

# Team cures diabetes in mice without side effects

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Bruno Doiron, Ph.D., left, and Ralph DeFronzo, M.D., of UT Health San Antonio co-invented a patented technique that has cured diabetes in mice for one year without side effects. Credit: UT Health San Antonio

A potential cure for Type 1 diabetes looms on the horizon in San Antonio, and the novel approach would also allow Type 2 diabetics to stop insulin shots.

The discovery, made at The University of Texas Health Science Center, now called UT Health San Antonio, increases the types of [pancreatic cells](#) that secrete [insulin](#).

UT Health San Antonio researchers have a goal to reach human clinical trials in three years, but to do so they must first test the strategy in large-animal studies, which will cost an estimated \$5 million.

Those studies will precede application to the U.S. Food and Drug Administration for Investigational New Drug (IND) approval, Bruno Doiron, Ph.D., a co-inventor, said.

The scientists received a U.S. patent in January, and UT Health San Antonio is spinning out a company to begin commercialization.

The strategy has cured diabetes in mice.

"It worked perfectly," Dr. Doiron, assistant professor of medicine at UT Health, said. "We cured mice for one year without any side effects. That's never been seen. But it's a mouse model, so caution is needed. We want to bring this to large animals that are closer to humans in physiology of the endocrine system."

Ralph DeFronzo, M.D., professor of medicine and chief of the Division of Diabetes at UT Health, is co-inventor on the patent. He described the therapy:

"The pancreas has many other cell types besides [beta cells](#), and our approach is to alter these [cells](#) so that they start to secrete insulin, but

only in response to glucose [sugar]," he said. "This is basically just like beta cells."

Insulin, which lowers [blood sugar](#), is only made by beta cells. In Type 1 diabetes, beta cells are destroyed by the immune system and the person has no insulin. In Type 2 diabetes, beta cells fail and insulin decreases. At the same time in Type 2, the body doesn't use insulin efficiently.

The therapy is accomplished by a technique called gene transfer. A virus is used as a vector, or carrier, to introduce selected genes into the pancreas. These genes become incorporated and cause digestive enzymes and other cell types to make insulin.

Gene transfer using a viral vector has been approved nearly 50 times by the U.S. Food & Drug Administration to treat various diseases, Dr. DeFronzo said. It is proven in treating rare childhood diseases, and Good Manufacturing Processes ensure safety.

Unlike beta cells, which the body rejects in Type 1 diabetes, the other cell populations of the pancreas co-exist with the body's immune defenses.

"If a Type 1 diabetic has been living with these cells for 30, 40 or 50 years, and all we're getting them to do is secrete insulin, we expect there to be no adverse immune response," Dr. DeFronzo said.

## **Second-by-second sugar control**

The therapy precisely regulates blood sugar in mice. This could be a major advance over traditional insulin therapy and some diabetes medications that drop blood sugar too low if not closely monitored.

"A major problem we have in the field of Type 1 diabetes is

hypoglycemia (low blood sugar)," Dr. Doiron said. "The [gene transfer](#) we propose is remarkable because the altered cells match the characteristics of beta cells. Insulin is only released in response to glucose."

People don't have symptoms of [diabetes](#) until they have lost at least 80 percent of their beta cells, Dr. Doiron said.

"We don't need to replicate all of the insulin-making function of beta cells," he said. "Only 20 percent restoration of this capacity is sufficient for a cure of Type 1."

Provided by University of Texas Health Science Center at San Antonio

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