

Principles of Anti-microbial Therapy



Antimicrobial drugs

An antimicrobial drug is any substance of natural, semisynthetic, or synthetic origin either to **kills** or **inhibits** the growth of a microorganism, BUT causes little or no host damage.....

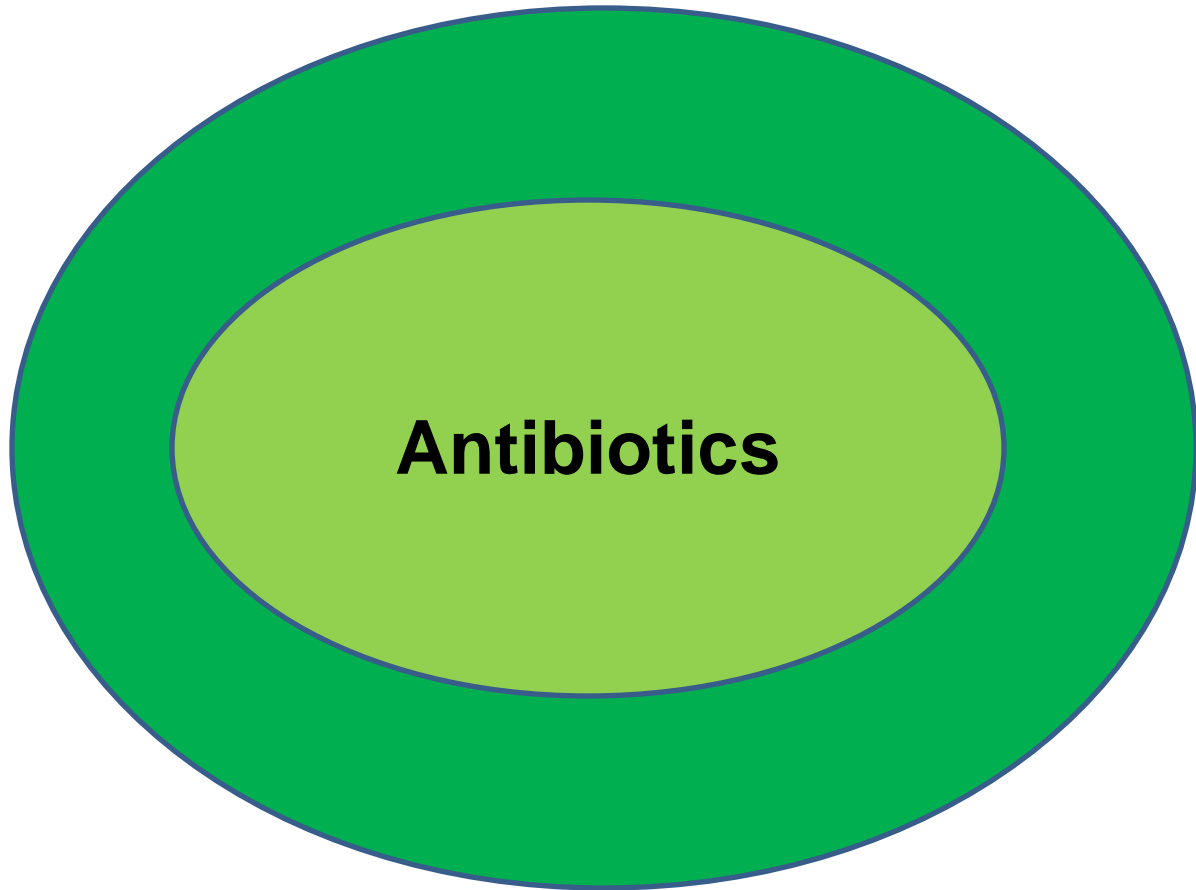
Ex:

(Quinolones, Sulfonamides)

Antibiotics (ATB)

- Substances produced from certain of microorganisms (M.O) either bacteria or fungi to act against other M.O.
- To suppress (inhibit) the growth of M.O or to destroy (kill) them.

Ex:- (penicillins, tetracyclines)



Antibiotics

❖ The most imp thing that these drugs should do no harm to the host cell.

❖ (Highly selectivity drugs against M.O rather than human cells)

❖ From where did this selectivity of these drugs come????

Differences between bacteria & human cells

Bacterial cells	Mammalian cells
Cell wall	No cell wall
Small Ribosome (70s)	Large Ribosome (80s)
Folic acid come from :- Conversion of Para AminoBenzoic Acid (PABA) to folate	Folic acid come from Dietary folate

- the selective toxicity is relative rather than absolute, requiring that the concentration of the drug be carefully controlled to attack the M.O while still being tolerated by the host



Classification of antimicrobial agents

1. According to the **CAUSTIVE AGENTS** (The type of organism) against which they are active.

(bacteria, virus, fungi) ` either (antibacterial, antiviral, antifungal & so on)

2. According to their **structure** (Macrolides, Aminoglycosides, Tetracyclines)

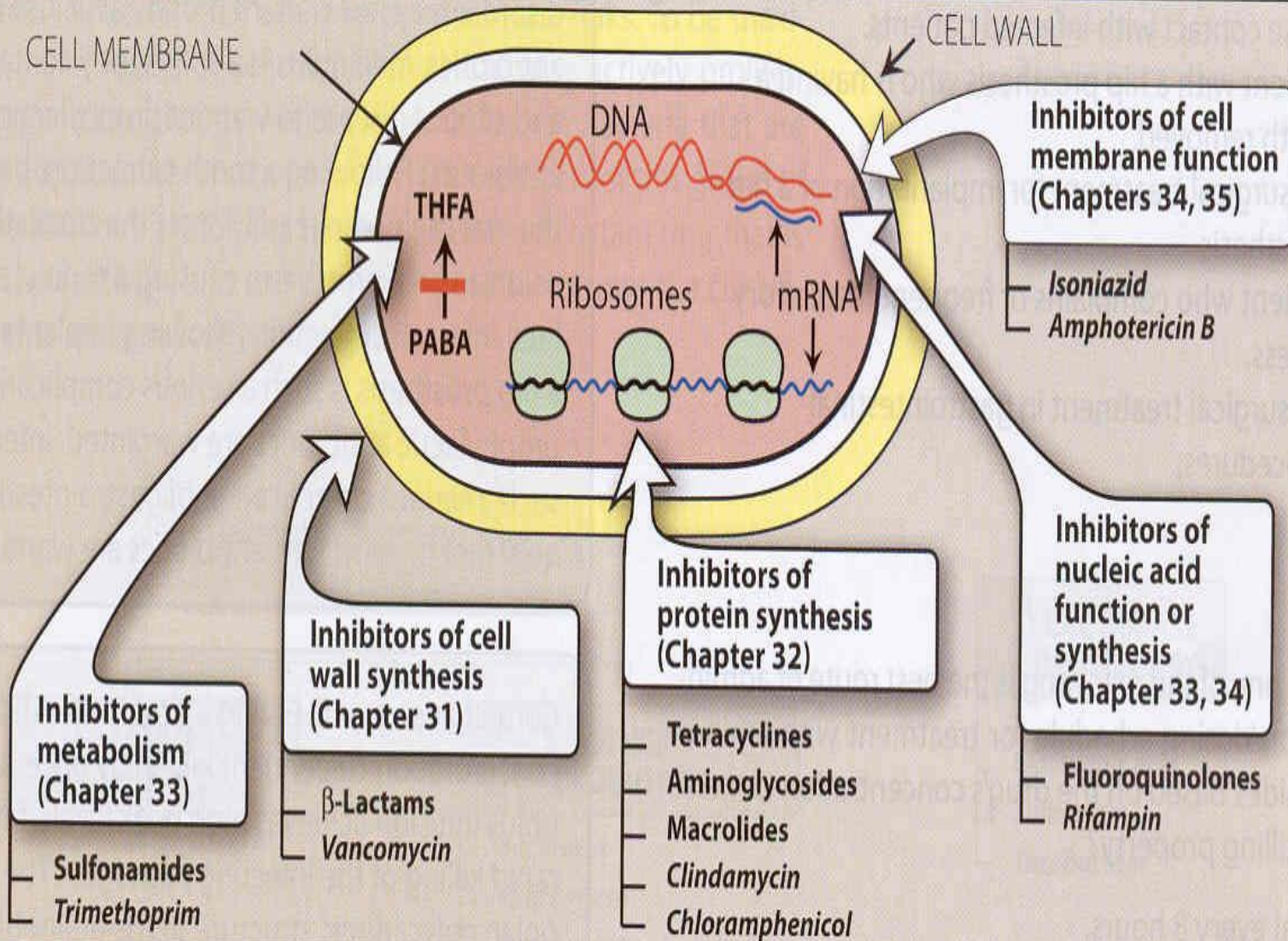
3. According to their **mechanism of action**:

- a. Inhibition of cell wall synthesis
(e.g. **Penicillins, Cephalosporins**).
- b. Inhibition of protein synthesis
(e.g. **Tetracyclines, Macrolides**).
- c. Inhibition of nucleic acid synthesis or function
(e.g. **Fluoroquinolones**)
- d. Inhibition of metabolism
(e.g. **Sulfonamides**)
- e. Inhibition of cell membrane function
(e.g. **Isoniazid**)

Classification of some antimicrobial agent by their site of action..

