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 Editorial Policies and the Editorial Policy Checklist.

Statistics	
For all statistical a	analyses, confirm that the following items are present in the figure legend, table legend, main text, or Methods section.
n/a Confirmed	
☐ ☐ The exac	ct sample size (n) for each experimental group/condition, given as a discrete number and unit of measurement
☐ X A statem	nent on whether measurements were taken from distinct samples or whether the same sample was measured repeatedly
The stati	istical test(s) used AND whether they are one- or two-sided mon tests should be described solely by name; describe more complex techniques in the Methods section.
A descrip	ption of all covariates tested
A descrip	ption of any assumptions or corrections, such as tests of normality and adjustment for multiple comparisons
A full de	scription of the statistical parameters including central tendency (e.g. means) or other basic estimates (e.g. regression coefficient) iation (e.g. standard deviation) or associated estimates of uncertainty (e.g. confidence intervals)
	hypothesis testing, the test statistic (e.g. F , t , r) with confidence intervals, effect sizes, degrees of freedom and P value noted lues as exact values whenever suitable.
For Baye	esian analysis, information on the choice of priors and Markov chain Monte Carlo settings
For hiera	archical and complex designs, identification of the appropriate level for tests and full reporting of outcomes
Estimate	es of effect sizes (e.g. Cohen's d , Pearson's r), indicating how they were calculated
1	Our web collection on <u>statistics for biologists</u> contains articles on many of the points above.
Software ar	nd code
Policy information	n about <u>availability of computer code</u>
Data collection	We collected data with GraphPad Prism: an analysis and graphing software. Version: GraphPad Prism 9
Data analysis	We analyzed data with GraphPad Prism: an analysis and graphing software. Version: GraphPad Prism 9
	ng 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 y 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

The data that support the findings of this study are available from the corresponding author upon reasonable request.

Research involving human participants, their data, or biological material

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

Reporting on sex and gender

Our findings applied only to female.

No comparison with male was performed because the present study is exploratory.

Reporting on race, ethnicity, or other socially relevant groupings

Please specify the socially constructed or socially relevant categorization variable(s) used in your manuscript and explain why they were used. Please note that such variables should not be used as proxies for other socially constructed/relevant variables (for example, race or ethnicity should not be used as a proxy for socioeconomic status).

Provide clear definitions of the relevant terms used, how they were provided (by the participants/respondents, the researchers, or third parties), and the method(s) used to classify people into the different categories (e.g. self-report, census or administrative data, social media data, etc.)

Please provide details about how you controlled for confounding variables in your analyses.

Population characteristics

- Healthy female volunteer, age: 20 and 40 years, BMI: 20 and 26 kg/m², height: 158 cm and 180 cm,
- Regular menstrual cycles and cycles lasting between 20 and 35 days.
- Without oestroprogestative contraception (i.e., oral progestative contraception, IUDs, implants or absence of contraception are allowed).
- Fitness level assessment:
- . if age < 35 years: 35 ml/min./kg < VO2max < 55 ml/min./kg,
- if age > 35 years: 30 ml/min./kg < VO2max < 55 ml/min./kg,
- Active and free from any orthopaedic, musculoskeletal, and cardiovascular disorders,
- Non active smokers, no alcohol, or drug addiction, and no medical treatment (with the exception of the aforementioned accepted means of contraception),

Non-inclusion criteria:

- Past record of orthostatic intolerance, arterial hypertension, and cardiac rhythm disorders,
- Chronic back pains, vertebral fracture, scoliosis or herniated disc, history of knee problems, or joint surgery/broken leg,
- History of hiatus hernia or gastro-oesophgeal reflux, thyroid dysfunction, renal stones, diabetes, glaucoma, and migraines,
- $\textit{Past record of thrombophlebitis, family history of thrombosis, or positive \textit{response} in the \textit{thrombosis screening procedure,} \\$
- Abnormal result for lower limbs in Doppler ultrasound,
- Bone mineral density: T-score < -1.5, osteosynthesis material, presence of metallic implants,
- Poor tolerance to blood sampling and having donated blood (> 8 mL/kg) in a period of 8 weeks or less before the stard of the experiment.

