# Respiratory Block PNEUMONIA

Dr. Wajd Althakfi, MD Consultant Histopathology KSU-KKUH





### **Objectives**

- Understand that pneumonia is an inflammatory condition of the lung characterized by consolidation (solidification) of the pulmonary tissue
- Is aware of the pathogenesis of pneumonia and its classification which principally include bronchopneumonia, lobar pneumonia and atypical pneumonia
- Can appreciate the etiology and pathogenesis of lung abscess

### **PULMONARY INFECTIONS**

- Pulmonary infections in the form of pneumonia are responsible for one sixth of all deaths in the United States
- Pneumonia defined as any infection in the lung
- Normally, the lung parenchyma remains sterile because of a number of highly effective immune and non- immune defense mechanisms that extend throughout the respiratory system from the nasopharynx to the alveolar air spaces
- · The vulnerability of the lung to infection despite these defenses is not surprising because
  - 1. many microbes are airborne and readily inhaled into the lungs
  - 2. nasopharyngeal flora are regularly aspirated during sleep, even by healthy individuals
  - 3. lung diseases often lower local immune defenses
- The vast majority are upper respiratory tract infections caused by viruses, but bacterial and fungal infections can also occur
- Pneumonias are classified by the specific etiologic agent, which determines the treatment, or, if no pathogen can be isolated, by the clinical setting in which the infection occurs

### Who's at risk ?

- Loss or suppression of the cough reflex, as a result of coma, anesthesia, neuromuscular disorders, drugs, or chest pain (may lead to aspiration of gastric contents)
- Injury to the mucociliary apparatus, by either impairment of ciliary function or destruction of ciliated epithelium, due to cigarette smoke, inhalation of hot or corrosive gases, viral diseases, or genetic defects of ciliary function (e.g., the immotile cilia syndrome)
- Accumulation of secretions in conditions such as cystic fibrosis and bronchial obstruction
- Interference with the phagocytic or bactericidal action of alveolar macrophages by alcohol, tobacco smoke, anoxia, or oxygen intoxication
- Pulmonary congestion and edema







# Pathogenesis & classification

#### Portal of entry for most pneumonias is

- Inhalation of air droplets
- Aspiration of infected secretions or objects
- Hematogenous spread from one organ to other organs can occur

#### • Classification of pneumonia can be made according to causative agent

#### TABLE 15–8 The Pneumonia Syndromes

#### COMMUNITY-ACQUIRED ACUTE PNEUMONIA

Streptococcus pneumoniae Haemophilus influenzae Moraxella catarrhalis Staphylococcus aureus Legionella pneumophila Enterobacteriaceae (Klebsiella pneumoniae) and Pseudomonas spp.

#### COMMUNITY-ACQUIRED ATYPICAL PNEUMONIA

#### Mycoplasma pneumoniae Chlamydia spp. (C. pneumoniae, C. psittaci, C. trachomatis) Coxiella burnetii (Q fever)

Viruses: respiratory syncytial virus, parainfluenza virus (children); influenza A and B (adults); adenovirus (military recruits); SARS virus

#### HOSPITAL-ACQUIRED PNEUMONIA

- Gram-negative rods, Enterobacteriaceae (Klebsiella spp., Serratia marcescens, Escherichia coli) and Pseudomonas spp.
- Staphylococcus aureus (usually penicillin resistant)

#### ASPIRATION PNEUMONIA

Anaerobic oral flora (Bacteroides, Prevotella, Fusobacterium, Peptostreptococcus), admixed with aerobic bacteria (Streptococcus pneumoniae, Staphylococcus aureus, Haemophilus influenzae, and Pseudomonas aeruginosa)

#### CHRONIC PNEUMONIA

#### Nocardia

Actinomyces

Granulomatous: Mycobacterium tuberculosis and atypical mycobacteria, Histoplasma capsulatum, Coccidioides immitis, Blastomyces dermatitidis

#### NECROTIZING PNEUMONIA AND LUNG ABSCESS

- Anaerobic bacteria (extremely common), with or without mixed aerobic infection
- Staphylococcus aureus, Klebsiella pneumoniae, Streptococcus pyogenes, and type 3 pneumococcus (uncommon)

#### PNEUMONIA IN THE IMMUNOCOMPROMISED HOST

Cytomegalovirus Pneumocystis jiroveci Mycobacterium avium-intracellulare Invasive aspergillosis Invasive candidiasis "Usual" bacterial, viral, and fungal organisms (listed above)

SARS, severe acute respiratory syndrome.

