





# **Respiratory Fungal Infections**

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# Respiratory fungal infections

- Respiratory System
- Rout of infection?
- Oral Cavity, any role?
- Respiratory fungal infections are less common than viral and bacterial infections.
- Are opportunistic infections
  - Diseases in immunocompromised mainly , rarely in healthy hosts
- Have significant difficulties in diagnosis and treatment.

## **Inducing Immunosuppression**

AIDS

Bone marrow/ organ transplantation

Cancer: Leukemia, lymphoma etc

Drugs: Cytotoxic drugs, steroids etc

Endocrine related: Diabetes

Failure of organs: multi-organ

### **Other factors**

Increased survival of premature neonates

.More elderly pts

Long Stay in hospital/ ICU

Surgery

Devices

# Respiratory fungal infection - Etiology

## ➤ YEAST

- Candidiasis (*Candida* and other yeast)
- Cryptococcosis (*Cryptococcus neoformans*, *C. gattii*)

## ➤ Pneumocystosis (*Pneumocystis jiroveci*)

## ➤ Mould fungi

- Aspergillosis (*Aspergillus* species)
- Zygomycosis (*Zygomycetes*, e.g. *Rhizopus*, *Mucor*)
- Other mould

## ➤ Dimorphic fungi

- *Histoplasma capsulatum*
- *Paracoccidioides brasiliensis*
- *Blastomyces dermatitidis*
- *Coccidioides immitis*

Opportunistic

Primary infections

# Primary Systemic Mycoses

Infections of the respiratory system

Dissemination seen in immunocompromised hosts

Common in North America and to a lesser extent South America. Not  
.common in other parts of the World

Etiologies are dimorphic fungi. In nature found in soil of restricted  
.habitats

:They include

,Blastomycosis

,Histoplasmosis

,Coccidioidomycosis

Paracoccidioidomycosis

# Aspergillosis

Aspergillosis is a spectrum of diseases of humans and animals caused by members of the genus *Aspergillus*

These include

mycotoxicosis (1)

Allergy (2)

invasion and extension ) in preformed cavities Colonization (without (3)

Invasive, inflammatory, granulomatous, necrotizing disease of lungs (4)

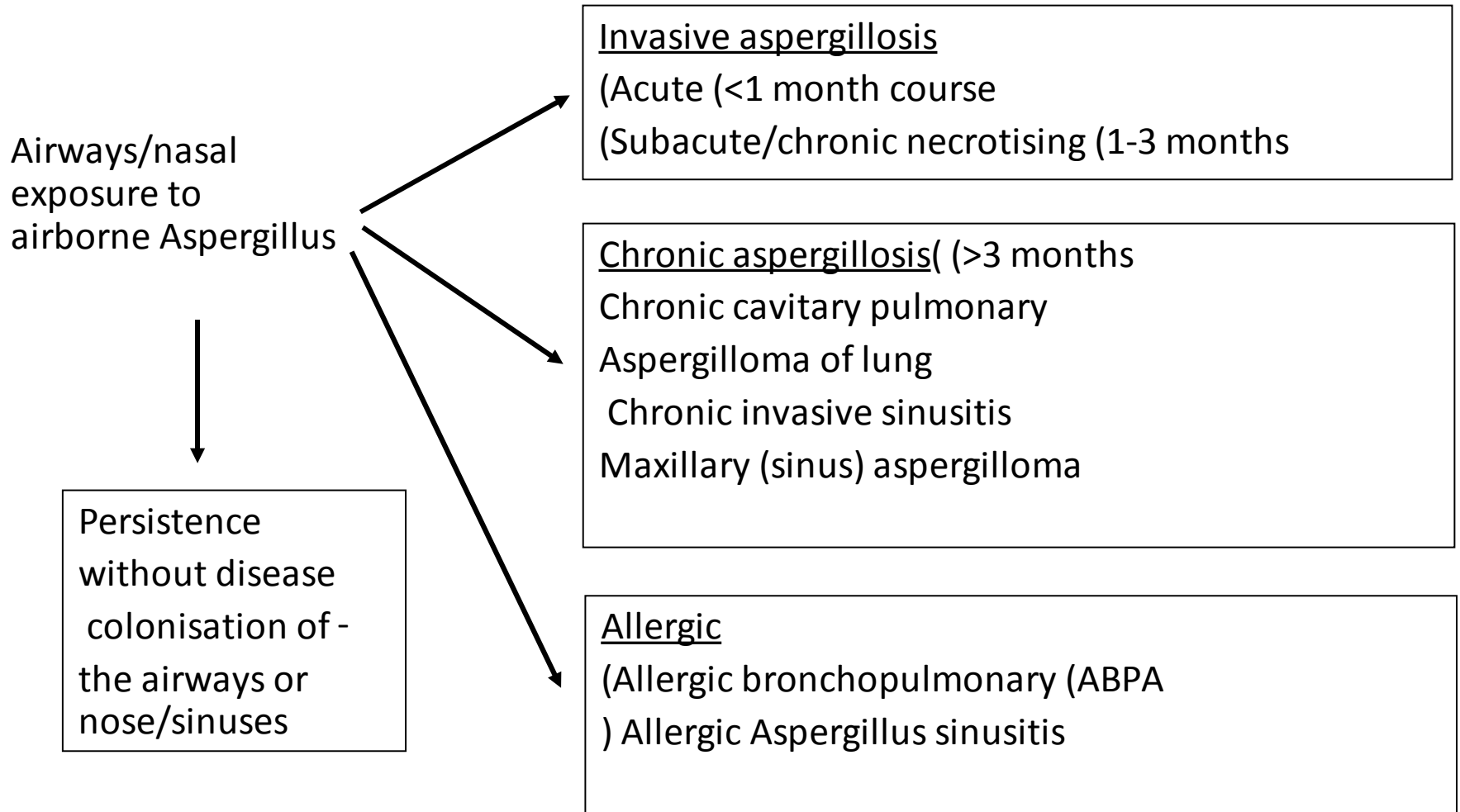
.systemic and disseminated disease (5)

