

# Breaking up the superbugs' party

August 13 2013

---

The fight against antibiotic-resistant superbugs has taken a step forward thanks to a new discovery by scientists at The University of Nottingham.

A multi-disciplinary research team at the University's Centre for Biomolecular Sciences has uncovered a new way of inhibiting the toxicity and virulence of the notorious superbug, *Pseudomonas aeruginosa*.

This bacteria produces an armoury of [virulence factors](#) and is resistant to many conventional antibiotics. It is almost impossible to eradicate *P. aeruginosa* from the lungs of people with [cystic fibrosis](#) and is therefore a leading cause of death among sufferers. The bug also causes a wide range of infections particularly among hospital patients.

The new discovery concerns the [bacterial cells'](#) ability to 'talk' to each other by producing and sensing small [chemical signal](#) molecules. This is called 'quorum sensing' (QS) and enables a population of individual bacteria to act socially rather than as individuals. QS allows a population of bacteria to assess their numerical strength and make a decision only when the population is 'quorate'.

The mechanism through which QS signals work is by activating gene expression upon interaction of a QS signal molecule with a receptor protein. In many disease-causing bacteria, QS controls genes which are essential for infection. These genes code for virulence factors such as toxins which cause damage to host tissues and the immune system. Interfering with the QS signalling process blocks [bacterial virulence](#) and

renders bacteria unable to cause infection. Consequently QS systems are [molecular targets](#) for the development of new anti-infective drugs which do not kill bacteria but instead block their ability to cause disease.

In a study published in the journal, *PLOS Pathogens*, the Nottingham team has described how they solved the 3D structure of a receptor protein called PqsR used by *P. aeruginosa* to sense alkyl quinolone QS signal molecules so that they could visualize the shape of the QS signal molecule-binding site within the PqsR protein.

Professor of Molecular Microbiology, Paul Williams, said: "We were able to synthesize and screen a library of chemical compounds which could fit within the PqsR binding site and block receptor activation by the QS signal molecules. The active compounds were screened for their ability to inhibit QS and through a process of chemical refinement some novel potent QS inhibitors were discovered which were tested biologically on *P. aeruginosa* and shown to block virulence gene expression."

Professor of Macromolecular Crystallography, Jonas Emsley, added: "This ground-breaking work establishes a platform for the future evaluation and further development of these new QS inhibitor compounds as potential drugs for the treatment of *P. aeruginosa* infections."

Provided by University of Nottingham

Citation: Breaking up the superbugs' party (2013, August 13) retrieved 27 April 2024 from <https://phys.org/news/2013-08-superbugs-party.html>

This document is subject to copyright. Apart from any fair dealing for the purpose of private study or research, no part may be reproduced without the written permission. The content is

provided for information purposes only.