

# Supporting Information

## Design and Engineering of Biomimetic Aloe vera Sponges via Recombination of Functionalized Peel and Gel for Enhanced Wound Healing

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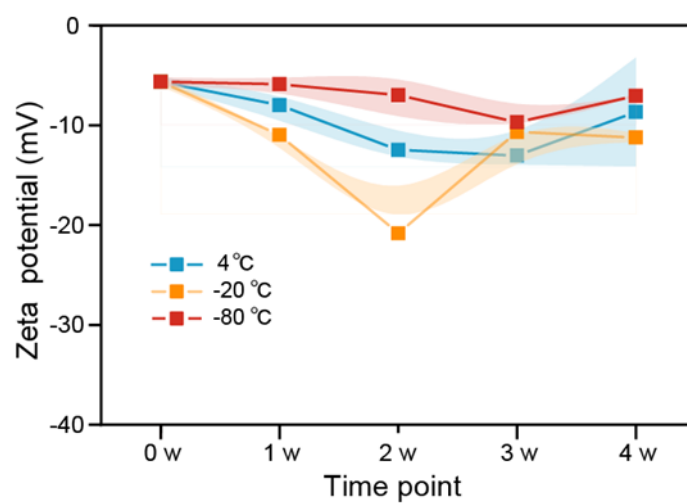
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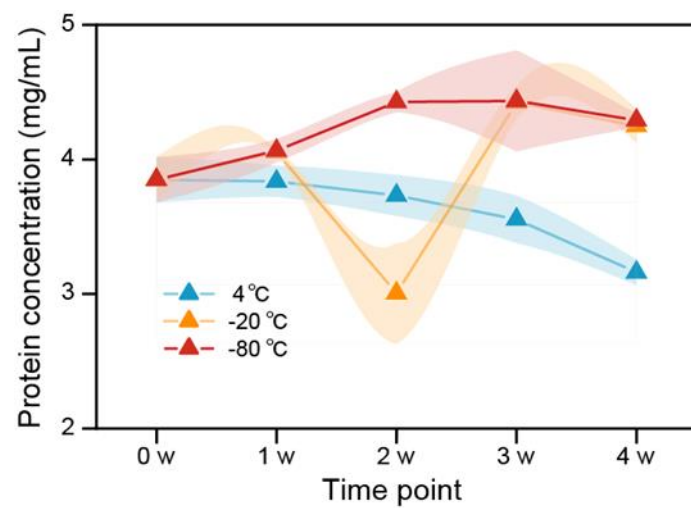
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(X. Zhang).*

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## 1. Stability evaluation of APNs

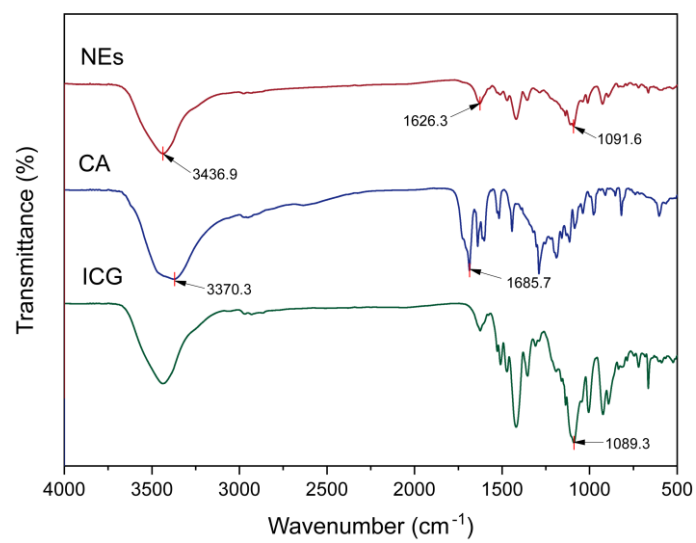


**Fig. S1.** Zeta potential of APNs under different storage temperatures (4, -20 and -80 °C) for 4 weeks (n = 3).



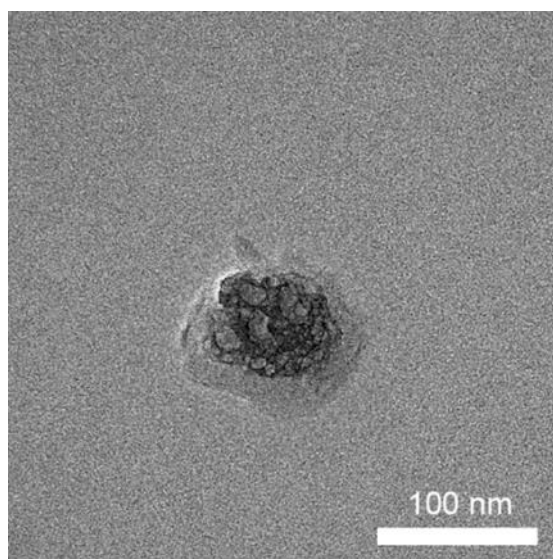
**Fig. S2.** Protein content of APNs under different storage temperatures (4, -20 and -80 °C) for 4 weeks (n = 3).

## 2. FTIR spectrum of NEs



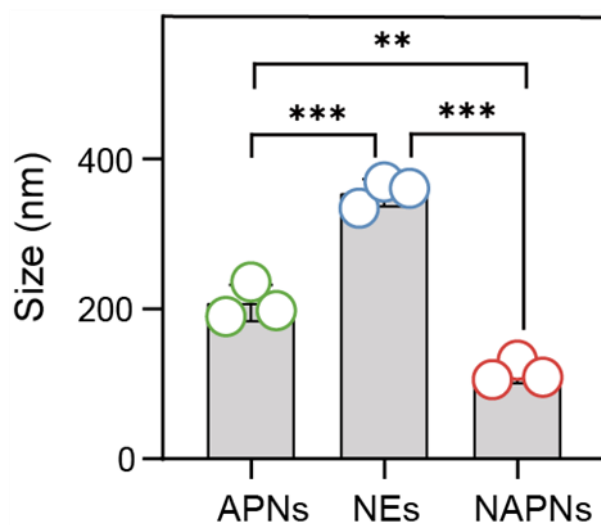
**Fig. S3.** FTIR spectra of CA, ICG and NEs.

