

# **A Comparative Study of CRISPR-Cas9 Treatment and Conventional Diabetes Treatment**

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# Introduction

Overview of diabetes and why it matters

# Purpose statement



## What is Diabetes?

- Chronic disease affecting blood sugar regulation
- Caused by lack of insulin or insulin resistance
- Leads to long-term health complications



## Type 1 diabetes

- Autoimmune destruction of pancreatic beta cells
- Body stops producing insulin
- Requires lifelong insulin therapy



## Why it matters.

- 537 million adults worldwide have diabetes
- Type 1 diabetes increasing 3-5% per year
- Over 1 million children and teens affected



**3.9 million (9.7%)**

Of people aged 1 year and older live with diagnosed diabetes  
(Canada, 2023–2024)

[\\*https://health-infobase.canada.ca/diabetes/\\*](https://health-infobase.canada.ca/diabetes/)

02

**Current**

**Diabetes**

**Treatments**

How diabetes is managed today

# Insulin Replacement Therapy (Basal-Bolus)

- **Overview:**
  - Multiple daily injections or insulin pump
  - Basal insulin: long/intermediate-acting
  - Bolus insulin: rapid/short-acting, taken before meals
- **Safety:**
  - Hypoglycemia risk
  - Possible weight gain
  - Does not cure diabetes
- **Cost & Inconveniences:**
  - Moderate-high cost
  - Daily injections, frequent monitoring
  - Lifelong dependency
- **Future Potential:**
  - Could restore natural insulin production
  - May reduce or eliminate injections

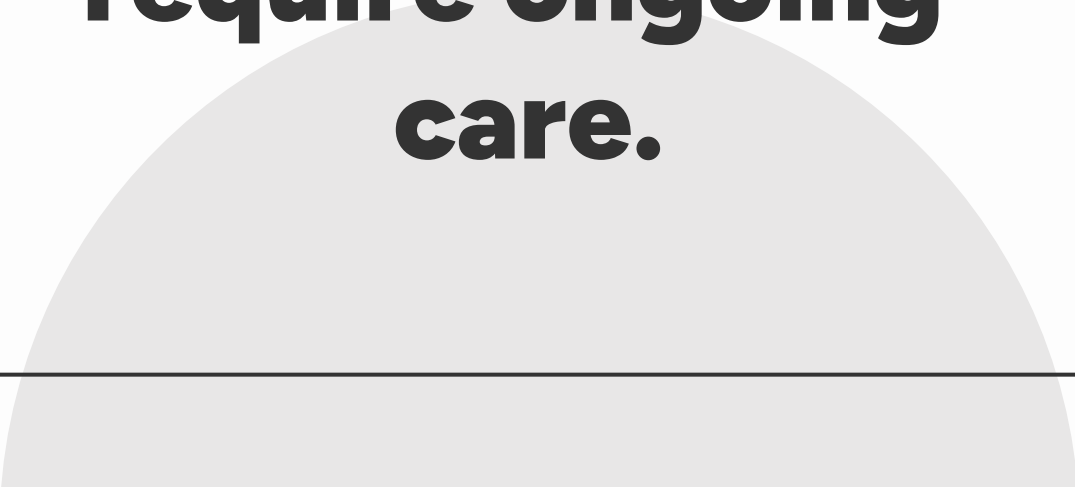
# Artificial Pancreas Systems

- **Overview:**
  - Continuous glucose monitoring (CGM) + insulin pump
  - Automated insulin delivery based on glucose readings
  - Mimics pancreatic feedback
- **Safety:**
  - Reduces hypoglycemia compared to manual injections
  - Possible device malfunctions
  - Skin irritation possible
- **Cost & Inconveniences:**
  - High upfront cost + ongoing supplies
  - Must wear device constantly
  - Device alarms, calibration, and technical learning curve
- **Future Potential:**
  - Gene-edited  $\beta$ -cells could replace devices
  - Reduces dependency on technology

