

Researchers discover genes behind antibiotic resistance in deadly superbug infections

September 12 2023



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Australian researchers have uncovered new genetic insights into



Staphylococcus aureus, revealing what makes the bacterium so dangerous when it enters the blood.

While common, Staphylococcus aureus infections—known as Golden staph—can be life-threatening if the bacteria enter the bloodstream, causing sepsis. Golden staph is notorious for its ability to become resistant to antibiotics, making it hard to treat, which can lead to adverse health outcomes for patients infected with a drug-resistant form of the bacteria.

In one of the most comprehensive studies of its kind, published in <u>Cell</u> <u>Reports</u>, researchers, led by the Peter Doherty Institute for Infection and Immunity (Doherty Institute), analyzed the unique genetic profiles of more than 1,300 Golden staph strains.

By combining this data with patient and antibiotic information, the researchers found that, while patient factors are critical in determining mortality risks, <u>specific genes</u> are linked to <u>antibiotic resistance</u>, along with the bacteria's ability to linger in the blood, evading antibiotics and the immune system.

University of Melbourne Dr. Stefano Giulieri, a Clinician-Researcher at the Doherty Institute and first author of the paper, said the findings highlighted the diagnostic power of integrating clinical and genomic data.

"To the best of our knowledge, this is one of the first times that the method we used, called a <u>genome-wide association study</u> (GWAS), has been applied to delve into the role of <u>bacterial genomes</u>, host factors and antibiotics on the course of staphylococcal sepsis," said Dr. Giulieri.

"In GWAS, scientists scan the genome of a big collection of bacteria to look for tiny changes (mutations) that show up more often in strains with



a certain characteristic, such as antibiotic resistance. Mutations with a strong statistical link are precious clues to figure out how bacteria acquire attributes that are important for patient outcomes."

"Our study uncovered a deeper understanding of the intricate genetic dynamics underlying severe Golden staph infections. It highlights the potential of combining bacterial <u>whole-genome sequencing</u>, <u>clinical data</u> and sophisticated statistical genomics to discover clinically relevant bacterial factors that influence infection outcomes."

University of Melbourne Professor Ben Howden, Director of the Microbiological Diagnostic Unit (MDU) Public Health Laboratory at the Doherty Institute and co-senior author of the paper, said that this work represents a significant advancement in <u>medical research</u> as it reshapes our strategies against complex health challenges like Golden staph infections.

"By revealing the genes responsible for antibiotic resistance in Golden staph, our GWAS is pointing the scientific community to clearer targets for the development of effective solutions to treat Golden staph bloodstream infections," said Professor Howden.

"This knowledge has the potential to shape and enhance our ability to tackle these persistent infections. As bacterial genomes become increasingly available in the clinical routine, we inch closer to customized therapeutic strategies, where treatments will be tailored to the unique genetic makeup of the infecting strain, rather that treating everyone in the same way."

More information: A statistical genomics framework to trace bacterial genomic predictors of clinical outcomes in Staphylococcus aureus bacteraemia, *Cell Reports* (2023). DOI: 10.1016/j.celrep.2023.113069



Provided by The Peter Doherty Institute for Infection and Immunity

Citation: Researchers discover genes behind antibiotic resistance in deadly superbug infections (2023, September 12) retrieved 29 April 2024 from <u>https://phys.org/news/2023-09-genes-antibiotic-resistance-deadly-superbug.html</u>

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