

Deletion of FAT10 gene reduces body fat, slows down aging in mice

March 24 2014



A single gene appears to play a crucial role in coordinating the immune system and metabolism, and deleting the gene in mice reduces body fat and extends lifespan, according to new research by scientists at the Jean Mayer USDA Human Nutrition Research Center (USDA HNRCA) on Aging at Tufts University and Yale University School of Medicine. Their results are reported online today in the *Proceedings of the National Academy of Sciences*.

Based on gene expression studies of [fat tissue](#) conducted at the USDA HNRCA, the Tufts University researchers initiated studies of the role of FAT10 in adipose tissue and metabolism. "No one really knew what the FAT10 gene did, other than it was 'turned on' by inflammation and that it seemed to be increased in gynecological and gastrointestinal cancers."

said co-author Martin S. Obin, Ph.D., an adjunct scientist in the Functional Genomics Core Unit at the USDA HNRCA at Tufts University. "Turning off the FAT10 gene produces a variety of beneficial effects in the [mice](#), including reduced body [fat](#), which slows down aging and extends lifespan by 20 percent."

Typically, mice gain fat as they age. The authors observed that activation of the FAT10 gene in normal mice increases in fat tissue with age. Mice lacking FAT10 consume more food, but burn fat at an accelerated rate. As a result, they have less than half of the fat tissue found in normal, aged mice. At the same time their skeletal muscle ramps up production of an immune molecule that increases their response to insulin, resulting in reduced circulating insulin levels, protection against type 2 diabetes and longer lifespan.

The authors note that eliminating FAT10 will not fully address the dilemma of aging and weight gain. "Laboratory mice live in a lab under ideal, germ-free conditions," said Obin, who is also an associate professor at the Friedman School of Nutrition Science and Policy at Tufts University. "Fighting infection requires energy, which can be provided by stored fat. Mice without the FAT10 gene might be too lean to fight infection effectively outside of the laboratory setting. More research is needed to know how to achieve that balance in mice and then hopefully, at some point, people."

The possibilities for future research of FAT10 are exciting. Recent high-profile studies reported that FAT10 interacts with hundreds of other proteins in cells. Now the Tufts and Yale researchers have demonstrated that it impacts immune response, lipid and glucose metabolism, and mitochondrial function.

"Now there is dramatic road map for researchers looking at all of the proteins that FAT10 gets involved with," said co-first and corresponding

author Allon Canaan, Ph.D., an associate scientist in the Department of Genetics at Yale. "Blocking what FAT10 does to coordinate immunity and metabolism could lead to new therapies for metabolic disease, metabolic syndrome, cancer and healthy aging, because when we knock it out the net result is mice live longer."

Canaan and colleagues initially developed the FAT10-deficient mouse to study the role of FAT10 in sepsis. In an attempt to increase sensitivity for sepsis, Canaan aged the FAT10 knockout mice and made the discovery that mice lacking the gene were lean and aged more slowly. The mice appear younger and more robust than comparably-aged normal mice, have better muscle tone, and do not develop age-related tumors.

More information: Canaan A; Defuria J; Perelman E; Schulz V; Seay M; Tuck D; Flavell R; Snyder M; Obin M; and Weissman S. "Extended Lifespan and Reduced Adiposity in Mice Lacking the FAT10 Gene." *Proceedings of the National Academy of Sciences*. Published online ahead of print March 24, 2014.

www.pnas.org/cgi/doi/10.1073/pnas.1323426111

Provided by Tufts University

Citation: Deletion of FAT10 gene reduces body fat, slows down aging in mice (2014, March 24) retrieved 2 May 2024 from

<https://medicalxpress.com/news/2014-03-deletion-fat10-gene-body-fat.html>

<p>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.</p>
--