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GLOBAL EDUCATION



Bacteriology of Tuberculosis

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Outline

1. *Mycobacterium tuberculosis* complex
2. Classification of mycobacteria within the genus
3. Bacteriological characteristics of tuberculosis
4. TB Diagnostic tools

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Mycobacterium tuberculosis Complex (MTBC)

Robert Koch identified the organism of the MTBC

- *Mycobacterium tuberculosis*
 - *Mycobacterium bovis*
 - *Mycobacterium bovis* BCG
 - *Mycobacterium microti*
 - *Mycobacterium africanum*
-
- ➔ Acid-fast organisms
 - ➔ Slow growing organisms
 - ➔ Only one group in the Genus *Mycobacteria*
 - ➔ Major public health implications



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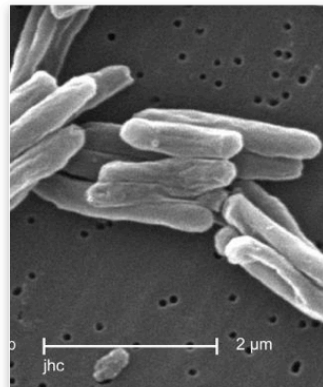
Genus Mycobacteria

Almost 100 different species

- 50 have potential for disease

Runyon classification system

- morphology
- growth rate
- pigmentation



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Nontuberculous mycobacteria

Rapidly growing: < 7 days	Intermediately growing: 7-10 days	Slowly growing: > 7 days on agar
<i>M. fortuitum</i> complex	<i>M. marinum</i> – pigmented 30°C	<i>M. avium</i> complex
(<i>M. fortuitum</i> , <i>M. peregrinum</i> , <i>M. fortuitum</i> third biovariant complex, <i>M. mucogenicum</i>)	<i>M. goodii</i> - pigmented 35°C	(<i>M. avium</i> , <i>M. intracellulare</i> , <i>M. avium</i> complex "xvar") <i>M. kansasii</i> , <i>M. xenopi</i> , <i>M. simiae</i> , <i>M. szulgai</i> , <i>M. malmoense</i> , <i>M. terrae</i> / <i>M. nonchromogenicum</i> complex, <i>M. haemophilum</i> , <i>M. genavense</i>
<i>M. chelonae</i> / abscessus group		Others:
(<i>M. chelonae</i> , <i>M. abscessus</i> , <i>M. immunogenium</i>)		<i>M. interjectum</i> , <i>M. confluentens</i> , <i>M. lentiflavum</i> etc
<i>M. smegmatis</i> – pigmented / nonpigmented		
Others pigmented: (<i>M. phlei</i> , <i>M. aurum</i> , <i>M. flavescens</i> , <i>M. neoaurum</i> , <i>M. vaccae</i> , <i>M. thermoresistibile</i>)		

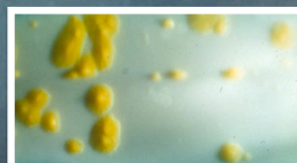
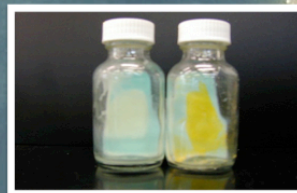
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TB Bacteriology

Growth rates and pigmentation

Runyon groups

- ✓ slowly growing *photochromogens*
- ✓ slowly growing *scotochromogens*
- ✓ slowly growing non-chromogens
- ✓ rapid growers



TB Bacteriology

Clinical classification

Human pathogens

- ✓ *M. tuberculosis*, *M. leprae*, *M. ulcerans*

Animal pathogens

- ✓ *M. bovis*, *M. avium*, *M. marinum*

Environmental mycobacteria

- free living in water or soil
- Classification is not strict
- *M. tuberculosis* infection is acquired primarily by person to person transmission
- Infections with non-tuberculous mycobacteria (NTM) are acquired from the environment and are not transmitted person to person

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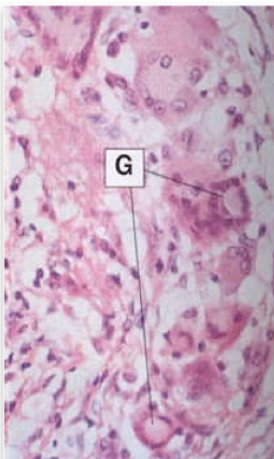
Tuberculosis transmission and disease (I)

