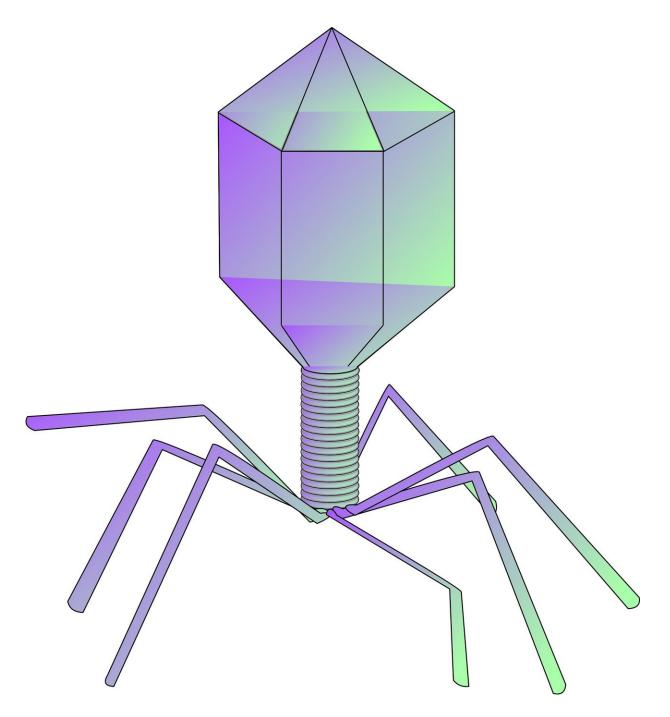


Researchers surprised to find bacterial parasites behind rise of 'super bugs'

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For the first time ever, researchers from the University of Pittsburgh School of Medicine discovered that phages—tiny viruses that attack



bacteria—are key to initiating rapid bacterial evolution leading to the emergence of treatment-resistant "superbugs." The findings were published today in *Science Advances*.

The researchers showed that, contrary to a dominant theory in the field of evolutionary microbiology, the process of adaptation and diversification in bacterial colonies doesn't start from a homogenous clonal population. They were shocked to discover that the cause of much of the early adaptation wasn't random point mutations. Instead, they found that phages, which we normally think of as bacterial parasites, are what gave the winning strains the <u>evolutionary advantage</u> early on.

"Essentially, a parasite became a weapon," said senior author Vaughn Cooper, Ph.D., professor of microbiology and molecular genetics at Pitt. "Phages endowed the victors with the means of winning. What killed off more sensitive bugs gave the advantage to others."

When it comes to bacteria, a careful observer can track evolution in the span of a few days. Because of how quickly bacteria grow, it only takes days for <u>bacterial strains</u> to acquire new traits or develop resistance to antimicrobial drugs.

The researchers liken the way bacterial infections present in the clinic to a movie played from the middle. Just as late-arriving moviegoers struggle to mentally reconstruct events that led to a scene unfolding in front of their eyes, physicians are forced to make treatment decisions based on a static snapshot of when a patient presents at a hospital. And just like at a movie theater, there is no way to rewind the film and check if their guess about the plot or the origin of the <u>infection</u> was right or wrong.

The new study shows that bacterial and <u>phage</u> evolution often go hand in hand, especially in the early stages of bacterial infection. This is a



multilayered process in which phages and bacteria are joined in a chaotic dance, constantly interacting and co-evolving.

When the scientists tracked changes in genetic sequences of six bacterial strains in a skin wound infection in pigs, they found that jumping of phages from one bacterial host to another was rampant—even clones that didn't gain an evolutionary advantage had phages incorporated in their genomes. Most clones had more than one phage integrated in their genetic material—often there were two, three or even four phages in one bug.

"It showed us just how much phages interact with one another and with new hosts," said Cooper. "Characterizing diversity in early bacterial infections can allow us to reconstruct history and retrace complex paths of evolution to a clinical advantage. And, with growing interest in using phages to treat highly resistant infections, we are learning how to harness their potency for good."

More information: "Rampant prophage movement among transient competitors drives rapid adaptation during infection" *Science Advances* (2021). <u>advances.sciencemag.org/lookup1126/sciadv.abh1489</u>

Provided by University of Pittsburgh

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