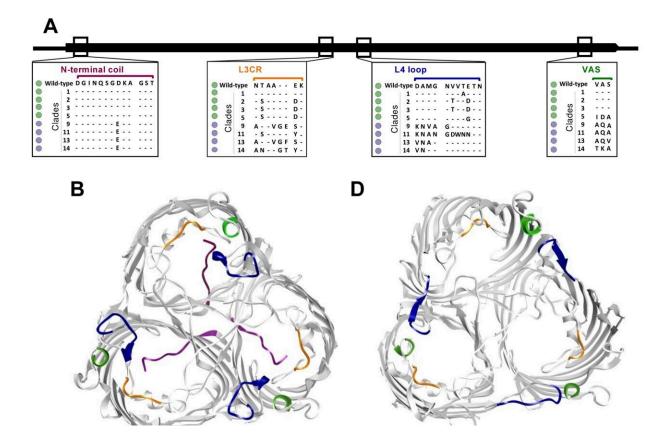


Scientist uncovers roots of antibiotic resistance

May 5 2023, by Suhtling Wong



Comparative visualization of domains within OmpU associated with bile resistant phenotypes. (A). Multiple sequence alignment of ompU alleles resistant and sensitive to whole bile. Conserved residues are indicated by dots and those absent are indicated by gray boxes. (B-E) Comparative architecture of OmpU domains associated with the bile resistant phenotype. Domains are color coded and visualized in OmpU N16961 and GBE1114. Top and slabbed view of (B, C) N16961 and (D, E) GBE1114, respectively. The identified domains are colored as follows: NTC (purple), L3R (orange), L4 (blue) and VAS (green). Credit:



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Bacteria naturally adapt to various environmental stimuli and as they mutate, these changes can make them resistant to drugs that would kill or slow their growth.

In a recent article published in *PLoS Genetics*, UCF College of Medicine microbiologist Dr. Salvador Almagro-Moreno uncovers the evolutionary origins of antimicrobial resistance (AMR) in <u>bacteria</u>. His studies on the bacterium that causes cholera, Vibrio cholerae, provide insight into deciphering what conditions must occur for infectious agents to become resistant.

"How AMR occurs in bacterial populations and the pathways leading to these new traits are still poorly understood," he said. "This poses a major public health threat as antimicrobial resistance is on the rise."

Dr. Almagro-Moreno studied genetic variants of a protein found in bacterial membranes called OmpU. Using computational and molecular approaches, his team found that several OmpU mutations in the cholera bacteria led to resistance to numerous antimicrobial agents.

This resistance included <u>antimicrobial peptides</u> that act as defenses in the human gut. The researchers found that other OmpU variants did not provide these properties, making the protein an ideal system for deciphering the specific processes that occur to make some bacteria resistant to antimicrobials.

By comparing resistant and antibiotic sensitive variants, the researchers were able to identify specific parts of OmpU associated with the emergence of antibiotic resistance. They also discovered that the <u>genetic</u>



<u>material</u> encoding these variants, along with associated traits, can be passed between <u>bacterial cells</u>, increasing the risk of spreading AMR in populations under antibiotic pressure.

By understanding how mutations occur, researchers can better understand and develop therapeutics to combat resistant infections. Dr. Almagro-Moreno is also looking at <u>environmental factors</u> such as pollution and warming of the oceans, as possible causes of resistant bacteria. "We are studying the <u>genetic diversity</u> of environmental populations, including coastal Florida isolates, to develop a new approach to understanding how antimicrobial resistance evolves," he explained.

Understanding the bacteria that causes cholera, an acute diarrheal illness linked to infected water and foods, has global implications. The disease sickens up to 4 million people worldwide and severe cases can cause death within hours.

More information: Trudy-Ann Grant et al, Allelic diversity uncovers protein domains contributing to the emergence of antimicrobial resistance, *PLOS Genetics* (2023). DOI: 10.1371/journal.pgen.1010490

Provided by University of Central Florida

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