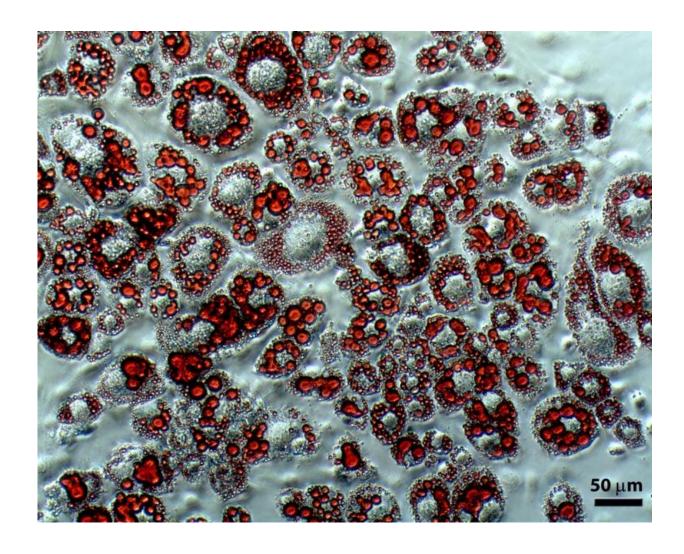


## New fat cell metabolism research could lead to new ways to treat diabetes and obesity

November 16 2015



Microscopy image depicting fat cells (or adipocytes) after differentiation. The cells are stained with Oil Red O, which highlights lipid or fat droplets that accumulate with the fat cells. The metabolic studies described here indicated that fat cells produce these fatty acids, in part, from essential amino acids rather than sugar only. Credit: Metabolic Systems Biology lab, UC San Diego Jacobs School



of Engineering

Researchers at the University of California, San Diego report new insights into what nutrients fat cells metabolize to make fatty acids. The findings pave the way for understanding potential irregularities in fat cell metabolism that occur in patients with diabetes and obesity and could lead to new treatments for these conditions. The researchers published their findings online in the Nov. 16 issue of *Nature Chemical Biology*.

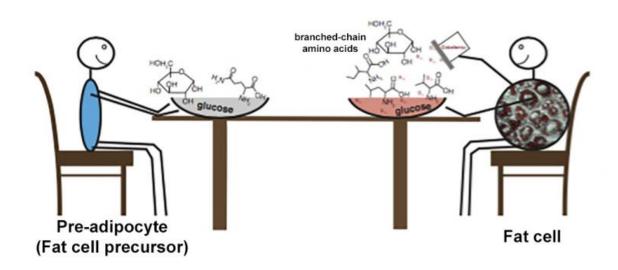
"This study highlights how specific tissues in our bodies use particular nutrients. By understanding fat cell metabolism at the molecular level, we are laying the groundwork for further research to identify better drug targets for treating diabetes and obesity," said Christian Metallo, a bioengineering professor in the Jacobs School of Engineering at UC San Diego and senior author of the study. Metallo is affiliated with the Institute for Engineering in Medicine, the Moore's Cancer Center, and the CHO Systems Biology Center, all at UC San Diego.

In the new study, researchers discovered that as <u>fat cells</u> develop, they change what types of nutrients they metabolize to produce fat and energy. Pre-adipocytes, which are precursors to fat cells, preferentially consume glucose, a simple sugar, to grow and make energy. But when pre-adipocytes become fat cells, researchers found that they metabolize not just glucose, but also branched-chain amino acids, a small set of the <u>essential amino acids</u> for humans.

This finding is important because it shows that fat cells play an important role in regulating the body's levels of branched-chain amino acids—which are typically elevated in individuals with diabetes and obesity.



"We've taken a step towards understanding why these <u>amino acids</u> are accumulating in the blood of diabetics and those suffering from obesity," said Courtney Green, a bioengineering Ph.D. student at UC San Diego and first author of the study. "The next step is to understand how and why this metabolic pathway becomes impaired in the fat cells of these individuals."



Cartoon illustrating the difference between the metabolic intake of fat cell precursors (left; glucose) versus differentiated fat cells (right; glucose and branched-chain amino acids). Credit: Metabolic Systems Biology lab, UC San Diego Jacobs School of Engineering

Metallo and his team studied the metabolism of fat cells from the preadipocyte stage throughout the <u>fat cell differentiation</u> process. They induced pre-adipocytes to differentiate into fat cells and cultured the cells in media containing nutrients enriched with carbon-13 isotopes, a form of carbon atoms that are used as metabolic tracers in cells, animals, and people. Through this method, researchers were able to trace what



carbon-based nutrients the cells metabolized and what they produced at different stages of the cell differentiation process.

"We are curious about how different cells in our body, such as fat cells, consume and metabolize their surrounding <u>nutrients</u>. A better understanding of how these biochemical pathways are used by cells could help us find new approaches to treat diseases such as cancer or diabetes," said Metallo.

**More information:** Courtney R Green et al. Branched-chain amino acid catabolism fuels adipocyte differentiation and lipogenesis, *Nature Chemical Biology* (2015). DOI: 10.1038/nchembio.1961

## Provided by University of California - San Diego

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