### 3. TEM of NAPNs

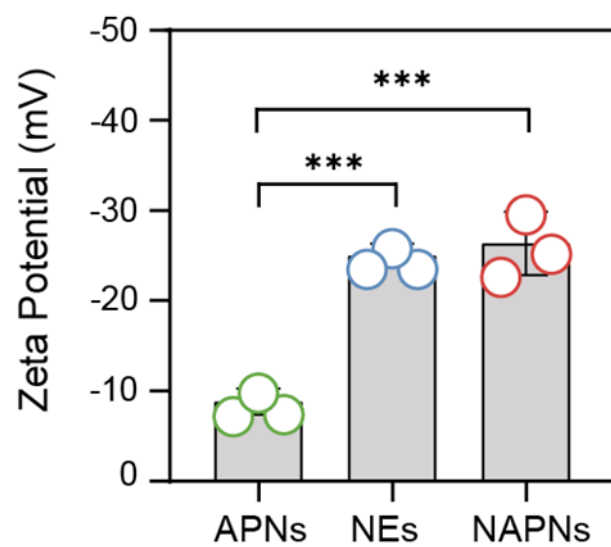


**Fig. S4.** The membrane structure and black contents of NAPNs. Scale bar: 100 nm.

#### 4. Characterization of particle size and zeta potential

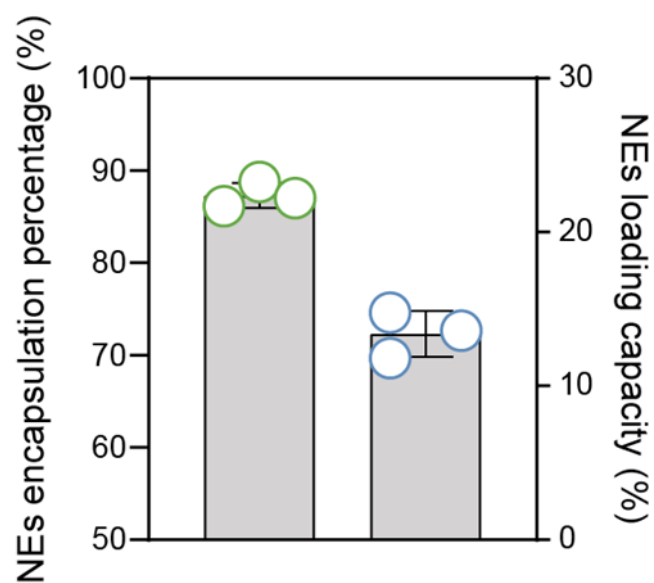


**Fig. S5.** Size distribution of APNs, NEs and NAPNs ( $n = 3$ ).  $**p < 0.01$ ,  $***p < 0.001$ .



**Fig. S6.** Zeta Potential of APNs, NEs and NAPNs (n = 3). \*\*\* $p < 0.001$ .

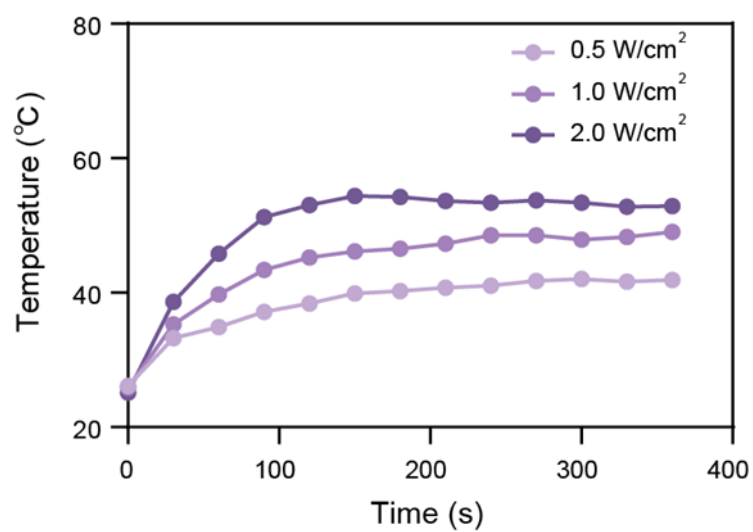
## 5. Encapsulation efficiency and drug loading of NAPNs



**Fig. S7.** Encapsulation efficiency and loading capacity of NAPNs (n = 3).

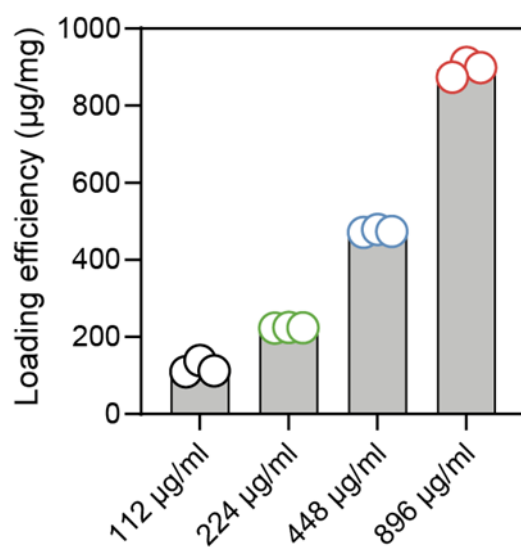


## 6. Temperature variations at different powers of NAPNs



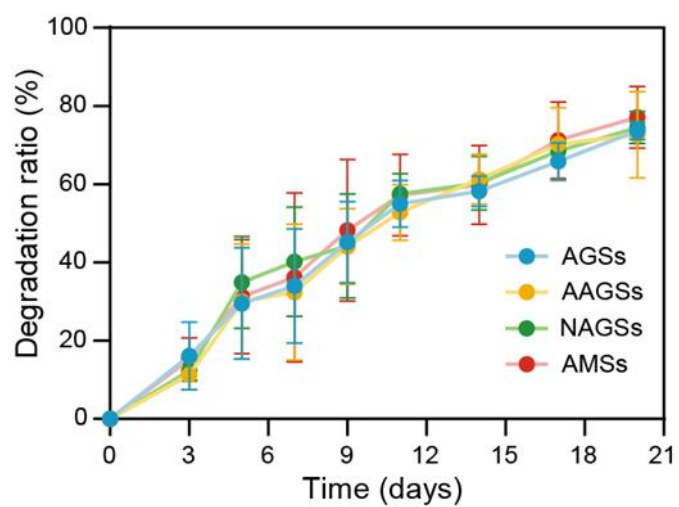
**Fig. S8.** Photothermal heating curves of NAPNs under different laser powers.

## 7. The loading efficiency



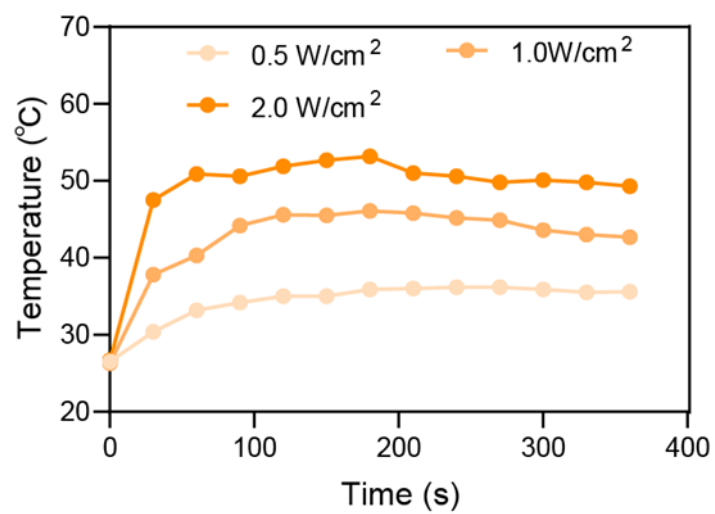
**Fig. S9.** The loading efficiency of AGSs for NAPNs (n = 3).