### **Anatomic classification of pneumonia**

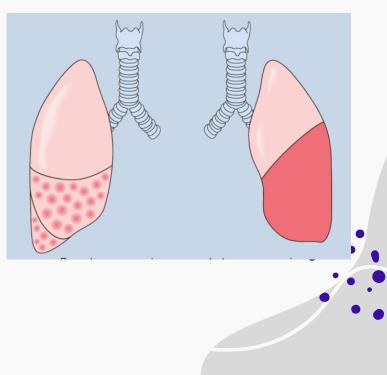
Classification of pneumonia also can be made according gross anatomic distribution of the disease:

#### 1. Alveolar

- Bronchopneumonia: (Streptococcus pneumoniae, Haemophilus influenza, Staphylococcus aureus) Represent an extension from preexisting bronchitis or bronchiolitis. Extremely common tends to occur in two extremes of life
- Lobar pneumonia: (Streptococcus pneumoniae) Acute bacterial infection of a large portion of a lobe or entire lobe. Classic lobar pneumonia is now infrequent
- Note: Overlap of the two patterns often occur

#### 2. Interstitial:

o Influenza virus (children), Mycoplasma pneumoniae

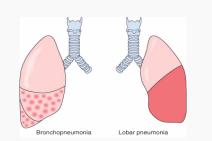


#### Community-Acquired Acute Pneumonia

- Usually, Bacterial
- Can follow URT infection
- It can be lobar or bronchopneumonia
- **Clinical features:** Sudden onset of high fever, chills, pleuritic chest pain and productive cough, may be with hemoptysis
- Reduced air entry and dullness by percussion
- Cause:
  - The most common cause of Community-Acquired Acute Pneumonia is *Streptococcus pneumoniae*
  - **Other common causes:** Haemophilus influenzae, Moraxella catarrhalis, Staphylococcus aureus, Legionella pneumophila, Klebsiella pneumoniae and Pseudomonas aeruginosa spp.
  - In intravenous drug abuser: Staphylococcus aureus
  - It is more common in:
    - Underlying chronic disease e.g., DM, COPD, and congestive heart failure
    - Congenital or acquired immune deficiency
    - Decreased or absent splenic function

#### Bronchopneumonia

- most common agents are:
- Streptococcus pneumoniae,
- Haemophilus Influenza, in COPD
- Pseudomonas Aeroginosa in CF
- coliform bacteria.
- staphylococci



#### Lobar pneumonia

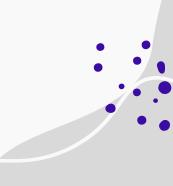
- 90-95% are caused by pneumococci(Streptococcus pneumoniae)
- Rare agents: *K. pneumoniae,* staphylococci, streptococci, *H. influenzae,* Pseudomonas and Proteus





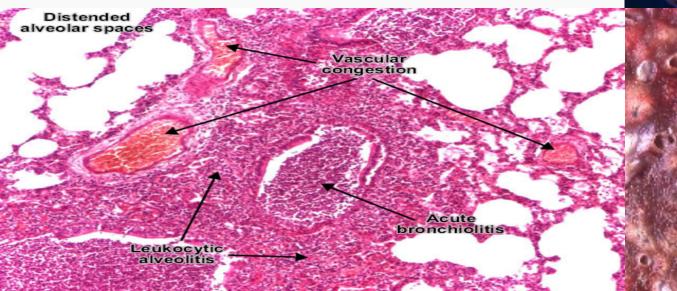
### Bronchopneumonia

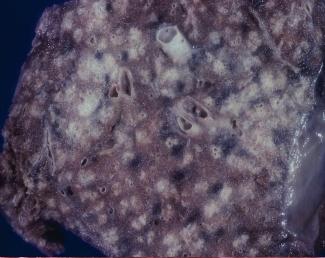
- Are focal/patchy areas of consolidated acute suppurative inflammation in one or more lobes
- Usually it involves lower lobes (basal) bilaterally because there is a tendency of the secretions to gravitate into the lower lobes
- Well developed lesions are 3 to 4 cm dry grey red ill defined nodules