The type of disease and severity depends upon the physiologic state of the host and the species of *Aspergillus*. causing the disease

**Aetiological Agents:** ,*Aspergillus species*

.common species are *A. fumigatus*, *A. flavus*, *A. niger*, *A. terreus* and *A. nidulans*

# CLASSIFICATION OF ASPERGILLOSIS





# Aspergillosis

**(Chronic Aspergillosis (Colonizing aspergillosis**

(Aspergilloma OR Aspergillus fungus ball)

signs include: Cough, hemoptysis, variable fever

Radiology will show mass in the lung , radiolucent crescent ■

**Invasive pulmonary Aspergillosis**

Signs: Cough , hemoptysis, Fever, Pneumonia, Leukocytosis

# Aspergillosis

:*Aspergillus sinusitis*

The most common cause is *Aspergillus flavus*  
in addition *Aspergillus*, there are other fungi that can cause fungal sinusitis

:Clinical

Nasal polyps – sinusitis

(Could disseminate to – eye → cranium (Rhinocerebral

Other diseases caused by :*Aspergillus*

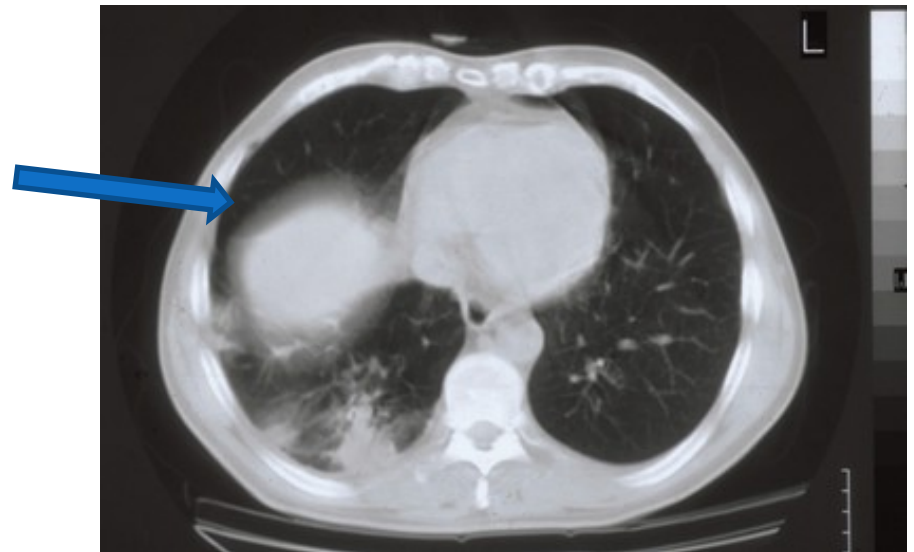
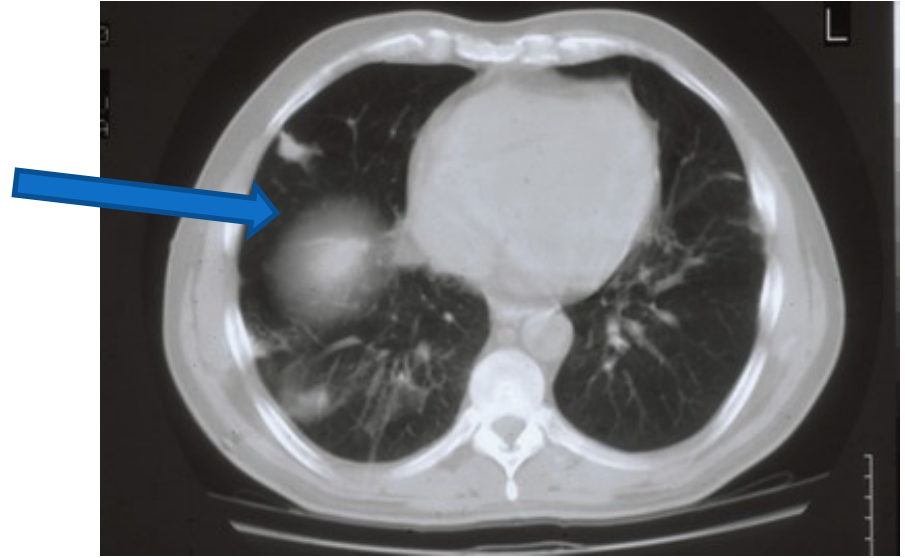
Corneal ulcer – endophthalmitis