## 8. *In vitro* degradation



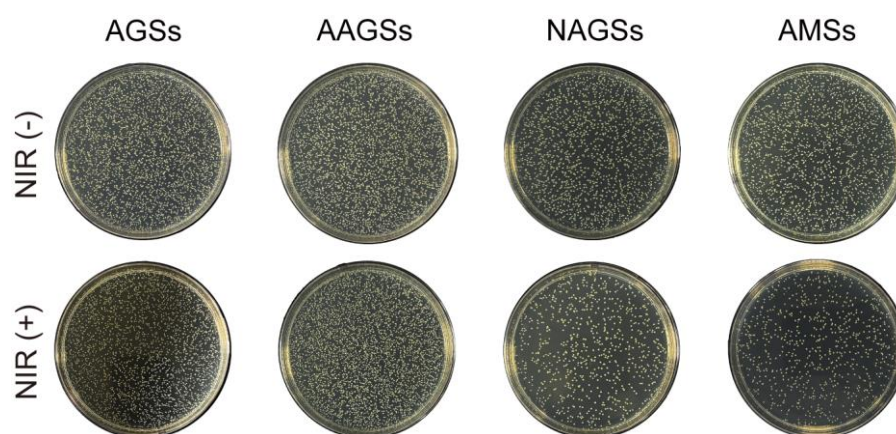
**Fig. S10.** *In vitro* degradation of the sponges in PBS over 21 days (n = 3).

## 9. Temperature changes under different power levels of AMSs

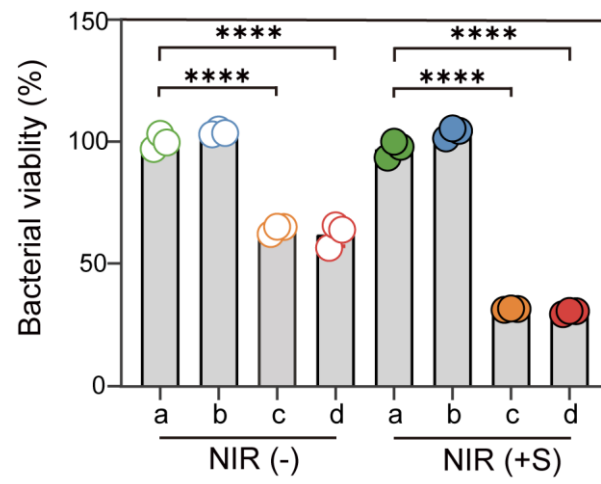


**Fig. S11.** Photothermal heating curves of AMSs under different laser powers.

## 10. Anti MRSA effects of AMSs in Vitro

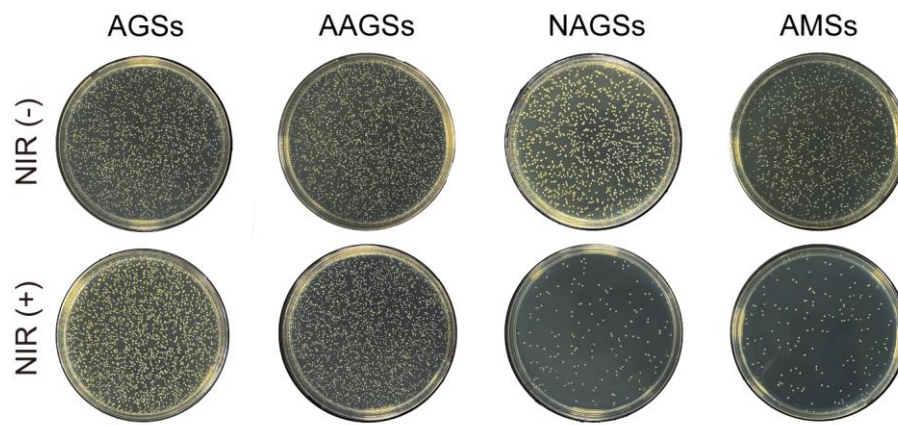


**Fig. S12.** Agar plate images showing MRSA growth following different treatments.

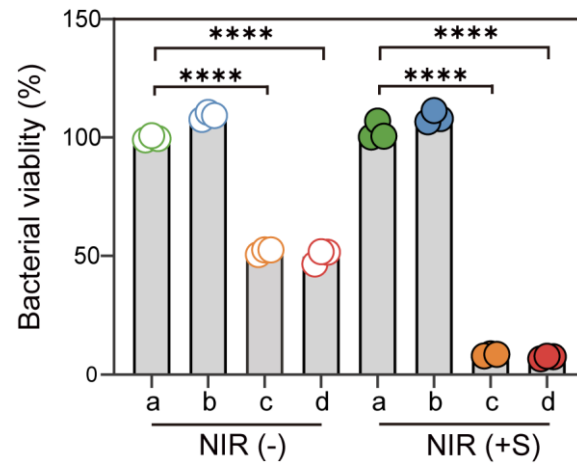


**Fig. S13.** Corresponding quantitative analysis of bacterial colonies. The groups are labeled as follows: a - AGSs, b - AAGSs, c - NAGSs, d - AMSs. Data were presented as mean  $\pm$  s.d.

\*\*\*\* $p < 0.0001$ .

