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# **Definition:**

- Diphtheria is an acute, toxin-mediated disease caused by toxigenic *Corynebacterium diphtheriae*
- It's a very <u>contagious and potentially life-</u> <u>threatening</u> bacterial disease.
- It's an infectious disease, which usually attacks the <u>throat and nose mucous</u>
   membrane.

### Diphtheria

- Greek diphtheria (leather)
- Recognized by Hippocrates in 5th century
- Epidemics described in 6th century
- C. diphtheriae described by Klebs in 1883
- Toxoid developed in 1920s



### **Etiology:**

<u>C. diphtheriae</u> is an aerobic gram-positive bacillus.

Pleomorphic, club-end
 Non-spore-forming
 Non-acid-fast
 Non-motile



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### Infectious agent:

 Toxin production (toxigenicity) occurs only when the bacillus is itself infected (lysogenized) by a specific virus (bacteriophage) carrying the genetic information for the toxin (toxogene).

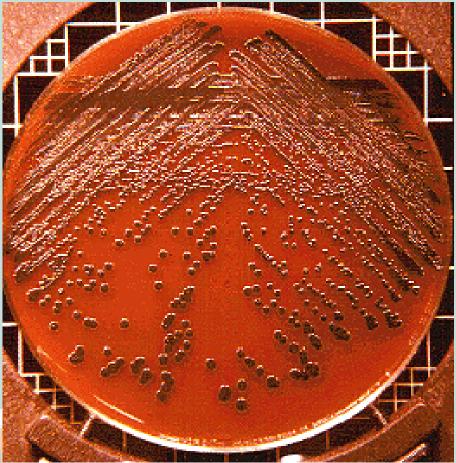


• C. diphtheriae has four biotypes—gravis, intermedius, mitis and belfanti.

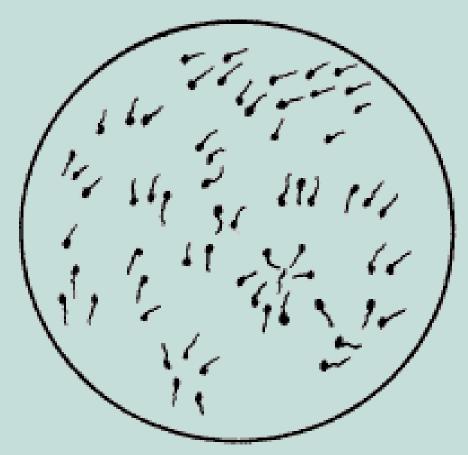
• All strains may produce toxin and can cause severe disease.

• All isolates of **C. diphtheriae** should be tested for toxigenicity.

#### **Gram +ve Bacilli and Colonies**



Corynebacterium diphtheriae, mitis Chocolate tellurite agar







# **Pathogenesis and pathology:**

- Susceptible persons may acquire toxigenic diphtheria bacilli in the <u>nasopharynx, middle ear</u> or anterior nares ,skin.
- The organism produces a toxin that inhibits cellular protein synthesis and is responsible for local tissue destruction and pseudomembrane formation.

## **Pathogenesis and pathology:**

- The pseudomembrane consists of coagulated fibrin, inflammatory cells, destructed mucous tissues and bacteria.
- The pseudomembrane in **larynx**, trachea or bronchi may have the potential for **airway obstruction**.
- The toxin is responsible for the major complications of myocarditis and neuritis, and can also cause low platelet counts (thrombocytopenia) and protein in the prifice (proteinuria).

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# EPIDEMIOLOGY



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#### **Occurrence:**

- Diphtheria occurs worldwide and is more prevalent in winter months in temperate zones.
- It was eradicated from most of the developed world by mass vaccination in the mid-20th century.
- However, it's still **common** in developing countries where immunizations aren't given.

- In 1993 and 1994, thousands cases were reported during a serious outbreak of diphtheria in countries of the former Soviet Union.
- The epidemic has **declined** after **a peak in 1995** ,it was responsible for over **140 000 cases** and over **4000 deaths**.
- Over **70%** of cases were <u>aged 15 years or older</u>.

### **Transmission:**

- Transmission is most often <u>person-to-</u>
   <u>person spread</u> from the respiratory
   tract (by small droplet when coughing or sneezing).
- Rarely, transmission may occur from skin lesions or articles soiled with discharges from lesions of infected
   persons.