Recruitment

The selection was carried out in two phases:

- One preliminary screening based on the application files, comprising first in a questionnaire on the subject's way of life, his education and professional experience, and a physical activity questionnaire (IPAQ); and secondly a medical questionnaire on the subject's personal and family medical history.
- A selection session, including clinical and paramedical examinations, questionnaires, carried out where the investigations

Subject Aptitude:

Candidates was declared apt to participate in the experiment on the following conditions:

- After verification that the information document has been read carefully, candidates had the possibility to ask any additional question concerning the research project. After having obtained satisfactory answers to these questions, they signed the specific Information and Consent Form.
- After checking the tests results, verification was made that all inclusion criteria are present and that there are no noninclusion criteria.

Ethics oversight

Approval by both the CPP IIe de France II Ethics Committee (July 5, 2021) and the French Health Authority, ANSM (May 31,

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

Power-based calculation of the number of subjects is not directly applicable for such explorative studies. However, based on effect sizes obtained with previous MEDES DI in men, minimal sample to find statistically significant difference between Pre- and Post-DI, when there is one (at power 80% and alphal evel 0.05), could be estimated for example as n=5 for plasma volume evolution (effect size 2.01), n=11 for orthostatic tolerance (effect size 0.96), n=15 for glucose tolerance (effect size 0.8), n=19 for VO2max (effect size 0.69). Therefore, a total of 20 subjects was deemed necessary for this study.

Data exclusions

One subject could not complete the experiment due to technical issues and another subject was excluded for clinical trials regulatory reasons. Overall, 18 healthy women participated in the study.

The quantification of ganglion cell complex thickness was performed in 16 subjects due to technical glitches.

Replication

The quality of ophthalmological measurements is determined with a quality index provided by the device.

Randomization

Randomization is not relevant for our study. No allocation was necessary.

Blinding

Blinding is not relevant for our study. No allocation was necessary.

Behavioural & social sciences study design

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

Study description

Briefly describe the study type including whether data are quantitative, qualitative, or mixed-methods (e.g. qualitative cross-sectional, quantitative experimental, mixed-methods case study).

Research sample

State the research sample (e.g. Harvard university undergraduates, villagers in rural India) and provide relevant demographic information (e.g. age, sex) and indicate whether the sample is representative. Provide a rationale for the study sample chosen. For studies involving existing datasets, please describe the dataset and source.

Sampling strategy

Describe the sampling procedure (e.g. random, snowball, stratified, convenience). Describe the statistical methods that were used to predetermine sample size OR if no sample-size calculation was performed, describe how sample sizes were chosen and provide a rationale for why these sample sizes are sufficient. For qualitative data, please indicate whether data saturation was considered, and what criteria were used to decide that no further sampling was needed.

Data collection

Provide details about the data collection procedure, including the instruments or devices used to record the data (e.g. pen and paper, computer, eye tracker, video or audio equipment) whether anyone was present besides the participant(s) and the researcher, and whether the researcher was blind to experimental condition and/or the study hypothesis during data collection.

Timing

Indicate the start and stop dates of data collection. If there is a gap between collection periods, state the dates for each sample cohort.

Data exclusions

If no data were excluded from the analyses, state so OR if data were excluded, provide the exact number of exclusions and the rationale behind them, indicating whether exclusion criteria were pre-established.

Non-participation

State how many participants dropped out/declined participation and the reason(s) given OR provide response rate OR state that no participants dropped out/declined participation.

Randomization

If participants were not allocated into experimental groups, state so OR describe how participants were allocated to groups, and if allocation was not random, describe how covariates were controlled.

Ecological, evolutionary & environmental sciences study design

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

Study description

Briefly describe the study. For quantitative data include treatment factors and interactions, design structure (e.g. factorial, nested, hierarchical), nature and number of experimental units and replicates.

Research sample

Describe the research sample (e.g. a group of tagged Passer domesticus, all Stenocereus thurberi within Organ Pipe Cactus National Monument), and provide a rationale for the sample choice. When relevant, describe the organism taxa, source, sex, age range and any manipulations. State what population the sample is meant to represent when applicable. For studies involving existing datasets, describe the data and its source.