CELL MEMBRANE

CELL WALL



THFA

PABA

DNA

mRNA

Ribosomes

Inhibitors of metabolism (Chapter 33)

- Sulfonamides
- Trimethoprim

Inhibitors of cell wall synthesis (Chapter 31)

- β -Lactams
- Vancomycin

Inhibitors of protein synthesis (Chapter 32)

- Tetracyclines
- Aminoglycosides
- Macrolides
- Clindamycin
- Chloramphenicol

Inhibitors of cell membrane function (Chapters 34, 35)

- Isoniazid
- Amphotericin B

Inhibitors of nucleic acid function or synthesis (Chapter 33, 34)

- Fluoroquinolones
- Rifampin

4- According to the Spectrum of activity

Spectrum: the range of bacteria that a drug is effective against.

1. **Narrow spectrum** antibiotic: active against single or limited # of M.O. (ex Isoniazid)
2. **Extended spectrum** : is one that, as a result of chemical modification, affects additional types of bacteria, usually those that **are gram-negative**. active against types of bacteria G- & G+.
(ex Ampicillin)
3. **Broad spectrum**: active against a wide variety of microbial species.
(ex Tetracyclines, quinolones, clindamycin , Chloramphenicol).

❑ Using narrow spectrum effective drug clinically is much better than using broad spectrum.

So, if u know the causative M.O & u know their susceptibility to the drugs it's better to choose the narrowest drug.

Why???

❑ Coz wider spectrum means the drug will affect other M.O ex. Normal flora in respiratory tract, urinary tract and so on and we will get condition called superinfection which is one of complications of using broad spectrum.

5- By their action

1. Bacteriostatic drugs

- ❑ They **arrest/inhibit** (stops does not kill it) the growth & replication of the bacteria.
- ❑ **Given to pts who have good immunity.**

E.x.

Sulphonamides,
Trimethoprim, Tetracyclines,
macrolides, Clindamycin,
Chloramphenicol.

2. Bactericidal drugs

- kills** the bacteria that are rapidly dividing or multiplying.
- more preferable in pts who have low immunity. .(HIV patients, cancer patients, taking steroids)

E.x.

Vancomycin ,B-lactams,
Aminoglycosides,
Rifampicin

Sometimes (not a rule) when you combine two of these bacteriostatic in one drugs, the new drug will become bactericidal.

EXAMPLES:

Chloramphenicol
Erythromycin
Clindamycin
Sulfonamides
Trimethoprim
Tetracyclines



EXAMPLES:

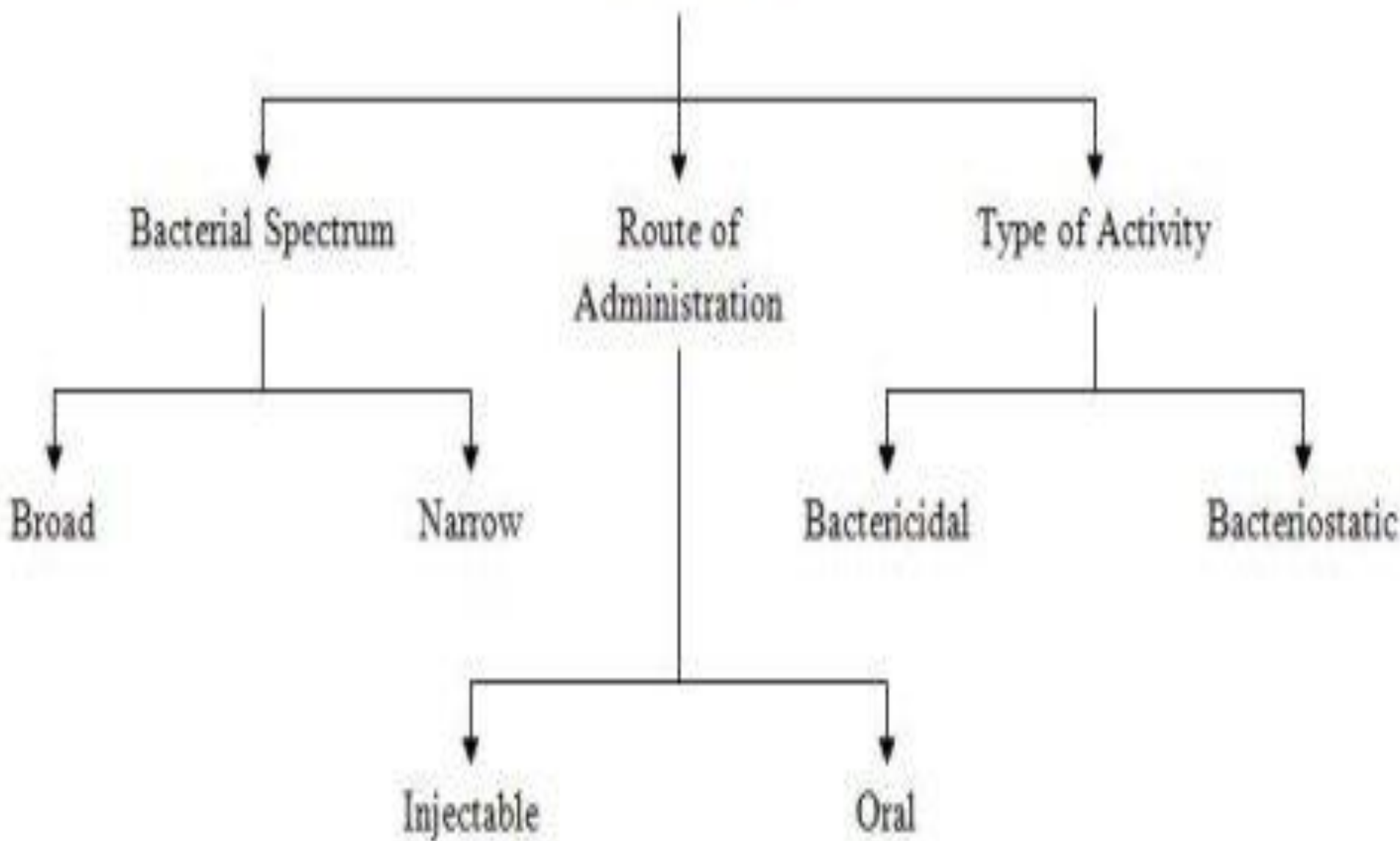
Aminoglycosides
Beta-lactams
Vancomycin
Quinolones
Rifampin
Metronidazole



Factor determine **statics** or **cidal**

- 1.The mechanism of action of the drug. For example, inhibition of cell wall synthesis in the bacteria is most likely to be bactericidal agents.**
- 2. The dose (serum level) of the drug at the site of action.**
- 3.The type of microorganism (sensitivity);if the microorganism is resistant to the drug it will be affected like the drug is bacteriostatic even if it's bactericidal.**

Antibiotics



Antibiotics

Sulfonamides

Sulfamethoxazole

Sulfadiazine

Sulfametho. +
Trimethoprim

Others

Quinolones Fluro-Quinolones

Nalidixic a
(1st Gen.)

Enoxacin

Norfluxacin

Ciprofluxacin

Ofluxacin

Ilevo-Fluxacin

B - Lactam

Penicillin

Ampicillin

Amoxycillin

Cephalosporins

Others

Tetracyclins

Doxycyclin

Others

Macrolides

Erythromycin

Azithromycin

Clarithromycin

Clindamycin

Aminoglycosides

Streptomycin

Neomycin

Gentamycin

Kanamycin

Others

M A S


Selection of Antimicrobial Agents



1- Making the diagnosis :

- ✓ To be sure that the pts is suffering from an bacterial infection.
- ✓ Know the site of infection(GI,RT,UT).
- ✓ Identify the organism.
- ✓ Take the required specimen from the patient.(blood , CSF, mid stream urine ,ear swap, vaginal discharge)

2-Remove the pathological barrier to cure (abscess ,obstruction)



Not my thing,
know what I mean...?

Antibiotics
DON'T WORK
ON COLDS...

... OR MOST COUGHS AND SORE THROATS. **NHS**

3-Select the best drug :So that it reach site of infection in the therapeutic conc.

➤ Drug properties: PK, TI.

➤ **Optimum dose & frequency**

➤ **the most appropriate route of administration**

4- The cost of therapy.

