

Infective Disorders Pathophysiology



Pharmacology 2



MENINGITIS PATHOPHYSIOLOGY

Meningitis is an acute inflammation of the protective covering of the CNS (brain & spinal cord), that known as meninges

→ Inflammation caused by pathogenic infection.

Types -

1) Viral Meningitis - Enterovirus category - 85% cases

↳ Coxsackie virus A & B, Echo virus

↳ other - Influenza, mumps, HIV, measles, herpes

2) Bacterial Meningitis → Contagious & fatal

→ S. pneumoniae - most common in infant & children, found in R.Tract, Sinus, & nasal cavity, Prevented by vaccine

→ N. meningitidis - highly contagious, common in upper R.tract infectn, after reach to blood Stream it can cause meningitis. "Vaccine"

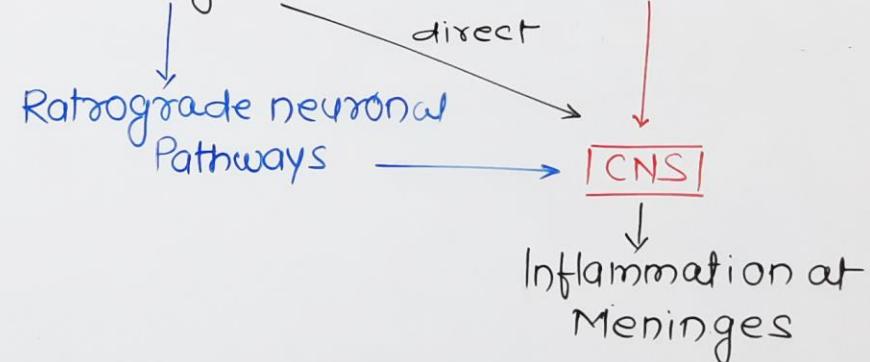
→ H. influenzae - type B - (HIB) - children

→ L. monocytogenes - found in unpasteurized cheeses, hot dog, & meats. Pregnant woman, newborn & older person.

3) Others → Fungal meningitis (Cryptococcal meningitis)
Mycobacterium tuberculosis

Symptoms - Fever, headache, stiff neck, confusion, seizure, No appetite or thirst, skin rashes, in baby - fever, crying, inactivity, poor feeding, flu like.

Pathology - Pathogens → Invasion of blood Stream



Treatments -

Bac. Meningitis - Antibiotics & Steroids

Viral Meningitis - Symptomatic Treatment

Fungal Meningitis - Antifungal Drugs

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Website/Notes

Typhoid Pathophysiology

Typhoid is a bacterial infection disease caused by "Salmonella typhimurium" which lives in intestine & blood. & spread through feces.

Transmission -

1. Fecal-Oral Transmission Route -

Infected person → Feces/Urine → Water/Food

↓
Spread Infection ← Taken by normal

2. Typhoid Carriers → Some people carries the bacteria in their intestine, gallbladder, often for years.

Symptoms →

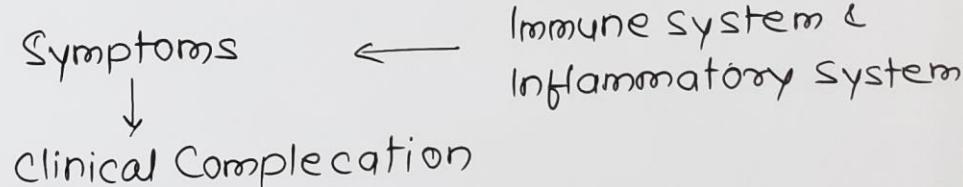
↳ Early → Fever, headache, weakness, fatigue, sweating, dry cough, abdominal pain, diarrhoea, rashes

↳ Later → Lie motionless, exhausted with your eyes half closed

Diagnosis → Detectn of S.typhi via blood, stool, urine

Pathophysiology

Infection → Intestine absorptn → blood stream
↓ +



Treatments - Antibiotics & Supporting therapy

↳ Antibiotic/AMAs → Ciprofloxacin, Azithromycin, Ceftriaxone, Co-trimoxazole, Ampicilline, Cefotaxime, etc