### Bronchopneumonia

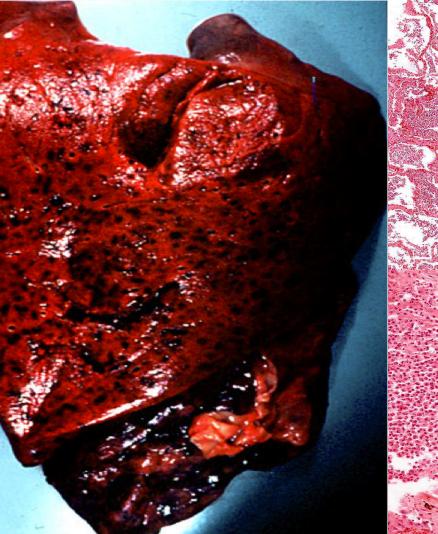
- **Grossly:** patchy consolidation of the lung, slightly elevated, dry, granular, gray-red to yellow, and poorly delimited at their margins
- **Micro:** suppurative, neutrophil-rich exudate that fills the bronchi, bronchioles, and adjacent alveolar spaces





### Lobar pneumonia

- It is widespread involvement of a large area and even an entire lobe of lung (widespread fibrinosuppurative consolidation)
- There are 4 stages:
- I. Stage I: Congestion: lung is heavy, boggy and red. The intra-alveolar space is filled with fluid, few scattered neutrophils and numerous bacterias
- II. Stage II: Red hepatization (solidification): alveolar spaces are filled with neutrophils, red cells (congestion) and fibrin. Grossly the lung is firm/solid red and liver-like
- III. Stage III: Gray hepatization: here the red cells are reduced but neutrophils and fibrin(fibrinopurulent/suppurative exudate) are still present. Grossly the lung is still firm/solid and liver-like but grey
- IV. Stage IV: Resolution: exudates within the alveoli are being enzymatically digested, resorbed, ingested by macrophages or coughed up



all the alveolar air spaces are distended by a cellular infiltrate

careful examination of this infiltrate shows it to consist of neutrophils.

Red hepatization Alveolitare filled with fibrin, RBC's and

Case 34: Lobar pneumonia

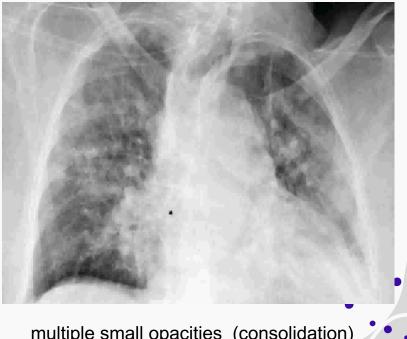
34: Lobar pneumonia

the alveolar septae are widened and also show infiltration by neutrophils.

neutrophils

### **Clinical features**

- Abrupt onset of high fever, shaking chills, cough • productive mucopurulent sputum occasional patients may have hemoptysis
- When fibrinosuppurative pleuritis is present, it is • accompanied by pleuritic pain and pleural friction rub
- Radiology: •
  - in lobar pneumonia there is a radio opaque (consolidation) well circumscribed lobe
  - in bronchopneumonia there are multiple small opacities usually basal and bilateral



multiple small opacities (consolidation)

# Complications

- Tissue destruction and necrosis, causing abscess formation (particularly common with type 3 pneumococci or *Klebsiella* infections)
- Spread of infection to the pleural cavity, causing the intrapleural fibrinosuppurative reaction known as **empyema**
- **Bacteremic dissemination** to the heart valves, pericardium, brain, kidneys, spleen, or joints, causing metastatic abscesses, endocarditis, meningitis, or suppurative arthritis









### COMMUNITY-ACQUIRED ATYPICAL (VIRAL AND MYCOPLASMAL) PNEUMONIAS

- Characterized by patchy inflammation in the lungs confined to the alveolar septae and pulmonary interstitium and therefore it is called interstitial pneumonitis
- The term *atypical* denotes the moderate amount of sputum, no physical findings of consolidation, only moderate elevation of white cell count, and lack of alveolar exudate
- It is caused by many organisms but the most common is *Mycoplasma pneumonia* (common in children and young adults)
- Others include:

• **Viruses** e.g. respiratory syncytial virus, influenza virus (children), influenza A and B (adults); adenovirus and SARS virus

• **Chlamydia** spp. (*C. pneumonia etc.*) and *Coxiella burnetti* (Q fever). Chlamydia is transmitted by inhalation of dried excreta of infected birds and causes ornithosis/psittacosis.

