# 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>.

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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.
$\boxtimes$	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. <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
$\times$	Estimates of effect sizes (e.g. Cohen's <i>d</i> , Pearson's <i>r</i> ), 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

AFM- Cypher, Asylum Research

microindenation - Hysitron TI 950 TriboIndenter

gross mechanical testing - MTS Insight II; Eden Prairie, MN; 250 N load cell

in vitro imaging - Nikon Eclipse Ti, Nikon A1R; Zeiss Pascal, Olympus DP70; Ziess LSM 5 Pascal

xray - (AJEX Meditech Ltd; J type stand unit)

microCT - Xradia 520 Versa (Carl Zeiss XRM, Pleasanton, CA, USA) and SCANCO µCT50 (Scanco Medical, Basserdorf, Switzerland)

in vivo Histology images - Axio Scan.Z1 (Zeiss)

Data analysis

mechanical testing - MATLAB (Mathworks, Natick, MA, R2019b)

xray - ImageJ (v 1.53c with Java 1.8.0\_172)

microCT - Dragonfly (version 4.1, Object Research Systems, Montreal, Canada)

in vivo Histology images - ImageJ (v 1.53c with Java 1.8.0\_172) Statistical analysis - GraphPad Prism (v 9.0.1, GraphPad Software)

Other - Adobe illustrator (2022 26.2.1)

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

The data included in this manuscript are available from the corresponding author upon request.	The data include	ed in this manuscrip	ot are available from th	ne corresponding author	upon request.
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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

The determination of treatment group sample size is based on data our lab previously generated on growth plate injuries in rabbits. Outcome measures include bony repair tissue by microCT, varus angulation (degrees), and bone length (cm). Taking the means and standard deviations from our data, we determined group sample size using SISA. With an alpha of 5% and power of 95%.

Data exclusions Indents exceeding 10 MPa were considered outside the soft tissue region of the growth plate and were discarded.

Replication

Randomization

Scaffolds and hydrogels were specifically designed and fabricated for their respective formulations, and thus no randomization was employed. Animals were randomly assigned to treatment groups with roughly equal distributions of males and females being placed in each group. However, due to lack of differences, sex was not ultimately evaluated as a covariate.

Blinding

Where possible samples were denoted as numbers rather than identified by group/treatment name.

### 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				
Antibodies	ChIP-seq				
Eukaryotic cell lines	Flow cytometry				
Palaeontology and archaeology	MRI-based neuroimaging				
Animals and other organisms	·				
Human research participants					
Clinical data					
Dual use research of concern					

#### **Antibodies**

Antibodies used Primary: aggrecan (Abcam, ab3778), collagen type II (Iowa hybridoma bank, CIIC1), collagen x (Abcam, ab49945), PEG (Academia Sinica, 6.3-PABG-A) Secondary: peroxidase conjugated goat anti-mouse IgG (Jackson ImmunoResearch 115-035-166) Goat anti-mouse IgG (H+L) Highly Cross-Absorbed secondary Antibody, Alexa Fluor 546 (Invitrogen, #A11030) Goat anti-mouse IgG (H+L) Cross-Absorbed secondary Antibody, Alexa Fluor 488 (Invitrogen #A-11001) Validation Each antibody was validated according to the manufacturer's protocols.

### Animals and other organisms

Policy information about studies involving animals; ARRIVE guidelines recommended for reporting animal research

Laboratory animals New Zealand White Rabbits, mix of male and female (n=36 and n=27 respectively), initial surgery at 6 weeks old

Wild animals No wild animals were used

Field-collected samples No field-collected samples were used

Ethics oversight All procedures complied with the Guide for the Care and Use of Laboratory Animals and were approved by the Institutional Animal

Care and Use Committee at the University of Colorado Denver.

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