# **Antimicrobial Effects of a Novel Combination Therapy Against Methicillin-Resistant** Staphylococcus aureus and Pseudomonas aeruginosa in a Porcine Wound Model

# Introduction

- Chronic wound infections contain various species of bacteria, primary among which are Staphylococcus aureus and Pseudomonas aeruginosa (PA)<sup>1,2</sup>
- The degree of microbial growth, especially biofilm formation, has a direct impact on wound healing<sup>3</sup>
- Therefore, limiting bacterial growth is an essential component of chronic wound care
- A novel technology has been designed to target components of wound healing in chronic or refractory wounds, regardless of pathology
- The combination therapy consists of formulations that address wound preparation, wound therapy (OCM<sup>IM</sup>), and skin integrity

# Objective

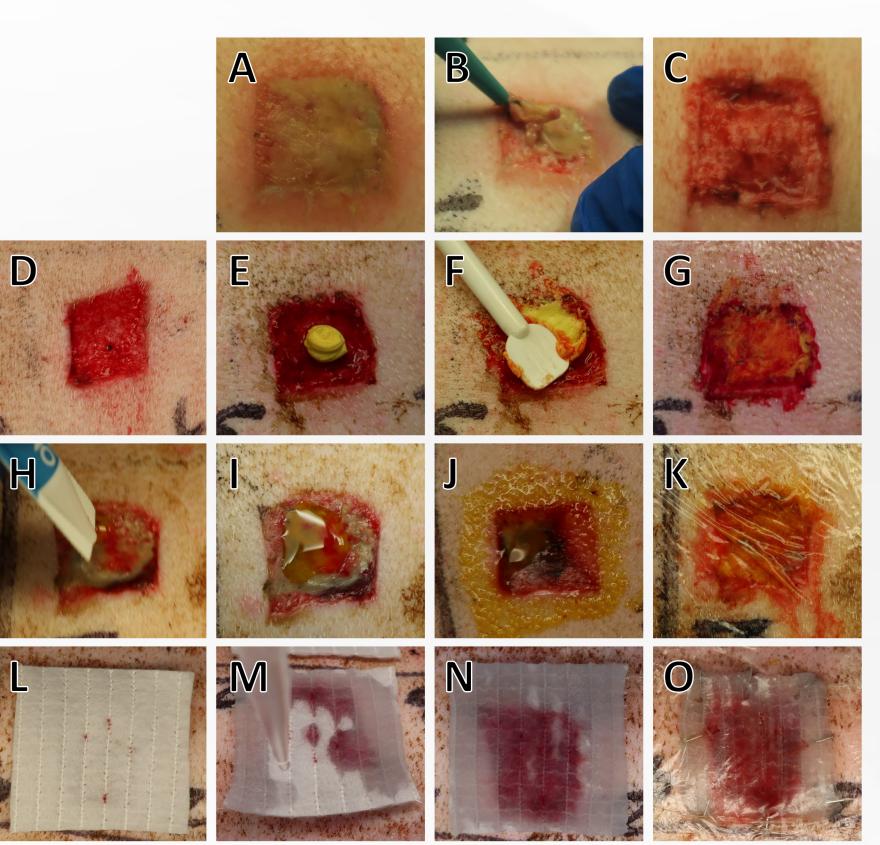
• To evaluate the antimicrobial and wound-healing effects of various wound care formulations against methicillin-resistant Staphylococcus aureus (MRSA) and PA using a porcine wound model