- MTB an obligate pathogen, requires host to survive
- Inhalation into alveolar spaces, ingestion by macrophages
- Development or elimination of disease depends on microbiocidal activity
- Degenerated macrophages sensitize local lymphocytes and mobilise more macrophages.
- Dynamic turnover of engulfment and degeneration



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Tuberculosis transmission and disease (II)



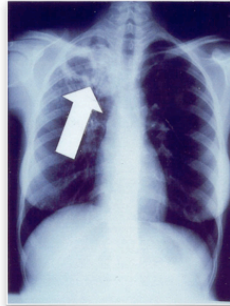
- Zone of **lymphocytes** and **mononuclears** surround centre of necrosis, development of capsule of fibrous connective tissue
- Extensiveness of disease due to virulence, route of infection, stage of infection, host factors

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TB Bacteriology

Oxygen dependence

- TB bacilli require oxygen
- Prefer top of lung or cavities where air circulation is best for multiplication
- TB bacilli can survive in a metabolically inactive form or "dormant" in the host
- Dormant bacilli can reactivate to cause active disease after years or decades and especially with deterioration of the host immune system



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Probability TB will be transmitted

- Infectiousness of person with TB
- Environment in which exposure occurred
- Susceptibility of the host
- Duration of exposure
- Virulence of the organism

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Disease progression

- Only some individuals infected will develop active disease
- Approx. 10% of infected persons with normal immune systems develop TB at some point in life
- HIV strongest risk factor for development of TB
- Risk of developing TB disease among persons with HIV is 7% - 10% each year
- Certain medical conditions increase risk that TB infection will progress to TB disease

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Conditions that increase the risk of progression to TB disease

- HIV infection
- Substance abuse
- Recent infection
- Diabetes mellitus
- Silicosis
- Immunosuppression (corticosteroids and other drugs, malnutrition)

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Cellular characteristics

Cell wall composition

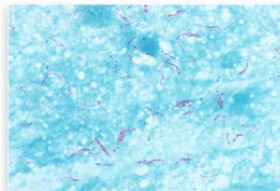
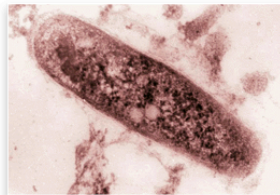
- layer of mycolic acid
- heat and phenol needed for the primary stain to penetrate the cell wall

Acid fast bacilli - AFB

- all mycobacteria are AFB (i.e resistant to decolorization with acid or acid alcohol)
- lack of specificity
- resistance to chemicals inc. disinfectants acids, alkali, detergents

Long generation time

- chronicity of disease



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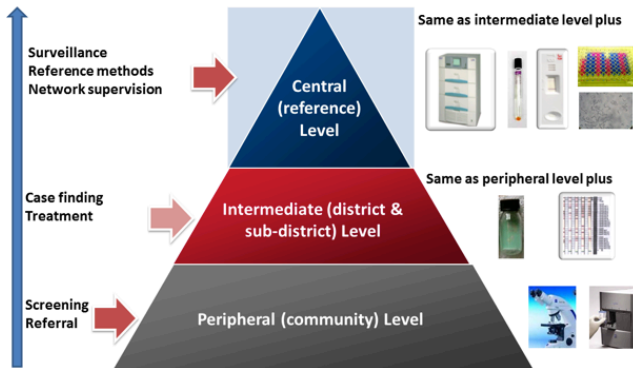
Resistance characteristics

Natural resistance to common antibiotics

- ✓ Inherent resistance to penicillins
- ✓ Intracellular bacteria are protected by the acid environment of macrophages
- ✓ Spontaneous mutations each million – 100 million divisions
- ✓ High selection pressure once mutants occur especially for the key mycobactericidal drugs (rifampicin and isoniazid)
- ✓ Exposure to a single drug
 - poor adherence to treatment / inappropriate prescription
 - irregular drug supply / poor quality

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WHO recommended diagnostics for use at different levels of laboratory sophistication



Available at: www.who.int/tb/dots/laboratory/policy/en WorldMEDSchool

Thank you!

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