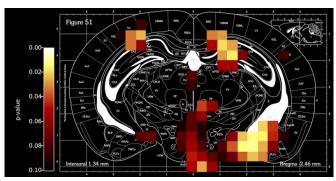


Fat tissue in energy saving mode

31 March 2016



Active regions of the AgRP neurons in mice brain

Scientists at the Max Planck Institute for Metabolism Research in Cologne have discovered brain cells that not only tell hungry mice to search for food, but also to limit blood sugar use by the brown fat tissue. This could ensure the survival of mice when they cannot find enough to eat.

What happens when we get hungry? How does the brain control energy expenditure? To find answers to these questions a research team led by Jens Brüning, director of the Max Planck Institute for Metabolism Research, analyzed the function of a specific class of nerve cells in the brain, called AgRP neurons. "These nerve cells are located in the hypothalamus, which can be seen as the brain command center controlling appetite", explains Sophie Steculorum, one of the authors of the study and associate of Brüning. "It has already been known for a few years that these cells control feeding behavior in hunger states".

In the recent study, researchers demonstrated that in mice AgRP neurons use an additional mechanism to regulate the sugar metabolism in the body. "These AgRP neurons tell the body to use less blood sugar when the mouse is hungry and cannot find food", says Johan Ruud, co-author of the study.

Reprogramming the brown fat tissue

The cologne scientists could show that the AgRP neurons are connected with the <u>brown adipose</u> <u>tissue</u>, also called brown fat. "When the neurons are activated, the cells in this brown fat tissue are reprogrammed – they produce a different set of proteins, for example high levels of myostatin", explains Ruud.

The protein myostatin is usually found in muscle cells and slows down muscle growth. Now the scientists could show for the first time that myostatin directly controls the sensitivity of the brown fat to insulin, which dictates how the body utilizes blood sugar.

AgRP neurons in human

AgRP neurons, myostatin and insulin are not only found in mice, but also in humans. Obesity and type-2-diabetes are likely associated with chronic activation of AgRP- neurons, at least in mice. The mechanism could explain why AgRP neurons are connected with those diseases. "Next we want to find out whether the cells also control the sensitivity of brown fat to insulin in humans", explains Steculorum.

More information: Sophie M. Steculorum et al. AgRP Neurons Control Systemic Insulin Sensitivity via Myostatin Expression in Brown Adipose Tissue, *Cell* (2016). DOI: 10.1016/j.cell.2016.02.044

Provided by Max Planck Society

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APA citation: Fat tissue in energy saving mode (2016, March 31) retrieved 16 May 2021 from https://medicalxpress.com/news/2016-03-fat-tissue-energy-mode.html

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