# Xenotransplantation

- **Overview:**
  - Transplant insulin-producing pig islet cells
  - Requires immune-suppressing drugs
- **Safety:**
  - High risk of immune rejection and infection
  - Ethical concerns about animal-to-human transplants
- **Cost & Inconveniences:**
  - Very high: surgery + lifelong immune-suppressing drugs
  - Major surgery with recovery risk
  - Ongoing monitoring to prevent rejection
- **Future Potential:**
  - Could make donor cells safer and more human-compatible
  - Still experimental, mostly preclinical research

**These treatments  
manage diabetes  
but don't cure it and  
require ongoing  
care.**



**03**

# **Gene Editing Technologies**

CRISPR, TALENs, and ZFN overview

# CRISPR Cas9

- **How it works:** RNA-guided enzyme edits DNA at specific sites; can be done in stem cells (ex vivo) or directly in the body (in vivo)
- **Safety:** Temporary off-target effects possible; newer versions designed to be safer
- **Cost:** Medium-high; may reduce long-term costs if curative
- **Patient Impact:** Experimental; requires specialized centers; careful monitoring; ethical concerns for inheritable changes
- **Benefits:** Fixes disease at the genetic level; may restore natural insulin; targets root cause
- **Applications:** Used in rare genetic disorders (e.g., CPS1 deficiency); disease modeling

# Transcription Activator-Like Effector Nucleases (TALENs)

- **How it works:** Protein-based system cuts specific DNA sequences; each protein recognizes one DNA base; done in stem cells.
- **Safety:** More precise than ZFNs, fewer off-target effects; can still incorrectly bind DNA; delivery into human cells challenging.
- **Cost:** High; custom protein design is time-consuming.
- **Patient Impact:** Experimental; complex lab work; not widely available clinically.
- **Benefits:** Can correct genetic mutations in stem cells; preclinical research shows potential.
- **Applications:** Research on genetic disorders; disease modeling in labs.

# Zinc-Finger Nucleases (ZFN)

- **How it works:** Protein-based system edits DNA using engineered zinc finger domains; first widely used tool
- **Safety:** Higher off-target risk; can damage DNA; may trigger cell death
- **Cost:** High; complex protein engineering needed for each target
- **Patient Impact:** Experimental; difficult to redesign; limited flexibility
- **Benefits:** Foundation for modern gene-editing tools; early stem cell gene correction
- **Applications:** Sickle cell therapy research; early gene therapy studies.