5-Patient factors:

1. Immune system.
2. Renal dysfunction.
3. Hepatic dysfunction.
4. Poor perfusion.
5. Age.
6. Pregnancy.
7. Lactation.
8. Concomitant medication.
9. Allergy.

Antibiotic Therapy

The principle in any infection the first thing you **should do is that you should take a sample for culture & sensitivity test to identify** the causative organism and treatment.

Test :- (Blood ,sputum,urine,pus,etc..)



- The first step is culture and sensitivity but while waiting for the culture and sensitivity test to come back that **minimally takes 48 hrs** unless there is a weekend ,we can't keep the pt without treatment for 4 days ,we treat him with what we call **empiric therapy**....
- After lab result if the organisms were similar to what u have diagnosed u continue with ur treatment..

□ Empiric therapy: is treating the pt without knowing the causative organisms & their **sensitivity** test.

- given BEFORE the lab results.

□ Definitive therapy :

treating exactly the causative agent depending on its **sensitivity** test (done after receiving the results of test)

□ Prophylactic therapy:

Used drugs to prevent an infection rather than to treat , to maintain health and prevent the spread of disease.

Before and after exposure to a disease entity.

Prophylactic antibiotics (chemoprophylaxis)

1. To prevention of opportunistic infection

- ❖ In bowel surgery to prevent peritonitis
- ❖ In dental manipulation to prevent bacterial endocarditis .

2. Prevention of spread among contacts.

- ❖ **Rifampicin** to prevent Meningitis.
- ❖ **Chloroquine** to prevent Malaria.

5- To prevent infection or disease :

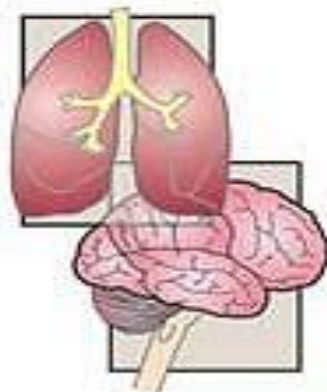
- To prevent recurrent UTI: Co-Trimoxazole twice per week
- to prevent Rheumatic fever : young man who is having recurrent tonsillitis, we start giving him a monthly interval long acting Benzathine penicillin to prevent the acute streptococcal infection from coming back.
- **To prevent hepatic coma -- Neomycin**

1

Prevention of streptococcal infections in patients with a history of rheumatic heart disease. Patients may require years of treatment.

**3**

Prevention of tuberculosis or meningitis among individuals who are in close contact with infected patients.

**2**

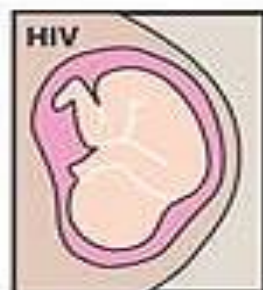
Pretreatment of patients undergoing dental extractions who have implanted prosthetic devices, such as artificial heart valves, to prevent seeding of the prosthesis.

**4**

Treatment prior to certain surgical procedures (such as bowel surgery, joint replacement, and some gynecologic interventions) to prevent infection.

**5**

Treatment of the mother with *zidovudine* to protect the fetus in the case of an HIV-infected, pregnant woman.



Combination of antimicrobials



Advantages

1. To delay or avoid the development of resistance.

(Ex. Tuberculosis)

2. To broaden the spectrum of activity. (Mixed infection, severe unknown infection,).

3. To obtain potentiation (synergistic effect).

B-lactams and aminoglycosides in endocarditis.

Co-trimoxazole.

- The antibacterial effects of all the B-lactam antibiotics are synergistic with the aminoglycosides.
- Because B- lactam facilitate the entry of aminoglycosides (inhibit protein synthesis) to gain access to intracellular target sites.
→ This can result in enhanced antimicrobial activity.
- these drug types **should never be placed in the same infusion fluid**, because on prolonged contact, the positively charged aminoglycosides form an inactive complex with the negatively charged penicillins.

Disadvantages:

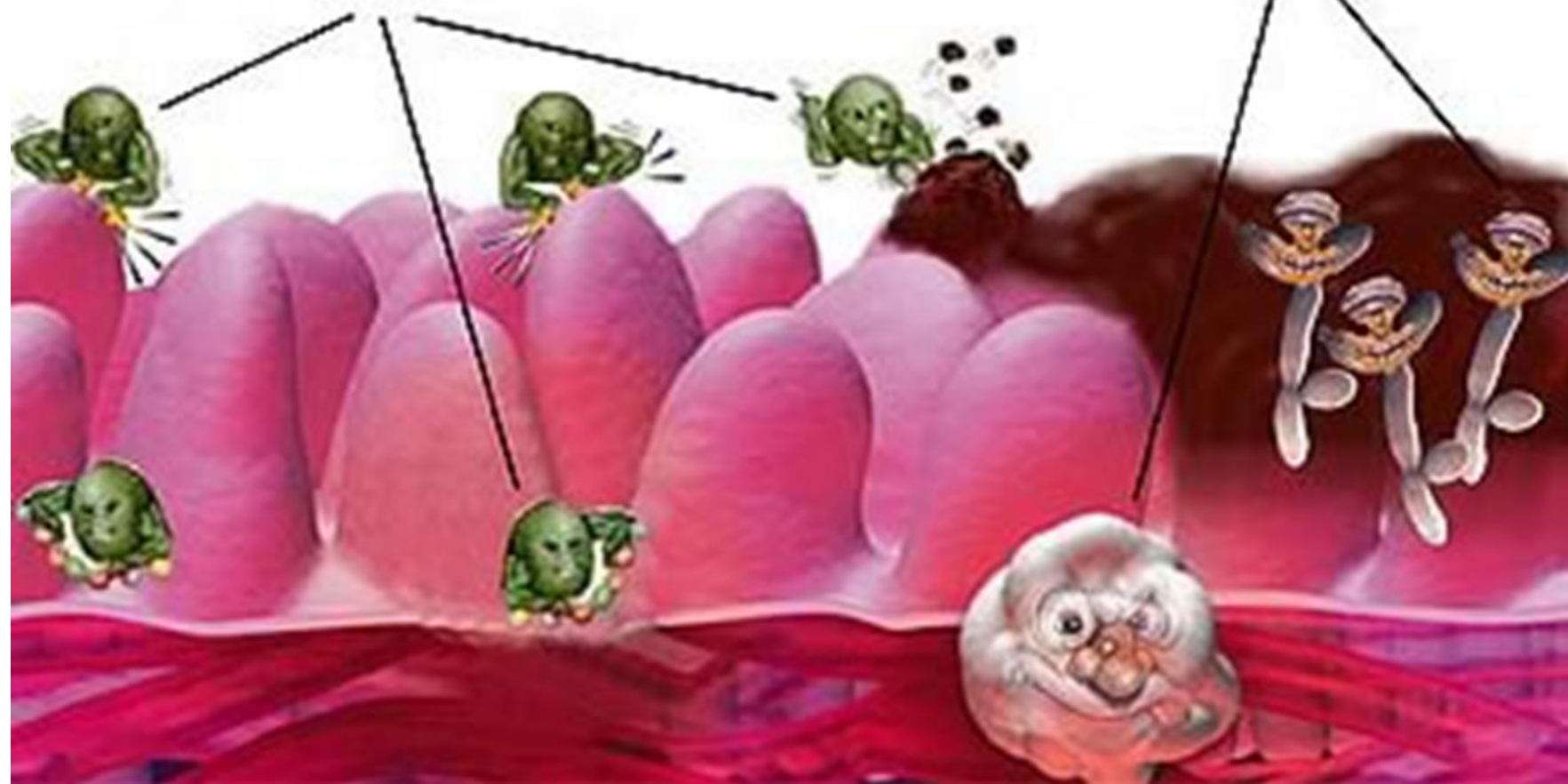
1. Concomitant administration of a second agent is usually bacteriostatic and may interfere with the action of the first drug that is bactericidal
2. Suppression of normal flora, so give higher chance for opportunistic infection (**superinfection**).
3. Increased incidence of adverse reactions.
4. Highly cost

Friendly Bacteria

L. acidophilus, *L. salivarius*,
L. casei, *L. thermophilus*,
B. bifidum, *B. longum*, etc.

Unfriendly Bacteria

Pathogenic bacteria & fungi,
such as *Candida albicans*, etc.



Problems with antimicrobial agents

1. Drug resistance.
2. Adverse effects
3. Drug-drug interaction

2. Adverse effects

a. Hypersensitivity; (not dose related)

e.g. Penicillin, cephalosporin.

b. Toxic effect (dose related)

High serum levels of certain antibiotics may cause Direct toxicity / Organ toxicity

e.g. Aminoglycosides(ototoxicity)

- Chloramphenicol (Aplastic anemia)

c. Superinfections: (clostridium difficile-colitis)

alterations of the growth of normal flora of intestine, genitourinary tracts.

