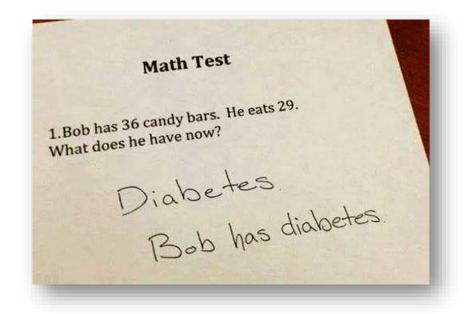


# Joe Bean, M.D. LLI Spring 2025

(Images are from a variety of sources and used only for educational purposes!)

# Diabetes

# When Sugar's Not So Sweet



### Not so long ago...

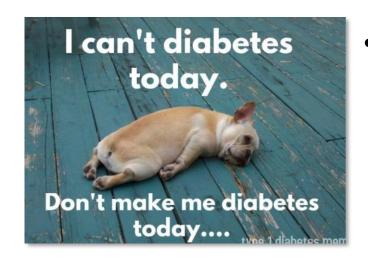


J. Bond Francisco, The Sick Child, 1893, oil on canvas, Smithsonian

 What was diabetes like for most people throughout history?



Cornelis de Bie, A Medical Practitioner Examining a Urine Flask, 1621



- Why is it important to understand this disease now?
- What will we talk about this afternoon?

#### Diabetes Statistics, U.S., 2021

Prevalence: 38.4 million (11.6%)

5% with type 1; 95% with type 2

10% undiagnosed

In seniors (65 and older) – 16.5 million

(29.2%)

New Cases: 1.2 million each year

**Diabetes** 13.5% of American Indians/Alaskan native adults

by race/ 12

12.1% of non-Hispanic Black adults

ethnicity: 1

11.7% of Hispanic adults

9.1% of Asian American adults

6.9% of non-Hispanic White adults

**Deaths:** 8<sup>th</sup> leading cause of

death in 2021 (103,000)

Cost: \$412.9 billion per year,

Average patient expense

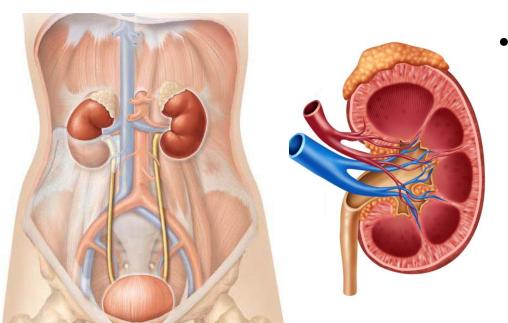
2.5x more than if they did

not have diabetes



#### What is Diabetes?

• Chronic disease named for its chief presenting symptom: Increased Urination



From Greek Dia "through"



Diabainen "to pass through"

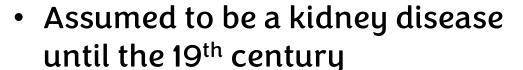


ADH probler

Diabetes <u>mellitus</u> vs. Diabetes <u>insipidus</u>











Ebers papyrus







#### A Disease with Ancient History...

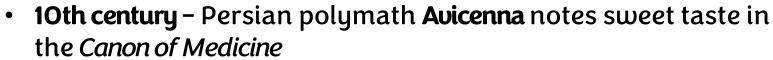
- 16<sup>th</sup> century BCE: descriptions on Egyptian medical papyri
- 5<sup>th</sup> century BCE Hindu physician Sushruta describes honeylike urine that attracts ants



3<sup>rd</sup> century BCE - Greek physician Apollonius of Memphis coins the term "diabetes"







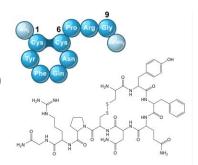




1674- English physician Thomas Willis differentiates types of "pissing evil" by taste

### The Other Diabetes...Diabetes Insipidus

 A failure to make or failure to use <u>antidiuretic hormone</u> (ADH)