(Otitis externa – otitis media (*A. niger*

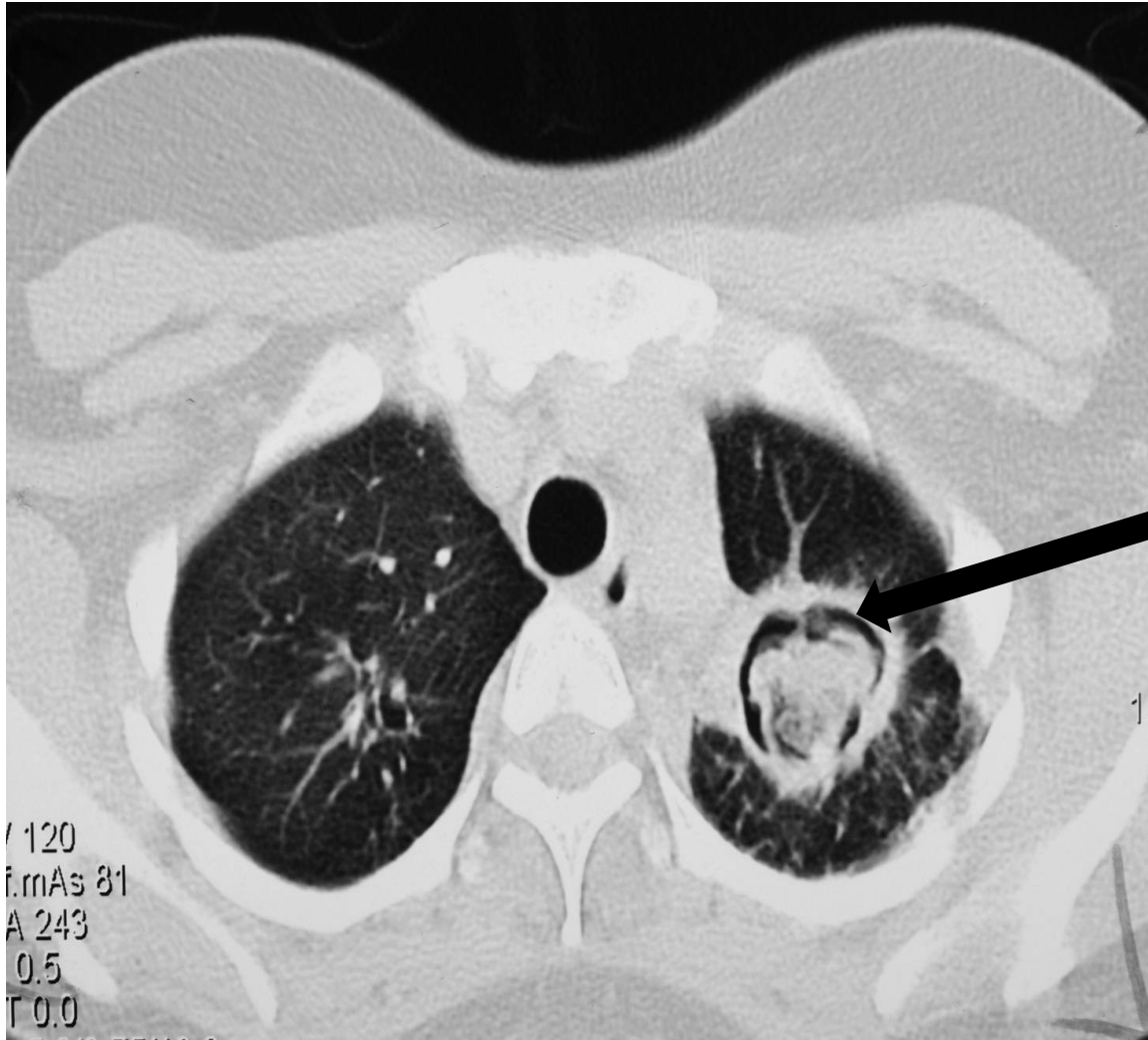
(Nail & skin infection (*A. niger*

# Invasive pulmonary aspergillosis in AIDS

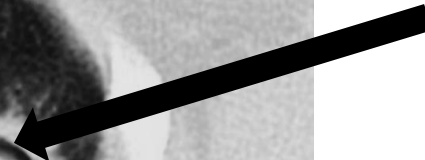
**Note the Halo sign**



# Simple (single) aspergilloma



**Note the Air  
crescent**



# ABPA

Hx Asthma  
Bronchial obstruction  
Fever, malaise  
Eosinophilia  
Wheezing +/-

Also:

Skin test reactivity to *Aspergillus*  
Serum antibodies to *Aspergillus*  
Serum IgE > 1000 ng/ml  
Pulmonary infiltrates



