral agents
	Enhance the virulence of a pathogen or render a nonpathogen virulent
x	Increase transmissibility of a pathogen
x	Alter the host range of a pathogen
x	Enable evasion of diagnostic/detection modalities
x	Enable the weaponization of a biological agent or toxin
X	Any other potentially harmful combination of experiments and agents

Plants

Seed stocks	
Novel plant genotypes	
Authentication	

ChIP-seq

Data deposition

\square	Confirm that both raw and final	processed data have been	deposited in a public data	abase such as <u>GEO</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. <u>UCSC</u>)	

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

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
Preprocessing	

F

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 Eklund et al. 2016)	
Correction	
Models & analysis	
n/a Involved in the study X Functional and/or effective connectivity X Graph analysis X Multivariate modeling or predictive analysis	S
Functional and/or effective connectivity	
Graph analysis	
Multivariate modeling and predictive analysis	

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