

## Supplementary data

### **Antibacterial activities against clinical isolates and *in vivo* efficacy of coralmycins**

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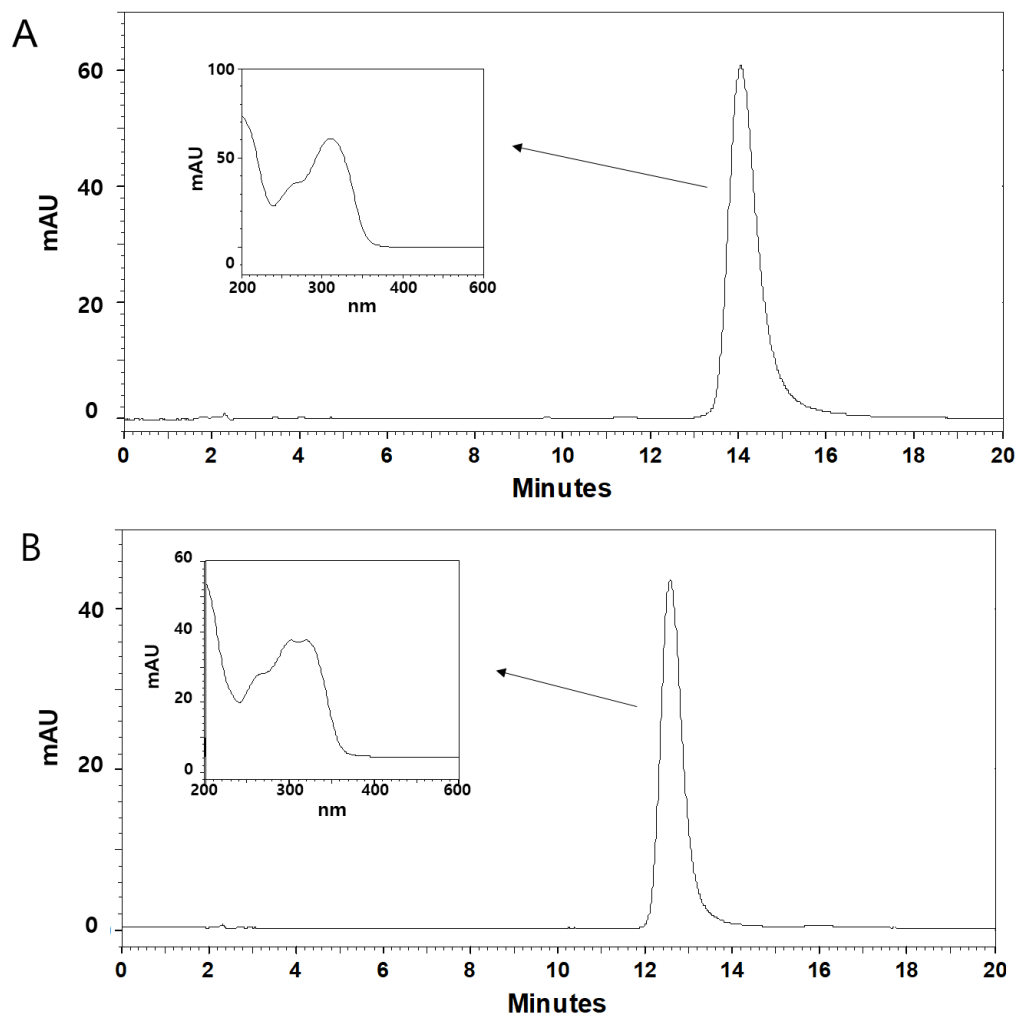
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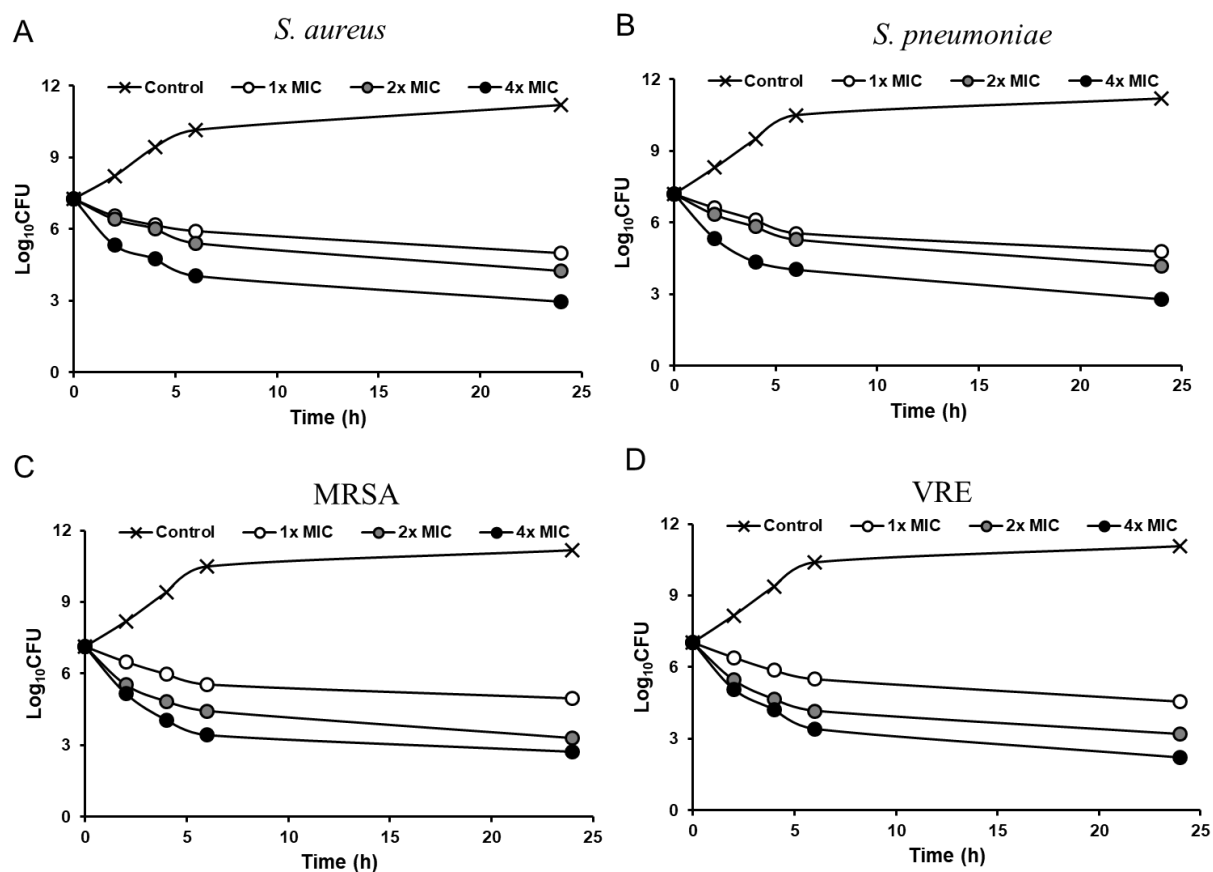
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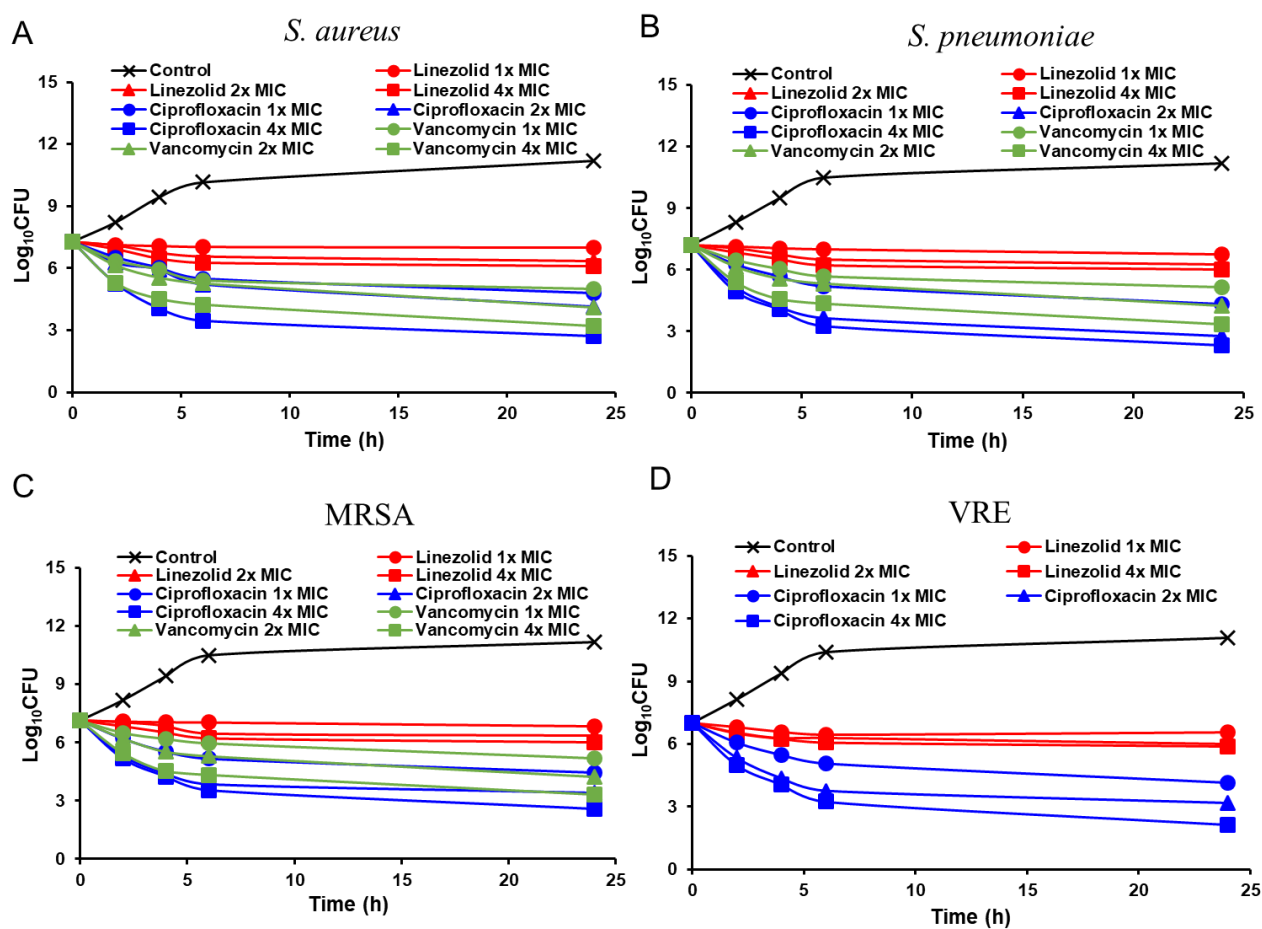
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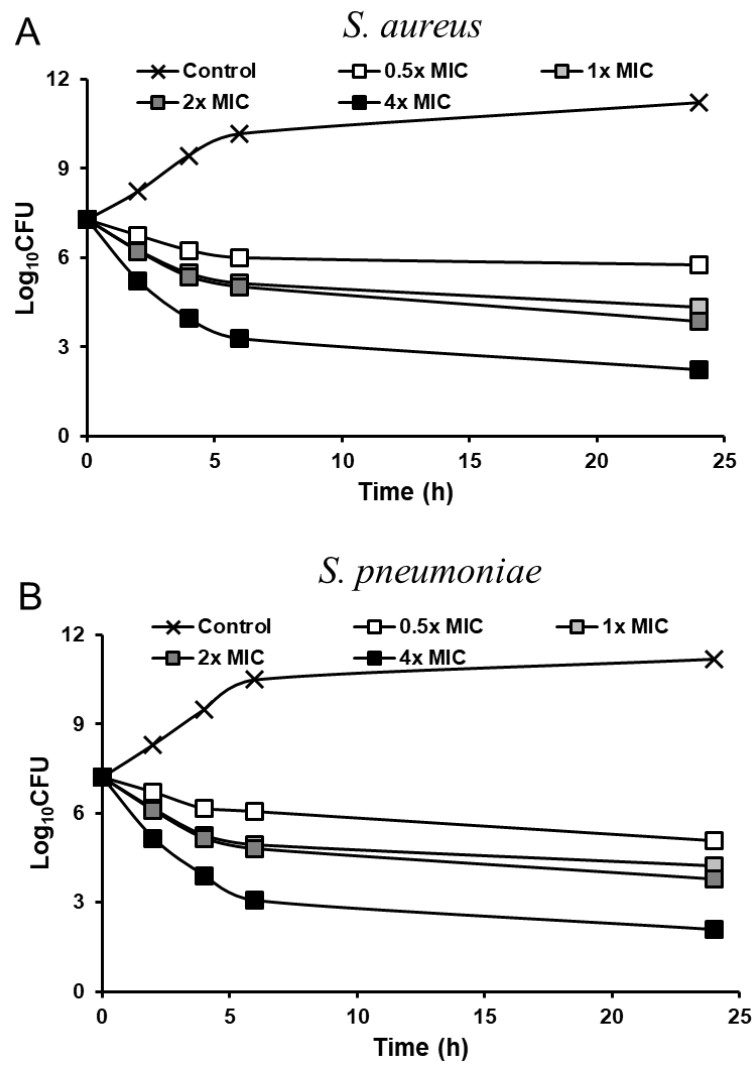
**Figure S1.** Purity test of coralmycin A and DH-coralmycin A isolated from a large-scale *C. coralloides* M23 culture. The coralmycin A (A) and DH-coralmycin A (B) purities were determined to be all over 95% at 225 nm using an analytical HPLC column (J'sphere ODS-H80,  $4.6 \times 150$  mm, S-4  $\mu$ m, YMC) with  $\text{CH}_3\text{CN}:\text{H}_2\text{O}$  (45:55) containing 0.01% TFA at a flow rate of 0.8 mL/min.



