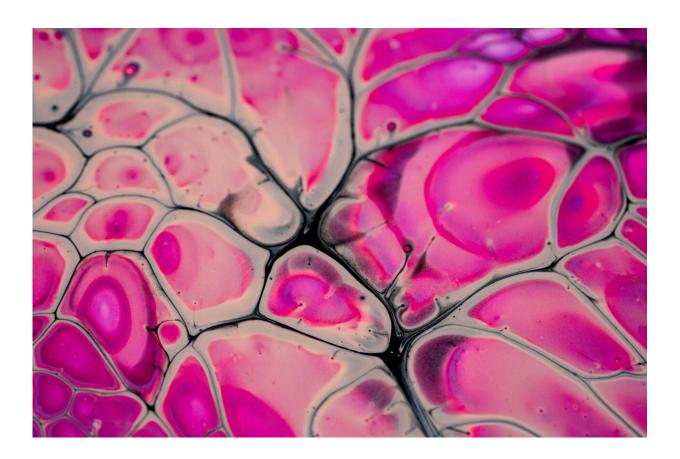


Detoxing body of two fat by-products could extend lifespan, researchers discover

March 9 2023



Credit: Unsplash/CC0 Public Domain

University of Virginia scientists have identified a promising approach to delay aging by detoxifying the body of glycerol and glyceraldehyde, harmful by-products of fat that naturally accumulate over time.



The new findings come from UVA researcher Eyleen Jorgelina O'Rourke, Ph.D., and her team, who are seeking to identify the mechanisms driving healthy aging and longevity. Their new work suggests a potential way to do so by reducing glycerol and glyceraldehyde's health-draining effects.

"The discovery was unexpected. We went after a very well-supported hypothesis that the secret to longevity was the activation of a cellrejuvenating process named autophagy and ended up finding an unrecognized mechanism of health and lifespan extension," said O'Rourke, of UVA's Department of Biology and the UVA School of Medicine's Department of Cell Biology.

"An exciting aspect of the discovery is that the key to switch on this longevity mechanism is the activation of two enzymes that are very well studied because of their role in ethanol detoxification. [Ethanol is the alcohol contained in beer and bourbon]. This existing knowledge greatly facilitates our search for drugs that can specifically activate this antiaging process."

Anti-aging discovery

In their search for the secrets to slow down aging, O'Rourke and her graduate student Abbas Ghaddar and postdoc Vinod Mony turned to <u>microscopic worms</u> called C. elegans. These soil dwellers share more than 70% of our genes and are an invaluable tool for <u>biomedical research</u>; two Nobel prizes in medicine were awarded to discoveries made using this worm exclusively.

Prior aging research in worms, mice and <u>human cells</u> made O'Rourke and others in the field suspect that the key to extending lifespan was to activate autophagy, a process that renews broken and old parts in our cells. But O'Rourke and her collaborators were surprised to find that



wasn't necessary—the scientists improved the worms' health and lifespan by 50% with no increase in autophagy at all.

They did this by capitalizing on a mechanism they discovered and named AMAR, the Sanskrit word for immortality. AMAR, in this instance, stands for "Alcohol and aldehyde-dehydrogenase Mediated Anti-aging Response." In short, the scientists found that they could prompt an anti-aging response by putting the spurs to a particular gene, adh-1.

Doing so prompted the gene to produce more of an enzyme, alcohol dehydrogenase, that prevented the toxicity caused by glycerol and, indirectly, glyceraldehyde. The result was that the worms lived longer, healthier lives.

Findings in lab models such as worms and mice don't always hold true in people, of course. So the researchers took several more steps to see if their lead was as promising as it appeared. First, they confirmed that the enzyme had similar beneficial effects on lifespan in another lab model, yeast.

Then they scoured through research looking at gene activity in creatures, including humans, who had undergone fasting or calorie restriction because both fasting and <u>calorie restriction</u> are known to extend healthspan and lifespan. Sure enough, the scientists found increased levels of the anti-aging enzymes in all the mammals tested, including in humans.

The scientists suspect that our levels of glycerol and glyceraldehyde naturally increase over time because they are toxic byproducts of fat, which we store more of as we age. Thus, AMAR may offer a way to head off the fat-derived toxicity, extend the number of years we live in good health, and maybe help us shed some extra pounds, too.



"We hope to attract interest in developing therapeutics that target AMAR," said O'Rourke, who is part of UVA's Robert M. Berne Cardiovascular Research Center.

"With <u>age-related diseases</u> currently being the major health burden for patients, their families and the healthcare system, targeting the process of aging itself would be most effective way to reduce this burden and increase the number of years of independent healthy living for all of us."

The researchers have published their findings in the scientific journal *Current Biology*.

More information: Abbas Ghaddar et al, Increased alcohol dehydrogenase 1 activity promotes longevity, *Current Biology* (2023). DOI: 10.1016/j.cub.2023.01.059

Provided by University of Virginia

Citation: Detoxing body of two fat by-products could extend lifespan, researchers discover (2023, March 9) retrieved 26 April 2024 from <u>https://phys.org/news/2023-03-detoxing-body-fat-by-products-lifespan.html</u>

This document is subject to copyright. Apart from any fair dealing for the purpose of private study or research, no part may be reproduced without the written permission. The content is provided for information purposes only.