↳ Supporting Therapy - Fluids, NSAIDs, Antigastric, Multivitamins



LEPROSY PATHOPHYSIOLOGY - HANSEN'S DISEASE

- # Leprosy, caused by "Mycobacterium leprae"
 - # It is infective dis. that cause sever, disfiguring skin sores & nerve damage in the arms & legs
 - # It has been considered incurable since ages & bears a social stigma. Now availability of drugs it is curable but deformities may not be reversed
- TYPES : → Based on Ridley-Jopling system, There are six categories (based on severity of symptoms)
1. Intermediate : - A few flat lesions that sometimes heal by themselves & may progress to more severe
 2. Tuberculoid : - Some large lesions, nerve damage can be healed itself & may progressive (T)
 3. Borderline Tuberculoid : - lesions are similar to tuberculoid but smaller & numerous; less nerve enlargement; persist; revert to tuberculoid or advance to another form (BT)
 4. Mid Borderline leprosy : - Redish plaques, moderate numbness, swollen lymph glands, may regress, persist, or progress to other forms. (BB)
 5. Borderline lepromatous : - Several types of lesions, persist, regress or progressive. (BL)
 6. Lepromatous leprosy - Many lesions with bacteria, hair loss, nerve involvement, limb weakness, disfigurements, doesn't regress (LL)

SYMPTOMS : - It is primarily a granulomatous disease of the peripheral nerves and mucosa of the upper respiratory tract.

- # Skin lesions - light & dark patches - External Sign
- # Progress → Permanent damage to Skin, Nerve Limbs, Eyes

Secondary Infection → Tissue loss, shortened and deformed toes & fingers.

CLINICAL SIGNS & FEATURES

- # Hypopigmented/Erythematous Skin Patches
- # Thickened Peripheral Nerve
- # Acid fast bacilli detected on Skin smear or biopsy material

TRANSMISSION : - # Skin, # Nasal mucosa (Resp. Tract)

INCUBATION PERIODS : - Variable (weeks to years)

- # Minimum → Few weeks
- # Maximum → 30 Years
- # Avg. → 3 - 10 years

RISK FACTORS : -

- # Inadequate Bedding
- # Contaminated Water & Foods
- # Insufficient Diet
- # Immunocompromised diseases (AIDS, TB)

TARGET CELL → Schwann Cells & Macrophages



PC

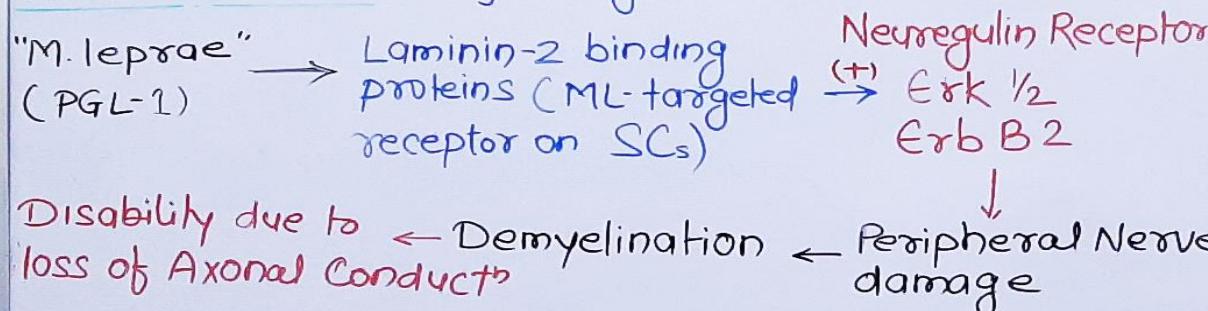
Pharmacology Concepts
By Ajesh Dhadhary

LAPROSY / HANSEN'S DISEASE

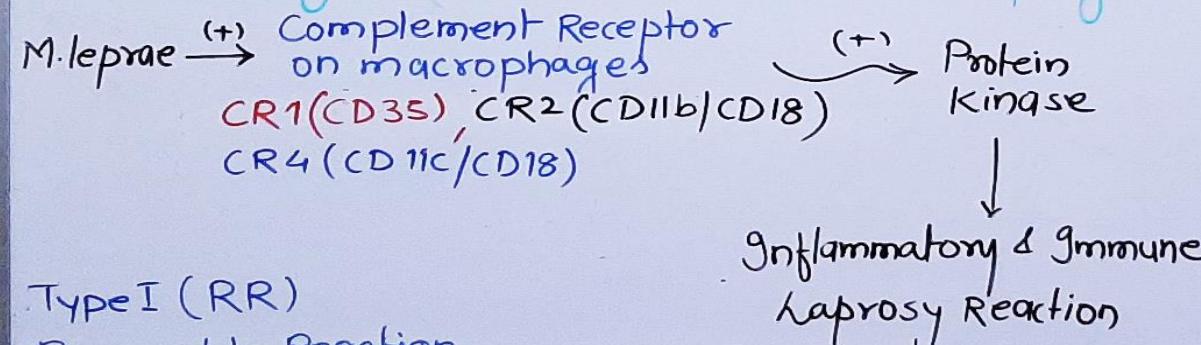
- ↳ *Mycobacterium leprae*, *M. lepromatosis*
- ↳ Tissue → Skin & Nerves
- ↳ G.H. Armaur Hansen - discovered *M. leprae*, 1873

