

New link between diet, intestinal stem cells and disease discovered

November 19 2021





Intestinal stem cells in green, cell division in red. Credit: Helmholtz Munich / Anika Böttcher

The intestine is essential for maintaining our energy balance and is a master at reacting quickly to changes in nutrition and nutrient balance. It manages to do this with the help of intestinal cells that among other things are specialized in the absorption of food components or the secretion of hormones. In adult humans, the intestinal cells regenerate every five to seven days. The ability to constantly renew and develop all types of intestinal cells from intestinal stem cells is crucial for the natural adaptability of the digestive system. However, a long-term diet high in sugar and fat disrupts this adaptation and can contribute to the development of obesity, type 2 diabetes, and gastrointestinal cancer.

The <u>molecular mechanisms</u> behind this maladaptation are part of the research field of Heiko Lickert and his group at Helmholtz Munich and the Technical University of Munich. The scientists assume that <u>intestinal</u> <u>stem cells</u> play a special role in maladaptation. Using a <u>mouse model</u>, the researchers investigated the effects of a high-sugar and high-fat <u>diet</u> and compared it with a control group.

From high-calorie diet to increased risk of gastrointestinal cancer

"The first thing we noticed was that the small intestine increases greatly in size on the high-calorie diet," says study leader Anika Böttcher. "Together with Fabian Theis' team of computational biologists at Helmholtz Munich, we then profiled 27,000 <u>intestinal cells</u> from control diet and high fat/high sugar diet-fed mice. Using new machine learning techniques, we thus found that intestinal stem cells divide and differentiate significantly faster in the mice on an unhealthy diet." The



researchers hypothesize that this is due to an upregulation of the relevant signaling pathways, which is associated with an acceleration of tumor growth in many cancers. "This could be an important link: Diet influences metabolic signaling, which leads to excessive growth of intestinal stem cells and ultimately to an increased risk of gastrointestinal cancer," says Böttcher.

With the help of this high-resolution technique, the researchers have also been able to study rare cell types in the intestine, for example, hormonesecreting cells. Among their findings, they were able to show that an unhealthy diet leads to a reduction in serotonin-producing cells in the intestine. This can result in intestinal inertia (typical of <u>diabetes</u> mellitus) or increased appetite. Furthermore, the study showed that the absorbing <u>cells</u> adapt to the <u>high-fat diet</u>, and their functionality increases, thus directly promoting weight gain.

Important basic research for non-invasive therapies

These and other findings from the study lead to a new understanding of disease mechanisms associated with a high-calorie diet. "What we have found out is of crucial importance for developing alternative non-invasive therapies," says study leader Heiko Lickert, in summarizing the results. To date, there is no pharmacological approach to prevent, stop or reverse obesity and diabetes. Only <u>bariatric surgery</u> causes permanent weight loss and can even lead to remission of diabetes. However, these surgeries are invasive, non-reversible and costly to the healthcare system. Novel non-invasive therapies could happen, for example, at the hormonal level through targeted regulation of serotonin levels. The research group will examine this and other approaches in subsequent studies.

About the people



Heiko Lickert and Anika Böttcher conduct research at the Helmholtz Diabetes Center of Helmholtz Munich. They specialize in the development of regenerative treatment approaches for numerous widespread diseases associated with impaired gut function. Lickert heads the Institute for Diabetes and Regeneration Research and is a professor at the Technical University of Munich (TUM). Both are scientists at the German Center for Diabetes Research (DZD).

The current study has been published as a cover story in *Nature Metabolism*.

More information: Alexandra Aliluev et al, Diet-induced alteration of intestinal stem cell function underlies obesity and prediabetes in mice, *Nature Metabolism* (2021). DOI: 10.1038/s42255-021-00458-9

Provided by Helmholtz Association of German Research Centres

Citation: New link between diet, intestinal stem cells and disease discovered (2021, November 19) retrieved 5 May 2024 from <u>https://medicalxpress.com/news/2021-11-link-diet-intestinal-stem-cells.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.