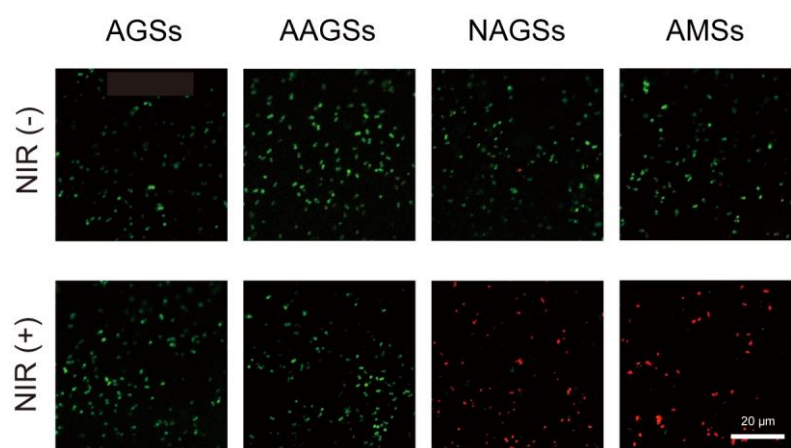


**Fig. S14.** Agar plate images of MRSA after treatments with NO donor addition.

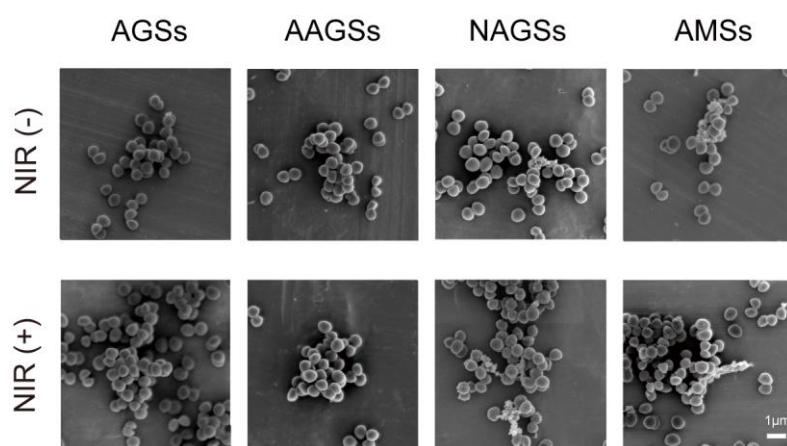


**Fig. S15.** Corresponding quantitative analysis of bacterial reduction ( $n = 3$ ). The groups are labeled as follows: a - AGSs, b - AAGSs, c - NAGSs, d - AMSs. Data were presented as mean  $\pm$  s.d. \*\*\*\* $p < 0.0001$ .



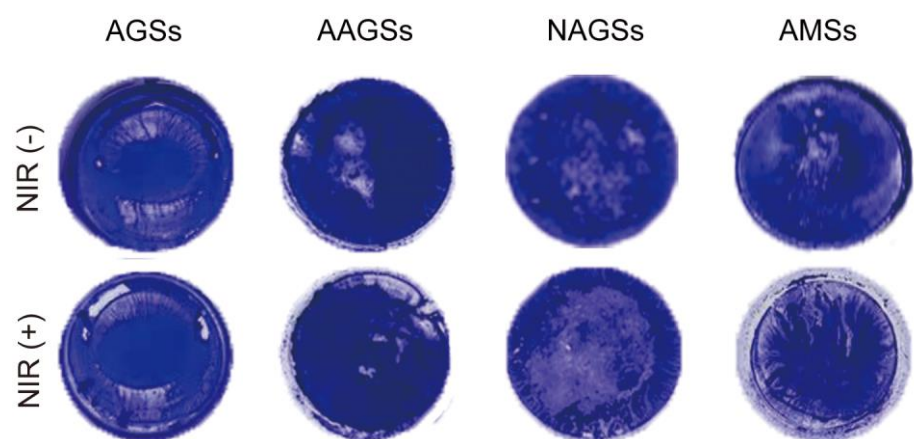


**Fig. S16.** Confocal fluorescence images of MRSA stained with Calcein-AM/PI under different treatment conditions. Scale bar: 20  $\mu\text{m}$ .

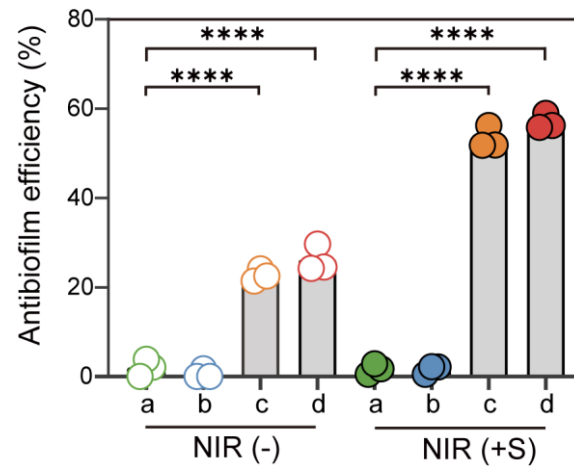


**Fig. S17.** SEM images of MRSA treated with various sponges, before and after laser irradiation.

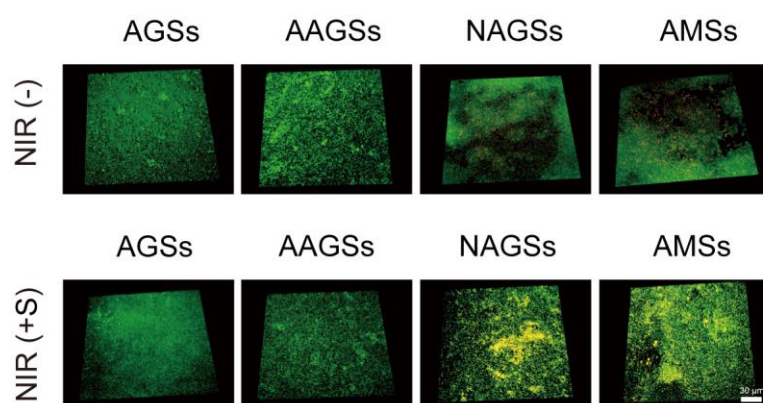
Scale bar: 1  $\mu\text{m}$ .



**Fig. S18.** Crystal violet-stained images of MRSA biofilms after different treatments.

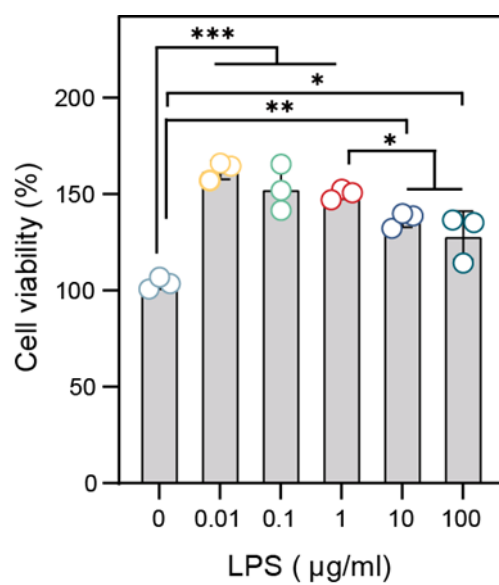


**Fig. S19.** Quantitative analysis of antibiofilm efficiency across treatments ( $n = 3$ ). The groups are labeled as follows: a - AGSs, b - AAGSs, c - NAGSs, d - AMSs. Data were presented as mean  $\pm$  s.d. \*\*\*\* $p < 0.0001$ .

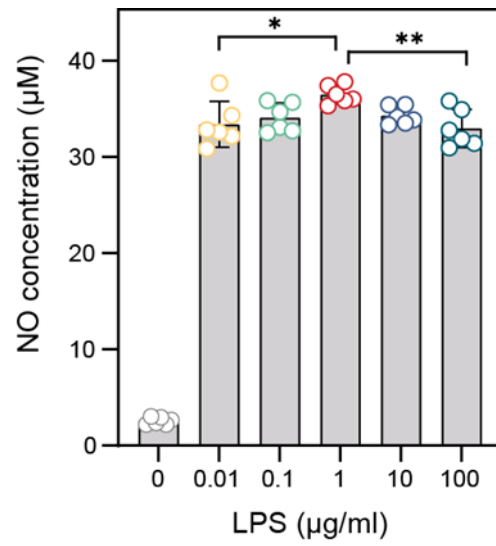


**Fig. S20.** 3D confocal fluorescence images of MRSA biofilm stained with Calcein-AM/PI under different treatment conditions. Scale bar: 30 μm.