PATHOGENESIS :-

1. **Schwan Cells** → Major Target area



2. **Macrophages (Monocyte derived macrophages)**



Type I (RR) → Delayed type Hypersensitivity React"

→ CD4+ lymphocyte → inflammatory React - skin, nerve
INFγ, TNF-β

Edema, Erythema, neuritis, sensory & motor loss, etc

Type II (ENL) → TNFα, IL-1, INFγ, C-reactive protein

↓
Inflammatory Response

Appearance of tender, Erythematous, nodules on Skin,
Systemic Symptoms - Fever, Enlarged lymph node, Edema,
weight loss, anorexia

TREATMENT = MDT by NLEP (1982)

MBL

Rifampin - 600 mg/month, supervised

600 mg/month, supervised

Dapsone 100 mg/day, self 100 mg/day, self

Clofazimine - 300 mg/month, supervised
+ 50 mg/day, self

PBL

Duration - 12 months 6 months

CHILD - Rifampin - 10 mg/kg/month

Dapsone - 2 mg/kg/day

Clofazimine - 1 mg/kg/day +
6 mg/kg/month

TUBERCULOSIS

TB is an infective disease caused by "Mycobacterium tuberculosis", & that most often affect the lungs.

* Symptoms → Coughing (with or without mucus & blood), chills, fever, fatigue, weight loss, appetite loss,

* Contagious → Spread through air by cough, sneeze or spit

* Risk Factors - Compromised immune system, HIV, Diabetes, starvation, Malnutrition, Smokers.

* Type of TB Forms -

1) Latent TB - Your immune system stops the spreading and inactivate them but still alive in your body & can become active.

2. Active TB - Microbes can multiply, spread & cause sick.

* Clinical Manifestation → spinal pain & joint destruction

↳ Infecting to brain cause **Meningitis**

↳ Infecting to Liver & kidney → functional loss

↳ Infecting to heart → Cardiac Tamponade

PATHOPHYSIOLOGY -

M. tuberculosis → Lung Tissue → Multiplicatⁿ (1000 - 10000)

Inflammatory Response ← + Immune Response

"Pulmonary Complication" → Spread to other organ

↓ Other Tissue Damage

Treatment → Specific dose regimen of diff. AMAs

↳ 1st line drugs - Isoniazid (H), Ethambutol (E), Rifampin (R), Pyrazinamide (Z), Streptomycin (S)

↳ 2nd line drug → Quinolones, P-amino Salicylate, etc

* Regimen based on RNTCP guideline 2016

New Patient = HRZE (2months) + HRE (4 month) = 6 months

Previously treated: HRZES(2) + HRZE (1) + HRE (5) = 8 months

Daily Single dose (mg/kg): H - 5 mg/kg, R - 10 mg/kg, Z - 25 mg/kg
E - 15 mg/kg, S - 15 mg/kg

URINARY TRACT INFECTION

UTI is an infection in any part of urinary system - kidneys, ureters, bladder, and urethra.

↳ Most of UTI occurs in lower urinary tract (urethra, and bladder)

↳ Urethra → Urethritis

↳ Kidneys - Pyelonephritis

↳ Bladder → Cystitis

* Women are at greater risk than men

Pathogen → "E. coli" - 90% cases

Symptom → ↳ Painful urination, burning sensation

↳ Strong, persistent urge to urination

↳ Pain at genital & pelvic region

↳ Cloudy & pink cola like urination

↳ Strong smell urine

↳ Fever, Vomiting

Risk Factors - ↳ Diabetes

↳ Weak Immune System

↳ Prostate hypertrophy

↳ Kidney stones

Diagnosis - Urine culture for bacteria

Urine analysis - Blood cells

CT Scan, ultrasound,

Cystoscopy

Treatments

↳ Urinary antiseptics - Nitrofurantoin

↳ Urinary Analgesics - Phenazopyridine

↳ AMAs & Antibiotics

Empirical AMAs (3-5 days) oral dose For Acute UTI

1. Ciprofloxacin → 250-500 mg/12h

2. Ofloxacin → 200-400 mg/12h

3. Cotrimoxazole → 960 mg/12h

4. Cephalexime → 250-500 mg/6h

5. Cefpodoxime → 200 mg/12h

6. Amoxycilline + Clavulanic acid

→ 500 mg + 125 mg / 8 h

7. Nitrofurantoin → 50 mg/8h or 100mg/12h

* For upper UTI (pyelonephritis) - same drugs can be given for 2 weeks except Nitrofurantoin