Appearance of a new infection while treating an original infection (multiply C.difficile).

Classification of Antimicrobial Agents

1) Inhibitors of cell wall synthesis

- **B-lactams** - **Vancomycin**

2) Inhibitors of Protein Synthesis

- **Tetracycline** – **Aminoglycosides** - **Macrolides**-
Clindamycin - **Chloramphenicol**.

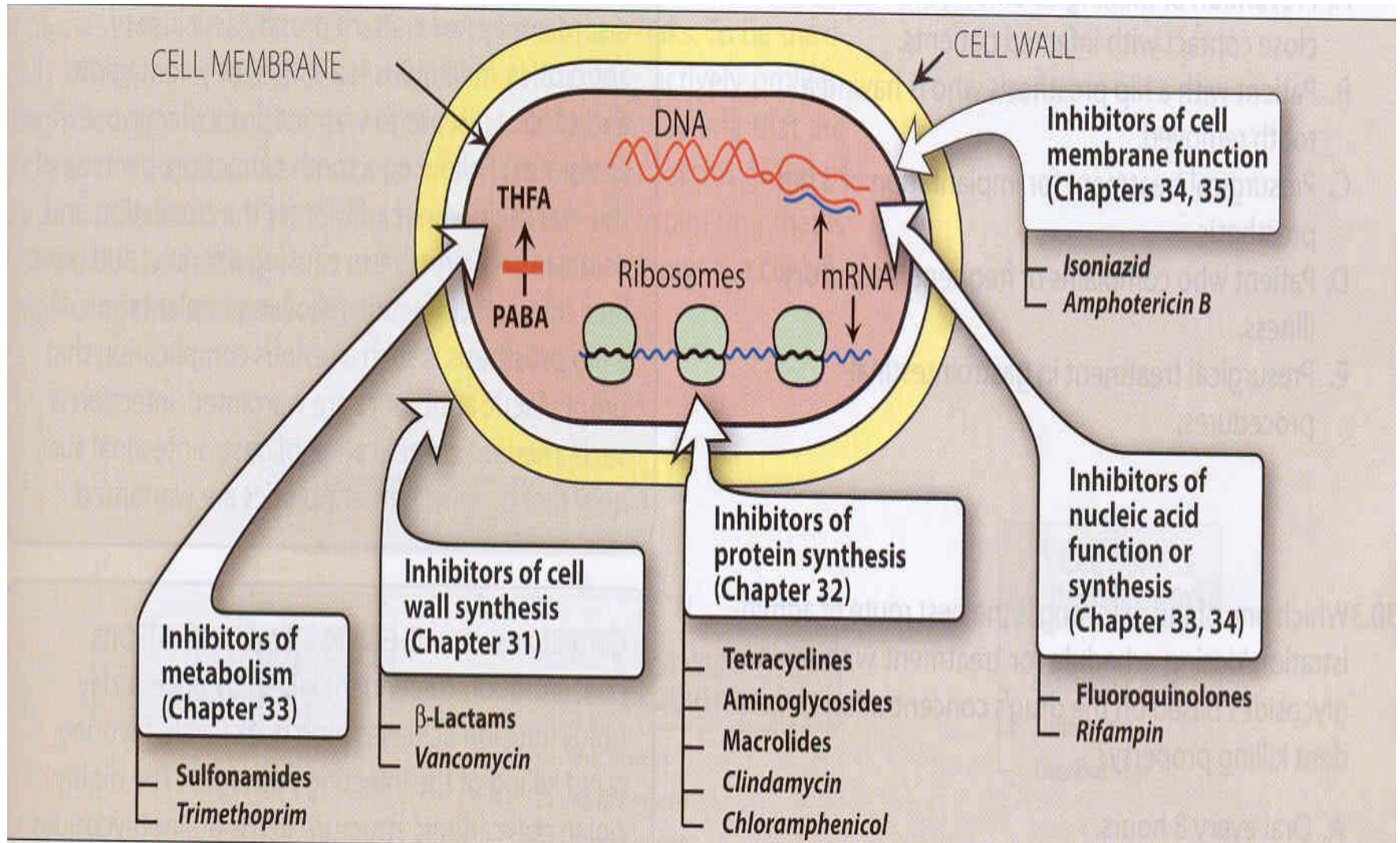
3) Inhibitors of Metabolism

- **Sulfonamides**- **Trimethoprim**

4) Inhibitors of Nucleic Acid function or synthesis

- **Fluoroquinolones**- **Rifampin**

Classification of some antimicrobial agent by their site of action..



A. Inhibitors of Cell Wall Synthesis

AGENTS AFFECTING THE CELL WALL

β-LACTAM ANTIBIOTICS

OTHER ANTIBIOTICS

β-LACTAMASE INHIBITORS

- Clavulanic acid
- Sulbactam
- Tazobactam

- Bacitracin
- Vancomycin
- Daptomycin

PENICILLINS

CEPHALOSPORINS

CARBAPENEMS

MONOBACTAMS

- Amoxicillin
- Ampicillin
- Dicloxacillin
- Indanyl carbenicillin
- Methicillin
- Nafcillin
- Oxacillin
- Penicillin G
- Penicillin V
- Piperacillin
- Ticarcillin

- Ertapenem
- Imipenem/cilastatin*
- Meropenem

- Aztreonam

1st GENERATION

2nd GENERATION

3rd GENERATION

4th GENERATION

- Cefadroxil
- Cefazolin
- Cephalexin

- Cefaclor
- Cefprozil
- Cefuroxime
- Cefoxitin

- Cefdinir
- Cefixime
- Cefotaxime
- Ceftazidime
- Ceftibuten
- Ceftizoxime
- Ceftriaxone

- Cefepime

CELL WALL SYNTHESIS INHIBITORS

1. β -LACTAM ANTIBIOTICS.

- I. Penicillins.
- II. Cephalosporins.
- III. Carbapenems.
- IV. Monobactams.

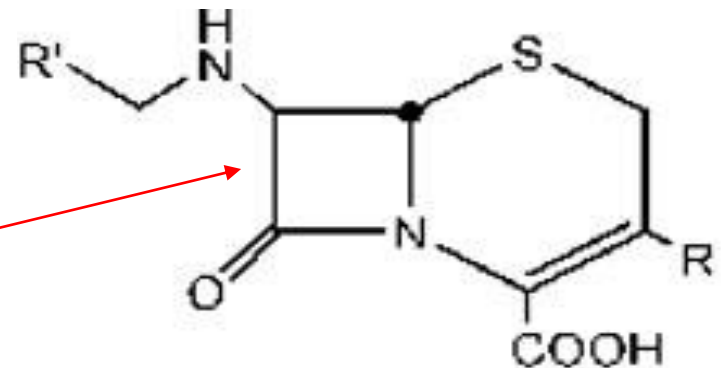
All these have one thing in common which is the β -lactam ring, which is very essential & imp for the activity of the drug against M.O.

2. OTHER ANTIBIOTICS.

- I. Vancomycin.

1- Beta-Lactam Antibiotics

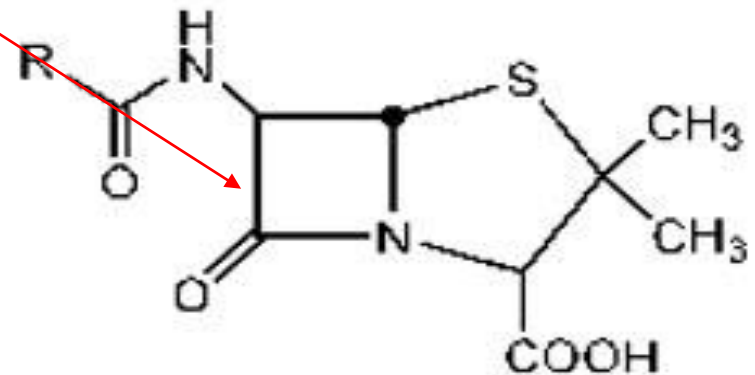
β -lactam ring



Cephalosporin

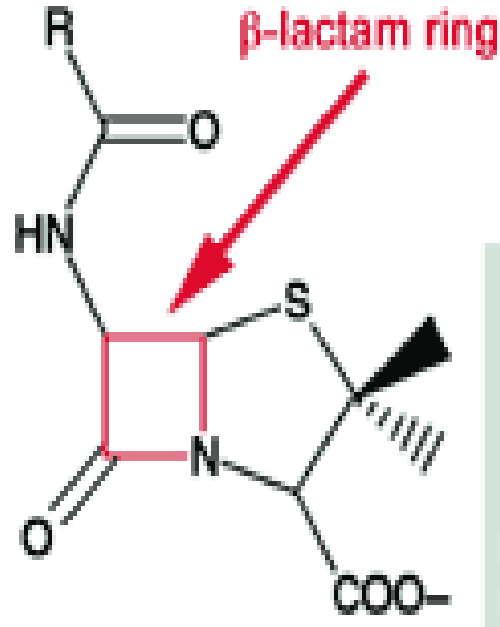
Penicillinases (Beta-lactamases)

enzymes that cleave the beta-lactam ring and thereby render penicillin and other beta-lactam antibiotics inactive against M.O.



