Viral Infections of the Respiratory System

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Objectives

- Introduction to respiratory viral infections
- Characteristics of respiratory viruses (*Orthomyxoviridae*, *Paramyxoviridae*)
- Mode of transmission
- Clinical features
- Lab diagnosis
- Treatment & prevention

Respiratory Tract Infections

- Are the commonest of human infections and cause a large amount of morbidity and loss of time at work (sick leave).
- Are common in both children and adults.
- Mostly caused by viruses.
- Mostly are mild and confined to the upper respiratory tract (URT).
- Mostly are self-limiting disease.
- ➤ URT-infection may spread to other organs causing more severe infection and death.

Clinical manifestations

- Common cold (rhinitis).
- Pharyngitis.
- Tonsilitis.
- Sinusitis & otitis media.
- Croup (acute laryngotracheobronchitis).
- Acute bronchitis.
- Acute bronchiolitis.
- Viral pneumonia.

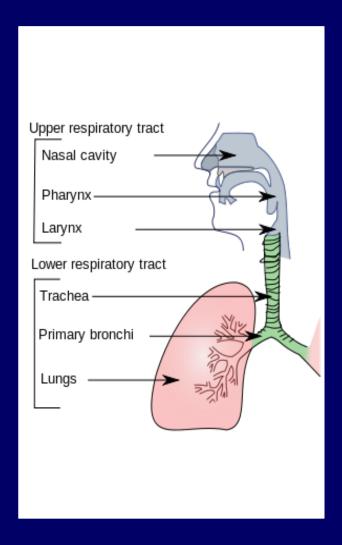


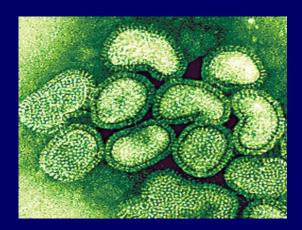
Table 22.1 Respiratory illnesses and their common viral causes*

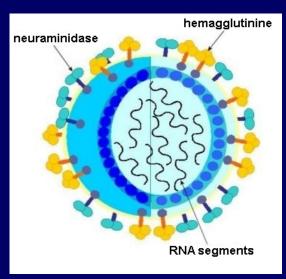
Respiratory illness	Main causal viruses
Rhinitis (common cold)	Rhinoviruses (100 serotypes) Coronaviruses (3 serotypes)
Pharyngitis	Influenza A and B viruses Parainfluenza virus (types 1–3) Adenoviruses B, C, E Coxsackie A virus
Laryngotracheobronchitis (Croup)	Influenza virus Parainfluenza virus (types 1 & 2)
Bronchitis	Respiratory syncytial virus Parainfluenza virus (usually type 3) Influenza virus
Bronchiolitis	Respiratory syncytial virus Human metapneumovirus Parainfluenza virus (usually type 3)
Pneumonia	Influenza virus Respiratory syncytial virus Human metapneumovirus Parainfluenza virus (usually type 3) Adenoviruses B, C, E

^{*}Other viruses can cause respiratory illness as part of a systemic infection, e.g. measles, pneumonia and pharyngitis in primary Epstein-Barr virus infection.

Influenza Virus

- > Family: Orthomyxoviridae.
- > Structural features: Enveloped virus with 2 projecting glycoprotein spikes:
 - Haemagglutinin (H)
 - Neuraminidase (N)
- ➤ Genome: 8 Segmented –ve polarity ssRNA.
- This virus is highly susceptible to mutations and rearrangements within the infected host.





Influenza viral proteins

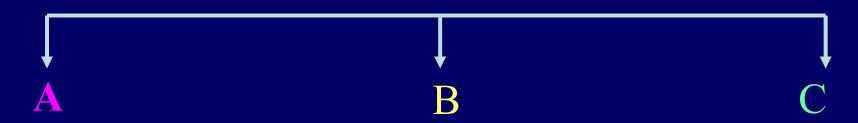
- Haemagglutinin (H):

- **Attachment to the cell surface receptors.**
- ❖ Antibodies to the HA is responsible for immunity.
- ❖ 16 haemagglutinin antigenic type, H1 H16...

- Neuraminidase (N):

- * Responsible for release of the progeny viral particles from the infected cell.
- 9 neuraminidase antigenic type, N1 N9.

