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Important! Doctor's Notes Only found in females' slides Only found in males' slides Extra Notes



This lecture is made by Physiology team

'I'm not telling you it's going to be easy. I'm telling you it's going to be worth it."

Objectives

- Discuss the epidemiology and pathophysiology of pneumonia and CAP
- Explain the different classifications of pneumonia
- Recognize clinical presentations associated with CAP
- Discuss the diagnosis and treatment of CAP
- Identify common etiological agents causing CAP and discuss their laboratory work up
- Discuss virulence factors and prevention of *Streptococcus pneumoniae*.

Definition

- Pneumonia: is an infection that leads to inflammation of the parenchyma of the lung (the alveoli) (consolidation and exudation).
- It may present as <u>acute</u>, <u>fulminant clinical</u> disease or as a chronic disease with a more prolonged course.

Epidemiology

- Overall the rate of CAP 5-6 cases per 1000 persons per year. Usually Its transmission by droplet.
- Mortality 23%
- High, especially in old people Almost 1 million annual episodes of CAP in adults > 65 yrs in the US.





Risk factors

- Age < 2 yrs, > 65 yrs
- Alcoholism
- Smoking
- Asthma and COPD
- Aspiration
- Dementia (a chronic mental disorder marked by memory loss, personality changes and impaired reasoning)
- Prior influenza
- HIV; Immunosuppression
- Institutionalization
- Recent hotel : Legionella
- Travel, pets, occupational exposures- birds (C. psittaci)



Defense mechanism of respiratory tract

- Filtration and deposition of environmental pathogens in the upper airways
- Cough reflux
- Mucociliary clearance
- Alveolar macrophages
- Humoral and cellular immunity
- Oxidative metabolism of neutrophils

Pathophysiology

- 1. Inhalation or aspiration(لما يدخل شيء للرئة بالغلط) of pulmonary pathogenic organisms into a lung segment or lobe.
- 2. Results from secondary bacteraemia from a distant source, such as Escherichia coli urinary tract infection and/or bacteraemia (less commonly).
- 3. Aspiration of oropharyngeal contents (multiple pathogens).

Pneumonia is classified according to:

Acquired environment

- community acquired pneumonia.
- hospital acquired pneumonia.
- nursing home acquired pneumonia.

Anatomy

- Lobar: entire lobe.
- Bronchopneumonia
 Interstitial.



Fungal Candida Aspergillosis Pneumocystis carn

Viral

- Adenoviruses

- Influenza virus

Pathogen

Parasite

Bacterial

- Typical: <u>Gram-positive</u> : Streptococcus pneumoniae(most common), Staphylococcus aureus, Group A hemolytic streptococci. <u>Gram-negative</u> : Klebsiella pneumoniae, Hemophilus influenzae, Moraxella catarrhal and Escherichia coli.

- Atypical:

Mycoplasma pneumoniae(most common), chlamydophila pneumoniae and legionella. Anaerobic bacteria



Community acquired pneumonia pathogens

•Strep pneumonia (the most)	48%
•Viral	23%
 Atypical orgs (MP,LG,CP) 	22%
•Haemophilus influenza	7%
 Moraxella catharralis 	2%
•Staph aureus	1.5%
•Gram –ive orgs	1.4%
•Anaerobes	

According to pathogens :

Thanks

to 436

team

1- Bacteria (dominant)				2- Fungal	3- Viral pneumonia	4- Others
Typical pneumonia			Atvnical pneumonia	pneumonia	common cause of pneumonia in children less than 5 vears	
Gram +	Gram -	Anaerobic	Atypical priedmonia			
1)Streptococcus pneumoniae (most common Typical pneumonia) 2)Staphylococcus aureus 3) Group A hemolytic streptococci	1)Klebsiella pneumoniae 2)Hemophils influenzae 3)Moraxella catarrhal 4)Escherichia coli		1)Legionnaies pneumonia(Legionella) 2) <u>Mycoplasma pneumoniae</u> (most common) 3) <u>Chlamydiophila</u> pneumoniae 4) <u>Chlamydophila</u> Psittaci 5) <u>Rickettsias</u> . 6) <u>Francisella</u> tularensis (tularemia)	1)Candida. 2)Aspergillosis. 3)Pneumocystis jiroveci (carinii), It causes PCP.	 1)Respiratory syncytial V. 2)Influenza V. 3)Adenoviruses. 4)Human metapneumovirus. 5)SARS and MERS CoV. 6)Cytomegalovirus. 7)Herpes simplex virus. 	1)Parasites 2)Protozoa 3)Chemical 4)Allergy

1)Streptococcus pneumoniae, H.influenzae and Moraxella: have cell wall therefore are gram stained and respond to Penicillin and B-lactam

2)Mycoplasma pneumoniae, Legionella and chlamydia: doesn't have cell wall (resistant to drugs that work on cell wall E.g (penicillin and B-lactam)

Actually legionella has cell wall but a little bit different

Clinical Manifestation

Typical Pneumonia Signs and symptoms are quite similar but Atypical pneumonia is less severe.