Sampling strategy

Note the sampling procedure. Describe the statistical methods that were used to predetermine sample size OR if no sample-size calculation was performed, describe how sample sizes were chosen and provide a rationale for why these sample sizes are sufficient.

Data collection

Describe the data collection procedure, including who recorded the data and how.

Timing and spatial scale

Indicate the start and stop dates of data collection, noting the frequency and periodicity of sampling and providing a rationale for these choices. If there is a gap between collection periods, state the dates for each sample cohort. Specify the spatial scale from which the data are taken

Data exclusions

If no data were excluded from the analyses, state so OR if data were excluded, describe the exclusions and the rationale behind them, indicating whether exclusion criteria were pre-established.

Reproducibility	Describe the measures taken to verify the reproducibility of experimental findings. For each experiment, note whether any attempts to repeat the experiment failed OR state that all attempts to repeat the experiment were successful.
Randomization	Describe how samples/organisms/participants were allocated into groups. If allocation was not random, describe how covariates were controlled. If this is not relevant to your study, explain why.
Blinding	Describe the extent of blinding used during data acquisition and analysis. If blinding was not possible, describe why OR explain why blinding was not relevant to your study.
Did the study involve fiel	d work? Yes No
Field work, collec	tion and transport
Field conditions	Describe the study conditions for field work, providing relevant parameters (e.g. temperature, rainfall).
Location	State the location of the sampling or experiment, providing relevant parameters (e.g. latitude and longitude, elevation, water depth).
Access & import/export	Describe the efforts you have made to access habitats and to collect and import/export your samples in a responsible manner and in compliance with local, national and international laws, noting any permits that were obtained (give the name of the issuing authority, the date of issue, and any identifying information).
Disturbance	Describe any disturbance caused by the study and how it was minimized.

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
\times	Antibodies	ChIP-seq
\times	Eukaryotic cell lines	Flow cytometry
\times	Palaeontology and archaeology	MRI-based neuroimaging
\boxtimes	Animals and other organisms	
	Clinical data	
\times	Dual use research of concern	
\boxtimes	Plants	
۸ ۵	tibadias	

Antibodies

Antibodies used

Describe all antibodies used in the study; as applicable, provide supplier name, catalog number, clone name, and lot number.

Validation

Describe the validation of each primary antibody for the species and application, noting any validation statements on the manufacturer's website, relevant citations, antibody profiles in online databases, or data provided in the manuscript.

Eukaryotic cell lines

Policy information about <u>cell lines and Sex and Gender in Research</u>

Cell line source(s)

State the source of each cell line used and the sex of all primary cell lines and cells derived from human participants or vertebrate models.

Authentication

Describe the authentication procedures for each cell line used OR declare that none of the cell lines used were authenticated.

Mycoplasma contamination

Confirm that all cell lines tested negative for mycoplasma contamination OR describe the results of the testing for mycoplasma contamination.

Commonly misidentified lines (See ICLAC register)

Name any commonly misidentified cell lines used in the study and provide a rationale for their use.

Palaeontology and Archaeology

Specimen provenance

Provide provenance information for specimens and describe permits that were obtained for the work (including the name of the issuing authority, the date of issue, and any identifying information). Permits should encompass collection and, where applicable, export.

Specimen deposition

Indicate where the specimens have been deposited to permit free access by other researchers.

Dating methods

If new dates are provided, describe how they were obtained (e.g. collection, storage, sample pretreatment and measurement), where they were obtained (i.e. lab name), the calibration program and the protocol for quality assurance OR state that no new dates are provided.

Tick this box to confirm that the raw and calibrated dates are available in the paper or in Supplementary Information.

Ethics oversight

Identify the organization(s) that approved or provided guidance on the study protocol, OR state that no ethical approval or guidance was required and explain why not.

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

Animals and other research organisms

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

Laboratory animals

For laboratory animals, report species, strain and age OR state that the study did not involve laboratory animals.

Wild animals

Provide details on animals observed in or captured in the field; report species and age where possible. Describe how animals were caught and transported and what happened to captive animals after the study (if killed, explain why and describe method; if released, say where and when) OR state that the study did not involve wild animals.

