

#### **EPFL** References

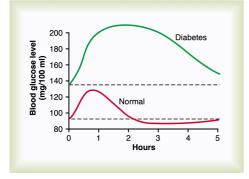
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# Introduction: insulin and it's metabolic effects

- Role of Insulin:
  - Hormone produced by the pancreas, critical for glucose regulation.
  - Promotes glucose uptake by cells for energy and storage as glycogen in the liver.
- Metabolic Effects:
  - High sugar level: secretion of insuline which lowers blood glucose by promoting storage in liver, muscle, and fat. Also inhibits glucose production (gluconeogenesis) in the liver.
  - Low sugar level: insuline secretion decrease; liver glycogen is split back into glucose
- Importance in Health:
  - Maintains blood glucose balance (hyperglycemia and hypoglycaemia)
  - Dysregulation linked to diabetes are a major threat to health

Physiology aspects



- Blood Glucose Regulation:
  - Maintained by the balance between insulin and glucagon.
  - Fluctuates based on food intake, physical activity, and hormonal changes.
- Normal Blood Glucose Levels
  - Fasting: 70-100 mg/dL.
  - Postprandial (after meals): 120-140 mg/dL.
- Impact of Dysregulation:
  - Hyperglycemia: Risk of long-term complications: loss of glucose in the urine, cell dehydration, tissue injury, increased utilisation of fats, metabolic acidosis and proteins
  - Hypoglycemia: Immediate risks like confusion, seizures, or loss of consciousness (brain, retina can only use glucose as nutrient).

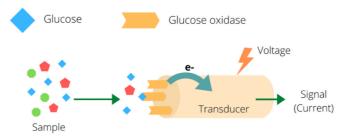
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### Technical description

- Components:
  - Biological Recognition Elements: Detect target molecules (e.g., enzymes, antibodies).
  - Transducer: Converts recognition events into measurable signals.
  - Signal Processing System: Converts signals into readable data.
- Transducer Types:
  - Electrochemical, optical, thermometric, piezoelectric, magnetic.
  - Electrochemical sensors dominate for glucose (sensitive, reproducible, low-cost).
- Electrochemical Sensors:
  - Types: Potentiometric, amperometric, conductometric.
  - Amperometric sensors: Most common, measure currents from electron exchange.
- Enzymes in Glucose Biosensors:
  - Common enzymes: Hexokinase, glucose oxidase (GOx), glucose-1-dehydrogenase (GDH).
  - GOx: High glucose selectivity, cost-effective, robust (handles pH, temperature, storage).

## EPFL Enzymatic Amperometric Sensors

- **Enzymatic**: Glucose Oxidase (GOx), generally present as a coat of the electrode, catalyses the oxidation of glucose in the blood producing hydrogen peroxide (H2O2). This reaction produces an electron transfer.
- Amperometric: a d.d.p. of 0.6-0.8V is present between anode and cathode, and the current flow is detected. The current is directly proportional to the concentration of the analyte that is oxidised at the electrode surface (H2O2).



### **EPFL** Main Categories





- Continuous Glucose Monitoring (CGM)
  - Real-time glucose data transmitted automatically
  - Consists of sensor, transmitter, and receiver/smartphone
  - Alarms for high/low glucose levels
  - Some integrate with insulin pumps for closed-loop systems
  - Subtype: flash CGM (requires manual scanning of sensor)
- Blood Glucose Meters (BGM)
  - Require multiple finger pricks to obtain blood samples
  - Limited ability to detect trends due to infrequent measurements
  - Require active user initiative

#### EPFL Dexcom G7









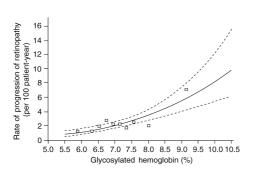
- Sensor life: 10 days
- Cost: \$378/month
- Real-time glucose readings through a smartphone app
- Standalone receiver (optional)
- FDA-approved for people with all types of diabetes ages 2 years and older
- Mean Absolute Relative Difference (MARD): 8.2%

### Abbott FreeStyle Libre 3



- Sensor life: 14 days
- Cost: \$132/month
- Real-time glucose readings through a smartphone app
- Standalone receiver (optional)
- FDA-approved for people with all types of diabetes ages 4 years and older
- Mean Absolute Relative Difference (MARD): 7.9%

### Medical Efficacy of CGM



- Result of Diabetes Control and Complications Trial show that improved glucose control, measured by a reduction in the fraction of glycosylated hemoglobin (HbA1c), leads to reduced long-term complications of diabetes (photo).
- Time in Range (TIR) improved from 18% with self-monitoring of blood glucose to 74% with CGM, exceeding the American Diabetes Association's recommended goal of 70% TIR.
- Modest but statistically significant declines in HbA1c among individuals with type 2 Diabetes.

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## Future development paths

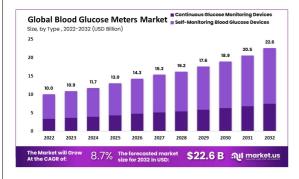
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- Non-invasive technologies (ex: Nemaura Medical's SugarBEAT and LifePlus' smartwatch) requiring no needle like optical or radio frequency sensors
- Implantable CGM Systems like the Swiss' Eversense solution
- Better algorithms (ML) for managing closed loop systems (insuline pumps)
- Improvements in CGM devices: longer duration between sensor swaps, better accuracy, smaller sensor footprint...

### EPFL Market considerations

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- The CGM market is projected to reach \$31.41 billion by 2031, growing at a CAGR (compound annual growth rate) of 15% from 2024 to 2031
- Market drivers
  - Increasing adoption of CGM devices in emerging markets
  - Technological advancements in glucose monitoring devices (like CGMs)
  - Rising diabetes prevalence globally
  - Rising awareness and detection

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