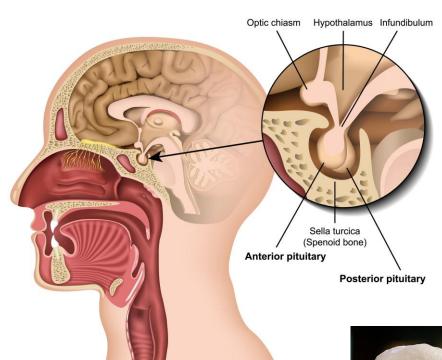


- Normally released from the pituitary gland
- Causes kidneys to keep water in the body





 Without ADH, too much water is lost in the urine



#### Diabetes Mellitus

Failure to make or failure to use insulin



The symptoms are great hair

The thegion of the left thioney, extreme

a better appetite then what is naturel

locather with a painfull sensation at the

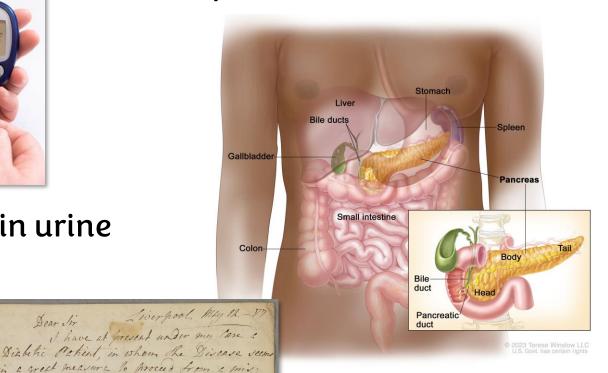
 Results in high blood sugar (hyperglycemia)





Leads to sugar in urine (glycosuria)

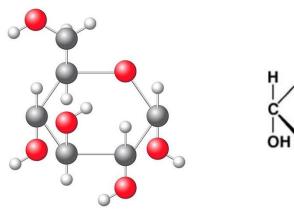
 1776 Matthew Dobson - boiled urine to dryness, crystallized "brown sugar" taste, subsequently found high glucose in blood  Normally made by the pancreas