Atypical Pneumonia

- The onset is acute (2-3 days)
- Prior viral upper respiratory infection
 (follows viral infection) (follows viral infection) المريض يصاب بالنومونيا
 يكون أصابه فيروس فيهيئ الاير وايز للنومونيا
- Fever
- Shaking chilis
- Cough with sputum production (rustysputum)
- Chest pain or pleurisy(inflammation of the pleura)
- Shortness of breath (dyspnea)

- Gradual onset
- Usually mild to moderate but in case of legionella it could be severe
- Headache, malaise and low grade fever
- Dry cough
- Arthralgia / Myalgia والعضلات
- In case of doctor examination, signs such as minimal, few crackles and rhonchi could notify.



- 2. X-ray (to determine whether the pneumonia is lobar, lobular or interstitial)
- 3. Laboratory
- CBC test shows leukocytosis
- Sputum either gram stain(low sensitivity due to normal flora) or culture

4. Blood culture not always but usually in case of pneumococcal pneumonia

5. Culture of pleural fluid if there is effusion

- 2. X-ray (to determine whether the pneumonia is lobar, lobular or interstitial)
- 3. Laboratory
- CBC test shows mild elevation of WBC
- LFTs show elevation of ALT & Alk Phos
- Sputum culture in case of Legionella
- Detection of antibodies
- DNA detection
- 4. Urine antigen for Legionella

Gram Positive diplococci - Alpha hemolytic - Catalase negative - Normal flora of UPT in 20-40% of people. Can cause respiratory infections such as **pneumonia, sinusitis and otitis**. Also, can cause non respiratory infections such as meningitis and bacteremia.

Virulence Factors:

- **Capsule** "most important virulence factor"
- There are more than 90 types of capsule.

important

- Pneumolysin (pore forming toxin, also stimulates cytokines and disrupts the cilia of respiratory epithelial cells released on lysis of organism by autolysin)
 Autolysin
 باللاب أول يومين ينمو بشكل طبيعي بعدها يكون شكل العينة مسطح وكانها ماتت
- Neuraminidase(cleaves sialic acid) Prevention: There is vaccination for S.pneumonia but it does not cover all the types of capsule.

Gram + diplococci Neutrophils





-Alpha hemolytic consist of two broad groups S.Pneumonia and S.Viridans.
- S.Pneumonia sensitive to Optochin unlike S.Viridans which is resistance."1"
- S.Pneumonia lysed"soluble" by bile unlike S.Viridans which insoluble in bile."2"

This two tests differentiate between S.pneumonia & S. Viridans



Atypical Pneumonia

General Principles

•Approximately 15% of all CAP.

- •Not detectable on gram stain.
- •Won't grow on standard media.
- •Most don't have a bacterial cell wall "Don't respond to β-lactams ".
- Usually less severe than Typical Pneumonia with some exceptions.

Causative Agents

- Chlamydia pneumonia
- Mycoplasma pneumonia
- •Legionella spp
- Psittacosis (Chlamydia psittaci)
- •Q fever (Coxiella burnettii)
- •Viral (Influenza, Adenovirus, Rhinovirus)
- Pneumocystis Jiroveci

Atypical(interstitial) pneumonia

Symptoms

- Insidious onset
- Mild to severe
- Headache
- Malaise
- Fever
- Dry cough
- Arthralgia myalgia

Signs

- Minimal
- Low grade fever

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عشان يوضح أننا

بدينا

بالأثبيكال

- Crackles
- Rhonchi (similar to wheezing)

Diagnosis and treatment

Diagnosis:

- X-ray
- CBC
- U&E test (urea and electrolytes) Low serum Na (legionella)
- LFT (Liver function test)
- ▲ALT ▲ ALK phos
- Sputum culture on special media(BCYE)
- Urine antigen for legionella
- Serology for detecting antibodies
- DNA detection

Treatment:

- Macrolides
- Quinolones
- Tetracycline
- B lactams have no effect

Because there is no cell wall to be destroyed

• Treat for 10-14 days

-CBC is general infection detector -X-ray because it's pulmonary and we need to see the pattern (lobar,interstitial...) -We culture to see the microorganisms - organisms specific signs like ALT We use every group that we have study except B-lactam

