

Antibiotics

Penicillins & Cephalosporins

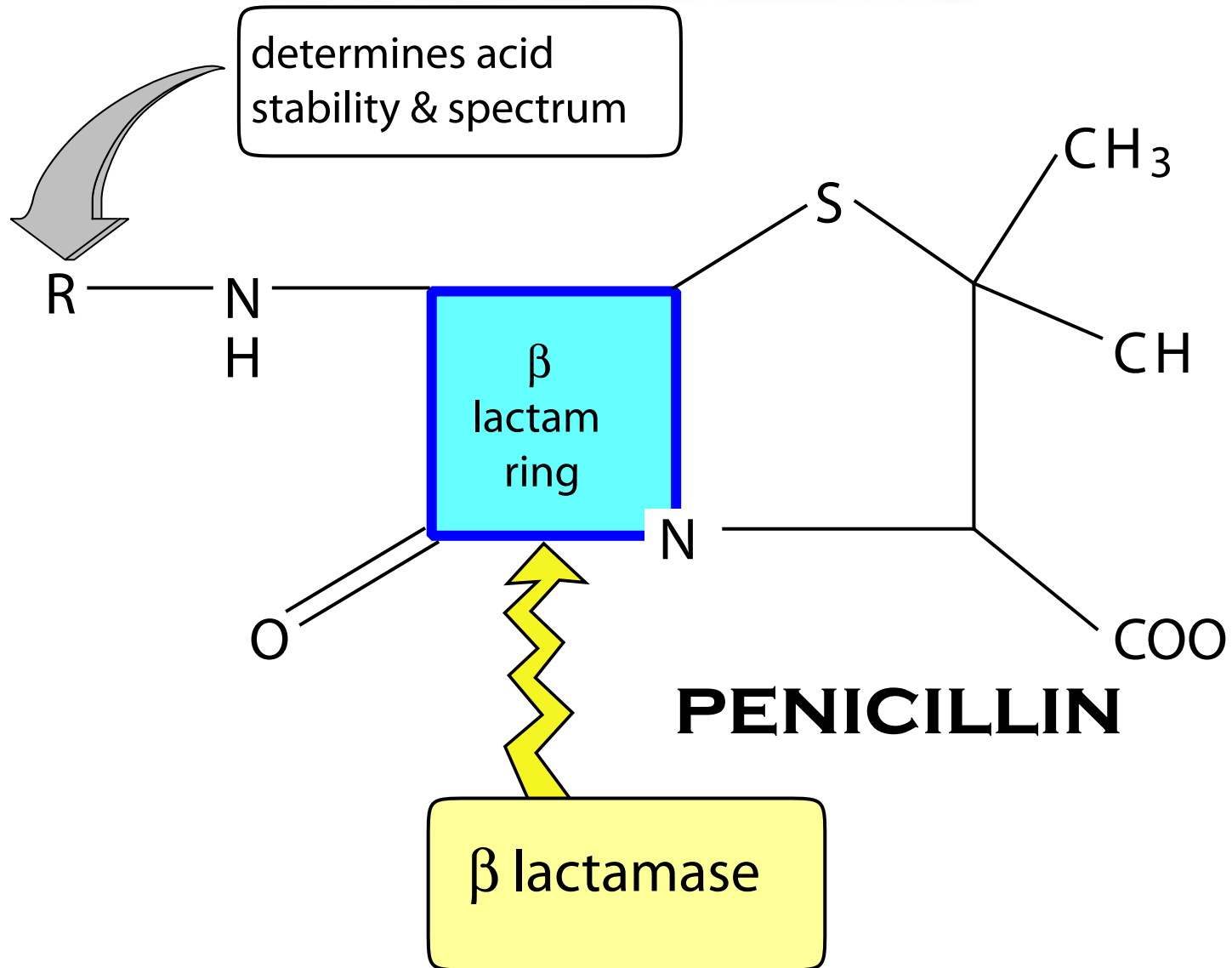


What would you do?