Reporting on sex

Indicate if findings apply to only one sex; describe whether sex was considered in study design, methods used for assigning sex. Provide data disaggregated for sex where this information has been collected in the source data as appropriate; provide overall numbers in this Reporting Summary. Please state if this information has not been collected. Report sex-based analyses where performed, justify reasons for lack of sex-based analysis.

Field-collected samples

For laboratory work with field-collected samples, describe all relevant parameters such as housing, maintenance, temperature, photoperiod and end-of-experiment protocol OR state that the study did not involve samples collected from the field.

Ethics oversight

Identify the organization(s) that approved or provided guidance on the study protocol, OR state that no ethical approval or guidance was required and explain why not.

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

Clinical data

Policy information about clinical 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

NCT05043974

Study protocol

https://classic.clinicaltrials.gov/ct2/show/NCT05043974

Data collection

 $All \ subjects \ underwent \ a \ complete \ bil \ ateral \ ophthal mologic \ examination \ 5 \ days \ before \ and \ 5-6h \ after \ dry \ immersion.$

Outcomes

The primary objective of the present study is to investigate the physiological effects of 5 days of dry immersion in 20 healthy female subjects, and to obtain Dry Immersion-in-Women Reference Dataset. A set of measurements will assess the changes in the cardiovascular, neuro-ophthalmological, hematological, metabolic, sensorimotor, immune, muscle and bone systems.

The secondary objectives are:

- to reveal/understand the particularities of the mechanisms of DI-induced changes in women,
- to compare changes induced by a 5-day dry immersion in women with those from men dry immersion studies, bedrest studies, and flight studies, using already published results.

Dual use research of concern

Policy information about <u>dual use research of concern</u>

Hazards

in the manuscript, pose a		to:
No Yes		
Public health		
National security		
Crops and/or livest	ock	
Ecosystems		
Any other significar	nt area	
Experiments of concer	n	
Does the work involve and	y of the	ese experiments of concern:
No Yes		
		er a vaccine ineffective
		peutically useful antibiotics or antiviral agents pathogen or render a nonpathogen virulent
Increase transmissi		
Alter the host range	e of a pa	athogen
Enable evasion of c	diagnost	ic/detection modalities
Enable the weapon	nization	of a biological agent or toxin
Any other potentia	lly harm	ıful combination of experiments and agents
Plants		
Seed stocks	Report	on the source of all seed stocks or other plant material used. If applicable, state the seed stock centre and catalogue number. If
		pecimens were collected from the field, describe the collection location, date and sampling procedures.
Novel plant genotypes	gene ei numbe	be the methods by which all novel plant genotypes were produced. This includes those generated by transgenic approaches, diting, chemical/radiation-based mutagenesis and hybridization. For transgenic lines, describe the transformation method, the or of independent lines analyzed and the generation upon which experiments were performed. For gene-edited lines, describe itor used, the endogenous sequence targeted for editing, the targeting guide RNA sequence (if applicable) and how the editor uplied.
a.		be any authentication procedures for each seed stock used or novel genotype generated. Describe any experiments used to the effect of a mutation and, where applicable, how potential secondary effects (e.g. second site T-DNA insertions, mosiacism, get gene editing) were examined.
ChIP-seq		
Data deposition	, and fi	nal processed data have been deposited in a public database such as <u>GEO</u> .
		sited or provided access to graph files (e.g. BED files) for the called peaks.
	. achos	
Data access links May remain private before public	cation.	For "Initial submission" or "Revised version" documents, provide reviewer access links. For your "Final submission" document, provide a link to the deposited data.
Files in database submissi	ion	Provide a list of all files available in the database submission.
Genome browser session (e.g. <u>UCSC</u>)		Provide a link to an anonymized genome browser session for "Initial submission" and "Revised version" documents only, to enable peer review. Write "no longer applicable" for "Final submission" documents.
Methodology		
Replicates	Describ	be the experimental replicates, specifying number, type and replicate agreement.
Sequencing depth		be the sequencing depth for each experiment, providing the total number of reads, uniquely mapped reads, length of reads and er they were paired- or single-end.
Antibodies	Describ lot nun	be the antibodies used for the ChIP-seq experiments; as applicable, provide supplier name, catalog number, clone name, and nber.