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Website/Notes

AIDS PATHOPHYSIOLOGY

AIDS - Acquired Immunodeficiency Syndrome

HIV - Human Immunodeficiency Virus

Virus → Retro Virus (RNA Virus)

Affected System → "Immune System"

Transmission - Unprotected Sex (STD), Placental, blood transfusion.

Symptoms - fever, chills, sore throat, joint pain, body aches, tiredness, weakness, weight loss.

* Late stage infection - Blurred vision, fever (100 F) dry cough, night sweat, shortness of breath, weight loss, esophagitis, infection on CNS, pneumonia, toxoplasmosis, TB, etc.

Treatment -

① Rev. Transcriptase inhibitors - Zidovudine, Lamivudine, Efavirenz, Nevirapine

② Protease Inhibitors - Indinavir, Ritonavir

③ Integrase Inhibitors - Raltegravir

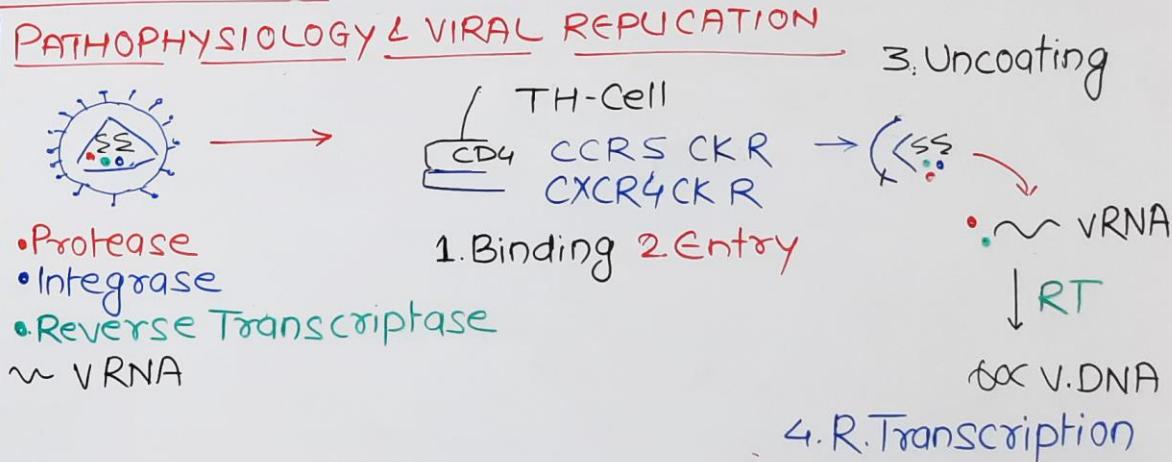
④ Vaccine

⑤ Entry blocker - Enfuvirtide, maraviroc

PATHOPHYSIOLOGY & VIRAL REPLICATION



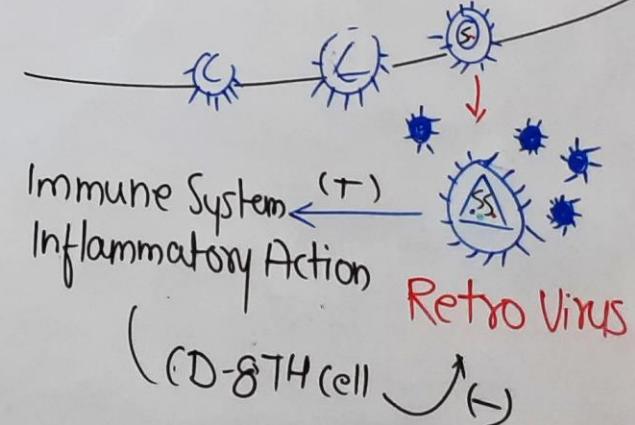
- Protease
 - Integrase
 - Reverse Transcriptase
- ~ vRNA



* ↓ CD4 count < 350

↓ Weaker the immunity

↑ Risk of 2^o infection



SYPHILIS

Syphilis is a bacterial infection, Sexually transmitted disease, caused by "*Treponema pallidum*".