Penicillin

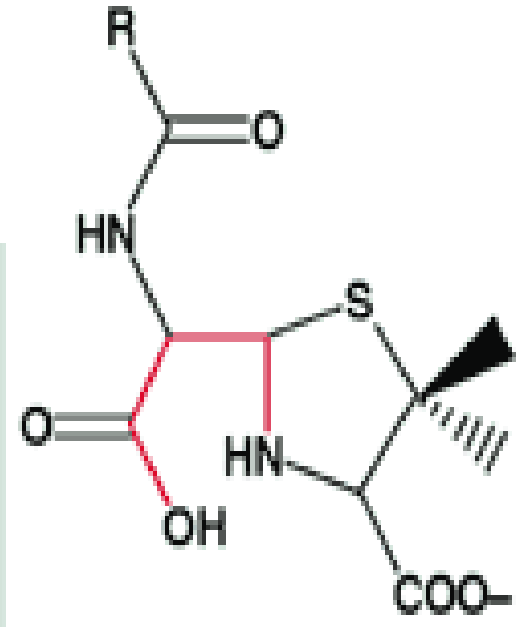
Penicillin Resistance



Penicillin

β -lactamase

β -lactamase breaks a bond in the β -lactam ring of penicillin to disable the molecule. Bacteria with this enzyme can resist the effects of penicillin and other β -lactam antibiotics.



Penicilloic acid

Inactive penicillin

B-lactamase inhibitors

Substances don't have antibacterial activity but they have the ability to **inhibit** the B-lactamase enzyme....

Ex. Clavulanic acid

Beta-Lactam Antibiotics

- Cell wall active agents
 - Prevent the final step in the synthesis of the bacterial cell wall.
- Range from very **narrow spectrum** to very **broad spectrum**
- **Only active on actively multiplying M.O. why???**
- The β -lactams are **BACTERICIDAL...WHY???**

KEVIN BROWN

PENICILLIN MAN

ALEXANDER FLEMING
AND THE ANTIBIOTIC REVOLUTION





I. Penicillins

- ✓ The most widely effective and use.
- ✓ Least toxic drugs.
- ✓ The safest drugs.
(we exclude the allergy rxn)
- ✓ All penicillins are excreted by the kidneys.

Classification of penicillin

1- Natural Penicillins

Penicillin G(parenteral)
Penicillin V(oral)

3- Anti-pseudomonal Penicillins

Piperacillin

2- The extended or broad Spectrum Penicillins

Aminopenicillins :
Ampicillin
Amoxicillin

4- Penicillinase Resistant Penicillins (anti-staphylococcal)

Cloxacillin.

Natural Penicillins

AND

**The extended or broad Spectrum
Penicillins**

ARE susceptible to inactivation by B-lactamases
(penicillinases)

1. Natural Penicillins

1- Penicillin G

(Benzylpenicillin)

- also called Crystalline penicillin.
- it is powder form.
- can be given IV (bolus or infusion) or IM.
- Has very short duration (1-2 hrs).
- **Destroyed by gastric juice if it is given orally so, NOT given orally.**
- Given in sever infection.
- Syphilis , acute Tonsillitis.**

2- Penicillin V

(Phenoxymethylpenicillin)

- Given orally (every 4h).
- Penicillin V is more acid-stable than penicillin G.
- **acute Tonsillitis.**
- Mild to moderate infections.
- Commonly used in children.

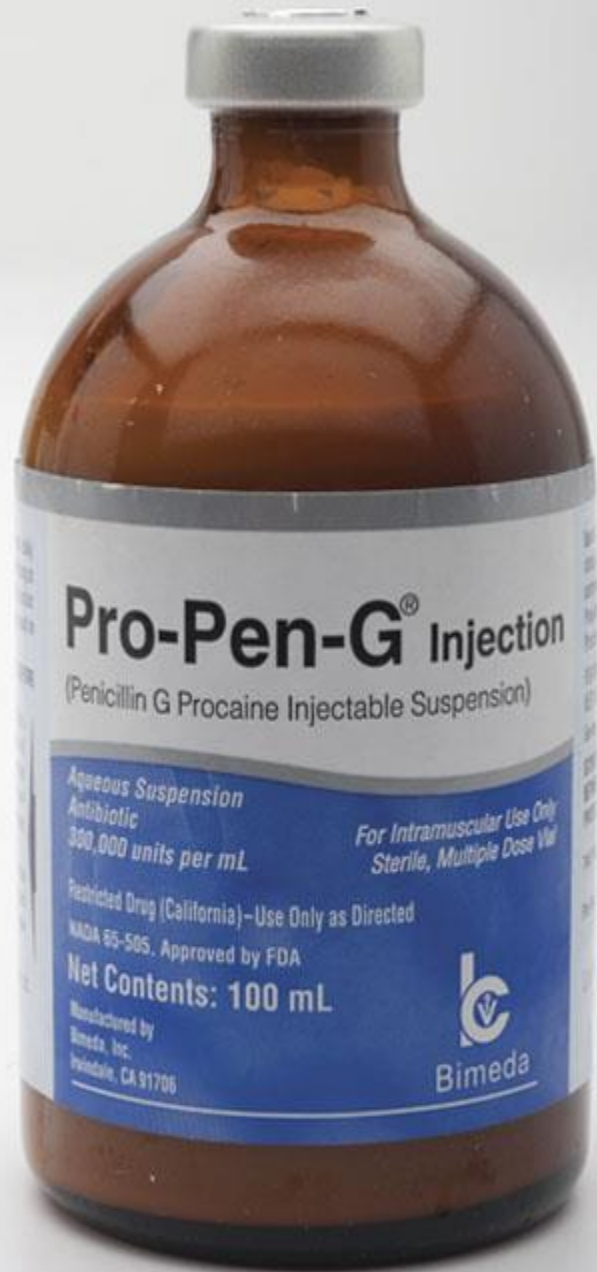
Derivatives of penicillin G

Long-acting forms:-

1- Procaine penicillin G (12 hrs) is given as every 12.

2- benzathine penicillin G (4 weeks) is given every month.

- Effective in treatment in syphilis.
 - Prophylaxis of endocarditis.
 - Prophylactic in rheumatic fever
- Both are administered **IM** and serve as depot forms..
 - they are **suspension** formulation that is never given by IV rout.



NDC 61570-148-10

Bicillin[®] L-A

(penicillin G benzathine
injectable suspension)

FOR DEEP IM INJECTION ONLY

2,400,000 units per 4 mL

Ten sterile single-dose disposable syringes
(4 mL size)

WARNING: NOT FOR INTRAVENOUS USE

R_x only



Monarch
Pharmaceuticals[®]



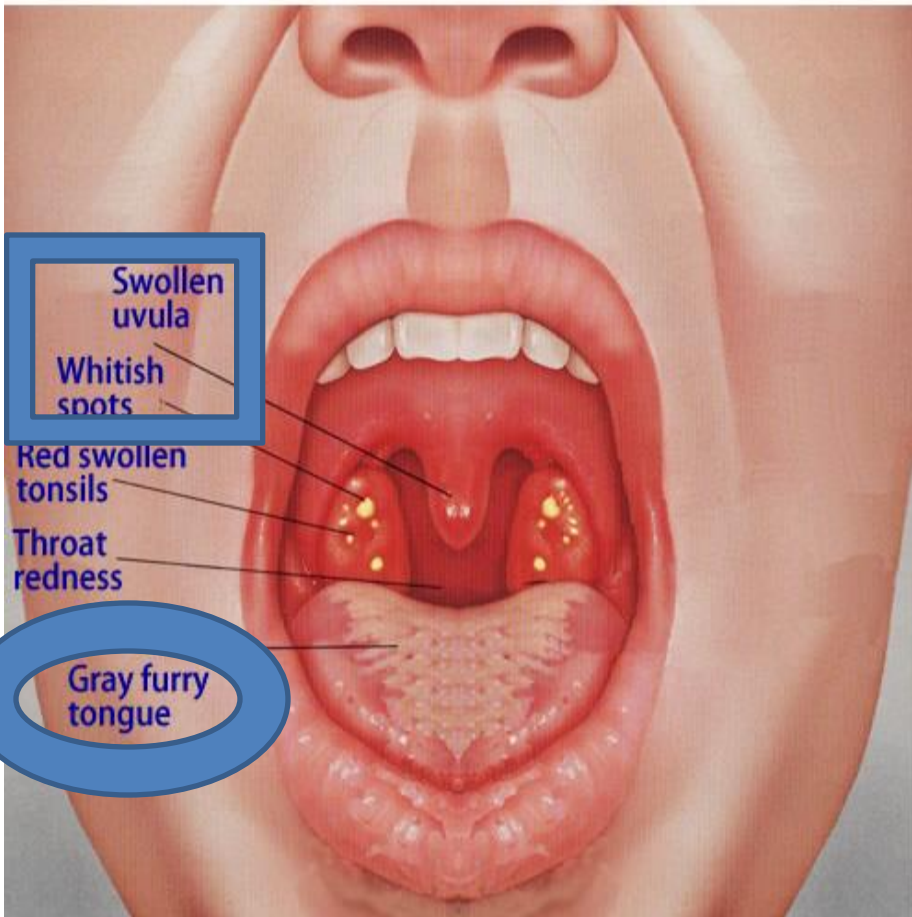
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Syphilis & acute severe tonsillitis :