#### **Source of infection:**

Patients and asymptomatic carriers

#### Reservoir:

Humans are the usual reservoir and carriers are usually asymptomatic.

#### **Incubation period:**

usually 2–5 days, occasionally longer (1–10 days).



### Period of communicability:

- Transmission may occur as long as <u>virulent</u> <u>bacilli are present in discharges</u> and lesions.
- The time is variable but is usually two weeks or less and seldom more than four weeks without antibiotics.
- Appropriate antibiotic therapy promptly terminates shedding. The rare chronic carrier may shed organisms for six months or more.

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#### **Susceptibility and resistance:**

- Infants born of immune mothers are relatively immune, but passive immunity is usually lost by six months of age.
- Lifelong immunity is usually, but not always, acquired after disease or inapparent infection.
- A primary course of toxoid vaccination provides long lasting but not lifelong immunity.
- Vaccinated individuals may become colonised by *C. diphtheriae* in the nasopharynx while still being
   protected from clinical disease.

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# **Risk factors**

- 1. Children under 5 and adults over 60 years old are particularly at risk.
- 2. People living in crowded or unclean conditions (low socio-economic status).
- 3. Undernutrition.
- 4. Children and adults who don't have up-todate <u>immunizations</u>.

# **Clinical Features:**

- Disease can involve almost any mucous membrane.
- For clinical purposes, it is convenient to classify diphtheria into a number of manifestations, depending on the anatomical site of disease.

### **Clinical features of diphtheria**

#### **Acute infection**

- Nasal infection
- Pharyngotonsillar infection
- Laryngeal infection
- Skin/wound/conjunctival infection (rare)



### **1.Anterior nasal diphtheria**

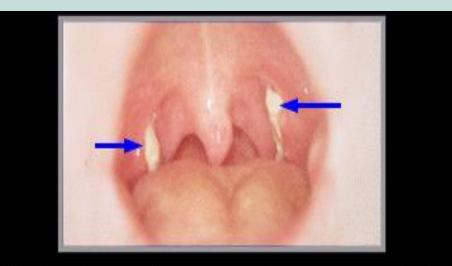
- The onset is indistinguishable from that of the <u>common cold</u> and is usually characterized by a mucopurulent nasal discharge (containing both mucus and pus) which may become blood-stained.
- A white pseudomembrane usually forms on the <u>nasal septum</u>.

#### 2. In pharyngotonsillar diphtheria

- Insidious onset of exudative pharyngitis
- Fever not high, but patient appears toxic
- Exudate spreads to form adherent "pseudomembrane"
- In early stages whitish membrane which can be wiped off easily over pharynx or tonsils.
- Later it becomes thick, white- blue to grey –black and adherent. It is difficult to remove if tried to
   remove it will result in bleeding.

#### Mucosal erythema around the membrane

# Edema of submandibular areaBull neck appearance



Diphtheria - notice the pseudomembrane in the posterior pharynx. It can become very large and may obstruct the airway.







#### Pharyngeal Diphtheria: Distant Complications

- Myocarditis
  - Cardiac arrhythmias in acute phase
  - Sudden death
- Neuropathy
  - Motor neuropathy
  - Paralysis of soft palate from 3rd week
    - Eye muslces, limbs, diaprhagm after 5th week.

# 3. Laryngeal diphtheria

- Laryngeal diphtheria can be either an extension of the pharyngeal form (often) or the only site involved (rarely).
- Symptoms include mild fever (with little absorption of toxin), dyspnea, hoarseness, and a barking cough.
- The pseudomembrane can lead to airway
   obstruction, coma, and death.



# Laryngeal diphtheria





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#### **4.Cutaneous and Other site diphtheria**

 <u>Skin infections</u> are quite common in the tropics and are probably responsible for the high levels of natural immunity found in these populations.

 Skin infections may be manifested by a scaling rash or by ulcers with <u>clearly demarcated edges</u> and pseudomembrane.

### **Cutaneous (skin) diphtheria**





10 y/o boy with severe diphtheria
conjunctivitis
pharyngeal membrane
bull neck
severe myocarditis
all vaccines contraindicated





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# **Diphtheria Complications**

Most attributable to toxin, severity generally related to extent of local disease

- Most frequent complications are myocarditis About 25% of survivors of the early toxaemia may later develop myocarditis with arrhythmias or cardiac failure.
- Neurological involvement occurs in 75% of cases.

