Bacterial Respiratory Tract Infections

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Respiratory module / 3rd year

• Contents:

-Streptococcus pneumoniae (pneumococcus) -Haemophilus influenzae (blood liking bacteria)

45 years old man, smoker. Sudden onset fever and chills Shortness of breath and pleuritic chest pain Productive rusty coloured sputum (blood stained)

Examination:
Vitals: RR 24 (normal 12-16), PR 110 (normal 60-100), T 39, B.P normal, O2 Saturation 90% (decreased)
Chest: decreased air entry, dull on percussion, decreased chest expansion
CXR: Right upper lobe consolidation
WBC 16000/mm3 (normal 4-11) mainly neutrophils

member of the oropharyngeal flora of 5-70% of the < population, with the highest isolation rate in children during .the winter months

:A gram positive diplococci catalase negative





:An important pathogen <

It primarily causes disease of the upper and lower * respiratory tract

May spread to other sites, such as the joints, peritoneum, * .endocardium, biliary tract and, in particular, the meninges

Virulence factors

1. Capsule

- The capsular polysaccharide is a crucial virulence factor.
- Antiphagocytic and anticomplement.
- A total of more than 90 different capsular serotypes have been identified.
- About 90% of cases of bacteraemic pneumococcal pneumonia and meningitis are caused by some 23 serotypes.

- 2. Lipotechoic acid and coline binding proteins:Adhesion molecule
- 3. IgA1 protease
- Pneumococci produce an extracellular protease that specifically cleaves human IgA1 in the hinge region.
- This protease enables these pathogens to evade the protective functions of the principal immunoglobulin isotype of the upper respiratory tract.

- 4. Pneumolysin (pore forming toxin)
- Pneumococci produce an intracellular membrane-damaging toxin known as pneumolysin, which is released by autolysis.
- Pneumolysin also suppresses organism-targeted immunity (Neutrophils, lymphocyte proliferation and immunoglobulin synthesis).
- Pneumolysin is immunogenic and might be suitable for a new pneumococcal vaccine.

5. Autolysin

- When this enzyme activated, the pneumococcal autolysin breaks the peptide cross-linking of the cell wall peptidoglycan, leading to lysis of the bacteria.
- Autolysis:
- Enables the release of pneumolysin
- Release of peptidoglycans from cell wall leading to massive inflammatory response and sepsis to these peptidoglycan fragments.



- EPIDEMIOLOGY
- Source:
- Humans are the reservoir of pneumococci, which are commonly found in the upper respiratory tract of healthy persons throughout the world.

Occurrence:

 Pneumococcal infections are among the leading causes worldwide of illness and death for young children, persons who have underlying debilitating medical conditions and the elderly.

- 1 million deaths yearly worldwide
- 6 million cases of otitis media in USA
- The estimated global annual incidence is 1-3 per 1000 of the population, with a > 5% case fatality rate.

Mode of Transmission:

 Pneumococci are transmitted from person-to-person by droplet spread, by direct oral contact and indirectly through articles freshly soiled with respiratory discharges.

- Period of Communicability:
- Communicability associated with respiratory infection likely persists while pneumococci are present in respiratory secretions.
- Healthy persons is the major source of transmission
- Treatment with an antibiotic to which the infecting organism is sensitive can be expected to terminate communicability within 24 hours.
- Incubation Period:
- The incubation period varies by type of infection and can be as short as 1-3 days.

Streptococcus pneumonia Clinical features:

- Predisposing factors:
- Most Str. pneumoniae infections are associated with various predisposing conditions.
- Pneumonia results from aspiration of pneumococci contained in upper airway secretions into the lower respiratory tract; for example:
 - Loss of consciousness: general anaesthesia, convulsions, alcoholism, epilepsy or head trauma
- respiratory viral infections, such as influenza, chronic bronchitis.

- Young and elderly people
- Immune suppressed people (e.g Chronic diseases, drugs, asplenia)
- Structural respiratory abnormalities.