**These gene-editing  
tools are powerful  
but mostly still  
experimental.**



**04**

# **Data Comparison**

Table comparing treatments and gene editing

**A Comparative Table of CRISPR Cas9 Treatment and Conventional Diabetes Treatment**

		<b>Factors</b>					
	<b>Treatment type</b>	<b>Drug Delivery</b>	<b>Safety</b>	<b>Cost</b>	<b>Patient Inconveniences</b>	<b>CRISPR Solution</b>	<b>Applications</b>
<b>Conventional Diabetes Treatment</b>	<b>Insulin Replacement Therapy (Basal-Bolus)</b>	<ul style="list-style-type: none"> <li>Multiple daily injections (MDI) or insulin pump</li> <li>Basal insulin: long-acting or intermediate-acting</li> <li>Controls fasting glucose and between meals</li> <li>Bolus insulin: rapid-acting or short-acting</li> <li>Taken 0–15 min before meals</li> <li>Controls post-meal glucose spikes</li> <li>Based on lifestyle and treatment goals</li> </ul>	<ul style="list-style-type: none"> <li>Risk of hypoglycemia</li> <li>Possible weight gain</li> <li>Requires precise timing and dose adjustments</li> <li>Long-term therapy does not cure diabetes</li> </ul>	<ul style="list-style-type: none"> <li>Moderate to high depending on insulin type</li> <li>Ongoing lifelong cost (insulin + monitoring supplies)</li> </ul>	<ul style="list-style-type: none"> <li>Daily injections (sometimes 4+ per day)</li> <li>Frequent blood glucose monitoring</li> <li>Careful diet (carbs)</li> <li>Lifelong dependency</li> </ul>	<ul style="list-style-type: none"> <li>Could restore natural <math>\beta</math>-cell insulin production</li> <li>Reduces need for lifelong insulin injections</li> <li>Potential long-term or permanent correction</li> </ul>	<ul style="list-style-type: none"> <li>Standard treatment for Type 1 Diabetes</li> </ul>
	<b>Artificial pancreas systems</b>	<ul style="list-style-type: none"> <li>Continuous glucose monitor (CGM) + insulin pump</li> <li>Automated insulin delivery based on glucose readings</li> <li>Closed-loop system recreates pancreatic feedback</li> </ul>	<ul style="list-style-type: none"> <li>Reduces hypoglycemia compared to manual injections</li> <li>Risk of device malfunction</li> <li>Skin infections at contact sites</li> </ul>	<ul style="list-style-type: none"> <li>High initial device cost</li> <li>Constant sensor and pump supply expenses</li> </ul>	<ul style="list-style-type: none"> <li>Must wear device continuously</li> <li>Device alarms and calibration</li> <li>Technical learning curve</li> </ul>	<ul style="list-style-type: none"> <li>Gene-edited <math>\beta</math>-cells could replace mechanical insulin delivery</li> <li>Eliminates reliance on wearable devices</li> </ul>	<ul style="list-style-type: none"> <li>Used in clinical practice</li> <li>Improves glycemic control and reduces A1C</li> </ul>
	<b>Xenotransplantation</b>	<ul style="list-style-type: none"> <li>Surgical transplantation of insulin-producing islet cells (often from pigs)</li> <li>Patient must take drugs to prevent immune system attack</li> </ul>	<ul style="list-style-type: none"> <li>High risk of immune rejection</li> <li>Risk of infection</li> <li>Ethical and cross-species concerns</li> </ul>	<ul style="list-style-type: none"> <li>Very high (surgical procedure + lifelong medicine)</li> </ul>	<ul style="list-style-type: none"> <li>Major surgery</li> <li>Lifelong immune suppression</li> <li>Risk of transplant failure</li> </ul>	<ul style="list-style-type: none"> <li>Gene-edit donor animals to reduce immune rejection</li> <li>Modify pig cells to make them more compatible to humans</li> </ul>	<ul style="list-style-type: none"> <li>Experimental and preclinical research</li> </ul>
<b>CRISPR Cas9 Treatment</b>	<b>CRISPR Cas9</b>	<ul style="list-style-type: none"> <li>RNA-guided nuclease system</li> <li>Cas9 enzyme guided by gRNA to specific DNA sequence</li> <li>Can be delivered ex vivo (edit stem cells outside body) or in vivo</li> </ul>	<ul style="list-style-type: none"> <li>Temporary off-target effects possible</li> <li>New variants reduce risk</li> <li>New versions are designed to be safer</li> </ul>	<ul style="list-style-type: none"> <li>Medium–High (specialized lab procedures)</li> <li>Potential long-term cost reduction if curative</li> </ul>	<ul style="list-style-type: none"> <li>Requires specialized medical centers</li> <li>Still experimental for diabetes</li> <li>Ethical concerns (especially germline editing)</li> <li>Careful monitoring required</li> </ul>	<ul style="list-style-type: none"> <li>Fixes the genetic cause of the disease</li> <li>Potential restoration of natural insulin production</li> <li>May prevent hereditary transmission</li> <li>Targets root cause, not just symptoms</li> </ul>	<ul style="list-style-type: none"> <li>Successfully used in genetic disorders (CPS1 deficiency case)</li> <li>Disease modeling and research</li> </ul>
	<b>Transcription Activator-Like Effector Nucleases (TALENs)</b>	<ul style="list-style-type: none"> <li>Ex vivo stem cell modification</li> <li>Special proteins that can be designed to cut specific DNA</li> <li>Each TALE recognizes a single DNA base (A, T, C, G)</li> <li>Requires protein engineering for each new target</li> </ul>	<ul style="list-style-type: none"> <li>More specific than ZFNs</li> <li>Fewer off-target effects than ZFNs</li> <li>Can still bind incorrect DNA sequences</li> <li>Delivery into human cells can be challenging</li> </ul>	<ul style="list-style-type: none"> <li>High (custom protein design required)</li> <li>Time-consuming development</li> </ul>	<ul style="list-style-type: none"> <li>Complex lab-based procedure</li> <li>Not widely available clinically</li> <li>Still largely experimental</li> </ul>	<ul style="list-style-type: none"> <li>CRISPR easier and faster to design (RNA-guided instead of protein redesign)</li> <li>More flexible and scalable</li> <li>Lower production cost</li> </ul>	<ul style="list-style-type: none"> <li>Stem cell gene editing</li> <li>Correction of genetic disorders in research settings</li> <li>Preclinical disease modeling</li> </ul>
	<b>Zinc-Finger nucleases (ZFN)</b>	<ul style="list-style-type: none"> <li>Ex vivo stem cell editing</li> <li>Older gene-editing tool that uses custom-built proteins to cut DNA</li> <li>First widely used gene-editing tool</li> </ul>	<ul style="list-style-type: none"> <li>Higher off-target mutation risk</li> <li>Can cause unintended DNA damage</li> <li>May trigger cell death if incorrect cuts occur</li> </ul>	<ul style="list-style-type: none"> <li>High due to complex protein engineering</li> <li>Custom-built for each DNA target</li> </ul>	<ul style="list-style-type: none"> <li>Requires specialized labs</li> <li>Difficult to redesign for new mutations</li> <li>Limited flexibility compared to CRISPR</li> </ul>	<ul style="list-style-type: none"> <li>RNA-guided system simpler and faster to program</li> <li>More precise with newer variants</li> <li>More cost-effective and adaptable</li> </ul>	<ul style="list-style-type: none"> <li>Sickle cell gene therapy research</li> <li>Early stem cell gene correction studies</li> <li>Foundation for development of modern gene-editing tools</li> </ul>