A link between airborne fungi and severe  
asthma

# Common airborne Aspergilli



*Aspergillus  
niger*



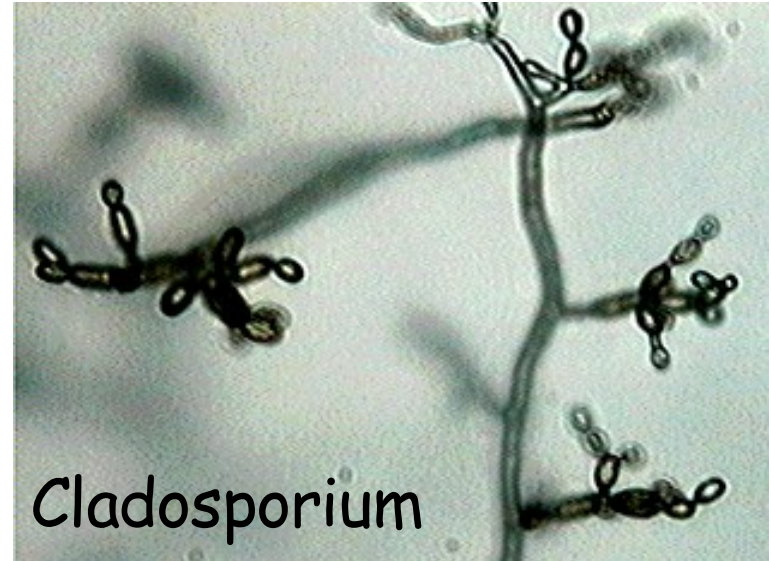
*Aspergillus  
fumigatus*



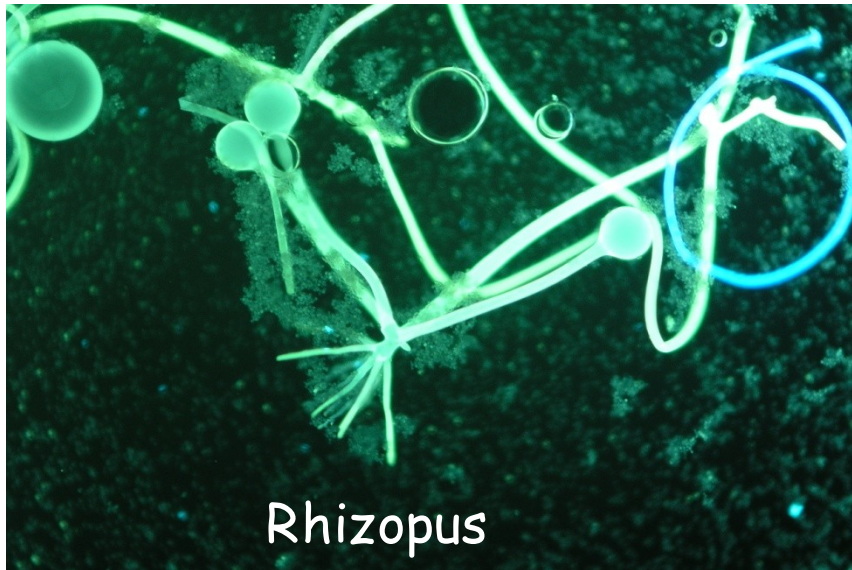
# Other important airborne fungi



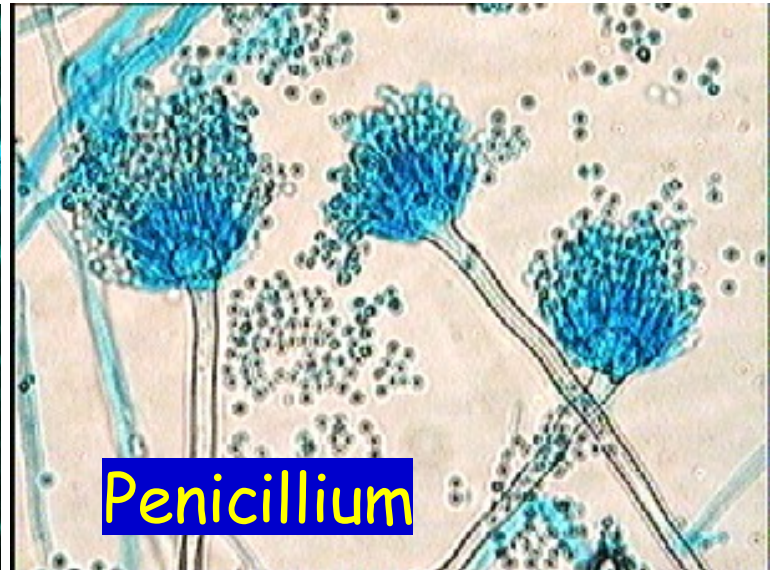
Alternaria



Cladosporium

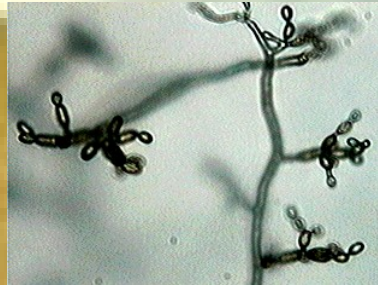
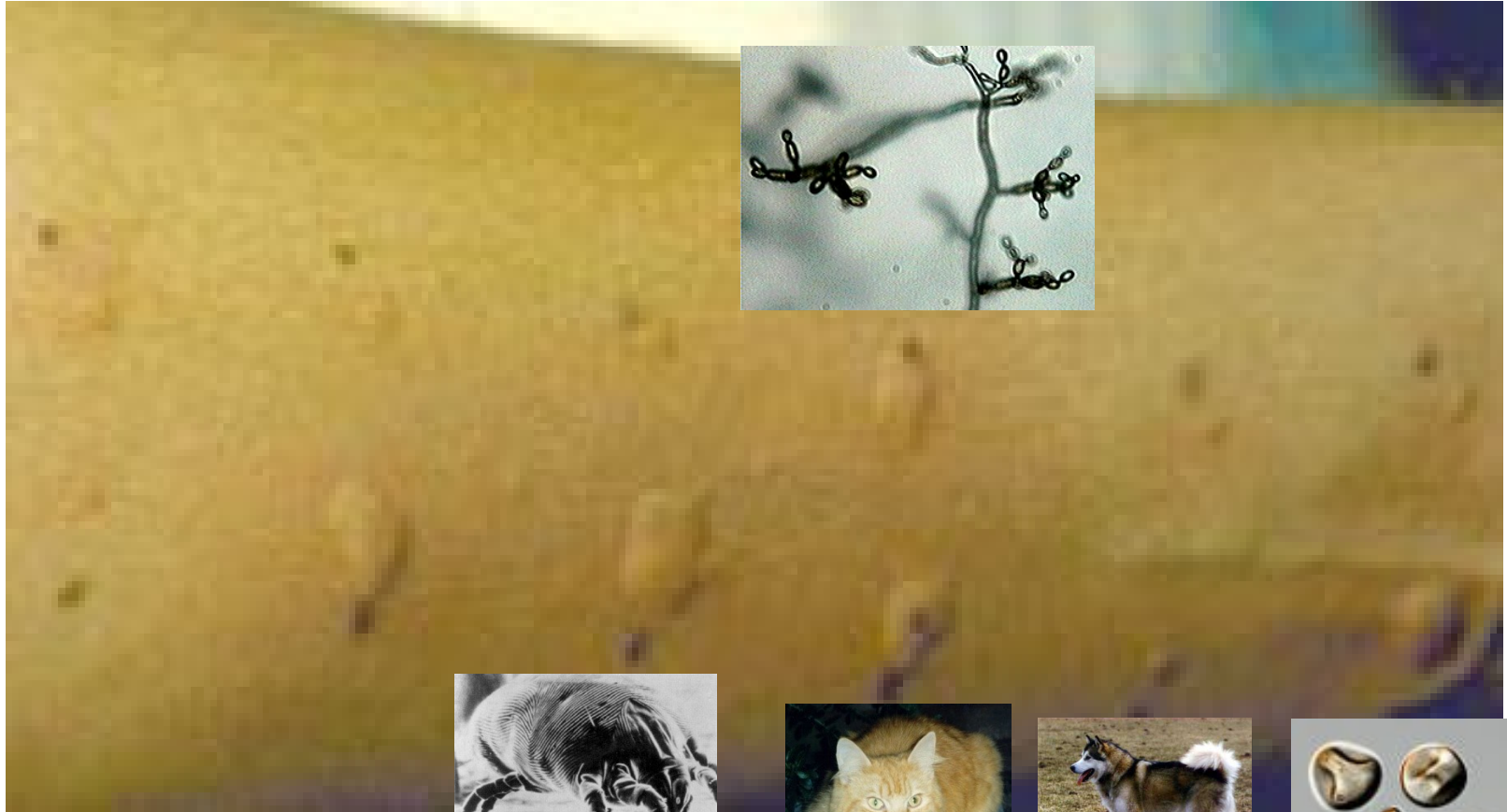


Rhizopus



Penicillium





# AIR QUALITY IN HOSPITALS

- Nosocomial pulmonary Aspergillosis
- > 500 cases of POST-OPARATIVE ASPERGILLOSIS



# Fungal sinusitis

# sinusitis after chemotherapy biopsy showed hyphal invasion



# Management of acute invasive *Aspergillus* sinusitis

diagnosis Requires both biopsy and preferably culture for mucormycosis, *Scedopsporium*– differential diagnosis = infection/*Fusarium*

tissue Requires systemic antifungal therapy to minimize eye, mouth and destruction, including spread to face, brain and cure

? Requires surgical removal

# Allergic Aspergillus sinusitis

Clinical features = nasal obstruction, recurrent sinus infections, loss of smell and nasal polyps

Aspergillus precipitins +ve in 85%

Surgical handling of specimen very important – mucus versus tissue: allergic or chronic invasive

# Chronic invasive *Aspergillus* sinusitis

Complications in immunocompromised:

- orbital apex syndrome
- generalised proptosis and blindness
- cavernous sinus thrombosis
- osteomyelitis of the base of the skull
  - cerebral aspergillosis