Mycoplasma pneumonia

- •Eaton's agent (1944)
- •No cell wall
- Common
- •Rare in children and in > 65
- •People younger than 40.
- •Crowded places like schools, homeless shelters, prisons.
- •Can cause URT symptoms
- •Usually mild and responds well to antibiotics.
- •Can be very serious

May be associated with

extra-pulmonary findings: skin rash, hemolysis, myocarditis, pancreatitis, encephalitis

Diagnosis:

Serology

NAAT Nucleic acid amplification tests. Done on throat or Np (nasopharynx) swab

Culture can be done but requires special media and slow grower (weeks)

Cold agglutinins test not specific but can be helpful

Al Rekabi notes: Hard to culture Can be detected by Complement fixation test But the better way is The cold agglutination test



Cx-ray

Chlamydia pneumonia

- •Obligate intracellular organism
- 50% of adults sero-positive
- Mild disease
- Sub clinical infections common
- •5-10% of community acquired pneumonia
- Diagnosis:SerologyNAAT

Al Rekabi notes: Causes three diseases: 1. Pneumonia 2. Urethritis 3. trachomatis

Psittacosis



- •Chlamydia psittaci
- •Exposure to birds
- Bird owners, pet shop employees, vets
- Parrots, pigeons and poultry
- Birds often asymptomatic.
- The disease/infection that come from animals are called zoonotic.
- Less common more severe

Q fever (Coxiella burnetti)



Drs note: know that its from animal like sheep Exposure to farm animals mainly sheep
Spread by inhalation of infected <u>animal birth</u> products

Pneumonia is acute form of infection

•Diagnosis: serology

Legionella pneumophila

Legionnaire's disease
<u>Serious outbreaks</u> linked to exposure to cooling towers
<u>Can be very severe and lead</u> to ICU admission.
Diagnosis:

Specimen: sputum(not accurate)

- -Culture on specialized media (BCYE)
- -DFA (low sensitivity)
- -NAAT

oUrine antigen testing

Can cause Hyponatraemia common (<130mMol)

Bradycardia WBC < 15,000 Abnormal LFTs Raised CPK Acute Renal failure

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الفنادق من

والمياة عشان كذا في البداية قلنا

Pontiac fever: Non pneumonic Influenza like illness Self limiting Related to exposure to environmental aerosols containing Legionella (potentially reaction to bacterial endotoxins)

Verv

important

Legionnaires in IC

Legionella presents on X-Ray as: interstitial pneumonia but can present as lobar or any other type

Antibiotic Treatment of CAP

- Factors to consider in selection of antibiotic:
 - Co morbidities
 - Previous antibiotic exposure in last 3 months
 - Severity
 - Out patient management vs requiring inpatient admission vs requiring ICU

Treatment

		Macrolides E.g. Erythromycin 	Doxycycline Like Tetracycline but wider range	Levoflo xacin From quinolo nes	B-lactam And Macrolide B-lactam includes Penicillin And cephalosporins	B-lactam And Levo To cover everythin 8
Outpatient, healthy patient with no exposure to antibiotics in the last 3 months	- <i>S. pneumoniae</i> -Atypical pathogens -Viral					
Outpatient, patient with comorbidity or exposure to antibiotics in the last 3 months	As above + Anaerobes S. aureus					
Inpatient : Not ICU	Same as above + coliforms					
Inpatient : ICU	Same as above + <i>Pseudomonas</i>					

Quiz

Which of the following most common causes of the community acquired pneumonia?

A-S.pneumonia

B-viral

C-H.influenza

D-M.catharralis

Which of the following is true about S.pneumonia? A-gram+ and beta hemolytic B-gram- and beta hemolytic C-gram+ and alpha hemolytic D-gram- and alpha hemolytic

Which of the following organism don't have cell wall? A-s.pneumonia B-s.aureus C-gram - organisms D-Mycoplasma pneumonia Which of the following organism is transmitted by exposure to birds ? A-Legionella B-Mycoplasma C-S.pneumonia D-Chlamydia psittaci

Which of the following culture is used for diagnosis legionella ? A-SDA B-BCYE C-Lowenstein-Jensen D-Blood agar

Which of the following antibiotics is best choice for legionella? A-Azithromycin B-clarithromycin C-cephalosporins D-penicillin

1-A 2-C 3-D 4-D 5-B 6-A (because legionella is gram - and azithromycin effect against gram - bacteria)

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