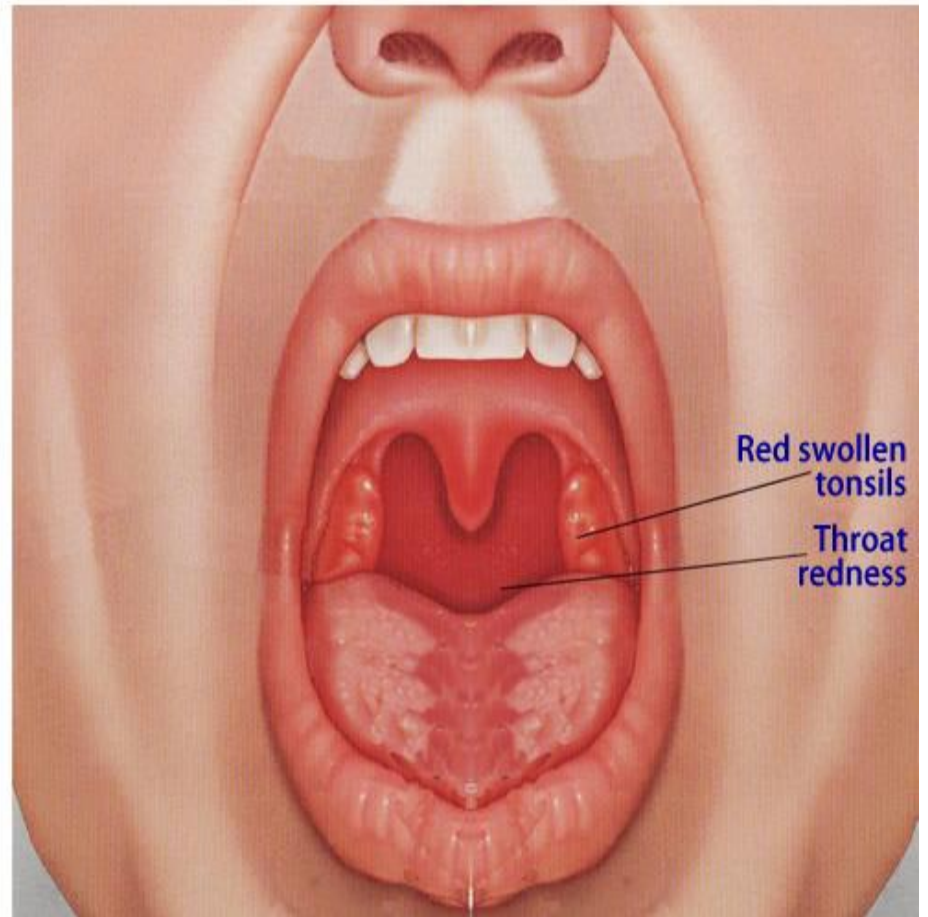
Pen G at the beginning...

apt with syphilis who requires a treatment for so many months (maintenance) can receive benzathine penicillin instead of giving him crystalline or procaine penc..

Bacterial



Viral

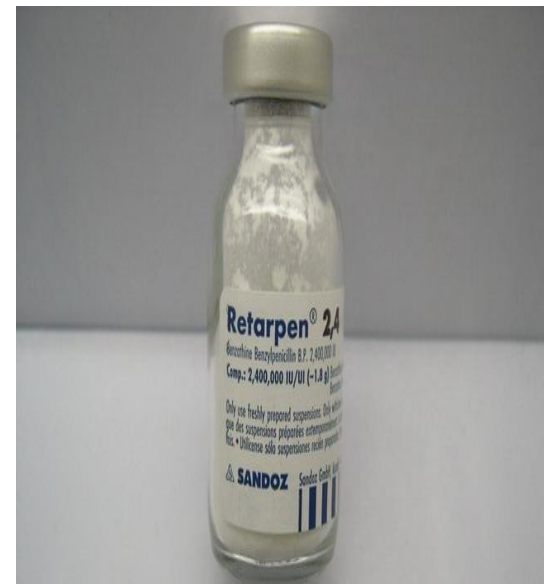




Pus in Tonsillar Crypts

Acute tonsillitis





Retarpen 240 萬 IU/Vial
瑞達平 注射劑

2. Extended-spectrum Penicillins

or

Aminopenicillin:

Ampicillin and amoxicillin

Ampicillin

❖ (IV, Oral) is given every 6h (4x1).

❖ must be taken on empty stomach.

Amoxicillin

❖ (IV, Oral) is given every 8h (3x1).

❖ better absorbed orally.

Ampicillin

- ❖ (IV, Oral) is given every 6h (4x1).
- ❖ must be taken on **empty stomach**.
- ❖ Is used in Bacillary Dysentery. **WHY???**
1g (4x1) for 5 days + fluid.
- ❖ Diarrhea is common side effect **WHY????**

Amoxicillin

- ❖ (IV, Oral) is given every 8h (3x1).
- ❖ better absorbed orally.
- ❖ Is employed prophylactically by dentists for patients with abnormal heart valves who are to undergo extensive oral surgery..
- ❖ used in treatment of peptic ulcer to eradicate H.Pylori.
- ❖ Otitis media.

B-Lactamase Inhibitors clavulanic acid.

- Has NO or lack or very weak antimicrobial effect.
- Binds irreversibly to B-lactmase → protect hydrolyzable pencyllins from inactivation by these enzymes.

Major S.E :- GI upset. (due to clavulanic acid)

Augmentin®:

consists of **Amoxicillin + Clavulanic acid.**



3- Anti staphylococcal penicillins

Also called anti-staph or penicillinase resistance penicillins.

❖ Strains of staphylococcus resistant to these drug called :
methicillin- resistant staphylococcus aureus (MRSA).
A serious source of nosocomial (hospital-acquired) infections.

❖ MRSA commonly respond to **vancomycin**.



3- Anti staphylococcal penicillins

Also called anti-staph or penicillinase resistance penicillins.

Ex. **Methicillin**, Flucloxacillin, Cloxacillin

- ❖ Given IV & orally. (every 4-6 hr)
- ❖ They are restricted to the treatment of infections caused by penicillinase-producing staphylococci.
- ❖ Because of its nephrotoxicity caused by **methicillin** nowadays this drug **is not** used clinically.
- ❖ Strains of staphylococcus resistant to these drug called :
methicillin- resistant staphylococcus aureus (MRSA).
A serious source of nosocomial (hospital-acquired) infections.



4- Anti pseudomonal Penicillins:

Ex. **piperacillin**

Ps.aeruginosa: G-ve bact Lacks porins → Making these organism resistant to many antimicrobial agents.

- ❖ **Pse.aeruginose** → very difficult to deal with & produce resistance easily.
- ❖ Given parentally not orally.
- ❖ **Piperacillin** is the most potent of these antibiotics.

Antipseudomonal penicillins

- *piperacillin with tazobactam*,

extends the antimicrobial spectrum to include penicillinase-producing organisms.

❧ All the penicillins cross the placental barrier, but **none has been shown to be teratogenic.**



❧ the inflamed meninges are more permeable to the penicillins.

❧ Penicillin levels in the prostate are insufficient to be effective against infections, penetration into bone, cerebrospinal fluid (CSF), is insufficient

❧ **Probenecid** inhibits the secretion of penicillins by competing for active tubular secretion via the organic acid transporter and, thus, can increase blood levels

Adverse reactions of penicillins

1-Hypersensitivity reaction :

- ✓ ranging from rash to angioedema & anaphylaxis.
 - ✓ Cross sensitivity with other β -lactam as cephalosporins.
 - ✓ Should be avoided if history is positive.
-
- To treat anaphylaxis:
 - ✓ **Maintain air way**
 - ✓ **Epinephrine** ,by which rout given??
 - ✓ **Antihistamine** .
 - ✓ **Hydrocortisone**.