Predisposing factors: malnutrition, alcoholism and any underlying debilitating disease

#### Gross:

- All causal agents produce essentially similar morphologic patterns
- The lung involvement may be quite patchy or may involve whole lobes bilaterally or unilaterally
- The affected areas are red-blue and congested. The pleura is smooth, and pleuritis or pleural effusions are infrequent

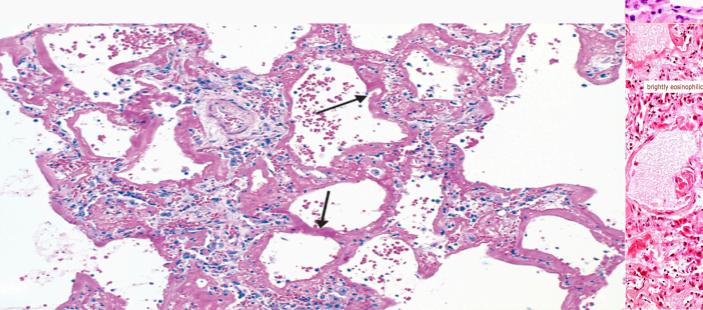






#### <u>Micro</u>:

- Predominantly there is inflammation in the interstitium/alveolar wall
- Alveolar septa are widened and edematous with mononuclear inflammatory infiltrate (and neutrophils in acute cases only)
- Sever cases: Intra-alveolar proteinaceous material with pink hyaline membrane lining the alveolar walls (diffuse alveolar damage)



# **Clinical course**

- The clinical course is extremely varied
- Many cases masquerade as severe upper respiratory tract infections or as chest colds
- Cough may be absent, and the major manifestations may consist only of fever, headache, muscle aches, and pains in the legs
- The ordinary sporadic form of the disease is usually mild with a low mortality rate, below 1%. Interstitial pneumonia (good prognosis)
- Secondary bacterial infection by staphylococci or streptococci is common in such circumstances



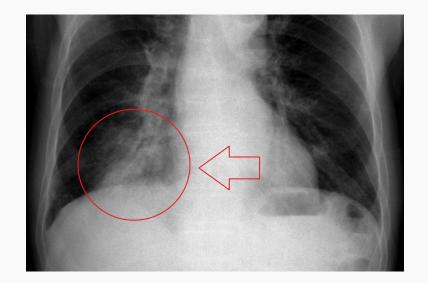
# HOSPITAL-ACQUIRED PNEUMONIA

- acquire terminal pneumonias while hospitalized (nosocomial infection)
- **Predisposing factor:** sever underlying conditions e.g. immunosuppression, prolonged antibiotic therapy, intravascular catheter and pt. with mechanical ventilator
- **Cause:** Gram-negative organisms like Klebsiella, Pseudomonas aeruginosa and E. coli have been implicated



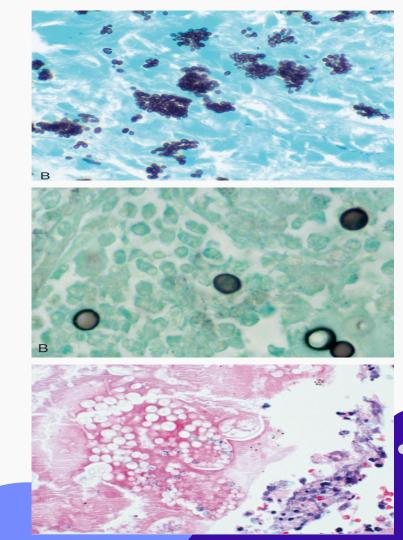
# Aspiration pneumonia

- Occurs in markedly debilitated patients or those who aspirate gastric contents either while unconscious (e.g., after a stroke) or during repeated vomiting
- These patients have abnormal gag and swallowing reflexes that predispose to aspiration
- The resultant pneumonia is partly chemical because of the extremely irritating effects of the gastric acid, and partly bacterial (from the oral flora)
- Typically, more than one organism is recovered on culture, aerobes being more common than anaerobes
- This type of pneumonia is often necrotizing, pursues a fulminant clinical course, and is a frequent cause of death. In those who survive, lung abscess is a common complication



# Chronic pneumonia

- Most often a localized lesion in an immunocompetent person, with or without regional lymph node involvement
- There is typically granulomatous inflammation
  - Bacteria (e.g., *M. tuberculosis*) or
  - Fungi (Histoplasma capsulatum, Coccidioides immitis, Blastomyces)
- In the immunocompromised, there is usually systemic dissemination of the causative organism, accompanied by widespread disease
- Tuberculosis is by far the most important entity within the spectrum of chronic pneumonias

