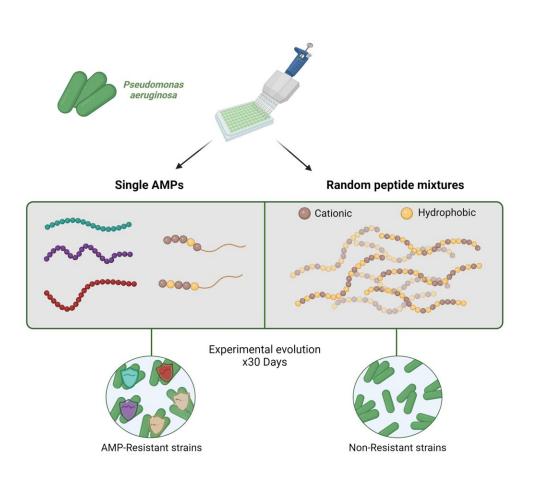


## Treatment with a mixture of antimicrobial peptides found to impede antibiotic resistance

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A common infection-causing bacteria was much less likely to evolve antibiotic resistance when treated with a mixture of antimicrobial peptides rather than a single peptide, making these mixtures a viable strategy for developing new antibiotic treatments. Jens Rolff of the Freie Universitat Berlin, Germany, and colleagues report these findings in a study published July 2 in *PLOS Biology*.

Antibiotic-resistant bacteria have become a major threat to <u>public health</u>. The World Health Organization estimates that 1.27 million people died directly from drug-resistant strains in 2019 and these strains contributed to 4.95 million deaths.

While bacteria naturally evolve resistance to antibiotics, misuse and overuse of these drugs has accelerated the problem, rendering many antibiotics ineffective. One emerging strategy to combat <u>antibiotic</u> resistance is the use of antimicrobial peptides, which are chains of amino acids that function as broad-spectrum <u>antimicrobial compounds</u> and are key components of the innate immune system in animals, fungi and plants.

In the new study, researchers investigated whether antimicrobial peptide mixtures synthesized in the lab could reduce the risk of the pathogen Pseudomonas aeruginosa from evolving antimicrobial resistance, compared to exposure to a single antimicrobial peptide.

They found that using antimicrobial peptide mixtures carried a much lower risk of the bacteria developing resistance. The mixtures also helped prevent the bacteria from developing cross-resistance to other antimicrobial drugs, while maintaining—or even improving—drug sensitivity.

Overall, the findings suggest that the use of antimicrobial peptide mixtures is a strategy worth pursuing in the search for new, longer-



lasting treatments for bacteria.

The researchers suspect that using a cocktail of multiple antimicrobial peptides creates a larger set of challenges for bacteria to overcome, which can potentially delay the evolution of resistance, compared to traditional antibiotics. Furthermore, these cocktails can be synthesized affordably, and previous studies have shown them to be non-toxic in mice.

Lead author Bernardo Antunes said, "Even after four weeks of exposure, a usual treatment duration for Pseudomonas infections, we could not find resistance against our new random peptide, but against other antimicrobials."

**More information:** Bernardo Antunes et al, The evolution of antimicrobial peptide resistance in Pseudomonas aeruginosa is severely constrained by random peptide mixtures, *PLoS Biology* (2024). <u>DOI:</u> <u>10.1371/journal.pbio.3002692</u>

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