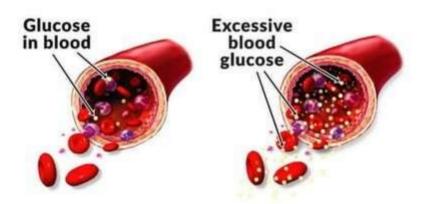


INTRODUCTION

 Diabetes mellitus (DM), is a group of metabolic diseases in which there are high blood sugar levels over a prolonged period.



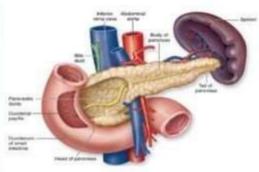
 Symptoms of high blood sugar include frequent urination, increased thirst, and increased hunger. o If left untreated, diabetes can cause many complications.



ketoacidosis, nonketotic hyperosmolar coma, or death.

 Serious long-term complications include <u>heart</u> <u>disease</u>, <u>stroke</u>, <u>chronic kidney failure</u>, <u>foot ulcers</u>, and <u>damage to the eyes</u>. Diabetes is due to either the <u>pancreas</u> not producing enough <u>insulin</u> or the cells of the body not responding properly to the insulin produced.

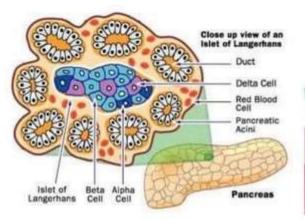




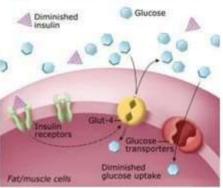
- There are three main types of diabetes mellitus:
 - □ Type 1 DM
 - ☐ Type 2 DM
 - Gestational Diabetes

o Type 1 DM

 Results from the pancreas's failure to produce enough insulin.



Type 1 Diabetes: Insufficient Insulin



 This form was previously referred to as "insulindependent diabetes mellitus" (IDDM) or "juvenile diabetes".

The cause is unknown.

o Type 2 DM

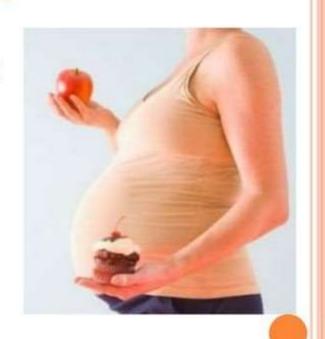
Begins with <u>insulin resistance</u>, a condition in which cells fail to respond to insulin properly.



- This form was previously referred to as "non insulindependent diabetes mellitus" (NIDDM) or "adult-onset diabetes".
- The primary cause is excessive body weight and not enough exercise.

o Gestational Diabetes

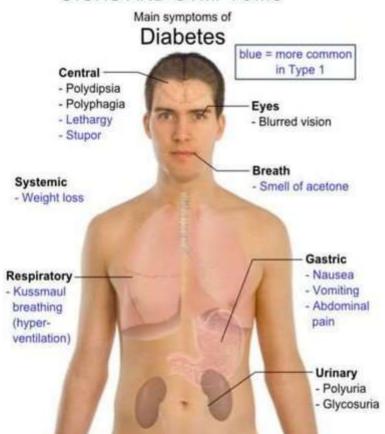
Is the third main form
and occurs in
pregnant women
without a previous
history of diabetes



COMPARISON OF TYPE 1 AND 2 DIABETES

Feature	Type 1 diabetes	Type 2 diabetes
Onset	Sudden	Gradual
Age at onset	Mostly in children	Mostly in adults
Body size	Thin or normal	Often obese
Ketoacidosis	Common	Rare
Autoantibodies	Usually present	Absent
Endogenous insulin	Low or absent	Normal, decreased or increased
Concordance in identical twins	50%	90%
Prevalence	~10%	~90%

SIGNS AND SYMPTOMS



SIGNS AND SYMPTOMS

- The classic symptoms of untreated diabetes are
 - weight loss
 - polyuria (increased urination)
 - polydipsia (increased thirst) and
 - polyphagia (increased hunger).



 Symptoms may develop rapidly (weeks or months) in type 1 DM, while they usually develop much more slowly and may be subtle or absent in type 2 DM.

SIGNS AND SYMPTOMS

In addition they also include:

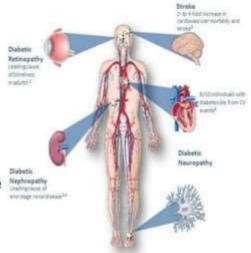
- Blurry vision
- Headache
- Fatigue
- Slow healing of cuts and
- Itchy skin.