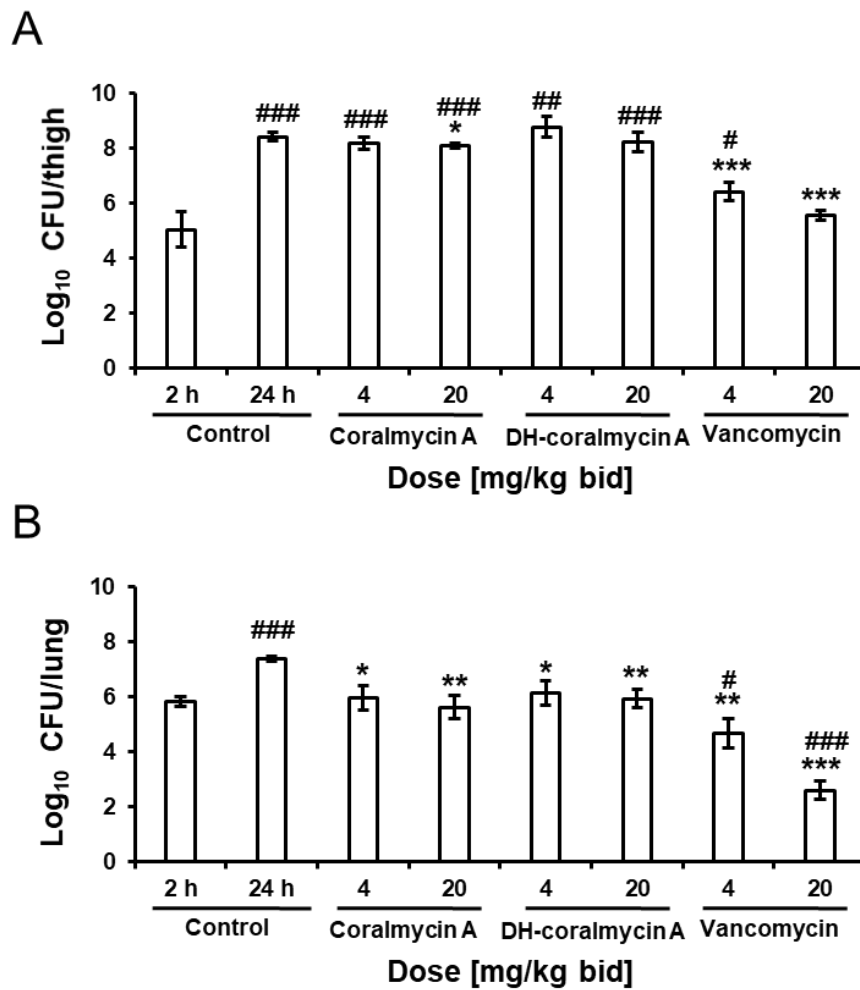
**Figure S2.** Time-kill curves of DH-coralmycin A. MIC values (mg/L) against *S. aureus* Giorgio (A), *S. pneumoniae* ATCC49619 (B), MRSA CCARM 3167 (C), and VRE 3 (D) were 0.06, 0.25, 0.25, and 2, respectively.



