# Supplementary Information

## A periodontal tissue regeneration strategy via biphasic

## release of zeolitic imidazolate framework-8 and FK506

## using a uniaxial electrospun Janus nanofiber

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#### Methods

#### **Preparation of ZIF-8 nanoparticles**

ZIF-8 nanoparticles (ZIF-8 NPs) was carried out at room temperature as reported, previously.<sup>1</sup> In brief, 1.17 g Zn (NO)<sub>3</sub>·6H<sub>2</sub>O, dissolved in 8 g of deionized (DI) water was added to 22.7 g 2-methylimidazole dissolved in 80 g DI water. The reaction solution was stirred at room temperature for ~ 5 min. After that, the produce was isolated via centrifugation, washed with deionized DI water and dried in a vacuum oven. The morphology of ZIF-8 NPs was obtained by transmission electron microscopy (TEM, JOEL JEM-201). The information on the crystalline phase structure of ZIF-8 NPs was obtained by X-ray diffractometer (XRD, SmartLab SE). The functional groups of ZIF-8 NPs were measured by Fourier transform infrared spectrometer (FTIR, FTIR-650).

#### Results

#### **Characterization of ZIF-8 NPs**

The morphology of ZIF-8 NPs was rhombic dodecahedral in shape and the size distribution was 110.58±9.39 nm (Fig. S1-S2). The XRD pattern of the synthesized ZIF-8 NPs were in good agreement with the simulated crystal image, suggesting a pure phase of the synthesized ZIF-8 NPs (Fig. S3). The FT-IR spectrum showed the peak positions and their assignments agreed well with those reported in the literature (Fig. S4).<sup>2</sup> These results showed the successful synthesis of ZIF-8 NPs and can be used without further purification.

### Figures



Fig. S1. TEM images of ZIF-8 NPs.



Fig. S2. Size distribution of the ZIF-8 NPs.



Fig. S3. XRD patterns of the ZIF-8 NPs.



Fig. S4. FT-IR spectra of the ZIF-8 NPs.



Fig. S5. The phase separation phenomenon in precursor solution.



Fig. S6. Proliferation of BMSCs on PP Janus nanofibers and PPZ Janus nanofibers with different concentrations of ZIF-8 NPs. Data are presented as means  $\pm$  SD. \* *P* < 0.05, \*\* *P* < 0.01, \*\*\* *P* < 0.001.



Fig. S7. Proliferation of BMSCs on PP Janus nanofibers, PPZ Janus nanofibers, and PPZF Janus nanofibers with different concentrations of FK506. Data are presented as means  $\pm$  SD. \* *P* < 0.05, \*\* *P* < 0.01, \*\*\* *P* < 0.001.



Fig. S8. H&E staining of vital organs including hearts, livers, spleens, lung and kidneys. Scale bar is 10  $\mu$ m.

### Table

primer	sequence (5' to 3')
rat- <i>8-actin</i> -F	GGAGATTACTGCCCTGGCTCCTA
rat- <i>θ-actin</i> -R	GACTCATCGTACTCCTGCTTGCTG
rat- <i>Runx2</i> -F	CATGGCCGGGAATGATGAG
rat- <i>Runx2</i> -R	TGTGAAGACCGTTATGGTCAAAGTG
rat-Osx-R	CATCCATGCAGGCATCTCA
rat- <i>Osx</i> -R	CTGCCCACCACCTAACCAA
rat- <i>Alp</i> -F	CATCGCCTATCAGCTAATGCACA
rat- <i>Alp</i> -R	ATGAGGTCCAGGCCATCCAG
rat- <i>Ocn</i> -F	TTATTGTTTGAGGGGCCTGGG
rat- <i>Ocn</i> -R	TGCTCCTACAAAGCTGTCTCC

**Table S1.** Primer Sequences of the Genes Involved in this Study

### References

1 Y. C. Pan, Y. Y. Liu, G. F. Zeng, L. Zhao, Z. P. Lai, *Chem. Commun.*, 2011, **47**, 2071-2073.

2 S. R. Venna, M. A. Carreon, J. Am. Chem. Soc., 2010, 132, 76-78.