

Shrinking to survive: Bacteria adapt to a lifestyle in flux

June 14 2021, by Marta Wegorzewska



E. coli cells shifted from nutrient-rich conditions (left) to nutrient-free conditions (right). These cells contain fluorescent protein fusions that make the cytoplasm green and the periplasm, the space between the cell membrane and the outer membrane, red. Credit: Huang laboratory

Summer picnics and barbecues are only a few weeks away. As excited as you are to indulge this summer, *Escherichia coli* bacteria are eager to feast on the all-you-can-eat buffet they are about to experience in your gut.



However, something unexpected will occur as *E. coli* <u>cells</u> end their journey through your digestive tract. Without warning, they will find themselves swimming in your toilet bowl, clinging to the last bits of nutrients attached to their bodies. How do these tiny organisms adapt to survive sudden starvation? Scientists at Washington University in St. Louis wondered.

Close examination of nutrient-deprived *E. coli* under the microscope—a routine process in a lab that studies bacterial cell size—revealed cells that looked different, and that these differences are related to their ability to survive.

"Their cytoplasm shrank. As it shrank, the <u>inner membrane</u> pulled away from the outer <u>membrane</u> and left a big space at one end of the cell," said Petra Levin, professor of biology in Arts & Sciences, whose postdoctoral scientist, Corey Westfall, and undergraduate student, Jesse Kao, first made the observation.

The space to which Levin refers, between the bacteria's inner and <u>outer</u> <u>membranes</u>, is called the periplasm. In collaboration with Kerwyn Casey Huang, professor of bioengineering and of microbiology and immunology at Stanford University, and his postdoctoral scientist, Handuo Shi, Levin found an unexpected developmental response to starvation—one that may be keeping *E. coli* alive until they find their next buffet.

The work is published this week in the *Proceedings of the National Academy of Sciences*.

The biologists showed that when *E. coli* cells lack nutrients, the cytoplasm becomes more dense as its volume decreases, probably because of water loss. At the same time, the periplasm increases in volume as the inner membrane pulls away from the outer membrane.



"Although we don't know for sure yet, we think that the cell is concentrating the nutrients in the cytoplasm so that it can keep running metabolism at a high rate," Levin said. "Perhaps this is an adaptation to *E. coli*'s constantly and rapidly changing lifestyle, in which it knows that each environment is temporary."

The shrinking is reversible, the scientists found. Once they transferred the starving bacteria into a nutrient-rich medium, the inner membrane and the cytoplasm expanded. The bacterial cells rapidly rebounded from starvation, especially when *E. coli* received their favorite carbon source, glucose. And, importantly, if the Tol-Pal system was intact.

The Tol-Pal system is a critical cellular machinery composed of proteins that connect the outer membrane to the inner membrane. But its function has been understudied. As the inner membrane expands, the Tol-Pal system helps to reconnect it with the outer membrane, the scientists speculate. When the Tol-Pal system was absent, the internal contents of the cells bled out.

"We speculate that Tol-Pal acts as the zipper slider, helping the inner membrane zip into the outer membrane coat during recovery," Levin said.

What happens to the transmembrane proteins, embedded in both the inner and outer membrane, when the inner membrane pulls away from the outer membrane? Do they get ripped apart? Levin and colleagues do not know yet and hope to answer these questions in the future.

More information: Handuo Shi et al, Starvation induces shrinkage of the bacterial cytoplasm, *Proceedings of the National Academy of Sciences* (2021). DOI: 10.1073/pnas.2104686118



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