

Young vessels rejuvenate aged insulinproducing beta cells

November 17 2014



Per-Olof Berggren, Ph.D., is a Professor of Experimental Endocrinology at Karolinska Institutet in Sweden. Credit: Ulf Sirborn

A recent study published in the journal *PNAS* shows that young capillary vessels rejuvenate aged pancreatic islets. The finding challenges



prevailing views on the causes of age-dependent impaired glucose balance regulation, a condition that often develops into diabetes type 2. The international research team behind the study now suggests that targeting inflammation and fibrosis in the small blood vessels of the pancreatic islets may offer a new way of treatment for age-dependent dysregulation of blood glucose levels.

"This is an unexpected but highly important finding, which we expect will have a significant impact on diabetes research in the future. The results indicate that beta cell function does not decline with age, and instead suggest that islet function is threatened by an age-dependent impairment of vessels that support them with oxygen and nutrients", says Per-Olof Berggren at the Rolf Luft Research Center for Diabetes and Endocrinology at Karolinska Institutet, who led the study together with Alejandro Caicedo at University of Miami Miller School of Medicine and Hong Gil Nam at DGIST in Republic of Korea.

Beta cells are located in the <u>pancreatic islets</u> and secrete the hormone insulin, that play a key role in regulating <u>blood glucose levels</u>. Ageing may lead to a progressive decline in glucose regulation which may contribute to diabetes development. Generally, it has been assumed that this is due to reduced capacity of the beta cell to secrete insulin or multiply.

"In the study we challenged the view that the age-dependent impairment in <u>glucose homeostasis</u> is solely due to intrinsic, dysfunction of <u>islet cells</u>, and hypothesized that it is instead affected by systemic aging factors", says first author Joana Almaca at the Diabetes Research Institute, University of Miami.

In marked contrast to the widely held notion that the insulin-producing pancreatic beta cell loses function with wear and tear, the researchers now show that mouse and human beta cells are fully functional at



advanced age. When replacing the islet vasculature in aged islet grafts with young capillaries, the investigators found that the islets were rejuvenated and glucose homeostasis fully restored.

"While expanding beta cell mass may still be desirable for future <u>diabetes</u> therapy, improving the local environment of the otherwise healthy aged beta cell could prevent age-associated deterioration in glucose homeostasis and thereby promote healthy ageing, which is conceptually novel and highly exciting", says Per-Olof Berggren.

More information: "Young Capillary Vessels Rejuvinate Aged Pancreatic Islets", Joana Almaça, Judith Molina, Rafael Arrojo, Rafael Arrojo e Drigo, Midhat H. Abdulreda, Won Bae Jeon, Per-Olof Berggren, Alejandro Caicedo, and Hong Gil Nam, *PNAS*, Online Early Edition the week of Nov 17-Nov 21, 2014. <u>www.pnas.org/cgi/doi/10.1073/pnas.1414053111</u>

Provided by Karolinska Institutet

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