

## Researchers gain insights into cellular processes associated with diabetes

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Credit: Diabetes

Two new studies from the Diabetes Center of Excellence at UMass Medical School investigate functions of the insulin-producing beta cells



that are destroyed in patients with type 1 diabetes, and a reliable method to measure beta cell replication in individuals.

"These studies have improved our understanding of the disease process in the pancreas and have implications for preventing and treating both type 1 and type 2 diabetes," said David Harlan, MD, the William and Doris Krupp Professor in Medicine, professor of medicine and codirector of the Diabetes Center of Excellence.

Harlan and Kent labs identify immunopathogenic pathway in type 1 diabetes

The labs of Dr. Harlan and Sally Kent, Ph.D., associate professor of medicine, led a collaboration that studied the aberrant expression of an immune system cell marker on beta <u>cells</u> in those with type 1 diabetes. The findings suggest that beta cells may interact directly with immune system cells to play a role in the development of type 1 diabetes.

"For still incompletely understood reasons, a big part of the disease process is that the individual's immune system somehow begins to attack and kill the insulin-producing cells in the pancreas," said Dr. Kent. "The greatest genetic risk for type 1 diabetes is driven by the expression of important immune genes called human leukocyte antigen class II (HLA class II) and yet human beta cells were not thought capable of expressing HLA class II and other important genes supporting its function. This new research definitively shows that beta cells from individuals with type 1 diabetes express these important gene products."

## Alonso lab confirms method to measure replication in beta cells

A study led by Rohit Sharma, Ph.D., also published in the journal *Diabetes*, found that a well-established method for quantifying replication in other cell types applies to beta cells as well.



"Research studying the pathways important for increasing beta cell number require a reliable method to assess beta cell replication. A major concern in beta cells was to distinguish DNA repair from DNA replication. We have now addressed this concern," said Dr. Sharma, instructor in medicine in the lab of Laura Alonso, MD, the George F. and Sybil H. Fuller Foundation Term Chair in Diabetes and associate professor of medicine. "Our findings suggest that measuring incorporation of the BrdU (thymidine analogue) nucleotide, an engineered-for-study structural unit of DNA, is a reliable method to assess beta cell replication. Quantifying beta cell replication provides a method for high throughput chemical screens to identify small chemical compounds capable of making beta cells replicate in order to overcome diabetes."

**More information:** Rohit B. Sharma et al. DNA Damage Does Not Cause BrdU Labeling of Mouse or Human  $\beta$ -Cells, *Diabetes* (2019). DOI: 10.2337/db18-0761

Mark A. Russell et al. HLA Class II Antigen Processing and Presentation Pathway Components Demonstrated by Transcriptome and Protein Analyses of Islet β-Cells From Donors With Type 1 Diabetes, *Diabetes* (2019). DOI: 10.2337/db18-0686

## Provided by University of Massachusetts Medical School

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