- Prolonged high blood glucose can cause glucose absorption in the <u>lens of the eye</u>, which leads to changes in its shape, resulting in vision changes.
- A number of skin rashes that can occur in diabetes are collectively known as <u>diabetic dermadromes</u>

COMPLICATIONS

- All forms of diabetes increase the risk of long-term complications. These typically develop after many years (10–20)
 - The major long-term complications relate to damage to <u>blood vessels</u>.
 - Diabetes doubles the risk of cardiovascular disease
- About 75% of deaths in diabetics are due to coronary artery disease.
 - Other <u>"macrovascular"</u> diseases (stroke)
 - o peripheral vascular disease.

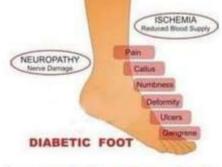


COMPLICATIONS

- The primary complications of diabetes due to damage in small blood vessels include damage to the eyes, kidneys, and nerves.
 - Damage to the eyes, known as <u>diabetic retinopathy</u>, is caused by damage to the blood vessels in the <u>retina</u> of the eye, and can result in gradual vision loss and <u>blindness</u>.
- Damage to the kidneys, known as <u>diabetic nephropathy</u>, can lead to tissue scarring, urine protein loss, and eventually <u>chronic kidney disease</u>, sometimes requiring <u>dialysis</u> or <u>kidney transplant</u>.
 - Damage to the nerves of the body, known as <u>diabetic</u> <u>neuropathy</u>, is the <u>most common</u> complication of diabetes.

COMPLICATIONS

The symptoms can include numbness, tingling, pain, and altered pain sensation, which can lead to damage to the skin.



o <u>Diabetes-related foot</u>
problems (such as <u>diabetic foot</u>
ulcers) may occur, and can be
difficult to treat, occasionally
requiring amputation.

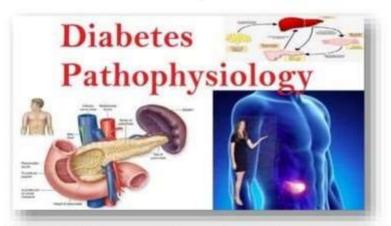


Additionally, <u>proximal diabetic</u>
 <u>neuropathy</u> causes painful <u>muscle</u>
 <u>wasting</u> and weakness – <u>Diabetic</u>
 <u>Amyotrophy</u>.



PATHOPHYSIOLOGY - GENERAL

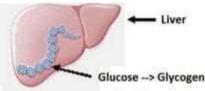
o <u>Insulin</u> is the principal hormone that regulates the uptake of <u>glucose</u> from the blood into cells of the body, especially liver, adipose tissue and muscle, except smooth muscle, in which insulin acts via the <u>IGF-1</u> (Insulin-like growth factor -1).



 Therefore, deficiency of insulin or the insensitivity of its <u>receptors</u> plays a central role in all forms of diabetes mellitus.

- The body obtains glucose from three main places:
 - The intestinal absorption of food
 - The breakdown of glycogen, the storage form of glucose found in the liver





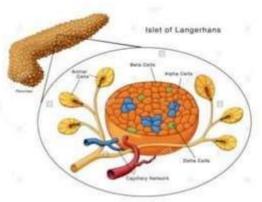
Gluconeogenesis, the generation of glucose from non-carbohydrate substrates in the body.



- Insulin plays a critical role in balancing glucose levels in the body:
 - It can inhibit the breakdown of glycogen or the process of gluconeogenesis.
 - It can stimulate the transport of glucose into fat and muscle cells.

It can stimulate the storage of glucose in the form of glycogen.

 Insulin is released into the blood by <u>beta cells</u> (β-cells), found in the <u>islets of</u>
 <u>Langerhans</u> in the pancreas, in response to rising levels of blood glucose, typically after eating.



Lower glucose levels result in decreased insulin release from the beta cells and results in the breakdown of glycogen to glucose.

This process is mainly controlled by the hormone glucagon, which acts in the opposite manner to insulin.

- If the amount of insulin available is insufficient
- If cells respond poorly to the effects of insulin
 - If the insulin itself is defective



 Then glucose will not be absorbed properly by the body cells



 The net effect is persistently high levels of blood glucose, poor protein synthesis, and break down of fat storage



 When the glucose concentration in the blood remains high over time, the <u>kidneys</u> will reach a threshold of <u>reabsorption</u> Glycosuria.