**Figure S3.** Time-kill curves of linezolid, ciprofloxacin, and vancomycin as comparators. MIC values (mg/L) against *S. aureus* Giorgio (A), *S. pneumoniae* ATCC49619 (B), MRSA CCARM 3167 (C), and VRE 3 (D) were 2, 2, 1, and 2, respectively, for linezolid; 0.06, 0.25, 4, and 16, respectively, for ciprofloxacin; 2, 0.5, 2, and 64, respectively, for vancomycin.



**Figure S4.** Time-kill curves of coralmycin A at concentrations below and above the MIC. MIC values (mg/L) against *S. aureus* Giorgio (A) and *S. pneumoniae* ATCC49619 (B) were 0.006 and 0.025, respectively.



**Figure S5.** Therapeutic efficacies of coralmycins in two mouse models of thigh and lung infection. (A) a mouse model of thigh infection induced by *S. aureus* Giorgio. The MICs for coralmycin A, DH-coralmycin A, and vancomycin against *S. aureus* Giorgio were 0.003, 0.06, and 2 mg/L, respectively. (B) a mouse model of lung infection induced by intranasal inoculation of *S. pneumoniae* ATCC49619. The MICs of coralmycin A, DH-coralmycin A, and vancomycin against *S. pneumoniae* ATCC49619 were 0.01, 0.25, and 0.5 mg/L, respectively. CFU in the thighs or lungs (n=4) of vehicle- and drug-treated mice were determined. The experiment shown is representative of two independent experiments. Data are expressed as the mean  $\pm$  SD (n = 4; \* $P$  < 0.01, \*\* $P$  < 0.001, and \*\*\* $P$  < 0.0001 versus 24 h control; # $P$  < 0.01, ## $P$  < 0.001, and ### $P$  < 0.0001 versus 2 h control).  $P$  values were obtained using unpaired Student's  $t$  test.