Stages of syphilis -

- 1) Primary] most 3) Latent - No symp. & contagious
- 2) Secondary] infectious 4) Tertiary - most destructive

1) Primary - after 3-4 weeks of infection, small & round sore (chancre), painless but infectious

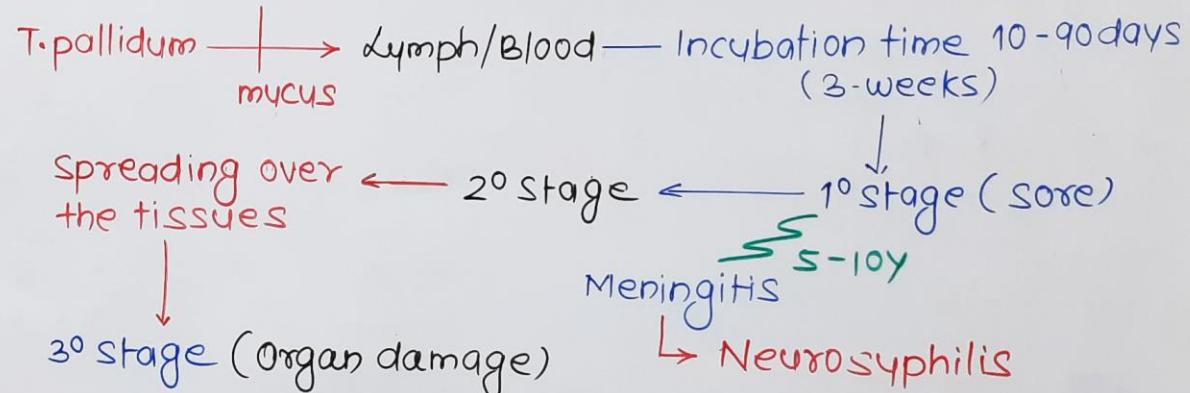
2) Secondary - Skin rash & Sore throat. Symptoms:- headache, fatigue, swollen lymph glands, fever, weight loss, hairfall, Joint pain

3) Latent → latent or hidden stage, No symptoms as 1^o & 2^o, but remain active for years, if no treatment progress to 3^o phase

4) Tertiary - late/last stage of Syphilis infection (15-30 % patients, without treatment), it can occurs years or decades after infections.
Life threatening clinical complications - blindness, deafness, mental illness (dementia), destruction of soft tissues & bone, Heart disease, neurosyphilis, etc.

Congenital Syphilis - Baby get infected from mother, symptoms - rash, deafness, teeth deformities, etc.

Pathophysiology - spread through unprotected unsafe sexual activity, blood transfusion, mother to baby, or direct contact with the infected lesions.



Delayed type hypersensitive reaction

Treatments

Early Infection → Penicillin G (1st choice), Doxycycline, tetracycline, Rifampin, Clindamycin, Ceftriaxone (Alternative drugs)

Late infection - Neurosyphilis (PenG, IV, large dose, 10 days)
- Ceftriaxone, Doxycycline, tetracycline → long duration

Pregnancy - Benzathine Penicillin-G (1m-2.4M unit)



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Website/Notes

GONORRHOEA

Gonorrhoea is a sexually transmitted disease, caused by bacterial infection (*Neisseria gonorrhoeae*)

Symptoms - Men - burning with urination, discharge from penis, testicular pain

Women - burning with urination, vaginal discharge, vaginal bleeding during periods, pelvis pain & pelvis inflammation

Transmission - Sexual contact with infected person, placental transmission,

Affected site - # Genital Organ

Rectum - Itching, pus discharge, bleeding

Eye - Pain, discharge

Throat - sore throat, swollen lymph

Joint - Pain, warm, swelling - (Inflammation)

Pathophysiology

N. gonorrhoeae, a gram -ve bacteria having pili which protect from phagocytic action of neutrophil and IgA protease enzyme which digest the IgA that present in urethra, fellopiant tube, and endo cervix allowing attachment to these surface.

Bacteria → Replicate & Grow at Site of Infectn → Inflammatory Reaction

Gonorrhoeal Complication ← Fibrosis (Fibrotic React^b)

Treatments - Antimicrobial Agents

↳ Amoxicillin/Ampicillin + Probenecid

↳ Ceftriaxone, Cefuroxime, azithromycin + Probenecid