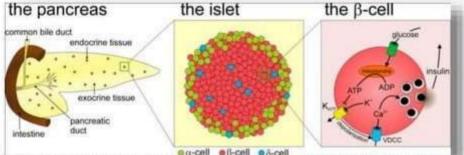
o This increases the osmotic pressure of the urine increased fluid loss



 Lost blood volume will be replaced osmotically from water held in body cells and other body compartments



 Type 1 diabetes mellitus is characterized by loss of the insulin-producing <u>beta cells</u> of the <u>islets of Langerhans</u> in the pancreas, leading to insulin deficiency.



- o This type can be rurmer crassined as immune-mediated or idiopathic.
 - The majority of type 1 diabetes is of the immunemediated nature, in which a <u>T-cell-</u> mediated <u>autoimmune</u> attack leads to the loss of beta cells and thus insulin.

- Most affected people are otherwise healthy and of a healthy weight when onset occurs.
- Sensitivity and responsiveness to insulin are usually normal, especially in the early stages.

oType 1 diabetes can affect children or adults, but was traditionally termed "juvenile diabetes" because a majority of these diabetes cases were in children.



 Type 1 diabetes is partly inherited, with multiple genes, including certain <u>HLA genotypes</u>, known to influence the risk of diabetes.



- In genetically susceptible people, the onset of diabetes can be triggered by one or more environmental factors, such as a viral infection or diet.
 - Among dietary factors, gluten may lead to type 1 diabetes, but the mechanism is not fully understood

- Type 2 DM is characterized by insulin resistance.
- The defective responsiveness of body tissues to insulin is believed to involve the <u>insulin receptor</u>.
 - o In the early stage of type 2, the predominant abnormality is reduced insulin sensitivity.
- Type 2 DM is due primarily to lifestyle factors and genetics.

- A number of lifestyle factors are known to be important to the development of type 2 DM, including
 - Obesity
 - lack of physical activity
 - poor diet
 - Stress
- Dietary factors also influence the risk of developing type 2
 DM such as
 - sugar-sweetened drinks
 - Type of <u>fats</u> in diet
 - a saturated fats and trans fatty acids increasing the risk
 - polyunsaturated and monounsaturated fat decreasing the risk
 - Eating lots of white rice also may increase the risk of diabetes.
 - A lack of exercise is believed to cause 7% of cases

- Hypoglycemia
 - Initial signs: mood changes, decreased spontaneity, hunger and weakness.
 - Followed by sweating, incoherence, tachycardia.
 - Results in unconsciousness, hypotension, hypothermia, seizures, coma, even death.

- 15 grams of fast-acting oral carbohydrate.
- Measured blood sugar.
- Loss of consciousness: 25-30ml 50% dextrose solution iv. over 3 min period.
- Glucagon 1mg.

- Severe hyperglycemia
 - A prolonged onset
 - Ketoacidosis may develop with nausea, vomiting, abdominal pain and acetone odor.
 - Difficult to different hypoglycemia or hyperglycemia.

 Hyperglycemia needs medical intervention and insulin administration.

- While emergency, give glucose first!
- Small amount is unlikely to cause significant harm.

DIAGNOSIS

- Can be diagnosed by demonstrating any one of the following:
 - □ Fasting plasma glucose level ≥ 7.0 mmol/l (126 mg/dl)
 - Plasma glucose ≥ 11.1 mmol/l (200 mg/dl) two hours after a 75 g oral glucose load as in a glucose tolerance test.
 - Symptoms of high blood sugar and casual plasma glucose ≥ 11.1 mmol/l (200 mg/dl)
 - Glycated hemoglobin (HbA_{1C}) ≥ 48 mmol/mol (≥ 6.5 DCCT %)

DIAGNOSIS

Oral Glucose Tolerance Test (OGTT)

- Measures the body's ability to metobolise glucose
- Most commonly done to check for <u>gestational</u> <u>diabetes</u>.
 - The patient is asked to take a glucose drink and their <u>blood glucose level</u> is measured before and at intervals after the sugary drink is taken.
 - For the standard glucose tolerance test, we should drink 75 grams or 100 grams.



MANAGEMENT

- Lifestyle
 - Good nutrition
 - Regular exercise
 - Diet control to maintain blood pressure.
- Medications
- Surgery
 - Pancreas transplant
 - kidney transplantation
 - Weight loss surgery