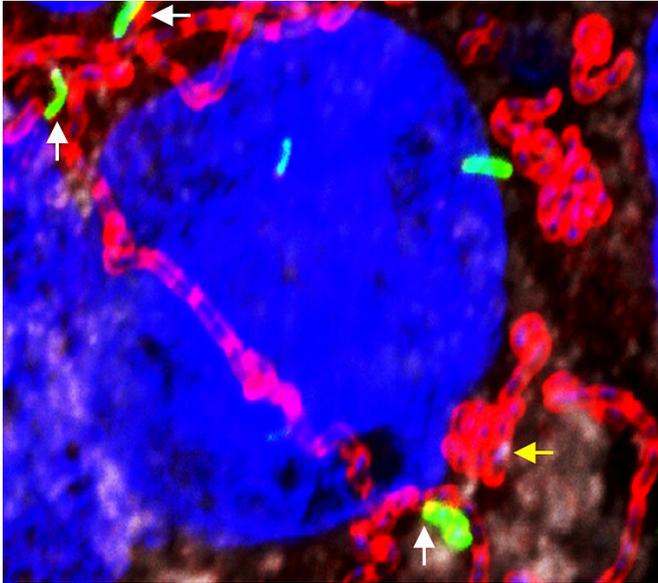


Bioengineered probiotic could prevent listeria infections

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Bioengineered lactobacillus (red) interacts with surface heat shock protein 60 (white, yellow arrows), blocking *Listeria monocytogenes* (green, white arrows) from crossing into the bloodstream. Credit: Rishi Drolia and Arun Bhunia

For pregnant women, the elderly and those with weakened immune systems, listeriosis is a serious foodborne illness often linked to deli meats, fresh produce and dairy products. Even with antibiotic treatment, listeriosis is fatal for about 20 percent of patients, resulting in thousands of deaths annually.

Purdue University's Arun Bhunia, a professor of food science, and postdoctoral researcher Rishi Drolia have developed a probiotic that could prevent infections in at-risk populations. A bioengineered version of *Lactobacillus*, a bacterium common in the [human gut](#), can block the pathway the *Listeria monocytogenes* bacteria use to cross intestinal wall [cells](#) into the bloodstream, his team reports in the journal *Nature*

Communications.

"The *Lactobacillus* bacteria we developed seeks out the same proteins as *Listeria monocytogenes* in the gut. When it attaches, it blocks the roadway for *Listeria*," Bhunia said. "This could be included in probiotic yogurts, capsules or gummies and used as a preventive measure to treat people who are at high risk of infection."

In a previous study, Bhunia's and Drolia's work showed how *Listeria* crosses the epithelial barrier, a wall of cells in the gut that generally protects the bloodstream from harmful pathogens. A protein in *Listeria*, called *Listeria* adhesion protein (LAP), interacts with [heat shock protein](#) in those epithelial cells and forces the cells apart. That gives *Listeria* access to the bloodstream.

Bhunias team has now isolated the *Listeria* adhesion protein from nonpathogenic strains of the *Listeria* bacterium and added it to the *Lactobacillus* to make a probiotic. When introduced to human gut cells and in mice, the *Lactobacillus* probiotic stably colonized the intestine and attached to epithelial cells on the heat shock protein. When pathogenic *Listeria* was introduced, it wasn't able to attach to those gut cells and invade the bloodstream.

"This novel approach of engineering a probiotic strain with an adhesion [protein](#) from a nonpathogenic bacterium to exclude a pathogen significantly enhances the prophylactic use of such bioengineered probiotic bacteria without raising serious health or regulatory concerns and, thus, their potential as preventive agents against listeriosis," the authors wrote. "This study, thus, provides the first and direct evidence that rational engineering of probiotic strains allows them to outcompete and diminish the colonization of the pathogens by competing for the receptor binding adhesion sites."

The *Lactobacillus* probiotic could also have

potential for other gut illnesses such as celiac disease or inflammatory bowel disease.

"We've seen evidence that the same proteins *Listeria* adheres to are overactive in these other illnesses," Drolia said. "The probiotic has an anti-inflammatory effect and colonizes the gut for a week or more at a time. We're interested in seeing how this could improve the lives of patients suffering from a variety of gut-related illnesses."

Bhunja has applied for a patent for the bioengineered *Lactobacillus* [probiotic](#) and envisions licensing the technology.

"We're also interested in using this model to look at other pathogenic bacteria such as *E. coli* or *Salmonella*," Bhunja said. "If we can engineer bacteria properly, we may be able to use this technology to prevent a wider range of foodborne illnesses."

More information: Rishi Drolia et al. Receptor-targeted engineered probiotics mitigate lethal *Listeria* infection, *Nature Communications* (2020). [DOI: 10.1038/s41467-020-20200-5](https://doi.org/10.1038/s41467-020-20200-5)

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