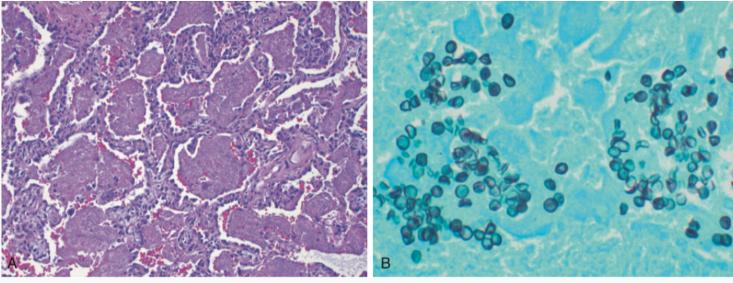


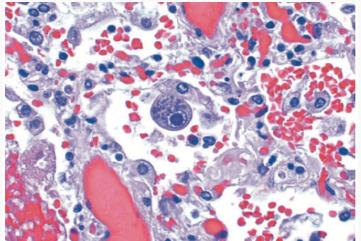
# Opportunistic pneumonia

- Infections that affect immunosuppressed patients (AIDS, cancer patients and transplant recipients)
- The appearance of a pulmonary infiltrate, with or without signs of infection (e.g., fever)
- Is one of the most common and serious complications in patients whose immune defenses are suppressed by disease, immunosuppressive therapy for organ transplants, chemotherapy for tumors, or irradiation

Diffuse Infiltrate	Focal Infiltrate
COMMON	
Cytomegalovirus <i>Pneumocystis jiroveci</i> Drug reaction	Gram-negative rods <i>Staphylococcus aureus</i> <i>Aspergillus</i> <i>Candida</i> Malignancy
UNCOMMON	
Bacteria <i>Aspergillus</i> <i>Cryptococcus</i> Malignancy	Cryptococcus Mucor Pneumocystis jiroveci Legionella pneumophila

TABLE 15–9 Causes of Pulmonary Infiltrates







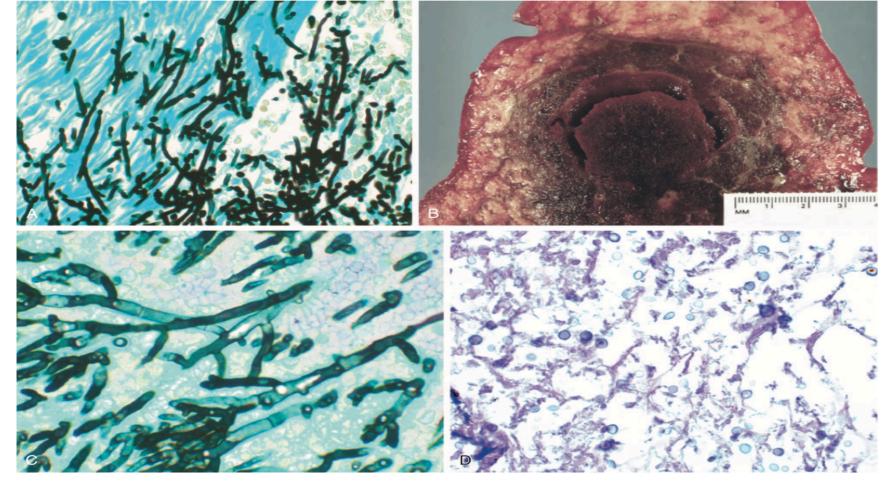
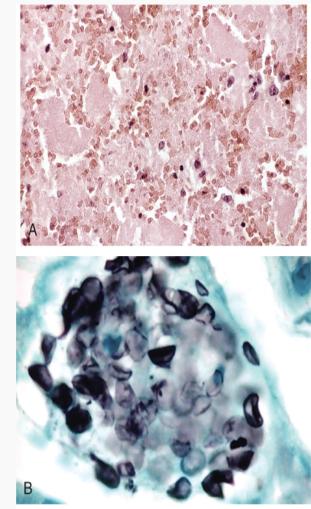


Fig. 13.42 The morphology of fungal infections. (A) Candida organism has pseudohyphae and budding yeasts (silver stain). (B) Invasive aspergillosis (gross appearance) of the lung in a hematopoietic stem cell transplant recipient. (C) Gomori methenamine-silver (GMS) stain shows septate hyphae with acute-angle branching, consistent with Aspergillus. (D) Cryptococcosis of the lung in a patient with AIDS. The organisms are somewhat variable in size. (B, Courtesy of Dr. Dominick Cavuoti, Department of Pathology, University of Texas Southwestern Medical School, Dallas, Texas.)