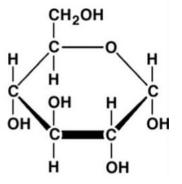


Type 1 us. Type 2

### What is Blood Sugar?

• Glucose - main energy source in cells





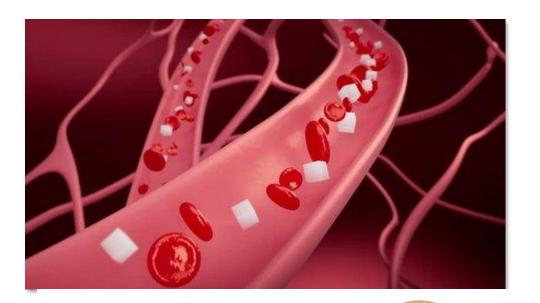
• Glycolysis and Cellular Respiration

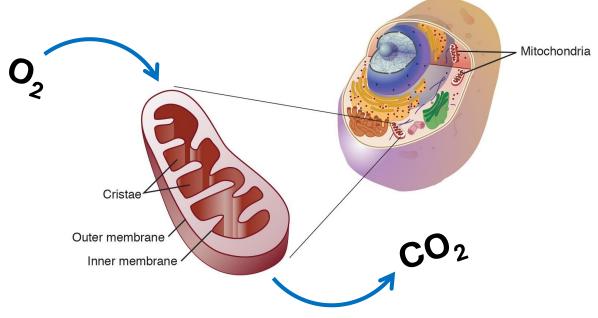




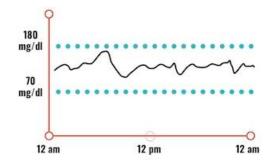
Breakdown of glucose

Occurs in <u>mitochondria</u>, uses oxygen and makes carbon dioxide





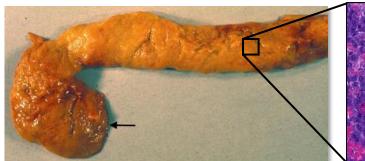
Energy is used to make <u>ATP</u>

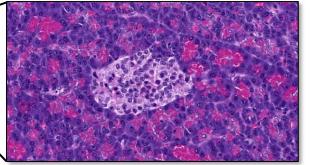


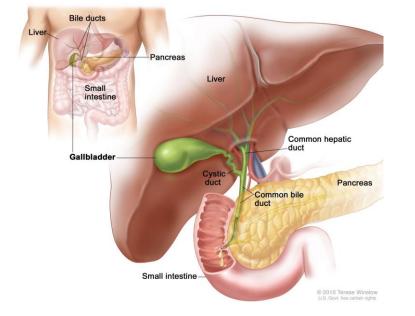
### Blood Glucose Regulation

Meet the pancreas!

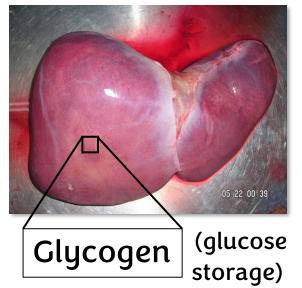
Pancreatic islets

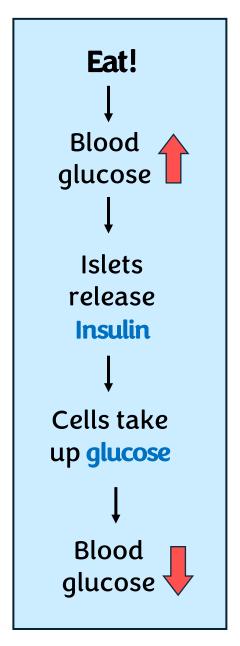


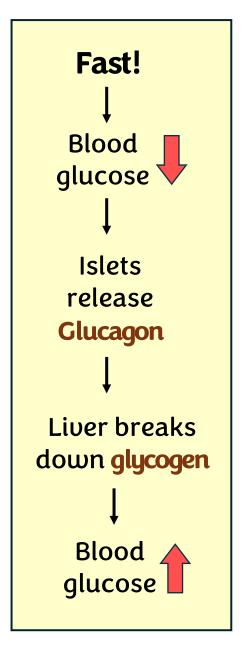




Meet the liver!







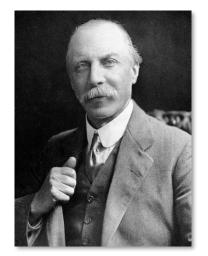




### Discovering the Problem

- 1889 Oskar Minkowski and Joseph von Mering
- Removed pancreas from dogs, causing fatal diabetes





- 1910 Edward Albert Sharpey-Schafer
- Hypothesized insulin, named from "insula" meaning island



- 1921 Frederick Banting and Charles Best
  - Discovered insulin!
  - Induced diabetes in dogs and cured it with purified pancreatic extract
  - Leonard Thompson, age 14, first person to receive insulin