# **Methods**

- Thirty-one deep reticular wounds (22 mm  $\times$  22 mm  $\times$  3 mm) were made across the paravertebral and thoracic areas on each of 6 specific pathogenfree pigs (Looper Farms, North Carolina)
- Pathogenic strains of MRSA (USA300) or PA (ATCC 27312) prepared as 10<sup>6</sup> CFU/mL inoculum suspensions were used to inoculate all wounds within 20 minutes after wounding
- Inoculated wounds were covered with polyurethane dressings (Tegaderm, 3M, USA) for 72 hours before being treated
- Treatment consisted of OCM alone, OCM plus skin protectant, or Aquacel Ag Advantage (positive control) or wounds left untreated (negative control)
  - Wounds treated with OCM alone were debrided before treatment and covered with polyurethane dressings (Figure 1, A-G)
  - Wounds treated with OCM plus skin protectant received a wound preparation formulation for 3 minutes before debridement, were debrided, were treated with OCM and skin protectant, and were covered with polyurethane dressing (Figure 1, H-K)
  - Wounds treated with Aquacel Ag Advantage were initially debrided. treated with Aquacel, and covered with polyurethane dressing (Figure **1, L-O**)
  - Untreated wounds were debrided then covered with polyurethane dressing
- All treatments (except wound preparation) were reapplied on Days 4 and 8
- Baseline wounds were biopsied before and after debridement, and baseline counts were obtained on Day 0; treated wounds were assessed on Days 4, 8, and 12 after treatment

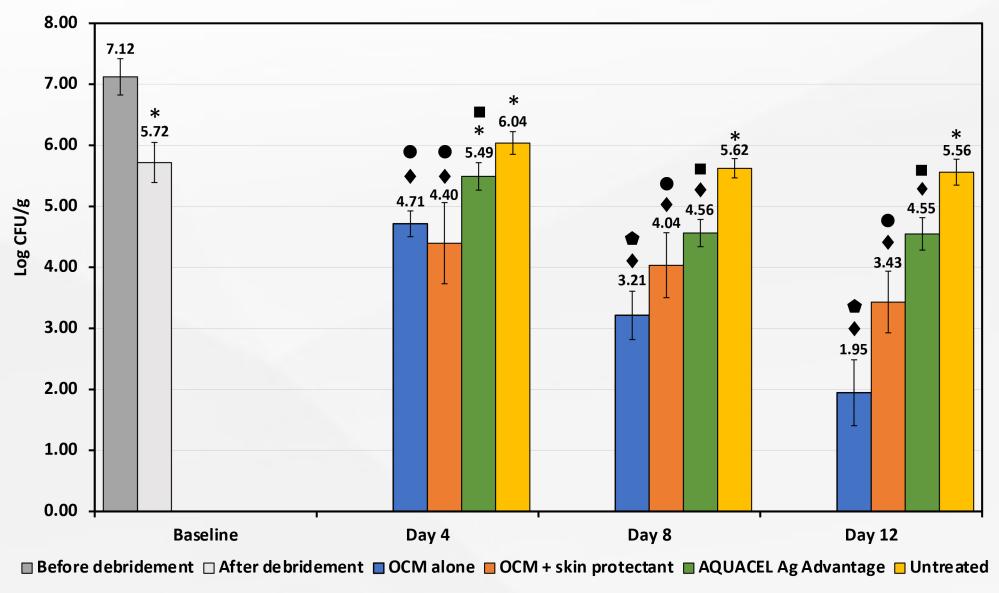
#### Figure 1. Wound preparation and application of OCM, OCM plus skin protectant, and Aquacel Ag Advantage