Types of influenza virus

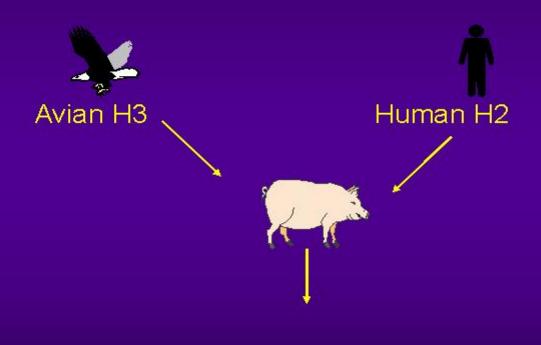


- Infects human and Animal
- Causes epidemic & pandemic
- Causes epizootic in animal
- Antigenic drift \rightarrow minor change
- Antigenic shift major change
- •Reassortment ,rearrangement

- Infects human only
- Causes outbreak
- Antigenic drift only

- Infects human only
- Causes mild illness

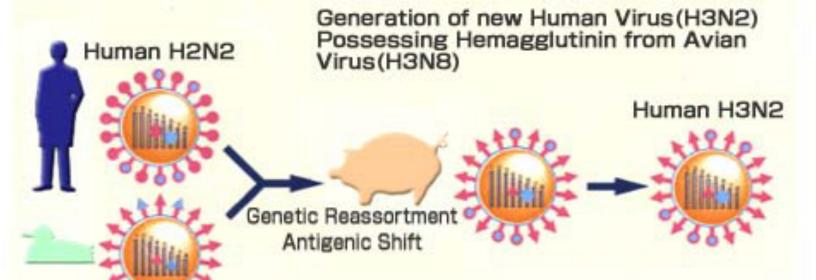
Reassortment



Human H3

Variation of Influenza Viruses

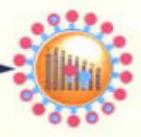
Glyco Word



Avian H3N8



Point Mutation of Hemagglutinin and Neuraminidase gene Antigenic Drift



Influenza Virus

- Divided into subtypes based on the haemagglutinine and neuraminidase proteins.
- ❖ The currently circulating strains are: H1N1 & H3N2.
- Pathogenesis: The virus infects the epithelial cells of the nose, throat, bronchi and occasionally the lungs.
- Transmission: Inhalation of infectious aerosol droplets.
- > I.P.: 1-4 days.
- Symptoms: Fever, malaise, headache, cough, chills, sore throat, and generalized pain.
- **Prognosis:** Usually self-limiting disease.

Continued...

- Complications: Primary influenza pneumonia
 2nd bacterial pneumonia
 Reye's syndrome [fatty degeneration of CNS and Liver (Aspirin)]
- Lab diagnosis: routine testing by Direct detection of Influenza A or B virus from sputum, nasopharyngeal swab, aspirate (NPA) or respiratory secretion by direct immunoflourecent assay (IFA).
- > Other detection methods: Cell culture, PCR.
- > Treatment:
- 1: Amantadine is effective against influenza A virus only.
- 2: Rimantadine, **Oseltamivir (Tamiflu)** or Zanamivir (Relenza) are effective against both influenza A & B viruses and can be used as treatment and prophylaxis.

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- > Prevention:
- ➤ Influenza vaccine: Two types of vaccines available:
- 1- The flu shot vaccine: Inactivated (killed vaccine).
- ❖ Given to people older than 6-months, including healthy people and those with chronic medical conditions.
- 2- The nasal spray flue vaccine (Flu mist): Live attenuated vaccine.
- ❖ Approved for use in healthy people between 5-49 years of age.
- ➤ Both vaccines contain two strains of the current circulating influenza A virus and the current circulating strain of influenza B virus.
- ➤ Vaccine should be given in October or November, before the influenza season begins.

Avian flu



Avian flu

- Viral etiology: Avian influenza type A virus (H5N1).
- Family: Typical orthomyxovirus.
- Epidemiology: Wild birds are the natural reservoir for the virus. They shed the virus in saliva, nasal secretion and feces.
- All domestic poultry are susceptible to infection.
- They become infected, when they eat food contaminated with secretion or excretion from infected bird.
- Avian influenza viruses do not usually infect human.
- High risk group includes those who working in poultry farms and those who are in close contact with poultry.

