Community acquired pneumonia and treatment options

•6 mechanisms in pathogenesis of pneumonia – (inhalation, aspiration, direct inoculation, reactivation, defects in pulmonary defenses, blunted cellular or humoral immune response)

•In general, clinical presentation: pleuritic chest pain, fever, cough, decreased breath sounds, dull to percussion, egophony

•Classified as "typical" or "atypical" or viral

•Typical = S. pneumoniae, H. influenza, S. aureus, enteric gram negative bacteria. More segmental or lobar infiltrate.

•Atypical = Mycoplasma, Legionella, Chlamydia pneumonia. More diffuse infiltrate. Sputum gram stain usually negative.

•Empiric treatment prior to susceptibility testing- cover S. pneumoniae, S. aureus, H. influenzae. Give amoxicillin-clavulanate, and 2nd or 3rd generation cephalosporin, or respiratory fluoroquinolone (3rd and 4th generation).

CAP patient characteristics

- Smokers: S. pneumo, H. influenzae, Moraxella
- Post viral bronchitis: S. pneumo
- No co-morbidity: atypicals, viral
- Alcoholic: S. pneumo, anaerobes
- IV drug user: S. aureus
- Epidemic: Legionnaire's
- Airway obstruction: anaerobes
- Animals: Psittacosis, Tularemia, Coxiella burnetii

Mechanisms of action of different antimicrobials

• **Penicillins-** Inhibits the transpeptidase enzyme step in peptidoglycan cell wall synthesis

1st generation:[ex Pen G] most streptococci, oral anaerobic coverage

2nd generation:[nafcillin] most streptococci, S. aureus (Penicillinase resistant)

3rd generation:[amoxicillin, ampicillin] most streptococci, basic G – coverage.

Amoxicillin is oral DOC for susceptible strains of S. pneumoniae

Amoxicillin with clavulanate treats beta lactamase producing bacteria like H. influenzae, Methicillin sensitive Staph aureus and anaerobes

4th generation:[piperacillin] extended spectrum, includes pseudomonas

• Cephalosporins-inhibits cell wall synthesis

Parenteral DOC for CAP caused by susceptible strains of S pneumoniae, H influenzae, Staph aureus. 1st generation: G+ (including Staph aureus), basic G-

- 2nd generation: G+, diminished Staph aureus, improved G- coverage, some anaerobic coverage
- 3rd generation: further diminished S. aureus, further improved G-, some Pseudomonal coverage and diminished G+ coverage
- 4th generation: same as 3rd plus coverage against Pseudomonas

**note that neither group is effective against "atypicals"

• Macroglides-inhibits 50s subunit of ribosome in protein synthesis

Treats mycoplasma, legionnaire's, chlamydial infections

Active against most common pathogens and atypical agents. Macroglide resistance is emerging to Strep pneumoniae

• Fluoroquinolones-inhibits DNA gyrase

- Broad spectrum against likely agents of CAP. Active against penicillin resistant Strep pneumoniae. Resistance developing.
- 1st generation: G- NOT pseudomonas, UTI only, NO atypicals
- 2nd generation: G- including pseudomonas, S. aureus, some atypicals NOT pneumococcus
- 3rd generation: G-, G+, expanded atypicals
- 4th generation: same as 3rd plus enhanced coverage of pneumococcus, decreased activity vs pseudomonas.

• **Tetracyclines-** inhibits 30s subunit of ribosome in protein synthesis

G+ and G-, aerobic & anaerobic bacteria, "atypicals"- mycoplasma, chlamydia and category A bioterrorism agents

Streptococcus pneumoniae (2/3 of CAP cases)

- Properties: G + diplococci "lancet shaped", α hemolytic.
 #1 cause CAP: "rusty sputum", single rigor, fever.
- Tx and dose:
- sensitive strains treat with penicillin G 250,000-400,000 units/kg/day IV divided q 4-6 or amoxicillin (oral) 1g PO tid.
- intermediate resistance strains are susceptible to 2nd/3rd generation parenteral cephalosporins and respiratory fluoroquinolones

Cefotaxime 1-2g IM/IV q6-8 or ceftriaxone1-2 g IM/IV q24h. Levofloxicin 500mgIV/po qd (others: alatrofox-, gati-, moxi-)

• **Prevention:** capsular vaccines (prevnar, pneumovax)

• H. influenzae

trimethoprim-sulfamethoxazole, Ampicillin IV, Amoxicillin po, azithro/clarithromycin, doxycycline

If severe, 3rd gen cephalosporin

Enteric gram negatives

Anaerobes: clindamycin or beta lactam/beta lactamase inhibitor

Staph aureus

Sensitive strains: Nafcillin or oxacillin (penicillinase resistant abx)

MRSA- Vancomycin

Atypicals

Legionella- fluroquinolone or azithromycin

Mycoplasma- erythro, azithro, clarithromycin, or fluoroquinolone

Chlamydia pneumoniae- doxycycline, erythrocyclin or floroquinolones

Duration of therapy

- S pneumoniae: until afebrile for 3-5d
- C pneumoniae:7-14 d
- M pneumoniae: not well established.
- Legionella: 10-21 d
- S aureus, P aeruginosa, Klebsiella, anaerobes >3weeks