# Diagnosis



## **:Specimen** ■

,Respiratory specimens: Sputum, BAL, Lung biopsy ■

(Other samples (depend on the site ■

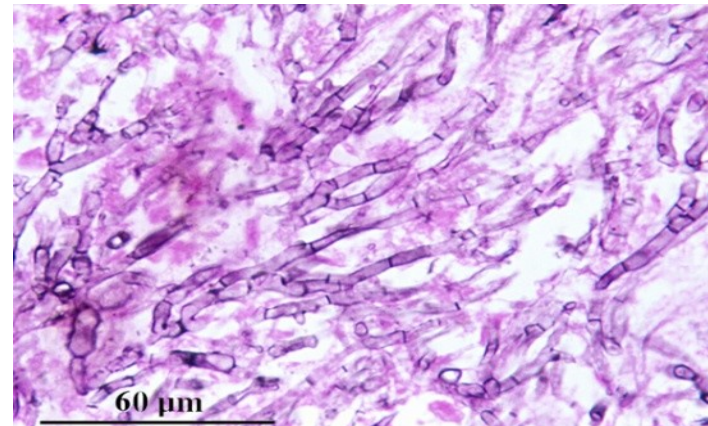
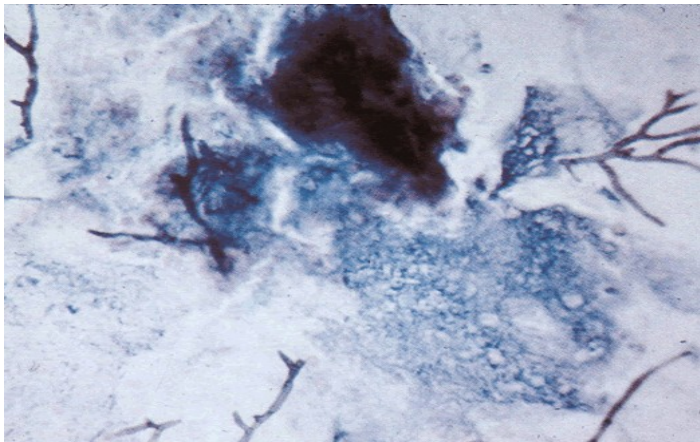
Blood ■

## **:Lab. Investigations**

### **:Direct Microscopy** ■

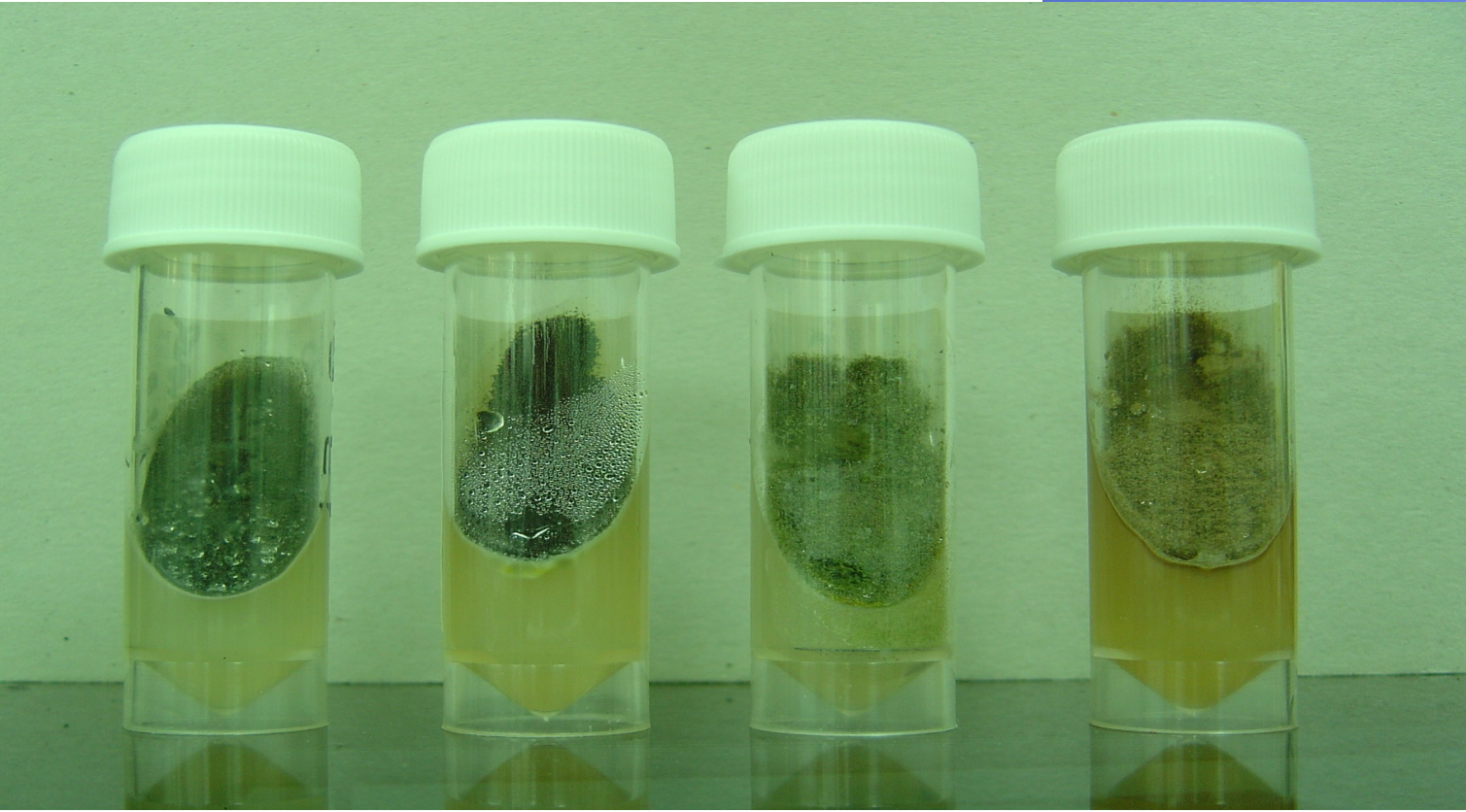
Stained smear : Periodic Acid Schiff (P.A.S); KOH, Giemsa, Greccott  
(methenamine silver stain (GMS