Continued...

- Symptoms in human:
- ☐ Ranges from typical flu to severe acute respiratory disease.
- ☐ Diarrhea, abdominal pain and bleeding from the nose have been reported.
- Treatment:
- ☐ Should be initiated within 48 hours.
- ☐ Oseltamivir and Zanamivir are used.
- Lab diagnosis: PCR, detection of the viral RNA in throat swap.

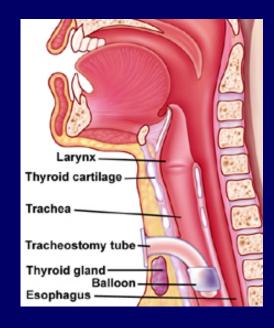
Swine flu



Parainfluenza Virus

- Family: Paramyxoviridae.
- Structural features: Enveloped virus with -ve polarity ssRNA genome, with 5 serotypes.
- Transmission: Inhalation of infectious aerosol droplets mainly in winter.
- Clinical syndrome:
 - a. Croup or acute laryngotracheobronchitis. PIV Type-I, II affect mainly in infants and young children. Fever, harsh cough, difficult inspiration can lead to airway obstruction which may require hospitalization and tracheostomy.
 - **b.** Bronchiolitis and Pneumonia: PIV Type-III in young children.





Continued...

- Lab diagnosis: routine testing by Direct detection of the virus from sputum, nasopharyngeal swab, aspirate (NPA) or respiratory secretion by direct immunoflourecent assay (IFA).
- > Other detection methods: tissue culture, PCR.
- Treatment and prevention: Supportive treatment, No specific treatment or vaccine available.

Respiratory Syncytial Virus (RSV) & Human metapneumovirus

- Family: Paramyxoviridae.
- > Structural features: Enveloped virus with –ve polarity ssRNA genome.
- Transmission: Inhalation of infectious aerosols mainly in winter.
- Clinical syndromes:
- **a.** Bronchiolitis: Life-threatening disease in infant especially under 6 month of life with respiratory distress and cyanosis can be fatal and can lead to chronic lung disease in later life.
- b. Pneumonia: can also be fatal in infant.

Continued...

- Lab diagnosis: routine testing by Direct detection of the virus from sputum, nasopharyngeal swab, aspirate (NPA) or respiratory secretion by direct immunoflourecent assay (IFA).
- ➤ Other detection methods: Isolated of virus by cell culture from N.P.A with multinucleated giant cell or syncytia as cytopathic effect (C.P.E); PCR.
- > Treatment and prevention: Ribavirin administered by inhalation for infants with severe condition.
- Vaccine: No vaccine available, but passive immunization immunoglobulin can be given for infected premature infants.

Measles Virus

- > Family: Paramyxovidae.
- > Structural features: Enveloped virus with –ve polarity ss-RNA genome.
- > Transmission: Inhalation of infectious aerosol droplets.
- Epidemiology: Measles virus infects human only.

 Most cases in preschool children, very infectious, infection occurs mainly in winter and spring.

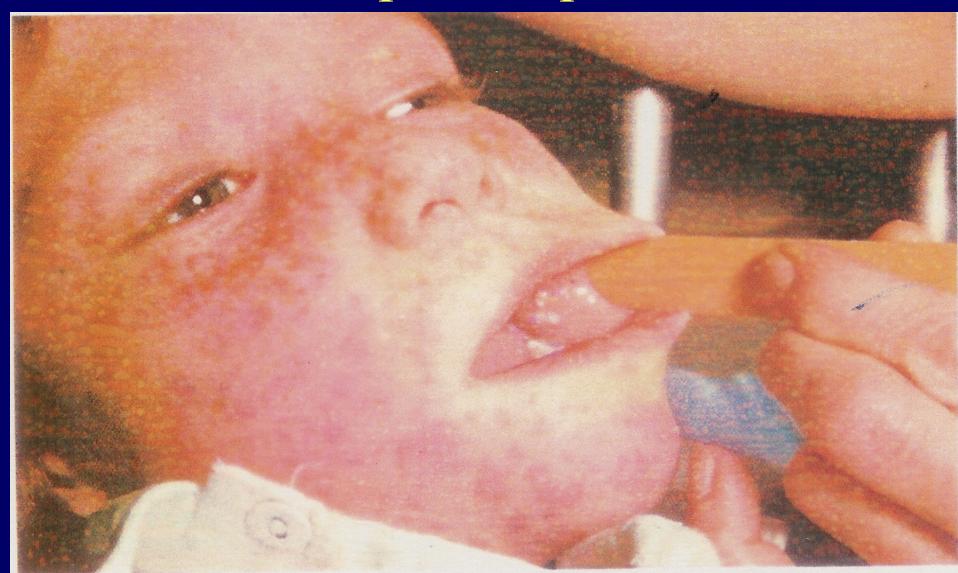
> Pathogenesis:

The virus infects first epithetical cells of upper respiratory tract, then the virus spread to **the blood** causing viremia and infect the endothelial cells of blood vessels. The virus reaches **the lymphoid tissue** where it replicates further and disseminates to the skin causing **maculopapular rash**.

> Clinical features:

- I.P.: 7- 14 days.
- Prodromal symptom: Fever, cough, conjunctivitis and running nose.
- **Koplik's spot:** Small red papules with white central dots appear mostly in buccal mucosa.
- Rash: Maculopapular rash first on face, trunk, extremities.
- The rash is red, become confluent, last 4 or 5 days, then disappears leaving brownish discoloration of the skin and final desquamation.
- Recovery complete in normal children with life long immunity.
- Complication also can occur.

Koplik's Spot



Measles



Complication:

- 1- Encephalitis: Acute or subacute sclerosing panencephalitis (SSPE)
- 2- Giant cell pneumonia: rare in immunocompromised children due to direct invasion of measles virus to the lung tissue.
- Lab diagnosis: Serology by detection of IgM Ab using ELISA, and in case of SSPE detection of measles antibodies in CSF or detection of viral NA using PCR.
- Treatment and prevention: No specific treatment, Prevention by giving the live attenuated vaccine (MMR) for Measles, Mumps and Rubella (given to all children 15 month and booster dose at school entry). Give excellent long last protection.

Mumps Virus

Mumps: is an acute benign viral parotitis.

Parotitis (painful inflammation and swelling of salivary gland and mainly parotid glands), it is a disease of children (5-15 years), but also can be seen in young adult with more complicated feature.

Virology Aspect:

- Family: Paramyxoviridae.
- > Structural features: Enveloped virus with polarity ss-ve RNA genome.
- ➤ The viral envelope is covered with two glycoprotein spikes, hemagglutinine and neuraminidase.
- > Transmission: Inhalation of infectious aerosol droplets during sneezing and coughing, direct contact with saliva.

> Epidemiology: Mumps virus infects human only.

- Highly infectious, peak in winter.
- Long incubation period 18-21 days.

Pathogenesis:

Infection started in the epithelial cells of upper respiratory tract, then virus spread by viremia to parotid gland mainly and to other organs as: testes, ovaries, pancreas and CNS.

Clinical Features:

- Classic mumps starts with moderate fever, malaise, pain on chewing or swallowing, particularly acidic liquids.
- Sudden onset of fever and painful swelling of parotid gland.
- Self-limiting disease resolve within one week.
- Solid and long life immunity developed.

- > Complications:
- Aseptic meningitis, Encephalitis, Pancreatitis, Thyroiditis.
- > after puberty:
- Orchitis: inflammation of one or both testicles. usually unilateral, rarely leads to sterility.
- Oophoritis: inflammation of ovaries.
- > Lab diagnosis:
- Serology by detection of **IgMAb using ELISA**, cell culture and isolation of the virus from saliva or detection of viral NA using PCR.
- > Treatment and prevention:
- No specific antiviral treatment.
- MMR: Live attenuated vaccine for Measles, Mumps and Rubella given to all children 15 month and booster dose at school entry. Give excellent long last protection.





Reference books

&the relevant page numbers

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By; Katherine N. Ward, A. Christine McCartney, and Bishan Thakker.

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Pages; 329-340.

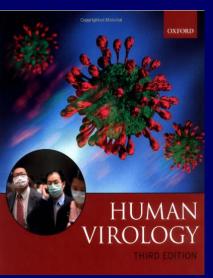
<u>Human Virology</u>

By; Leslie Collier and John Oxford.

(2006)

Pages, 71-95.





Thank you for your attention!

Questions?