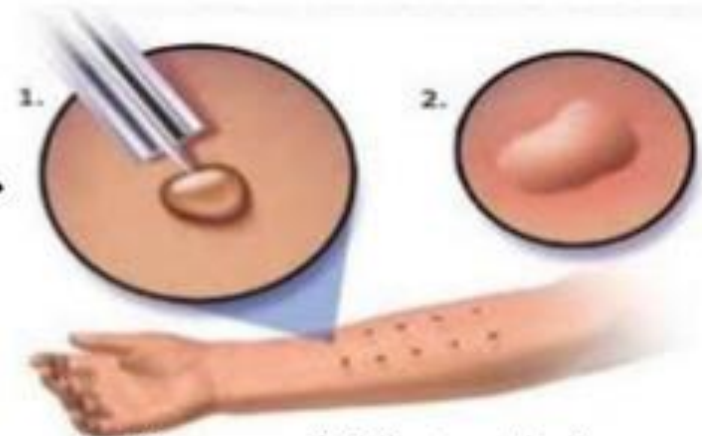
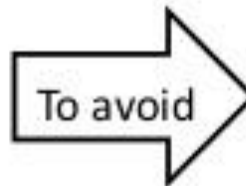


Hypersensitivity reaction

- All AMAs are capable to causing hypersensitive reaction, and this this reactions are unpredictable and unrelated to dose. E.g.: Penicillin induced anaphylactic shock (prick skin testing)



Inj. Penicillin induced anaphylactic shock



Perform sensitivity test before administering penicillin Inj.



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2-Diarrhea (most common): it is a common problem mainly with (Ampicillin).

Pseudomembranous colitis may occur.

❑ to treat Pseudomembranous colitis :

- **Stop the drug .**
- **Rehydration with electrolytes (fluid).**
- **Metronidazole → If no response Vancomycin.**

pseudomembranous colitis

- ❧ inflammation of the colon that occurs in some people who have taken antibiotics. it is sometimes called antibiotic-associated colitis or C. difficile colitis. It is almost always associated with an overgrowth of the bacterium *Clostridium difficile*.
- ❧ Severe pseudomembranous colitis can be life-threatening. However, treatment is usually successful.



CEPHALOSPORINS



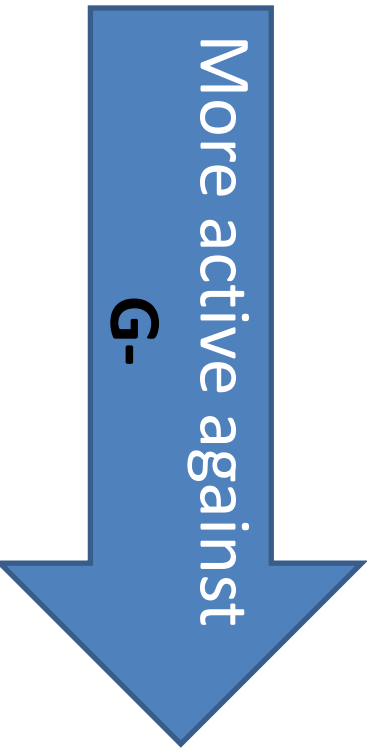
BY
V. SUSHIL

II. CEPHALOSPORINS

Suffix : Cef or Ceph

- A wider spectrum than penicillins.
- More resistance to B-lactmases enzyme.
 - Eliminated by kidney.
 - More expensive than penicillins.

Classification to four major groups



1st Generation

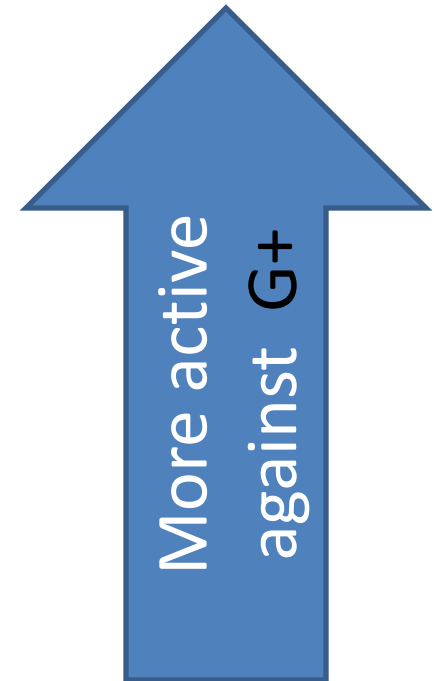
Cefazolin, Cefadroxil

2nd Generation

Cefoxitin

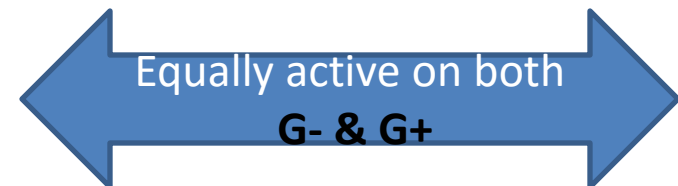
3rd Generation

Cefotaxime, Ceftriaxone



4th Generation

Cefepime



Cephalosporins

Gram + activity

1st Generation

β -lactamase sensitive

2nd Generation

Gram — activity

3rd Generation

β -lactamase resistant

4th Generation: good Gram + and Gram - activity;
more resistant to β -lactamase

(1st. Gen.)

(2nd. Gen.)

Cefazolin (IV)
Cefdroxil (oral)

Cefoxitin.(IM or IV)

cefazolin

Surgical prophylaxis
(orthopedic surgery) ???

Give 1g at induction of
anesthesia

because **cefazolin good
penetration to bone.**

Cefodroxil (2x1):
UTI,RI & sinusitis.

Cefoxitin

Surgical prophylaxis:

(Internal abdominal & pelvic
infection).

Why???

Against anaerobic M.O
(bacteroides fragilis).

Give 2g at induction of
anesthesia.

(3 rd . Gen.)	(4 th . Gen.)
<p>Cefotaxime, (IV) Ceftriaxone, (IV) Ceftazidime</p>	<p>Only parenteral Ex.Cefapime</p>
<p>The main adv of 3rd generation : their easy penetration to the CSF</p> <p><u>Ceftriaxone, Cefotaxime</u></p> <p>Due to Easy penetration to CSF</p> <p><u>DOC</u> in :- are effective in the treatment of neonatal and childhood meningitis caused by H.influenzae. meningococcal meningitis.</p> <p><i>Brain abscess</i> <i>Alternative to aminoglycoside</i></p> <ul style="list-style-type: none"> • <u>Ceftazidime active against P.A</u> • <u>Ceftriaxone</u> <p>Excretion: mainly in the bile = no need to adjust for renal insufficiency</p>	<p>Wider spectrum</p> <p>Used in Infection (organism) resistance to other cephalosporin.</p>

Adverse effects

- ✓ pain after injection.
- ✓ Diarrhea.
- ✓ Hypersensitivity reaction
(10% cross- sensitivity)

Adverse effects

- Allergic reaction: cross allergy with penicillin(5-15%).
- Some have anti-Vitamin K effect(bleeding).
- Some can cause a disulfiram – like reaction.
- False positive glucose test.
- Pain at site of injection.
- Diarrhea .
- neutopenia ,-----



NDC 63323-237-10 23710

CEFAZOLIN

FOR INJECTION, USP

1 gram*

FOR INTRAVENOUS OR INTRAMUSCULAR USE
Rx only

25 Single Use Vials







3-Carbapenems

E.x.. **Imipenem.**

administered IV infusion or IM every 6-8 hrs

✓ Very Broad-spectrum coverage.

Empiric therapy

Clinical application:-

life threatening infection .