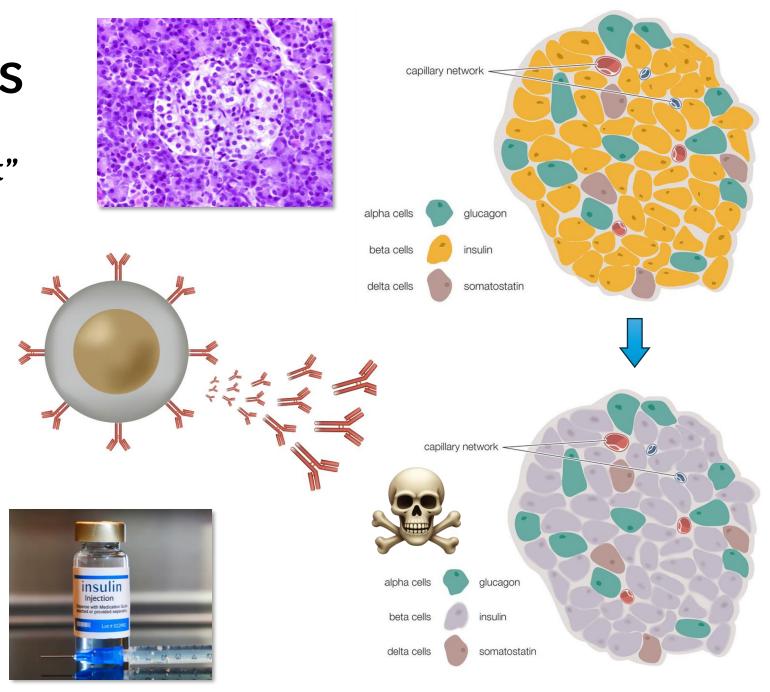


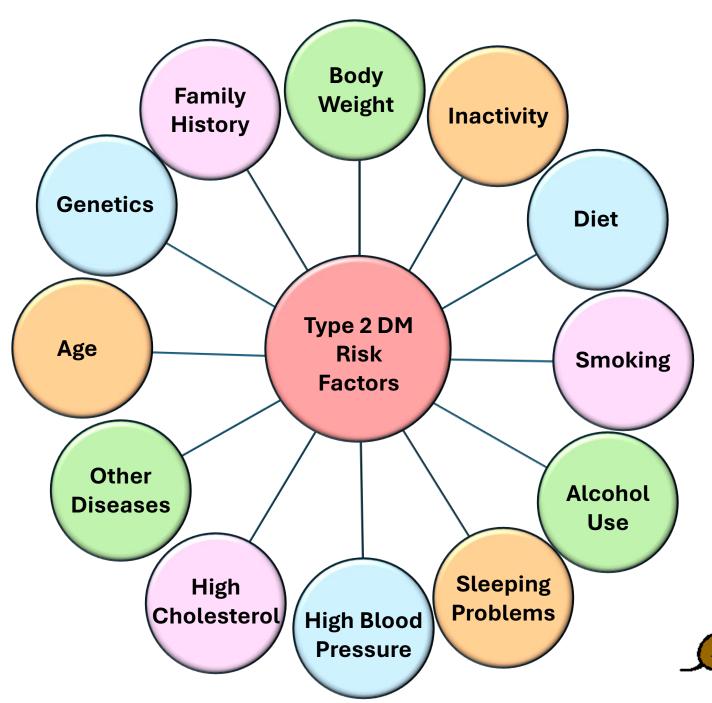
## Type I Diabetes

Formerly "childhood onset"



- Autoimmune destruction of pancreatic beta cells
- Insulin-dependent





# Type II Diabetes



Formerly "adult onset"

<u>Decreased insulin</u>
 <u>production</u>,
 pancreas "burnout"



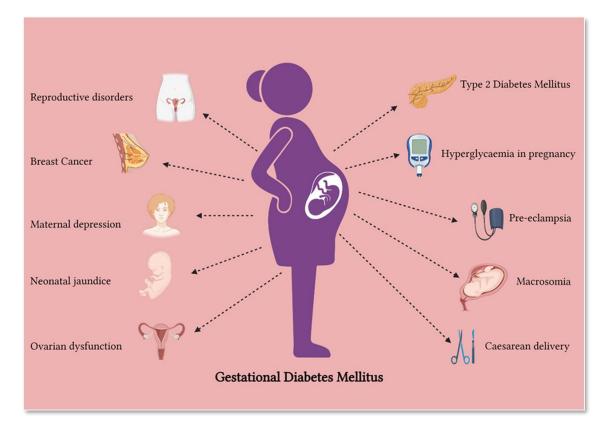
Insulin resistance, cells are not as responsive

#### **Gestational Diabetes**

- Diabetes appearing for the first time during pregnancy
- Cause unknown, excess weight prior to pregnancy increases risk

#### **Complications for baby**

- Higher birth weight
- Increased risk of preterm birth
- Higher risk of obesity and type II diabetes later in life