## 11. Screening of LPS concentration

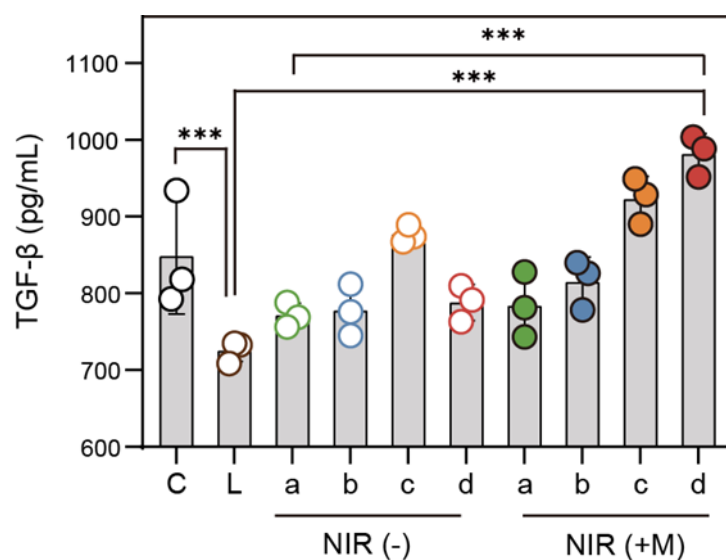


**Fig. S21.** The effect of LPS concentrations on RAW 264.7 (n = 3). \* $p < 0.05$ , \*\* $p < 0.01$ , \*\*\* $p < 0.001$ .



**Fig. S22.** NO production after stimulation with different concentrations of LPS (n = 6). \* $p < 0.05$ , \*\* $p < 0.01$ .

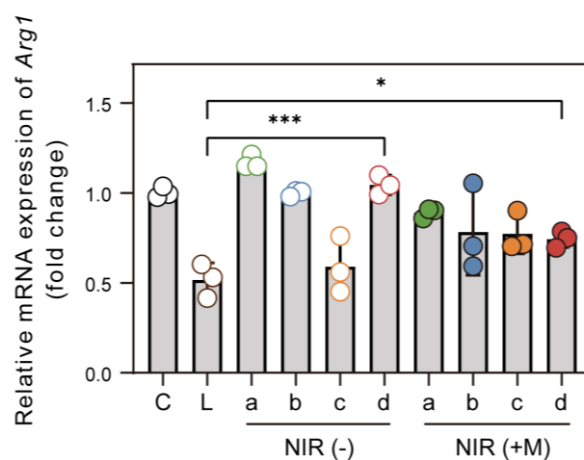
## 12. Quantitative analysis of TGF- $\beta$



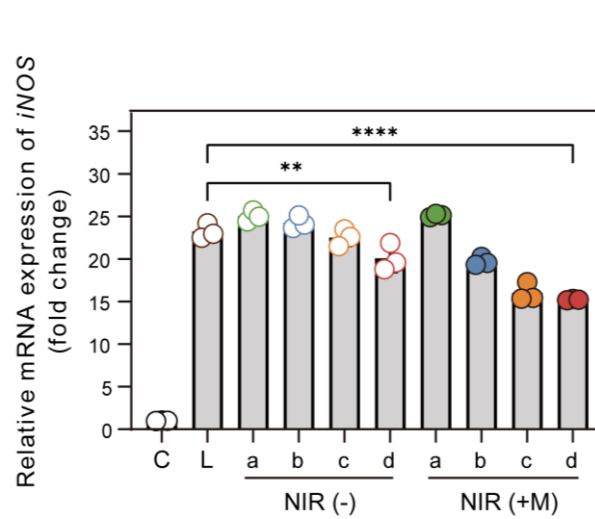
**Fig. S23.** Levels of TGF- $\beta$  in LPS-induced RAW 264.7 cells after treatments. The letters C, L, a, b, c and d represent the control group and the experimental groups treated with LPS, AGSs, AAGSs, NAGSs and AMSs, respectively ( $n = 3$ ). \*\*\* $p < 0.001$ .



### 13. RT-qPCR analysis

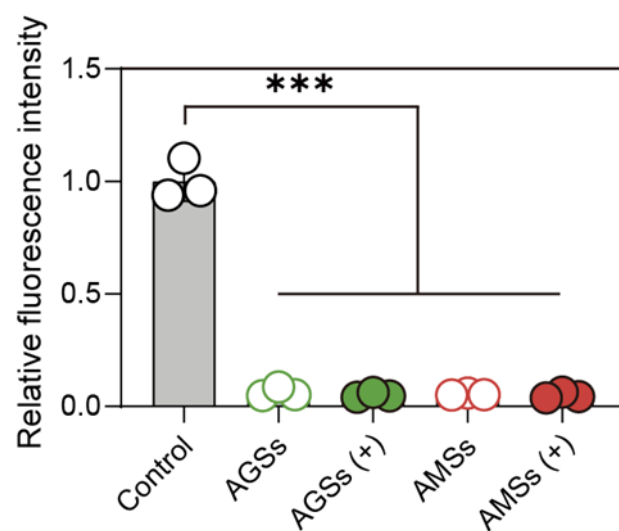


**Fig. S24.** The mRNA expression levels of *Arg1* in Raw 264.7 cells detected by RT-qPCR. Groups are labeled as follows: C - Control group, L - LPS only group, a - AGSs, b - AAGSs, c - NAGSs, d - AMSs. Data were presented as mean  $\pm$  s.d. \* $p < 0.05$ , \*\*\* $p < 0.001$ .

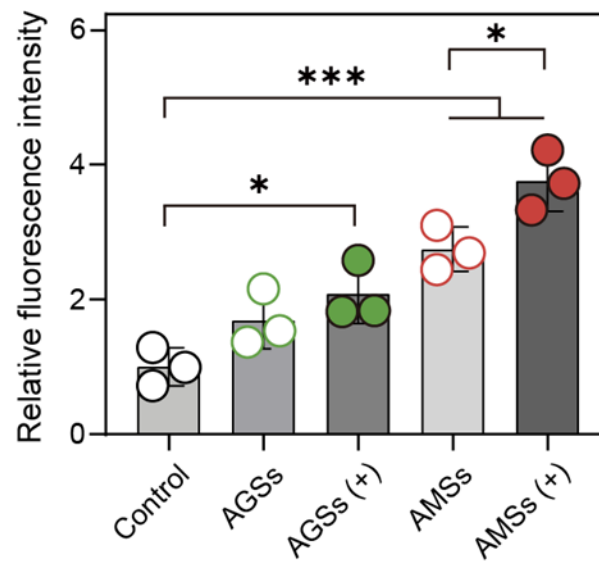


**Fig. S25.** The mRNA expression levels of *iNOS* in Raw 264.7 cells detected by RT-qPCR. Groups are labeled as follows: C - Control group, L - LPS only group, a - AGSs, b - AAGSs, c - NAGSs, d - AMSs. Data were presented as mean  $\pm$  s.d.  $**p < 0.01$ ,  $****p < 0.0001$ .

