

Supporting Information

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Metal-Organic Framework Functionalized Bioceramic Scaffolds with Antioxidative Activity for Enhanced Osteochondral Regeneration

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Supporting Information

Metal-organic Framework (MOF) Functionalized Bioceramic Scaffolds with Antioxidative Activity for Enhanced Osteochondral Regeneration

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Figure S1. (a) Digital photo of the larger-sized 17MOF-TCP scaffold. Vertical-section view (b)

and cross-section view (c) of the 3D reconstructed scaffold.



Figure S2. (a) The XRD patterns and (b) the compressive strength of the scaffolds functionalized with different concentrations of Zn/Co-MOF reaction solution (n=8). *p < 0.05



Figure S3. Element distribution on the surface of 17MOF-TCP scaffolds.



Figure S4. The release profiles of (a) Ca ions, (b) P ions, (c) Zn ions, and (d) Co ions from MOF-TCP scaffolds in Tris–HCl solution (n=5).



Figure S5. The color changes of ABTS free radicals after reacted with the β -TCP and MOF-TCP

scaffolds.



Figure S6. Multiple ROS-scavenging activities of MOF-TCP scaffolds. The absorbance spectra of (a) ABTS⁺, (b) DPPH⁻, (c) ONOO⁻, (d) O_2^{--} , and (e) H_2O_2 after treated with the β -TCP and MOF-TCP scaffolds. (f) O_2 produced from the H_2O_2 solution after treated with the β -TCP and MOF-TCP scaffolds.



Figure S7. The number of cells that adhered to different scaffolds with H_2O_2 stimulation (n=3). *p < 0.05, **p < 0.01, ***p < 0.001.



Figure S8. The promoting effects of MOF-TCP scaffolds on cell migration under oxidative stress. (a) The migration images of rBMSCs cultured with different scaffolds for 24 h under H_2O_2 stimulation. (b) Quantitative results of migration ratio of rBMSCs (n=4). (c) The migration images of chondrocytes cultured with different scaffolds for 24 h H_2O_2 stimulation. (d) Quantitative results of migration ratio of chondrocytes (n=4). *p < 0.05, **p < 0.01, ***p < 0.001.



Figure S9. Proliferation of (a) rBMSCs and (b) Chondrocytes cultured on the scaffolds under normal conditions (n=6). (c) Morphology of rBMSCs and Chondrocytes cultured on the scaffolds under normal conditions.



Figure S10. The semi-quantitative statistics of the fluorescence intensity of aggrecan protein in chondrocytes (n=4). *p < 0.05, **p < 0.01, ***p < 0.001.



Figure S11. ROS fluorescence images in rBMSCs after being treated with different types of

ROS and scaffolds.



Figure S12. ROS fluorescence images in chondrocytes after being treated with different types of ROS and scaffolds.



Figure S13. ROS fluorescence images in RAW 264.7 cells after being treated with different types of ROS and scaffolds.



Figure S14. Safranine O-fast green staining images and Van Gieson staining images of the osteochondral defects after 12 weeks of implantation.



Figure S15. Quantitative analysis of the Safranine-O sections based on the O'Driscoll grading system (n=10). **p < 0.01, ***p < 0.001.

| | A solution+H ₂ O/mL | B solution+H ₂ O/mL |
|-----------|--------------------------------|--------------------------------|
| 6MOF-TCP | 6.25+43.75 | 0.63+4.37 |
| 12MOF-TCP | 12.50+37.50 | 1.25+3.75 |
| 17MOF-TCP | 17.00+33.00 | 1.70+3.30 |
| 21MOF-TCP | 21.00+29.00 | 2.10+2.90 |
| 25MOF-TCP | 25.00+25.00 | 2.50+2.50 |
| 50MOF-TCP | 50.00+0.00 | 5.00+0.00 |

Table S1. Proportions of Zn/Co-MOF reaction solutions for different MOF-TCP scaffolds

Table S2. The primer sequences of osteogenic and chondrogenic genes used for RT-qPCR

| Gene | Forward primer | Reverse primer |
|----------|-----------------------|------------------------|
| GAPDH | TCACCATCTTCCAGGAGCGA | CACAATGCCGAAGTGGTCGT |
| OCN | CCGGGAGCAGTGTGAGCTTA | AGGCGGTCTTCAAGCCATACT |
| OPN | CACCATGAGAATCGCCGT | CGTGACTTTGGGTTTCTACGC |
| BMP2 | CGCCTCAAATCCAGCTGTAAG | GGGCCACAATCCAGTCGTT |
| RUNX2 | TCAGGCATGTCCCTCGGTAT | TGGCAGGTAGGTATGGTAGTGG |
| SOX9 | GGTGCTCAAGGGCTACGACT | GGGTGGTCTTTCTTGTGCTG |
| Aggrecan | AGGTCGTGGTGAAAGGTGTTG | GTAGGTTCTCACGCCAGGGA |
| COL- II | AACACTGCCAACGTCCAGAT | CTGCAGCACGGTATAGGTGA |

Table S3. The primer sequences of pro-inflammatory genes in chondrocytes used for RT-

qPCR

| Gene | Forward primer | Reverse primer |
|-------|------------------------|-------------------------|
| GAPDH | TCACCATCTTCCAGGAGCGA | CACAATGCCGAAGTGGTCGT |
| IL-1β | CAGGACCTGGACCTCTGCTGTC | GAGCCACAACGACTGACAAGACC |
| IL-6 | GAAAACACCAGGGTCAGCAT | CAGCCACTGGTTTTTCTGCT |
| TNF-α | CTCCTACCCGAACAAGGTCA | CGGTCACCCTTCTCCAACT |

Table S4. The primer sequences of anti-inflammatory and pro-inflammatory genes in

macrophages used for RT-qPCR

| GAPDH | AGAACATCATCCCTGCATCCAC | TCAGATCCACGACGGACACA |
|-------|--------------------------|--------------------------|
| IL-10 | GAGAAGCATGGCCCAGAAATC | GAGAAATCGATGACAGCGCC |
| Arg-1 | AACCTTGGCTTGCTTCGGAACTC | GTTCTGTCTGCTTTGCTGTGATGC |
| IL-1β | CTACCTGTGTCTTTCCCGTG | TTTGTTGTTCATCTCGGAGC |
| IL-6 | ATAGTCCTTCCTACCCCAATTTCC | GATGAATTGGATGGTCTTGGTCC |