#### **Complications for mom**

- Hypertension and preeclampsia
- Higher likelihood of C-section
- Future diabetes more likely

## Symptoms

Increased urination

Fatigue and weakness

Thirst

Blurred vision

Hunger

 Numbness and tingling of hands and feet

Weight loss

Fruity odor to breath

Nausea

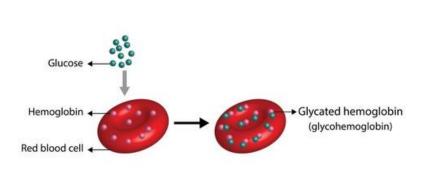
 Frequent infections, poorly healing wounds



# Diagnosis







**Tests** 



Normal

**Prediabetes** 

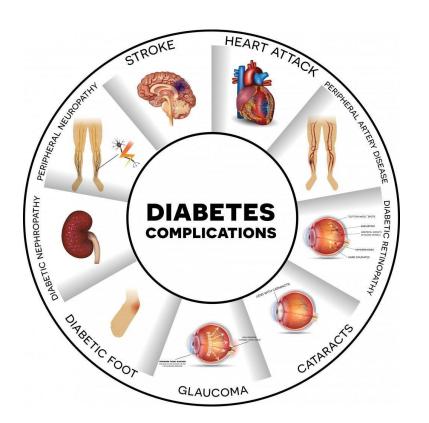
**Diabetes** 

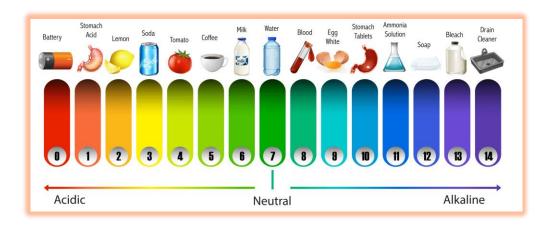
Fasting Glucose	Oral Glucose Tolerance Test	Hemoglobin A1C%
<100	<140 (2hrs)	5.6 or less
100-125	140-199	5.7-6.4
126+	200+	6.5+

### Diabetes Complications

#### **Acute problems**

- Ketoacidosis (diabetic crisis)
- Rapid fat breakdown creates acids





#### **Chronic problems**

- All long-term diabetic complications are due to BLOOD VESSEL damage
- The most highly vascular systems are most affected: brain, heart, nerves, eyes, kidneys, healing ability

#### Prevention is Key

Prediabetes is curable

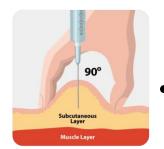


- Elimination of trans fats
- Listing calorie content in restaurants
- Public school cafeteria offerings
- Rethinking sugar-sweetened beverages
- Food label serving size realism, ingredient transparency

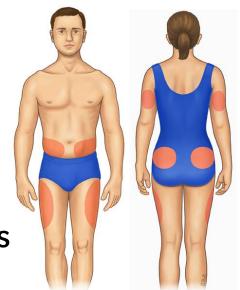
# Modern Therapy

#### **Treatments**

- Exercise and dietary changes
- · Glucose monitoring
- Insulin administration



Medications



- Preventative and multimodal treatment, stress avoidance
- Watch for hypoglycemia!



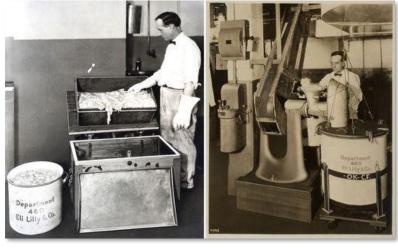
# The Story of Insulin

• Before insulin - fasting and diet modification

• 1920's - insulin purified from bovine and porcine pancreases

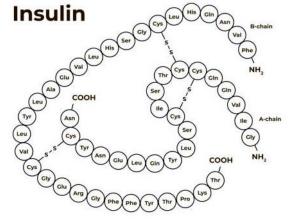
• 1950's - Frederick Sanger determines amino acid sequence