## 14. Quantitative analysis of immunofluorescence

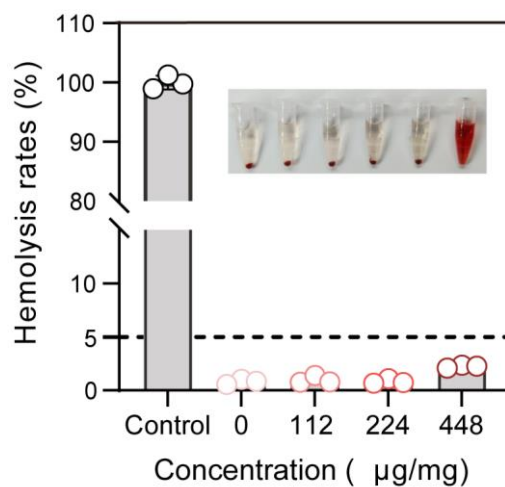


**Fig. S26.** Corresponding quantitative analysis of TNF- $\alpha$  in wound tissues ( $n = 3$ ). \*\*\* $p < 0.001$ .

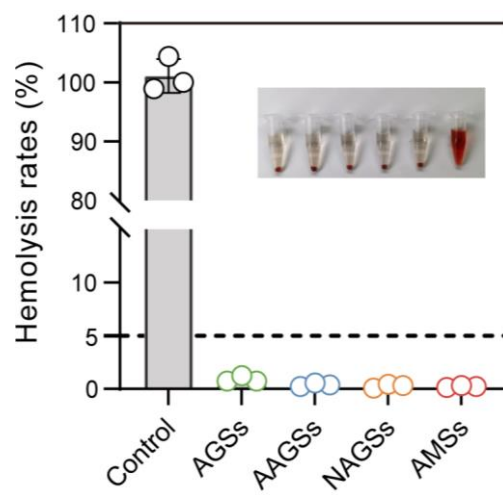


**Fig. S27.** Corresponding quantitative analysis of TGF- $\beta$  in wound tissues ( $n = 3$ ).  $*p < 0.05$ ,  $***p < 0.001$ .

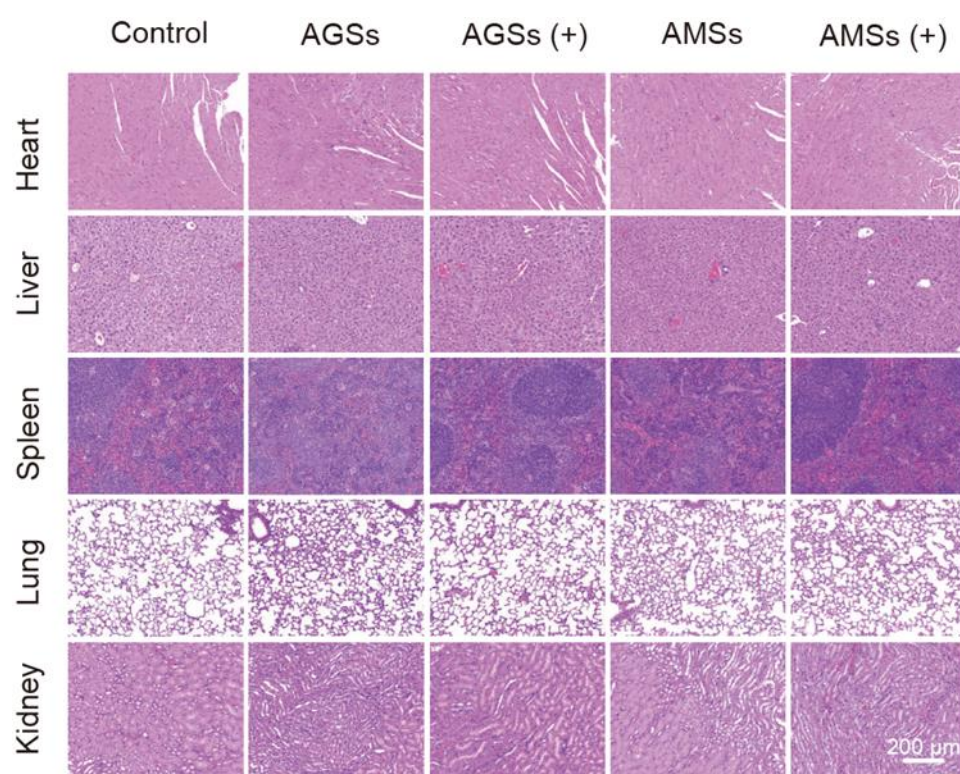
## 15. In vivo evaluation of biosafety of different sponges



**Fig. S28.** *In vitro* hemolysis results of different concentration of AMSs. The inserted image shows hemolysis of PBS, 0 µg/mg of AMSs, 112 µg/mg of AMSs, 224 µg/mg of AMSs, 448 µg/mg of AMSs and Water (from left to right).

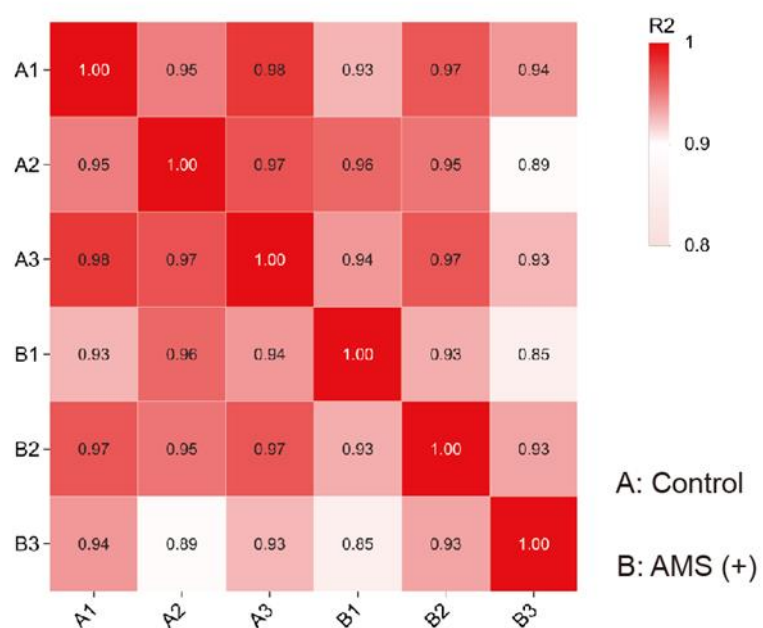


**Fig. S29.** *In vitro* hemolysis results of different sponges. The inserted image shows hemolysis of PBS, AGSs, AAGSs, NAGSs, AMSs and Water (from left to right).



