

## **Trust your gut: E. coli may hold one of the keys to treating Parkinson's**

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Escherichia coli. Credit: Rocky Mountain Laboratories, NIAID, NIH

*E. coli* usually brings to mind food poisoning and beach closures, but researchers recently discovered a protein in *E. coli* that inhibits the accumulation of potentially toxic amyloids—a hallmark of diseases such



as Parkinson's.

Amyloids are formed by proteins that misfold and group together, and when amyloids assemble at the wrong place or time, they can damage brain tissue and cause cell death, according to Margery Evans, lead author of the University of Michigan study, and Matthew Chapman, principal investigator and associate professor in U-M Molecular, Cellular, and Developmental Biology.

The findings could point to a new therapeutic approach to Parkinson's disease and a method for targeting amyloids associated with such neurodegenerative diseases.

A key biological problem related to patients with Parkinson's is that certain proteins accumulate to form harmful amyloid fibers in brain tissues, which is toxic to cells and causes <u>cell death</u>.

While these amyloids are a hallmark of Parkinson's and other diseases such as Alzheimer's, not all amyloids are bad. Some cells, those in *E. coli* included, assemble helpful amyloids used for cell function.

*E.. coli* make amyloid curli on the cell surface, where it's protective, rather than toxic. The curli anchor the bacteria to kitchen counters and intestinal walls, where they can cause infections and make us sick. These helpful amyloids that *E.. coli* produce do not form on the inside of the cell where they would be toxic.

"It means that something in *E. coli* very specifically inhibits the assembly of the amyloid inside the cell. Therefore, amyloid formation only occurs outside the cell where it does not cause toxicity," said Evans, a doctoral student in molecular, cellular, and <u>developmental biology</u>.

Evans and the U-M team went on a biochemical hunt to understand how



*E.*. *coli* prevented <u>amyloids</u> from forming inside <u>cells</u> and uncovered a protein called CsgC that is a very specific, effective inhibitor of *E. coli* amyloid formation.

U-M researchers have been collaborating with scientists from Umeå University in Sweden and Imperial College in London, and in the current study found that the CsgC protein also inhibits amyloid formation of the kind associated with Parkinson's.

Another implication of the research is that the curli could be a target for attacking biofilms, a kind of goo created by bacteria, which acts as a shield to thwart antibiotics and antiseptics. These bacteria can cause chronic infections, but treating these infections using molecules that block curli formation may degrade the biofilm and leave the bacteria more vulnerable to drug therapy.

The study, "The bacterial curli system possesses a potent and selective inhibitor of amyloid formation," is scheduled to appear Jan. 22 in the online edition of *Molecular Cell*.

Provided by University of Michigan

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