will show septate fungal hyphae with Dichotomous branching



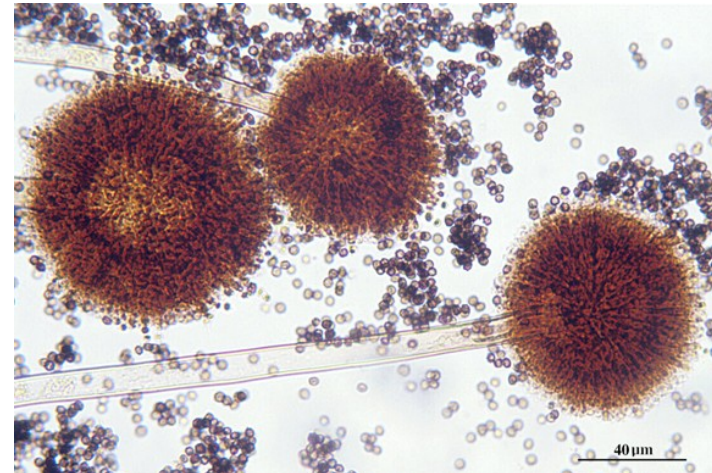
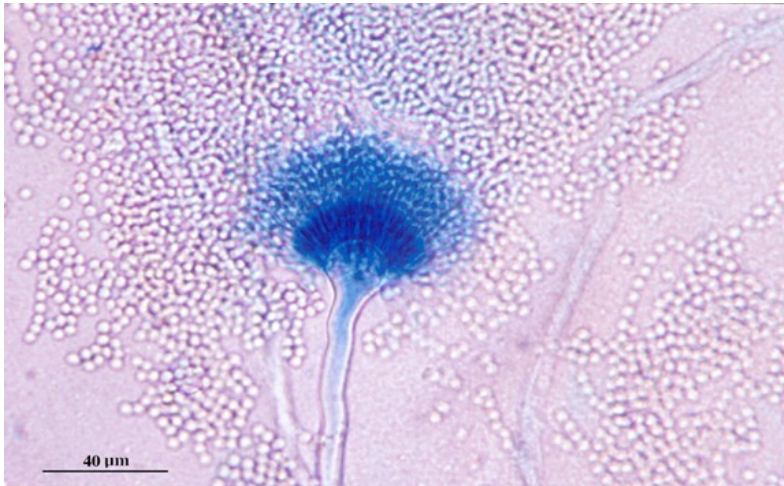
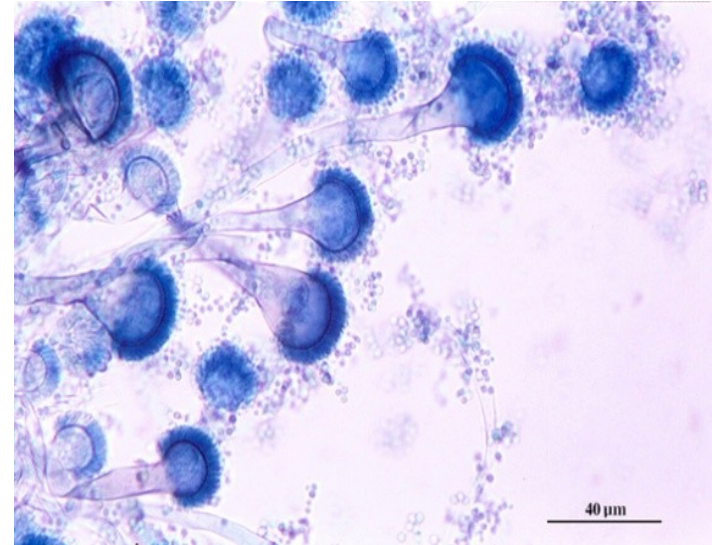
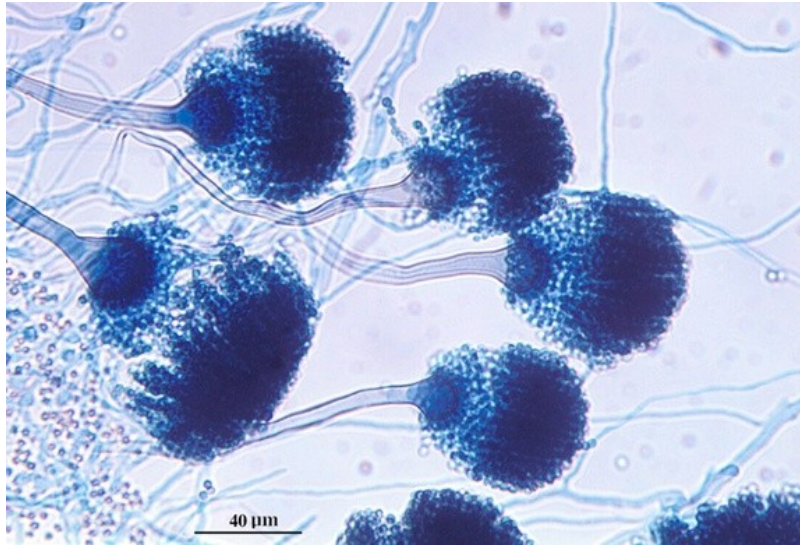
**:Culture** ■