 Death occurs in 5-10% .Death from acute circulatory failure may occur within the first 10 days.

# **Differential diagnosis**

- Membranous Tonsillitis
- Infectious mononucleosis
- Leukemia
- Aphthous ulcers
- Traumatic ulcer
- Foreign body (Nasal Diphtheria)



# **Diagnostic tests:**

- Specimens for culture should be obtained from the nose or throat or any mucosal or cutaneous lesion.
- Material should be obtained from beneath the membrane, or a portion of the membrane itself should be submitted for culture.



#### **Treatment:**

- Children and adults with diphtheria are treated in a hospital. After a confirmation of the diagnosis through a <u>throat culture</u>.
- The infected person receives a special antitoxin, given to neutralize the diphtheria toxin already circulating in the body.
- as well as antibiotics to kill the remaining diphtheria bacteria.

- Immediate hospitalization and strict isolation with early intervention allow most patients to recover from diphtheria.
- After the antibiotics and anti-toxin have taken effect, patient with diphtheria will need bed rest for a while (4 to 6 weeks, or until full recovery).
- Bed rest is particularly important if the person's heart has been affected by the disease. (Myocarditis)



## Treatment

- Antibiotic **not useful** in Acute infections.
- Antitoxin is a must.
- Antitoxin obtained from horse serum
   Mild 20,000 to 40,000 units
   Moderate 40,000 to 60,000 units
   Severe 80,000 to 1,00,000 units
- Commonly used antibiotics
  - **Penicillin** parentrally, Oral **Erythromycin**

- Those who have recovered should still receive a full course of the diphtheria vaccine to prevent a recurrence.
- because the occurrence of disease doesn't guarantee lifetime immunity.

#### **Prevention:**

- Preventing diphtheria depends almost completely on:
- immunizing children with the <u>diphtheria/pertussis/tetanus</u> (DPT) vaccine.
   non-immunized adults with the <u>diphtheria/tetanus vaccine (DT).</u>
- Most cases of diphtheria occur in people who haven't received the vaccine at all or haven't received the entire course.

### The immunization schedule:

- DPT vaccines at 2, 4, and 6 months of age.
- 1st booster dose given at 12 to 18 months.
- 2<sup>nd</sup> booster dose given again at 4 to 6 years (pre-school).
- booster shots given every 10 years after that to maintain protection.

#### **Diphtheria Toxoid**

- Formalin-inactivated diphtheria toxin
- Schedule Three doses + 2booster Booster shots every 10years
- Efficacy Approximately 95%
- Duration Approximately 10 years
- administered with tetanus toxoid .

#### Diphtheria and Tetanus Toxoids Adverse Reactions:

- Local reactions (erythema, induration)
- Exaggerated local reactions.
- Fever and systemic symptoms not common.
- Severe systemic reactions are rare.



# Diphtheria and Tetanus Toxoids contraindications and Precautions:

• Severe allergic reaction to vaccine component following a prior dose.

• Moderate or severe acute illness.



## **CONTROL MEASURES:**

- \***Prompt notification** to the local public health officials.
- \*Identification of close contacts of a person suspected to have diphtheria .



\*For close contact, regardless of their immunization status, the following measures should be taken :

1- Surveillance for 7 days for evidence of disease.

2-Culture for C. diphtheria.

3- Antimicrobial prophylaxes with oral erythromycin.

## **Treating Contacts**

 All contacts are advised to receive 500 mg
 Erythromycin 4 times a day.



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#### Follow up:

- <u>Pharyngeal culture</u> should be obtained from contacts proven to be carriers at a minimum of 2 weeks after completion of therapy.
- If cultures are positive , an <u>additional</u> 10- day therapy of erythromycin should be given .

## Outbreak measures:

- Outbreaks of diphtheria require immunising the largest possible proportion of the population involved.
- Emphasising the need for protection of infants and preschool children.
- In outbreaks amongst adults immunize groups that are at high risk.

- Repeat immunisations may be recommended after one month.
- Outbreak investigations involve enhanced case surveillance with laboratory confirmation of all suspected cases.
- Identification and appropriate management of close contacts and asymptomatic carriers.



## DIPHTHERIA is deadly-

#### **IMMUNISATION** is the ASK AT YOUR safeguard WELENRY CENTRE

