

# Poster abstract submission

**Approval Status**

Not Started

**Presenting author**

Meshack Omwega

**Presenting author's email**

meshacktweya2@gmail.com

**Further authors (if any)**

Lillian Musila  
Moses Gachoya  
Justin Nyasinga  
Martin Georges  
Erick Odoyo

**Affiliation(s)**

1. Microbiology Hub Kericho, Walter Reed Army Institute of Research – Africa, Kenya

**Country**

Kenya

**Type of organization**

Academic / research institution

**Poster title**

Systematic Combinatorial Optimization of Three-Phage Cocktails Against Multidrug-Resistant *Pseudomonas aeruginosa*

**Poster abstract****Background:**

Multidrug-resistant *Pseudomonas aeruginosa* poses a significant global health threat, necessitating alternative therapeutic strategies. Bacteriophage therapy shows promise, but single-phage use is prone to resistance. Bacteriophage cocktails can address this issue, but they are typically formulated empirically without a systematic evaluation of constituent phage interactions.

**Methodology:**

We employed a comprehensive combinatorial approach to optimize three-phage cocktails against clinical *P. aeruginosa* isolates. From 25 candidate bacteriophages, five were selected based on broad host range ( $\geq 60\%$  of 51 MDR *P. aeruginosa* clinical isolates). Ten possible three-phage cocktail combinations were systematically generated and evaluated using real-time Omnilog phenotypic microarray analysis. Phage interactions were quantified using the Highest Single Agent independence model to classify synergistic ( $\Delta > +5\%$ ), neutral ( $-5\% \leq \Delta \leq +5\%$ ), or antagonistic ( $\Delta < -5\%$ ) effects.

**Results:**

Individual phage inhibition efficiencies ranged from 35.4% to 75.4%. Cocktail performance varied dramatically (16.1–84.1% inhibition). Only one cocktail exhibited synergy, achieving 84.1% inhibition, an 8.7% improvement over the best individual phage. Four cocktails (40%) showed neutral interactions, while five (50%) demonstrated antagonistic effects. The optimal cocktail (Phages 1+3+4) demonstrated efficacy across six of eight diverse clinical isolates in biofilm inhibition assays (29–63% biomass reduction, 2.7–3.6  $\log_{10}$  CFU reduction) and achieved approximately 80% survival in *Galleria mellonella* infection models versus 10% in infected controls.

**Conclusions**

These findings demonstrate that phage cocktail optimization requires rational validation, as antagonistic interactions can occur, establishing a critical validation step before committing resources to in vivo

efficacy studies.

**Research topic**

Phage or phage products