Streptococcus pneumonia / Respiratory Presentation

1- Pneumonia (chest infection):

- Pneumonia is defined as an acute respiratory illness associated with recently developed radiological pulmonary shadowing which may be segmental, lobar or multilobar.
- *Str. pneumoniae* is a frequent cause of pneumonia where vaccination is not available.

Pneumonia

- Contiguous spread commonly results in complications such as:
- Inflammatory involvement of the pleura, Empyema and Pericarditis.
- Bacteraemia may complicate pneumococcal pneumonia in up to 15% of patients. This can result in metastatic involvement of the meninges, joints and, rarely, the endocardium.

Pneumonia

- Signs and symptoms:
- The patient rapidly becomes more ill with a high temperature (up to 39.5°C), pleuritic pain and a dry cough.
- A day or two later, rusty-coloured sputum is produced
- The patient breathes rapidly and shallowly, the affected side of the chest moves less, and signs of consolidation may be present together with a pleural rub.
- The mortality rate from pneumococcal pneumonia in those admitted to hospital is approximately 15-25%.

- 2- Otitis media and sinusitis
- Middle ear infections (otitis media) affect approximately half of all children between the ages of 6 months and 3 years
- Approximately one-third of cases are caused by *Str. pneumoniae*. Disease occurs after acquisition of a new strain to which there is no pre-existing immunity.
- The prevalence is highest among children attending primary school, where there is a constant exchange of pneumococcal strains.
- Pain, fever, ear discharge...

Chest X-ray confirms the area of consolidation (lobar), but * ;radiological changes lag behind the clinical course

So that X-ray changes may be minimal at the start of the illness. Conversely, consolidation may remain on the chest .X-ray for several weeks after the patient is clinically cured

The chest X-ray usually returns to normal by 6 weeks *



Streptococcus pneumonia LABORATORY DIAGNOSIS

Collection of specimens

- Throat swab, pus, blood cultures, expectorates, otitis media discharge or urine according to the site of infection.
- Blood cultures are of value in patients with invasive streptococcal infections. This is also the case in patients with suspected pneumococcal pneumonia, particularly when this is severe, as up to 15% of patients are bacteraemic.

- Cultivation and identification :
 - Blood or chocolate agar , 37°C, 5%CO2
 - Colonies of pneumococci are α-haemolytic and smooth dome shaped.
 - During prolonged incubation, autolysis of bacteria within the flat pneumococcal colonies results in a typical subsidence of the centre ('*draughtsman colonies*').
 - Gram positive cocci in pairs, alpha hemolytic, Catalase negative.
 - Optochin sensitive and bile soluble,



Alpha hemolysis



Optochin sensitive



Gram positive diplococci



Capsule:Typing :



- Pneumococci are typed on the basis of the differences in capsular polysaccharides, of which 90 have been described (in reference lab.
- Mixing a suspension of pneumococci with type-specific antisera increases the visibility of the capsule in the microscope, and is the basis of the *quellung reaction* or *capsular swelling test*.

□ TREATMENT

Follow the antibiotic guidelines

□ Vaccines:

- Protein Conjugated vaccine (PCV): protection for 7-13 ty
- 3 doses for those < 2 years age
- Non-conjugated polysaccharides : 23 polyvalent vaccine > 2 years who are at risk

- Carriage rate in respiratory tract:
 -Capsulated (types A-F):
 10% (50% are type B)
 -Non caps: 80%
- Gram negative small bacilli (old culture) or coccobacilli (young culture)
- Fastidious, Non motile non spore forming
- Catalase and Oxidase positive
- Facultative anaerobic (5% CO2)





Virulence factors:

- 1-Polysaccharides capsule (in 10% of strains): inhibits phagocytosis and complement activation
- six capsular types, designated a-f, which can be identified by a polymerase chain reaction (PCR) method
- The most important is type b, has a polymer of ribosyl ribitol phosphate capsule.