### **Pneumocystis** Pneumonia

- *P. jiroveci* (formerly *P. carinii*) is an opportunistic infectious agent considered as a fungus
- Seen in immunocompromised individuals especially
  AIDS
- Effective methods of diagnosis are:
  - Identify the organism in bronchoalveolar lavage fluids or in a transbronchial biopsy specimen.
  - Immunofluorescence antibody kits and PCR-based assays
- Microscopically:
  - characteristic intra-alveolar foamy, pink-staining exudate on H&E stains (A)
  - organism is trapped in the foamy material and can be seen on silver stain as oval cup shaped structures (B)



# Lung abscess

### Lung abscess

- Is localized suppurative necrotic process within
  the pulmonary parenchyma
- **Features:** tissue necrosis and marked acute inflammation. Abscess is filled with necrotic suppurative debris
- Organisms:
  - Staphylococci
  - Streptococci
  - Gram-negative organisms
  - Anaerobes
- Pathogenesis:
  - Can follow aspiration
  - As a complication of pneumonia
  - Septic emboli
  - Tumors
  - Direct infection





#### Lung abscess

#### **Clinical features**

- Prominent cough producing copious amount of foul smelling and bad-tasting purulent sputum
- Change in position evoke paroxysm of cough
- Fever malaise and clubbing of fingers
- Radiology shows fluid filled cavity

#### Complications

- Bronchopleural fistula and pleural involvement resulting in empyema
- Massive hemoptysis, spontaneous rupture into uninvolved lung segments
- Non-resolution of abscess cavity
- Bacteremia could result in brain abscess and meningitis

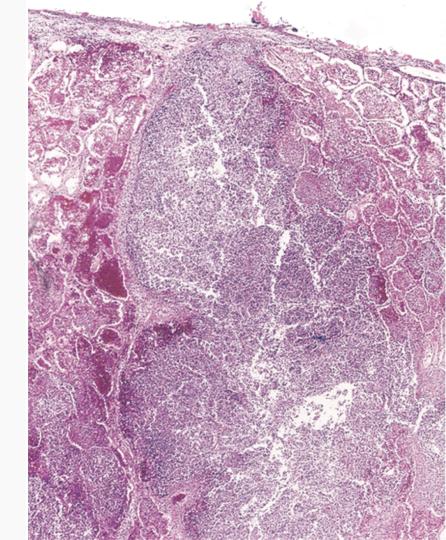
#### **Prognosis:**

• with antibiotic therapy 75% of abscess resolve



# Morphology

- Variable in size
- Single or multiple
- Pulmonary abscesses due to aspiration are more common on the right (because of the more vertical right main bronchus) and are most often single
- Abscesses that develop in the course of pneumonia or bronchiectasis are usually multiple, basal, and diffusely scattered
- The cardinal histologic change in all abscesses is suppurative destruction of the lung parenchyma within the central area of cavitation



#### **Summery (Acute pneumonia)**

- S. pneumoniae (the pneumococcus) is the most common cause of community-acquired bacterial pneumonia and usually has a lobar pattern of involvement.
- Morphologically, lobar pneumonias evolve through four stages: congestion, red hepatization, gray hepatization, and resolution.
- Other common causes of bacterial pneumonias in the community include H. influenzae and M. catarrhalis (both associated with acute exacerbations of COPD), S. aureus (usually secondary to viral respiratory infections), K. pneumoniae (observed in chronic alcoholics), P. aeruginosa (seen in individuals with cystic fibrosis, in burn victims, and in patients with neutropenia), and L. pneumophila, seen particularly in organ transplant recipients.
- Viral pneumonias are characterized by respiratory distress out of proportion to the clinical and radiologic signs, and by inflammation that is predominantly confined to alveolar septa, with generally clear alveoli.
- Common causes of viral pneumonia include influenza A and B, respiratory syncytial virus, human metapneumovirus, parainfluenza virus, and adenovirus.

# **Protect yourself**