**Fig. S30.** H&E staining images of major organs. Scale bar: 200  $\mu$ m.

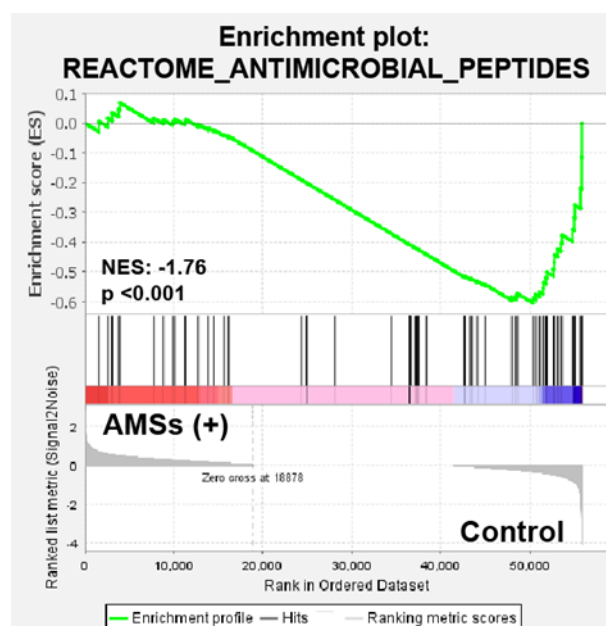
## 16. Transcriptome sequencing results of skin wound tissues



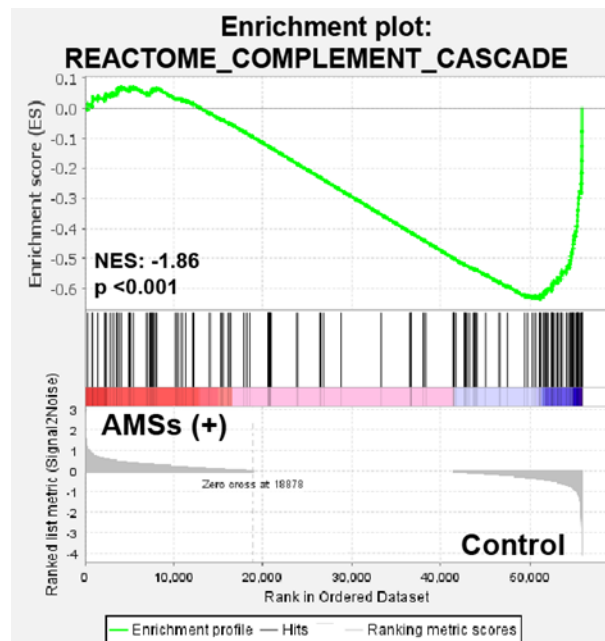
**Fig. S31.** The intergroup correlation of samples. Group A was the control group, Group B was the AMS (+) group.



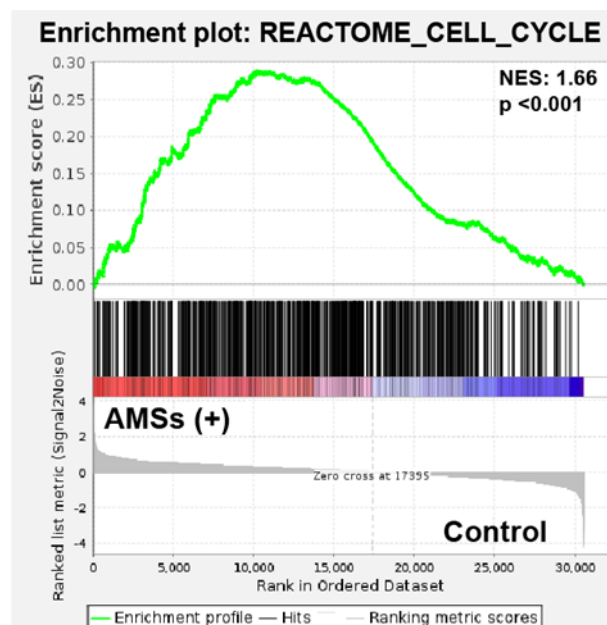
## 17. GSEA analysis of gene sets



**Fig. S32.** GSEA analysis showing the enriched pathway: antimicrobial peptides.

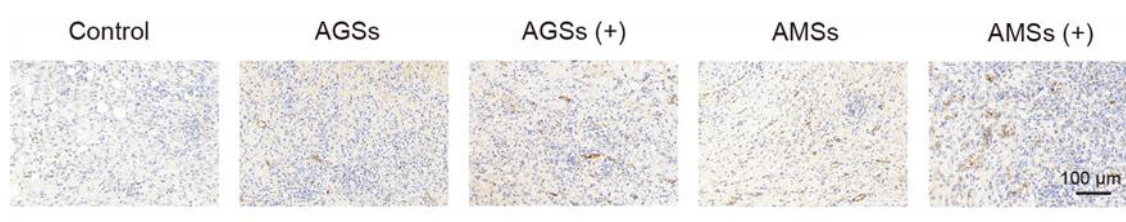


**Fig. S33.** GSEA analysis showing the enriched pathway: complement cascade.

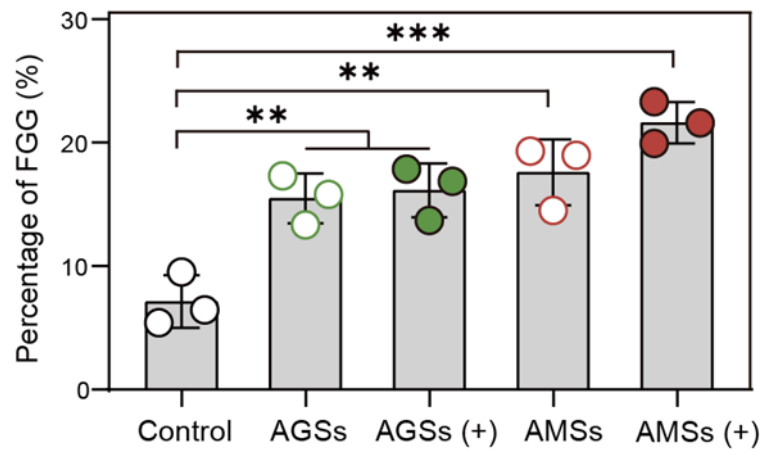


**Fig. S34.** GSEA analysis showing the enriched pathway: cell cycle.

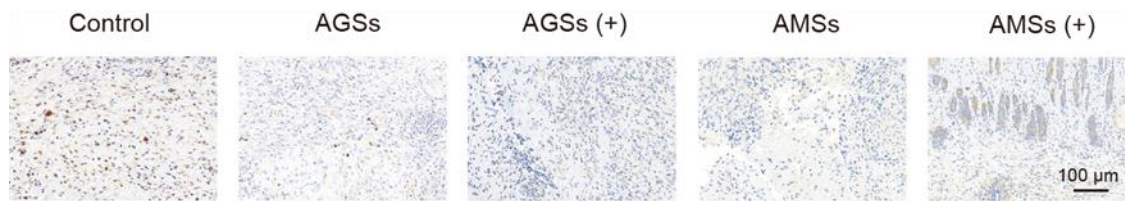
## 18. Images and quantitative analysis of immunohistochemical staining