## Figure 2. MRSA USA300 bacterial counts after treatment application at each assessment day.



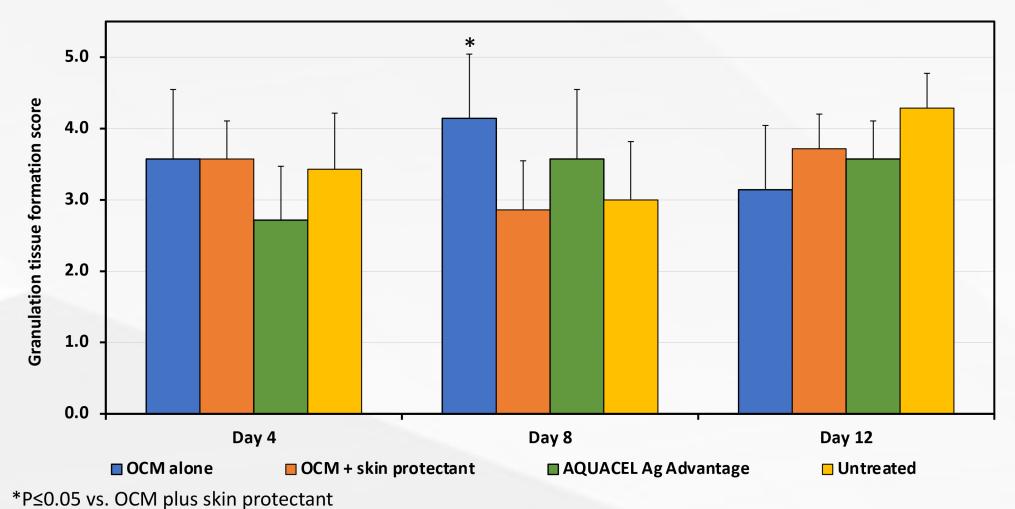


- On Days 8 and 12, MRSA USA300 counts were significantly lower in OCM alonetreated wounds versus all other treatments (Figure 2)
- On Days 4, 8, and 12, MRSA USA300 counts were significantly lower in wounds treated with OCM plus skin protectant versus those treated with the positive and negative controls (P≤0.05, all comparisons; **Figure 2**)
- On Days 4, 8, and 12, PA27312 counts were significantly lower in wounds treated with OCM alone or OCM plus skin protectant versus baseline before and after debridement (P≤0.05, all comparisons; **Figure 3**)
- Day 12 PA27312 counts were significantly lower with OCM alone versus all other treatments and with OCM plus skin protectant versus Aquacel and untreated control (P≤0.05, all comparisons; **Figure 3**)
- Among all treatments at all time points, the lowest MRSA USA300 and PA27312 counts occurred on Day 12 in wounds treated with OCM alone (Figures 2 and 3)
- On Day 8, increased granulation in MRSA USA300-infected wounds was observed with OCM alone compared with OCM plus skin protectant (Figure 4)
- Compared with Aquacel-treated wounds, MRSA USA300-infected wounds treated with OCM plus skin protectant showed increased re-epithelialization on Days 4 and 8 (Figure 5)
- In PA27312-infected wounds, increased re-epithelialization was observed with Aquacel compared with OCM alone at Day 4, and increased granulation was observed with Aquacel compared with OCM plus skin protectant at Day 8



\*P≤0.05 vs. Baseline before debridement ◆P≤0.05 vs. Baseline before/after debridement

#### Figure 4. Granulation tissue formation in MRSA USA300-infected wounds at each assessment day.



### Conclusions OCM alone was significantly better at halting proliferation in both MRSA USA300- and PA27312-infected wounds compared with baseline before and after debridement and compared with all other treatment groups • OCM and OCM plus skin protectant significantly reduced MRSA USA300 and PA27312 counts in this in vivo model, recording the lowest bacterial counts of any treatment in the study

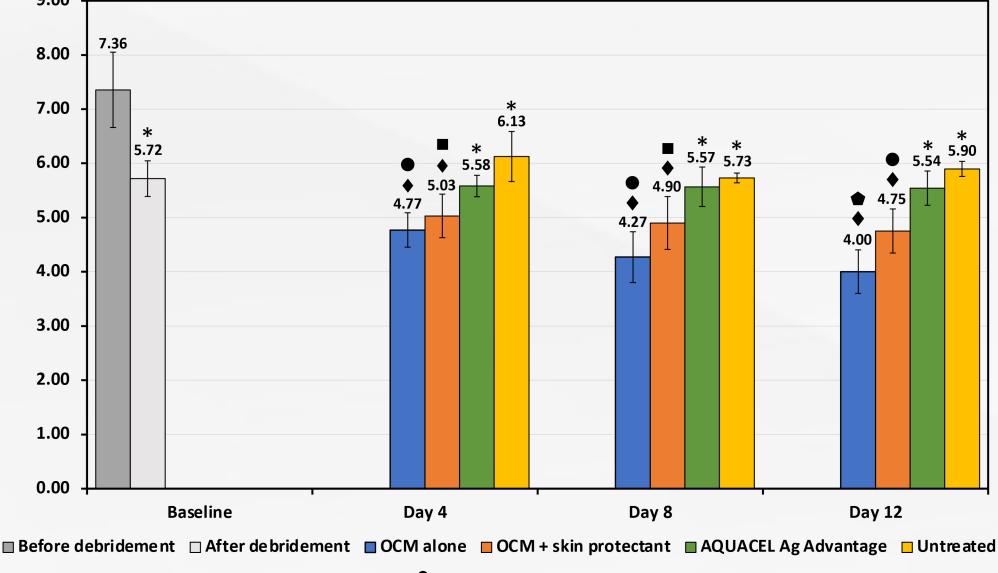
- Compared to the other treatments, OCM alone and OCM plus skin protectant showed significantly faster formation of new tissue in MRSA USA300-infected wounds
- These findings may have important clinical implications for the management of many wound etiologies, such as burns, diabetic foot ulcers, and pressure ulcers