Peak calling parameters	Specify the command line program and parameters used for read mapping and peak calling, including the ChIP, control and index files used.
Data quality	Describe the methods used to ensure data quality in full detail, including how many peaks are at FDR 5% and above 5-fold enrichment.
Software	Describe the software used to collect and analyze the ChIP-seq data. For custom code that has been deposited into a community repository, provide accession details.

Flow Cytometry

Plots	
Confirm that:	
The axis labels state the mark	er and fluorochrome used (e.g. CD4-FITC).
The axis scales are clearly visil	ble. Include numbers along axes only for bottom left plot of group (a 'group' is an analysis of identical markers).
All plots are contour plots wit	ch outliers or pseudocolor plots.
A numerical value for number	r of cells or percentage (with statistics) is provided.
Methodology	
Sample preparation	Describe the sample preparation, detailing the biological source of the cells and any tissue processing steps used.
Instrument	Identify the instrument used for data collection, specifying make and model number.
Software	Describe the software used to collect and analyze the flow cytometry data. For custom code that has been deposited into a community repository, provide accession details.
Cell population abundance	Describe the abundance of the relevant cell populations within post-sort fractions, providing details on the purity of the samples and how it was determined.
Gating strategy	Describe the gating strategy used for all relevant experiments, specifying the preliminary FSC/SSC gates of the starting cell population, indicating where boundaries between "positive" and "negative" staining cell populations are defined.
Magnetic resonance in	naging
Design type	Indicate task or resting state; event-related or block design.
Design specifications	Specify the number of blocks, trials or experimental units per session and/or subject, and specify the length of each trial or block (if trials are blocked) and interval between trials.
Behavioral performance measure	State number and/or type of variables recorded (e.g. correct button press, response time) and what statistics were used to establish that the subjects were performing the task as expected (e.g. mean, range, and/or standard deviation across subjects).
Acquisition	
Imaging type(s)	Specify: functional, structural, diffusion, perfusion.
Field strength	Specify in Tesla
Sequence & imaging parameters	Specify the pulse sequence type (gradient echo, spin echo, etc.), imaging type (EPI, spiral, etc.), field of view, matrix size, slice thickness, orientation and TE/TR/flip angle.
Area of acquisition	State whether a whole brain scan was used OR define the area of acquisition, describing how the region was determined.
Diffusion MRI Used	☐ Not used
Preprocessing	

Preprocessing software

Provide detail on software version and revision number and on specific parameters (model/functions, brain extraction, segmentation, smoothing kernel size, etc.).

Normalization	If data were normalized/standardized, describe the approach(es): specify linear or non-linear and define image types used transformation OR indicate that data were not normalized and explain rationale for lack of normalization.
Normalization template	Describe the template used for normalization/transformation, specifying subject space or group standardized space (e.g. original Talairach, MNI305, ICBM152) OR indicate that the data were not normalized.
Noise and artifact removal	Describe your procedure(s) for artifact and structured noise removal, specifying motion parameters, tissue signals and physiological signals (heart rate, respiration).
Volume censoring	Define your software and/or method and criteria for volume censoring, and state the extent of such censoring.
Statistical modeling & infe	rence
Model type and settings	Specify type (mass univariate, multivariate, RSA, predictive, etc.) and describe essential details of the model at the first and second levels (e.g. fixed, random or mixed effects; drift or auto-correlation).
Effect(s) tested	Define precise effect in terms of the task or stimulus conditions instead of psychological concepts and indicate whether ANOVA or factorial designs were used.
Specify type of analysis:	Whole brain ROI-based Both
Statistic type for inference	Specify voxel-wise or cluster-wise and report all relevant parameters for cluster-wise methods.
(See Eklund et al. 2016)	
Correction	Describe the type of correction and how it is obtained for multiple comparisons (e.g. FWE, FDR, permutation or Monte Car
Models & analysis n/a Involved in the study	
Functional and/or effective co	Report the measures of dependence used and the model details (e.g. Pearson correlation, partial correlation).
Graph analysis	Report the dependent variable and connectivity measure, specifying weighted graph or binarized graph, subject- or group-level, and the global and/or node summaries used (e.g. clustering coefficient, efficiency,

Specify independent variables, features extraction and dimension reduction, model, training and evaluation

Multivariate modeling and predictive analysis

metrics.