**combined with
cilastatin**

4-Monobactams

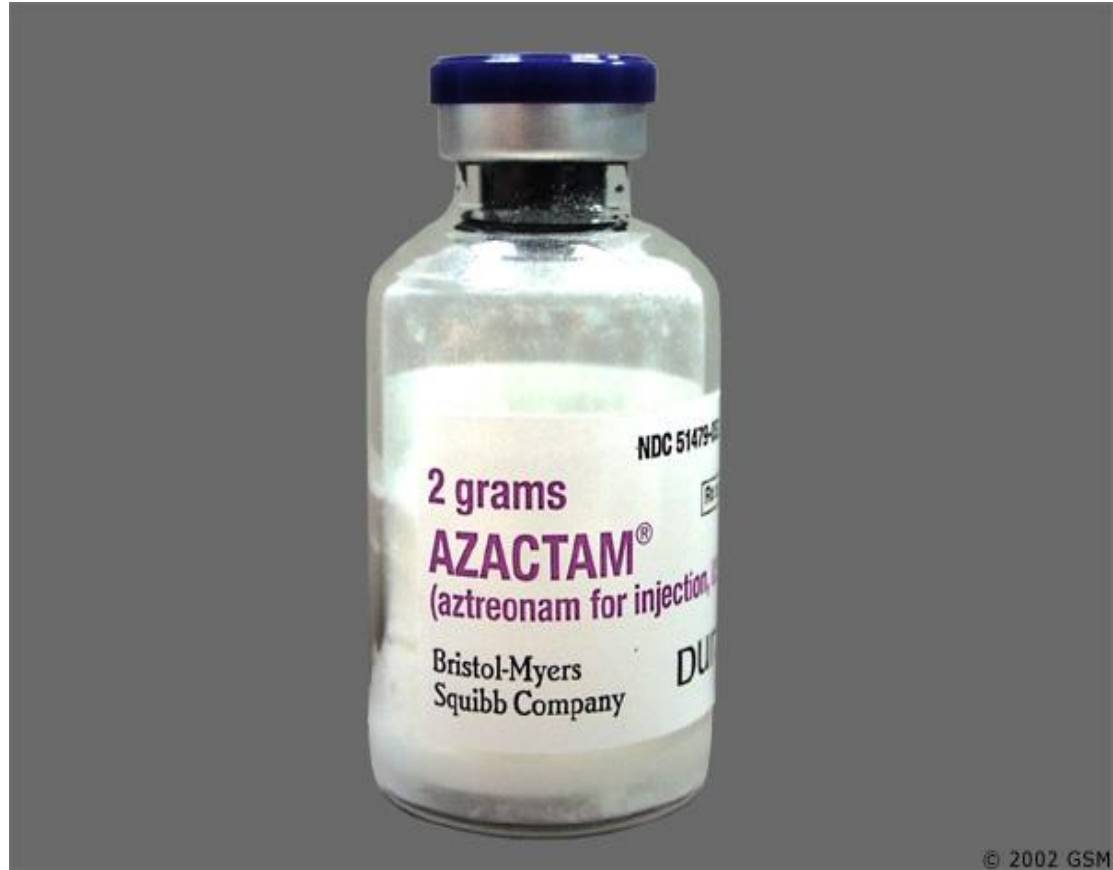
E.x. **Aztreonam.**

It is administered either IV or IM. Every 8 hrs

narrow spectrum.

- Only against G- aerobic rod.
- Aztreonam is resistant to the action of B-lactamases

❖ this drug **may offer a safe** alternative for treating patients who are allergic to penic &/or cephalosporins.



Carbapenem

- Imipenem , meropenem
- they Should be reserved to infection that are not responding to other antibiotic.
- These β -lactams antibiotics show cross sensitivity with Penicillins
- Imipenem/cilastatin and meropenem are the broadest-spectrum B-lactam antibiotic.
- Imipenem resists hydrolysis by most B-lactamases,
- Imipenem plays a role in empiric therapy .



**Other cell wall
inhibitor
vancomycin**

Vancomycin

Administered Oral , IV

Narrow spectrum

Bactericidal /not B-lactam

Orally:- every 6 hrs for refractory pseudomembranous colitis due to C. difficile.

Slow IV infusion (1-2 hrs) for treatment of systemic infections or prophylaxis.

✓ is effective against multiple drug-resistant organisms, such as MRSA. (DOC)

✓ Vancomycin in combination with A.G alternative regimen to treatment of enterococcal endocarditis.

S.E:-

1-Flushing (**red man syndrome**) with a **rapid** infusion. (**More common**)

2- phlebitis (inflammation of vein) at site of injection.

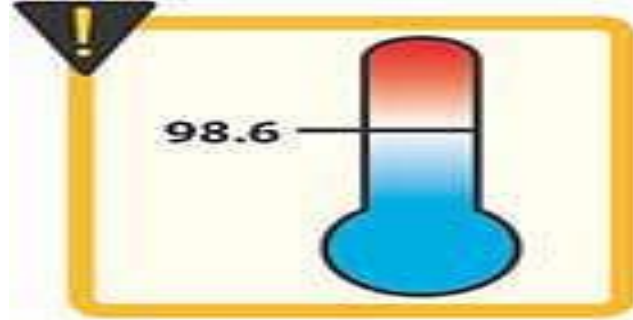
3- ototoxicity & nephrotoxicity (rare) but increased risk when administered with A.G.



Vancomycin

Clinical Uses

- Infections due to **methicillin-resistant staph** including bacteremia, empyema, endocarditis, peritonitis, pneumonia, skin and soft tissue infections, osteomyelitis
- Serious gram-positive infections in **β -lactam allergic patients**
- Infections caused by multidrug resistant bacteria
- Endocarditis or surgical prophylaxis in select cases
- Oral vancomycin for **refractory *C. difficile* colitis**



Fever



Chills

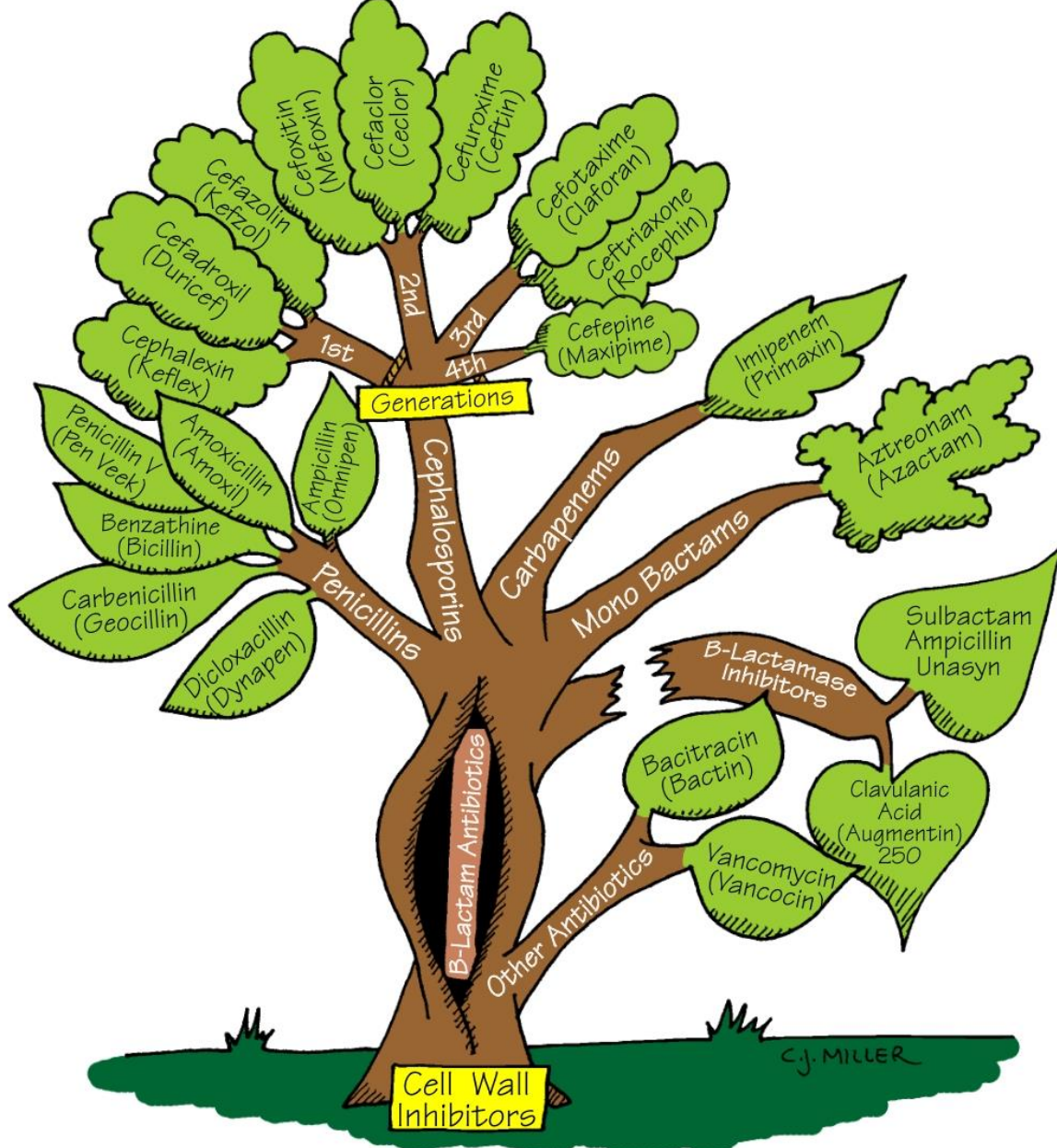


Flushing



Phlebitis





THE ANTIBIOTIC TREE