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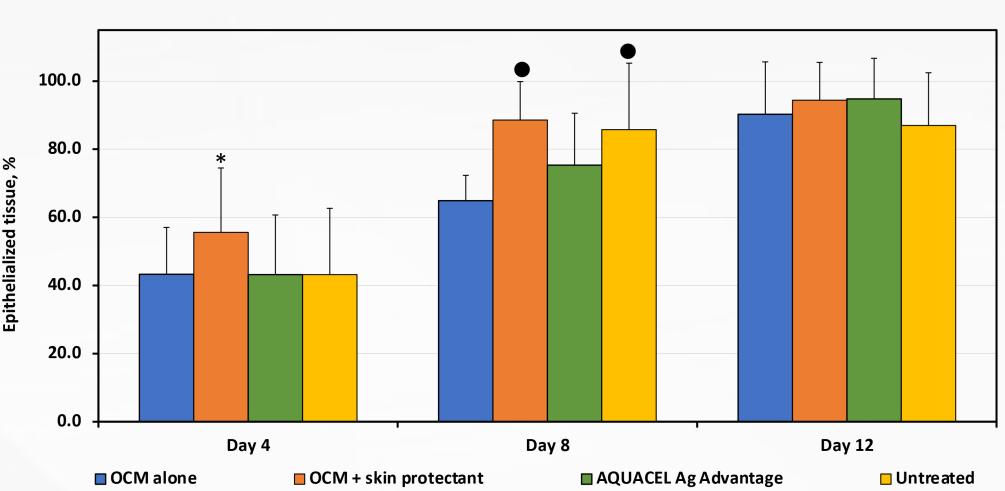
●P≤0.05 vs. Aquacel Ag Advantage and Untreated control <sup>●</sup>P≤0.05 vs. All treatments P≤0.05 vs. Untreated control

# 8.00 7.00 6.00 5.00



\*P≤0.05 vs. Baseline before debridement P≤0.05 vs. Aquacel Ag Advantage and Untreated control <sup>●</sup>P≤0.05 vs. All treatments ◆P≤0.05 vs. Baseline before/after debridement P≤0.05 vs. Untreated control

#### Figure 5. Re-epithelialization of MRSA USA300-infected wounds at each assessment day.



\*P≤0.05 vs. OCM alone

### **REFERENCES**

#### **DISCLOSURES**

SCD, JG, MS: Research support, Omeza, LLC. SJB, DB: Employees, Omeza.

### ACKNOWLEDGMENTS

The study was sponsored by Omeza, LLC (Sarasota, FL, USA). Medical writing and editorial assistance were provided by MedVal Scientific Information Services, LLC (Princeton, NJ, USA), and were funded by Omeza, LLC.

Presented at the SAWC Spring WHS 2024 Conference, May 15-17, 2024, Orlando, FL, USA.

Figure 3. PA27312 bacterial counts after treatment application at each assessment day.

●P≤0.05 vs. AQUACEL Ag Advantage

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