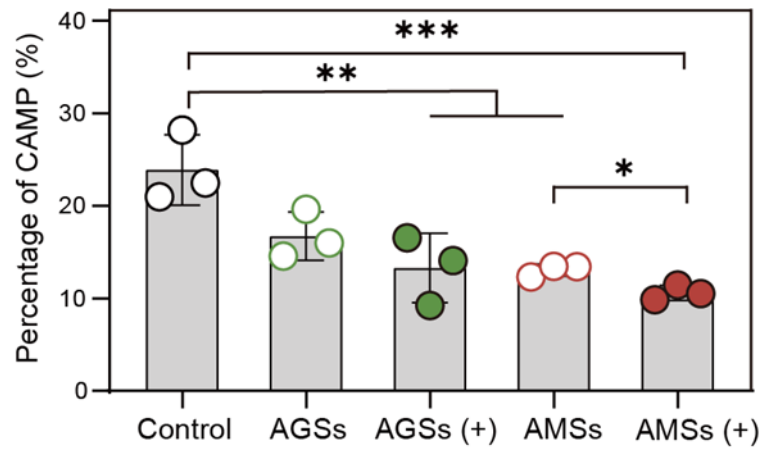
**Fig. S35.** Immunohistochemical diagram for FGG of infected skin wounds on day 10. Scale bar: 100  $\mu\text{m}$ .



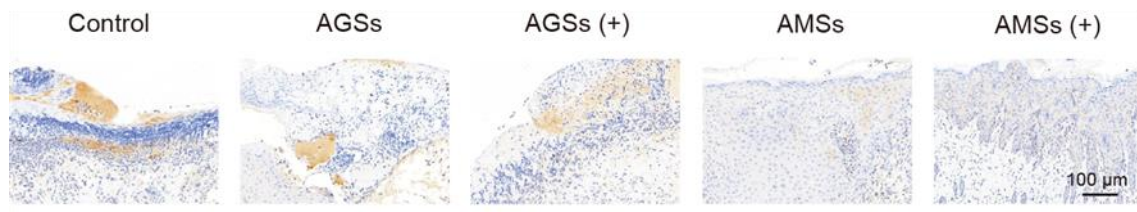
**Fig. S36.** Quantitative analysis of immunohistochemical staining for FG in wound tissues (n = 3). \*\* $p < 0.01$ , \*\*\* $p < 0.001$ .



**Fig. S37.** Immunohistochemical diagram for CAMP of infected skin wounds on day 10. Scale bar: 100 µm.

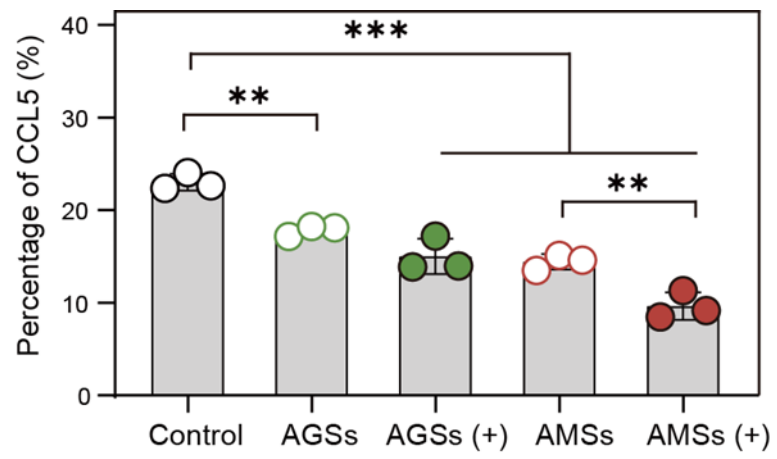


**Fig. S38.** Quantitative analysis of immunohistochemical staining for CAMP in wound tissues (n = 3). \* $p < 0.05$ , \*\* $p < 0.01$ , \*\*\* $p < 0.001$ .

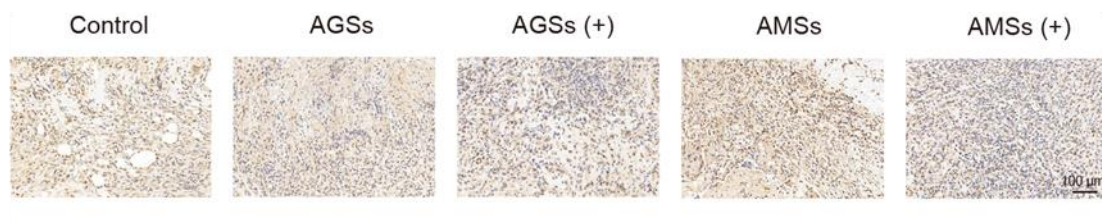


**Fig. S39.** Immunohistochemical diagram for CCL5 of infected skin wounds on day 10. Scale bar: 100  $\mu\text{m}$ .



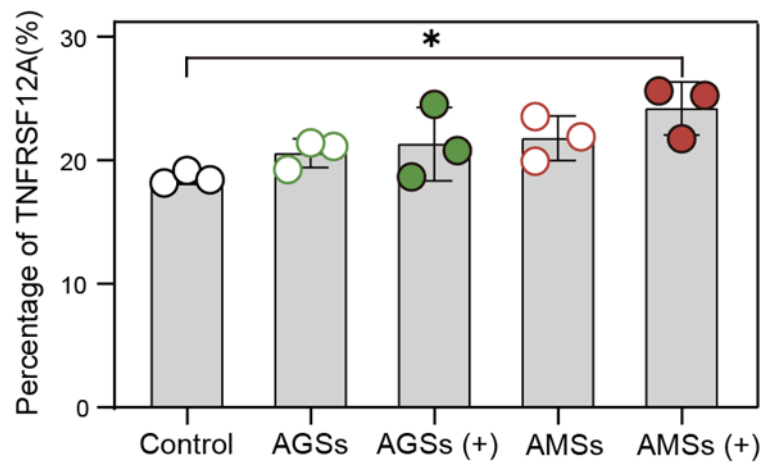


**Fig. S40.** Quantitative analysis of immunohistochemical staining for CCL5 in wound tissues (n = 3). \*\* $p < 0.01$ , \*\*\* $p < 0.001$ .



**Fig. S41.** Immunohistochemical diagram for TNFRSF12A of infected skin wounds on day 10.

Scale bar: 100  $\mu\text{m}$ .



**Fig. S42.** Quantitative analysis of immunohistochemical staining for TNFRSF12A in wound tissues (n = 3). \* $p < 0.05$ .