

Phage Therapy Against *Pseudomonas aeruginosa* Biofilms in Cystic Fibrosis: A Personalized Bacteriophage Cocktail Approach Guided by Host-Pathogen Genomics

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Abstract

Antibiotic resistance in *Pseudomonas aeruginosa* biofilms represents the leading cause of morbidity in cystic fibrosis (CF). We report a precision phage therapy protocol utilizing a cocktail of four lytic bacteriophages (Φ F1- Φ F4) selected based on patient-specific bacterial genotyping and CRISPR-spacer analysis. In a 24-month prospective cohort of 45 CF patients with multidrug-resistant infections, nebulized phage administration reduced bacterial load by 3.5 log₁₀ CFU/mL and improved forced expiratory volume (FEV1) by 18%. Whole-genome sequencing confirmed minimal phage resistance evolution, attributed to cocktail synergy and host immune modulation. This individualized phagetherapeutic framework offers a paradigm shift in managing recalcitrant respiratory infections.

Keywords: bacteriophage therapy, *Pseudomonas aeruginosa*, cystic fibrosis, biofilm disruption, personalized medicine, antimicrobial resistance



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