

New compound may stop bacteria from causing sickness

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Michael Federle, associate professor of medicinal chemistry and pharmacognosy in the UIC College of Pharmacy. Credit: UIC/Roberta Dupuis-Devlin

A study published in the *Journal of Biological Chemistry* is the first to describe a signaling pathway that affects communication—a process



called quorum sensing—between Streptococcus bacteria cells.

This type of bacterium is responsible for illnesses like <u>strep throat</u>, <u>scarlet fever</u>, and some cases of <u>soft tissue infections</u> and pneumonia. In extreme cases, or when bacteria become resistant to antibiotics, these common but serious infections can lead to death.

The University of Illinois at Chicago researchers who wrote the paper have been studying Streptococcus bacteria and have hypothesized that quorum sensing pathways are ideal drug targets for manipulating bacterial activity and suppressing virulence.

"Quorum sensing is how bacteria regulate their behavior as a collective," said Michael Federle, corresponding author on the study. "This social regulation may provide many benefits to bacteria and, in the case of bad bacteria, may help promote survival and help defend against the immune system."

That is why Federle and his team have developed a tool to help study the quorum sensing signaling pathways. Their technology allows them to easily monitor specific gene activities and whether chemicals or other conditions affect gene expression.

Using this technique, the UIC scientists have unlocked part of the communication process and identified a small molecule that promotes <u>quorum</u> sensing in Streptococci. This molecule stabilizes the chemical signal—called a pheromone—transmitted between cells. More specifically, the compound works by blocking the enzyme that would otherwise degrade the pheromone before it can deliver a message, thus promoting its ability to act as a signal.

"We identified a molecule that disrupts inhibition of signaling," said Federle, associate professor of medicinal chemistry and pharmacognosy



in the UIC College of Pharmacy. "Now that we identified this pathway and this chemical compound, we can look further for ways to manipulate <u>quorum sensing</u> and even silence the communication between cells."

By silencing this communication, Federle says, it might be possible "to convince bacteria to remain in a non-hostile state," something he considers vital in the face of rising antibiotic resistance in society.

"We're losing the war with bacteria," he said. "Every antibiotic we've come up with has some level of resistance in nature. We need a way to stop <u>bacteria</u> before they become a threat and require treatment."

Federle says this is analogous to finding a diplomatic solution or mediating a conflict before it causes harm.

More information: Tiara G. Pérez Morales et al, A novel chemical inducer of Streptococcusquorum sensing acts by inhibiting the pheromone-degrading endopeptidase PepO, *Journal of Biological Chemistry* (2017). DOI: 10.1074/jbc.M117.810994

Provided by University of Illinois at Chicago

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