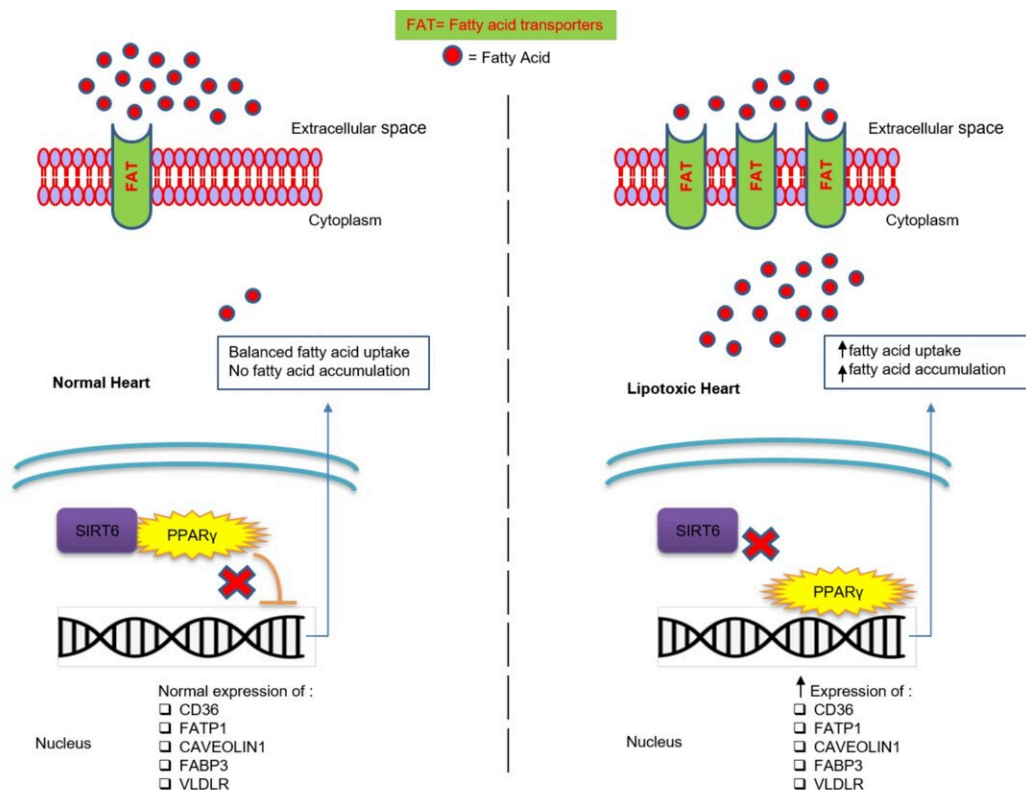


Regulating fatty acid uptake in cardiac muscle cells: A new insight

June 29 2021, by Sritama Bose



Schematic showing the mechanism by which SIRT6 regulates fatty acid uptake through PPAR γ transcription factor in the heart. Credit: Khan et al., 2021/Cell Reports

Fatty acids are formed when fat breaks down during digestion. While many of the body's organs use glucose as their primary energy source,

the heart derives most of its required energy (over 70%) from the oxidation of fatty acids. These are crucial for sustaining cardiomyocytes—cardiac muscle cells that control the rhythmic beating of the heart. However, accumulation of excess fatty acids in cardiomyocytes triggers harmful responses, often leading to severe cardiac diseases.

A recent study published in *Cell Reports* by a team of researchers from India and the US, led by scientists at the Indian Institute of Science (IISc), provides important insights into how fatty [acid](#) uptake is regulated in [cardiomyocytes](#).

"We identified a mechanism by which fatty acid [transport](#) (to cardiomyocytes) is critically regulated by a protein called SIRT6," says team lead Ravi Sundaresan, Associate Professor in the Department of Microbiology and Cell Biology, IISc. The study shows that SIRT6 could be a potential therapeutic target for treating several metabolic diseases affecting the heart.

Cardiomyocytes have several fatty acid transporters—specific proteins enhancing the uptake of [fatty acids](#) from the blood into the cells—to ensure sufficient supply. The authors say that this is the first study to show that SIRT6 regulates the genes responsible for the formation of these transporter proteins in cardiomyocytes.

The team observed that cardiomyocytes devoid of the SIRT6 protein had higher levels of fatty acid transporters, resulting in higher uptake and accumulation of fatty acids. They also showed that increasing the level of SIRT6 in cardiomyocytes lowered the levels of these transporters, thereby reducing fatty acid uptake and accumulation. The researchers carried out most of the studies in experimental mice models.

SIRT6 belongs to a family of proteins called sirtuins, which are

important biological enzymes that require specific molecules, called cofactors, to function. Surprisingly, the researchers found that SIRT6 neither functions as an enzyme nor does it require any co-factor to regulate fatty acid uptake in cardiomyocytes. Rather, it does so by binding with a specific protein involved in the production of fatty acid transporters. Sundaresan previously led another [study](#) which showed that SIRT6 can regulate [protein](#) synthesis without functioning as an enzyme.

"Fatty acids are very useful for the heart, if properly utilized," Sundaresan explains. However, accumulation of more fatty acids than the heart can use in cardiomyocytes causes lipotoxicity—a condition in which accumulation of lipids can eventually have grave consequences such as inflammation and cell death. Notably, diabetes and obesity are among certain pathological conditions in which levels of fatty acid transporters are reportedly higher. Under such conditions, activating SIRT6 and increasing its expression might be a beneficial therapeutic intervention, the researchers suggest.

The team also included Narayanaswamy Srinivasan, Professor in the Molecular Biophysics Unit, IISc, researchers from the Tata Institute of Fundamental Research, India, and Harvard Medical School, U.S..

More information: Danish Khan et al, SIRT6 transcriptionally regulates fatty acid transport by suppressing PPAR γ , *Cell Reports* (2021). [DOI: 10.1016/j.celrep.2021.109190](https://doi.org/10.1016/j.celrep.2021.109190)

Provided by Indian Institute of Science

Citation: Regulating fatty acid uptake in cardiac muscle cells: A new insight (2021, June 29) retrieved 25 April 2024 from <https://medicalxpress.com/news/2021-06-fatty-acid-uptake-cardiac-muscle.html>

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.