## **Supplementary Information**

## A composite dressing combining ultralong hydroxyapatite nanowire bio-paper and calcium alginate hydrogel accelerates wound healing

Yuankang Zhu<sup>1#</sup>, Liangshi Hao<sup>1#</sup>, Yurui Luo<sup>3</sup>, Jing Gao<sup>1</sup>, Fengming Xu<sup>1</sup>, Han Li<sup>1</sup>,

Changning Hao<sup>1</sup>, Chao-Po Lin<sup>3</sup>, Han-Ping Yu<sup>2\*</sup>, Ying-Jie Zhu<sup>2\*</sup>, Junli Duan<sup>1\*</sup>

<sup>1</sup> Department of Gerontology, Xinhua Hospital affiliated to Shanghai Jiaotong University School of Medicine, Shanghai 200082, P. R. China

<sup>2</sup> State Key Laboratory of High Performance Ceramics and Superfine Microstructure, Shanghai Institute of Ceramics, Chinese Academy of Sciences, Shanghai 200050, P. R. China

<sup>3</sup>School of Life Science and Technology, Shanghai Tech University, Shanghai, 201210,P. R. China

\* Correspondence:

J. L. Duan: duanjunli@xinhuamed.com.cn

Y. J. Zhu: <u>y.j.zhu@mail.sic.ac.cn</u>

H. P. Yu: yuhanping@mail.sic.ac.cn

# These authors contributed equally to this work.

Regents	Molecular weight	Purity	Source
Calcium chloride	110.08	٨P	Sinopharm Chemical
Calcium emoride	Calcium chioride 110.98 AK	AK	Reagent Co., Ltd.
Sodium hydroxide	40.00	ΔR	Sinopharm Chemical
Boardin nyaroxide	-0.00	AIX	Reagent Co., Ltd.
Sodium dihydrogen	156.01	ΔR	Sinopharm Chemical
phosphate dihydrate	150.01	AIX	Reagent Co., Ltd.
Alginic acid sodium	216.121	AR	Sigma-Aldrich
Oleic acid	282.46	ΔR	Aladdin Industrial
	202.40	AIX	Corporation
Mathanal	32.04	≥99.5%	Shanghai Lingfeng
Wethanor			Chemical Reagent Co., Ltd
Fthanol	46.07	>99 7%	Shanghai Lingfeng
Ethanor	40.07	<u>~</u> )).//0	Chemical Reagent Co., Ltd
Fetal bovine serum	/	/	LIFE ILAB BIO
Y-27632	320.3kDa	≥95%	Calbiochem
CHIR99021	465.34kDa	≥98%	<b>Tocris Bioscience</b>
BMP-4	34kDa	>95%	R&D Systems
VEGF-A	38.2kDa	$\geq$ 98%	Peprotech
Matrigel	/	/	Corning
SteamPro34	/	/	Gibco
FGF-2	16.4kDa	97%	Miltenyi Biotec
DAPI	350.25kDa	≥98%	Sigma
anti-human CD31	130kDa	/	DAKO
anti-Phalloidine	469kDa	/	ThermoFisher
anti-CD31	120kDa	/	Servicebio
anti-VEGF-A	32kDa	/	Affinity
anti-p-eNOS Ser1177	133kDa	/	Cell Signaling Technology
anti-AKT	56kDa	/	Cell Signaling Technology
anti-p-AKT Ser473	56kDa	/	Abclonal
anti-β-actin	42kDa	/	Abclonal

**Supplementary Table S1** List of detailed information for chemicals and reagents used in the experiments

Primer	Forward	Reverse	
S	rorwaru	KUVU SU	
Col-Ia	TGTTGGTCCTGCTGGCAAGAA	GTCACCTTGTTCGCCTGTCTCAC	
	TG	STEACETISTICOCCIOTETCAC	
Col-	AGTCGGAGGAATGGGTGGCT	CAGGAGATCCAGGATGTCCAG	
IIIa	ATC	AGG	
β-actin	GCAGGAGTACGATGAGTCCG	ACGCAGCTCAGTAACAGTCC	

Supplementary Table S2 Primer sequence



**Supplementary Figure S1.** Density and porosity of the HAP bio-paper consisting of ultralong HAP nanowires.



**Supplementary Figure S2.** Weights of the HAP bio-paper after water absorption and subsequently dried under the ambient environment.



Supplementary Figure S3. TG curves of the freeze-dried HAP-Alg<sub>x</sub> composite dressings and the HAP bio-paper.



Supplementary Figure S4. Bright-field images of H1 differentiation into blood vessel organoids on day 8 indicate that the aggregates tend to form extensive radial sprouting, especially under the stimulation of the HAP-Alg<sub>x</sub> composite dressing. The control group was treated without the HAP-Alg<sub>x</sub> composite dressing. Scale bar =  $400 \mu m$ .



**Supplementary Figure S5. Human vascular organoids (BVOs) with different samples.** (A) Number of master junctions of BVOs. (B) Number of master segments of BVOs. (C) Number of branches of BVOs. (D) Total branching length of BVOs. (E) Relative CD31 fluorescence intensity of BVOs to the control. Significant p-values are indicated: \*p<0.05, \*\*p<0.01 and #p<0.001 compared with the control group.



Supplementary Figure S6. Flou-4 AM and MitoSOX red staining. (A) Representative images of Flou-4 AM (Green), MitoSox Red (Red) and Hoechst (Blue). (B) Quantification of Fluo-4 AM fluorescence intensity. (C) Quantification of MitoSox Red fluorescence intensity. Significant p-values are indicated: p<0.05, p<0.01 compared with the control group. Scale bar = 5  $\mu$ m.