(on SDA (no cycloheximide) ■





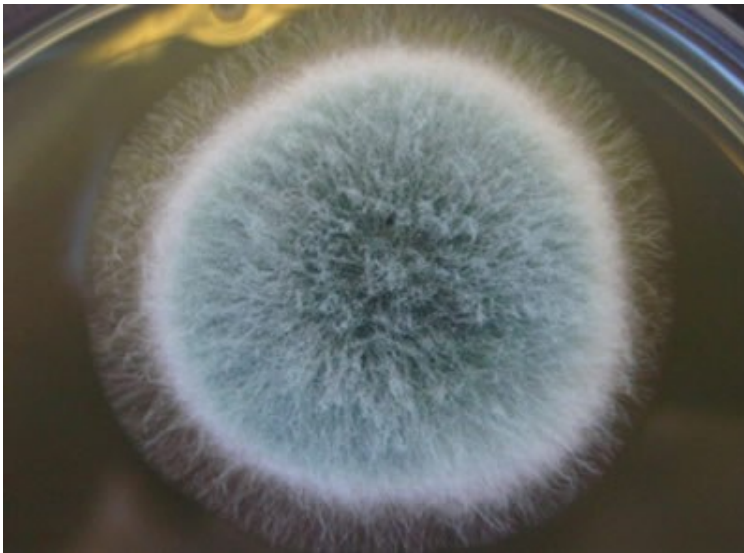
# Microscopy



LPCB preparations

# Cultures for *Aspergillus* from sputum and BAL

Yield in Invasive Aspergillosis from BAL and sputum ~40%



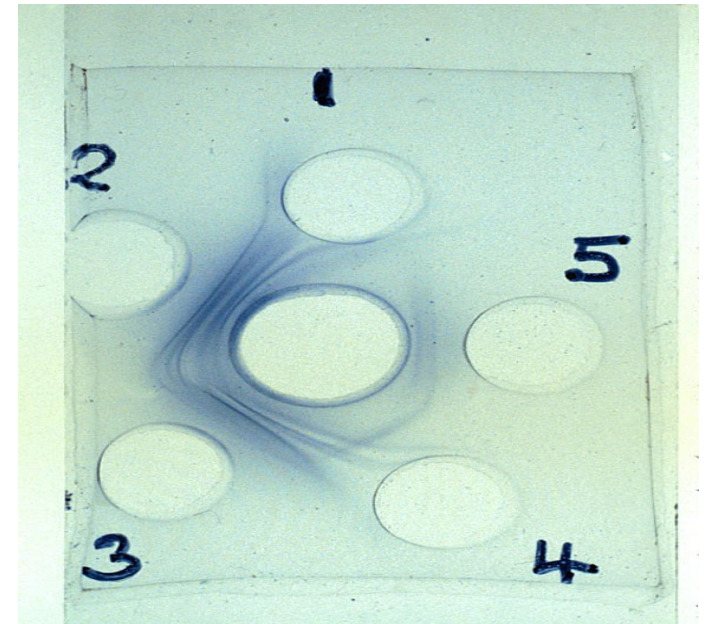
Bacteriological media inferior to fungal media – 32% higher yield on fungal media

# Diagnosis

## :Serology

Test for Antibody ■

(Using I.D (Immunodiffusion



Immunodiffusion

## Test for Antigen

ELISA test for galactomannan Antigen is available with a better sensitivity



# Diagnosis- PCR

**MycAssay™: Aspergillus**

**Real-time molecular based *in vitro* diagnostic tests for *Aspergillus* spp.**



# Choice of antifungal for aspergillosis

(Voriconazole (unless drug interaction

(Amphotericin B (AmBisome

OR

,Caspofungin

(Posaconazole (oral only

# Zygomycosis

- Pulmonary zygomycosis
- Rhinocerebral zygomycosis

## Risk factors

Diabetic ketoacidosis

Granulocytopenia

Corticosteroid therapy

Malignancy

HSCT

Many others



# Zygomycosis

## **Etiology:**

Zygomycetes

Non-septate hyphae

Mainly of the order Mucorales (Rarely Entomophthorales)

e.g. Rhizopus, Mucor, Absidia

- Angioinvasion, Thrombotic invasion of blood vessels
- Pulmonary infarctions and hemorrhage
- Rapid evolving clinical course
- High mortality

# Zygomycosis

Acute

Fever, pulmonary infiltrates refractory to antibacterial therapy.

Consolidation , nodules, cavitation, pleural effusion, hemoptysis

Infection may extend to chest wall, diaphragm, pericardium.

Early recognition and intervention are critical

# Zygomycosis

## Diagnosis

**:Specimen** ■

,Respiratory specimens: Sputum, BAL, Lung biopsy ■

:Other samples ■

## **:Lab. Investigations**

**:Direct Microscopy** ■

Periodic Acid Schiff (P.A.S); KOH, Giemsa, Grecoth methenamine silver (stain (GMS

**will show broad non- septate fungal hyphae**

**Culture:** (on SDA (no cycloheximide ■

**No Serology** available ■

# Zygomycosis

## Treatment:

Amphotericin B ,

Posaconazole (Other azoles are not effective)

?Surgery



**Thank you**