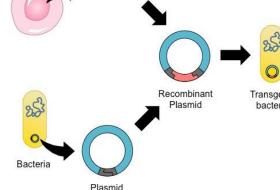












• 1980's - Biosynthetic human insulin using recombinant DNA technology

### Glucose Monitoring

• 1908 – Stanley Benedict develops copper reagent for urine glucose





• 1965 - First blood glucose test strip developed (Dextrostix)

 1980s – Glucose meters become available for home glucose monitoring,



 Steady improvement over decades – less blood, less pain, more accuracy

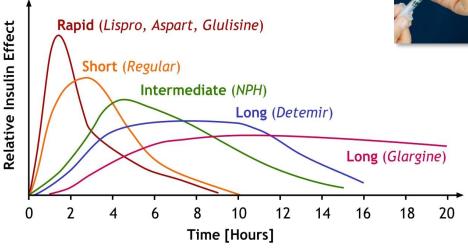


 2000's - Continuous glucose monitoring



#### Injected insulin

Several subtypes





#### **Oral medications**



 10 major classes of drugs with many mechanisms...

- Increase insulin secretion
- Decrease glucose release from liver
- Make cells more responsive to insulin
- Decrease glucose absorption in the gut
- Block the kidney from retaining sugar
- Alter carbohydrate metabolism in cells
- Promote weight loss And more!

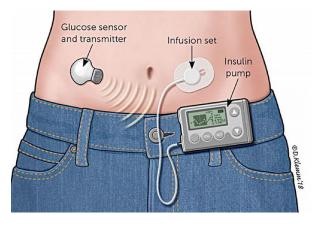




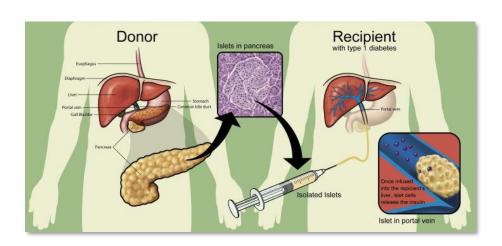
Pens, patches, pods and pumps



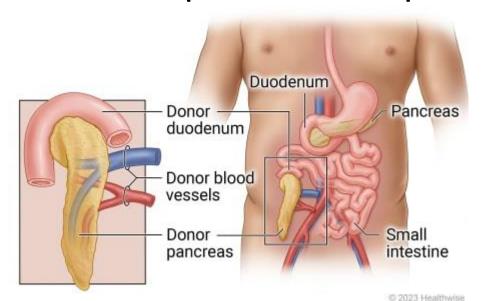
Artificial pancreas



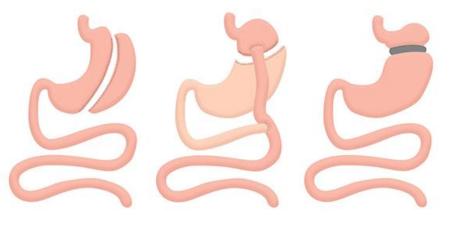
### Surgical Treatments



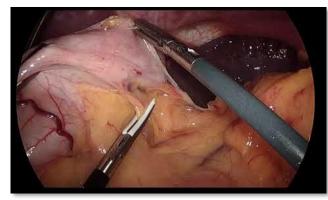
• Islet cell or pancreas <u>transplant</u>



- **Bariatric** surgery
  - **Gastric bypass**
  - Sleeve gastrectomy
  - Lap band
  - **Balloons**









**Gastric Sleeve** 

**Gastric Bypass** 

**Gastric Band** 

Balloons

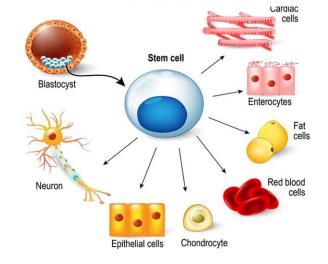


New medications

#### **Future Treatments**



Stem cell therapies





Transplant improvements

Immunotherapy



Artificial pancreas improvements













# Thank you!



Questions?