- 2- Fimbriae: which assist attachment to epithelial cells
- 3- Immunoglobulin (Ig) A proteases, which are also involved in colonization
- 4- Outer membrane proteins and lipopolysaccharide, which may contribute to invasion at several stages

- Growth requirements:
- Growth depends on a requirement for two factors, termed X and V (found on chocolate agar):
- X factor (haemin) is required for the synthesis of cytochrome c and other iron-containing respiratory enzymes.
 - Unlike most bacteria, haemin-dependent haemophili cannot synthesize protoporphyrin
- ✓ V factor is nicotinamide adenine dinucleotide (NAD),
 - It is essential for oxidation-reduction processes in cell metabolism.

:Clinically

- Host and organism factors?
- Respiratory route direct and indirect
- Common age: 5m-5years (capsulated) / elderly (mainly non-capsulated)
- □ invasive infections and non-invasive infections
- 1- Invasive: (encapsulated types)

Meningitis (50-60%), epiglottitis, pneumonia and septicarthritis, acute exacerbation of COPD Commonly caused by encapsulated types mainly serotype b-

2- Noninvasive (Non encapsulated types)

 localized disease of the respiratory tract including bronchitis and otitis media

-Can also sometimes cause invasive diseases mainly pneumonia and bacteremia

-More in elderly

-Is becoming commoner since the admission of Hib vaccine



History and ExaminationLab:

Samples:

- Respiratory samples
- Blood for culture in invasive conditions and in epiglottitis
- Blood cultures are usually positive in epiglottitis.
- throat swabs in patients with suspected acute epiglottitis should NOT be carried out, as attempts to obtain the sample may precipitate complete airway obstruction

H. Influenzae / diagnosis

- The viability of *H. influenzae* in clinical specimens declines with time, particularly at 4°C > immediate transfer
- 1. Antigen detection
- The detection of type b polysaccharide antigen in body fluids or pus is useful, particularly in patients who received antibiotics before specimens were obtained.
- A rapid latex agglutination test with rabbit antibody to type b polysach. Capsular antigen is used most commonly.
- □ Cross reactivity with *pneumococcus and E.coli*

H. Influenzae / diagnosis

2. Culture on chocolate agar - incubate in an aerobic atmosphere enriched with 5-10% carbon dioxide.(or on blood agar with staph streak - satellitism)

- On chocolate agar the colonies are smooth, mucoid, grey or colorless
- Gram stain: Gram negative CB
- Catalase and oxidase tests are positive
- Antibiotic sensitivity

H. Influenzae / diagnosis



H. Influenzae / treatment

- Untreated invasive infection: Mortality rate of 90%
- Start empirically until you get sensitivity results:
- I. Non-invasive: Clav. Acid / Amoxicillin or Macrolides orally
- (to overcome the beta lactamase production by the bacteria)
- II. Invasive: Cephalosporins IV

III. Skilled medical and nursing care is also vital in the management of acute epiglottitis, where maintenance of a patent airway is crucial.

H. Influenzae / prevention

- Conjugate vaccines for type b :
- polysaccharide capsule is covalently coupled to proteins such as a non-toxic variant of diphtheria toxin or *Neisseria meningitidis* outer membrane protein
 - 3 doses separated by a month 2,3,4 months age and a booster at 12 months of age

H. Influenzae / prevention

- Immunization of infants significantly reduces pharyngeal carriage of Hib, but has no effect on the carriage of other capsular types or non-capsulate strains.
- Conjugate Hib vaccine is recommended for children and adults with splenic dysfunction, sickle cells disease, malignancies because they are at increased risk of invasive Hib infection.

Other haemophilus

- H. parainfluenza:
- 🖌 As H. influenza
- Usually less serious
- Requires factor V only
 - H. Ducreyi:
- Causes sexually transmitted chancroid
- Requires factor X only

The End