# Table

Compares  
conventional vs.  
CRISPR-based  
diabetes treatments

Conventional: manage  
blood sugar, daily  
routines, risks, high  
cost

CRISPR/TALENs/ZFNs:  
edit genes to fix root  
cause, make more  
convenient potential  
long-term cure

Mostly experimental,  
expensive now

Could transform  
future diabetes care

Promising for  
hereditary and Type 1  
diabetes

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# Conclusion

Future potential of CRISPR for diabetes + citations

# CRISPR Cas9 Impact

- Targets the root cause of Type 1 diabetes, no symptoms.
- Potential to permanently restore insulin production by fixing faulty genes.
- Example: CPS1 deficiency treatment in a 7-month-old infant.
  - Base editing changes one DNA letter without cutting both strands → safer.
  - Delivered via lipid nanoparticles with custom gRNA and mRNA.
  - Result: lower ammonia, better protein tolerance, reduced medication, improved growth



## **Key Takeaway:**

Personalized CRISPR therapies can be safe and effective

# Future Diabetes Treatments

## Goal:

Restore insulin production and regulation, aiming for long-term insulin independence.



- Type 1 diabetes cases are rising worldwide; insulin therapy manages but does not cure.
- Promising strategies for the future:
  - Artificial pancreas systems
  - Stem cell-derived beta cells
  - Xenotransplantation
  - Combination therapies

# Why Research CRISPR?

- CRISPR Cas9 is still experimental but shows strong potential.
- Could revolutionize diabetes care by addressing the root genetic causes.
- Benefits:
  - Fewer lifelong injections
  - Reduced patient burden
  - Potential permanent treatment instead of symptom management



## **Conclusion:**

Continued research is crucial for safer, more effective gene-editing therapies.

**While insulin therapy remains a temporary treatment, the emerging field of gene editing, specifically with the CRISPR Cas9 should continue to be researched and studied, because of the strong success rate of it, and potential that would permanently change the healthcare system.**

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**Thank you  
for your  
time.**

