

Supplementary Materials: Enhanced Treatment Effects of Tilmicosin Against *Staphylococcus aureus* Cow Mastitis by Self-Assembly Sodium Alginate-Chitosan Nanogel

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Table S1. Factors and levels of the L9 (3^4) orthogonal design.

| Variable | Level | | |
|-------------------|-------|-------|--------------|
| | 1 | 2 | 3 |
| Type (A) | PVA | PVP | Poloxamer188 |
| Concentration (B) | 1% | 2% | 3% |
| Volume (C) | 15 mL | 20 mL | 25 mL |

Notes: PVA: Poly vinyl alcohol; PVP: Polyvinylpyrrolidone.

Table S2. The optimization of emulsifier by orthogonal experiment ($L_9 3^4$).

| Sample | Type (A) | Concentration (B) | Volume (C) | LC (%) | Size | PDI |
|---------|----------|-------------------|------------|------------------|-------------------|-------------------|
| 1 | 2 | 2 | 3 | 16.02 ± 0.85 | 1.339 ± 0.012 | >0.5 |
| 2 | 3 | 1 | 3 | 16.52 ± 0.70 | 1.677 ± 0.009 | >0.5 |
| 3 | 2 | 3 | 1 | 18.99 ± 1.97 | 5.847 ± 0.073 | >0.5 |
| 4 | 1 | 3 | 3 | 20.90 ± 0.65 | 0.336 ± 0.005 | 0.368 ± 0.024 |
| 5 | 3 | 3 | 2 | 17.61 ± 0.95 | 5.347 ± 0.036 | >0.5 |
| 6 | 3 | 2 | 1 | 17.04 ± 1.65 | 2.451 ± 0.013 | >0.5 |
| 7 | 1 | 1 | 1 | 17.19 ± 0.98 | 0.481 ± 0.013 | 0.424 ± 0.032 |
| 8 | 1 | 2 | 2 | 15.47 ± 0.98 | 0.311 ± 0.005 | 0.306 ± 0.018 |
| 9 | 2 | 1 | 2 | 15.91 ± 1.23 | 1.419 ± 0.005 | >0.5 |
| k1 | 17.85 | 16.54 | 17.74 | | | |
| k2 | 16.64 | 15.84 | 16.33 | | | |
| k3 | 17.05 | 19.16 | 17.81 | | | |
| R | 1.21 | 3.32 | 1.48 | | | |
| Optimum | A1 | B3 | C3 | | | |

Notes: k1, k2, and k3 are the average of grade for 3 levels in each factor; R is the different value between the max and min of k1, k2, and k3 in each level. Abbreviations: LC: load capacity; PDI: poly dispersion index. Optimum formulation of TIL-SLN: PVA, 3%, 25 mL.