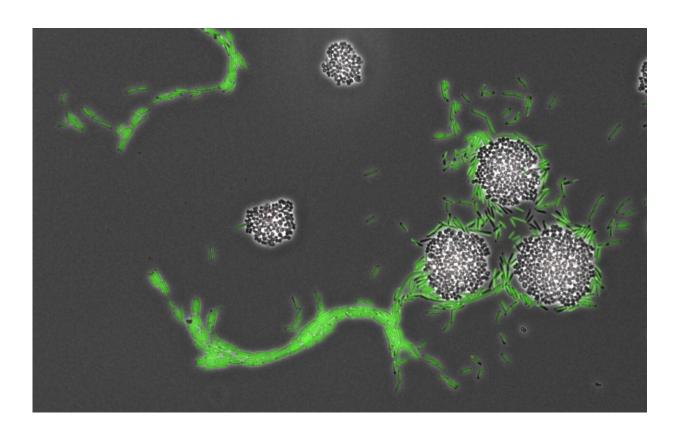


Study reveals 'bug wars' that take place in cystic fibrosis

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P. aeruginosa bacteria (green rods) respond to secreted interspecies factors and mount attack behaviours against *S. aureus*. Credit: Limoli et al. (CC BY 4.0)

Scientists have revealed how common respiratory bugs that cause serious infections in people with cystic fibrosis interact together, according to a new study in *eLife*.



The results provide insights into how bacterial pathogens wrestle each other for territory that could open avenues for new antibacterial treatments.

Studies of microbes from mouths, intestines, chronic wounds and chronic respiratory infections show that interactions between bacteria in these communities influence survival of the bugs and progression of disease. For example, infection with two bacterial species called *Staphylococcus aureus* (*S. aureus*) and *Pseudomonas aeruginosa* (*P. aeruginosa*) in the airways of cystic fibrosis patients is linked to decreased lung function and a shorter lifespan.

"One strategy to improve outcomes for people with cystic fibrosis carrying multiple infections is to block harmful interspecies interactions before they begin," explains lead author Dominique Limoli, Assistant Professor of Microbiology and Immunology at the University of Iowa, US. "We designed a system to visualise the early interactions between two microbes that cause infections in people with cystic fibrosis to follow their behaviour over time."

Limoli and her team used a co-culture system to grow both microbes together and then studied them using time-lapse microscopy to create videos of the bacteria's movements. They found that, on their own, *P. aeruginosa* cells multiply and form groups that appear like 'rafts' under a microscope. By contrast, when grown along with *S. aureus*, the *P. aeruginosa* cells move as single cells and accelerate their movement in the direction of the *S. aureus* colony. Studying the cells in more detail revealed that once the single *P. aeruginosa* cells reach the *S. aureus* cells, they enter the colony and dismantle it.

The researchers proposed that *P. aeruginosa* behaves in this way because *S. aureus* produces substances into the environment that trigger the change in movement of *P. aeruginosa*. To prove this, they took the liquid



that *S. aureus* had been growing in and placed it in the bottom of a petri dish, before using it to grow *P. aeruginosa*. In this environment, *P. aeruginosa* was much more mobile—moving across a greater surface area than on the normal petri dish.

So, does this hold true for other types of bacteria? Of three pathogens commonly found in patients with cystic fibrosis, the team found one that enhanced the movement of *P. aeruginosa*. However, when they looked at a broader group of bacteria, including Salmonella, Escherichia coli and Bacillus subtilis, they found a significant increase in the movement of *P. aeruginosa* in the presence of these communities.

"By acquiring a fundamental understanding of how bacteria sense and respond to life with each other, we move closer to learning how to rationally manipulate these behaviors during infection and in the environment," says Limoli. "For <u>cystic fibrosis</u> patients, this may mean preventing *P. aeruginosa* and *S. aureus* physical interactions. In other instances, we might bring together <u>bacterial species</u> that synergise to produce a beneficial compound."

More information: Dominique H Limoli et al, Interspecies interactions induce exploratory motility in Pseudomonas aeruginosa, *eLife* (2019). DOI: 10.7554/eLife.47365

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