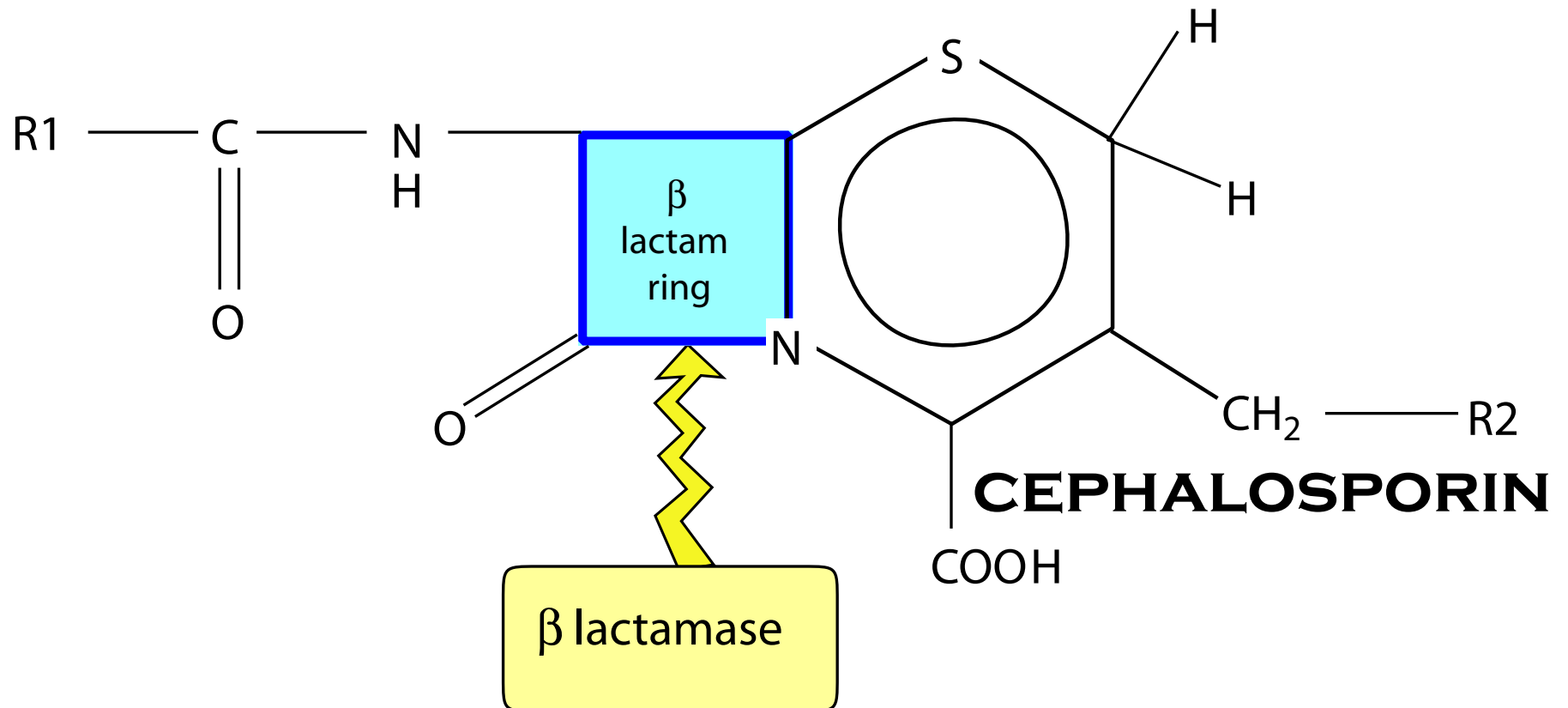
- calved 10 days ago
- all quarters painful and swollen
- clots & blood in milk



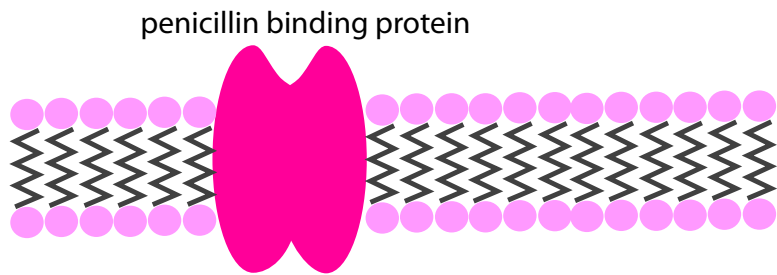
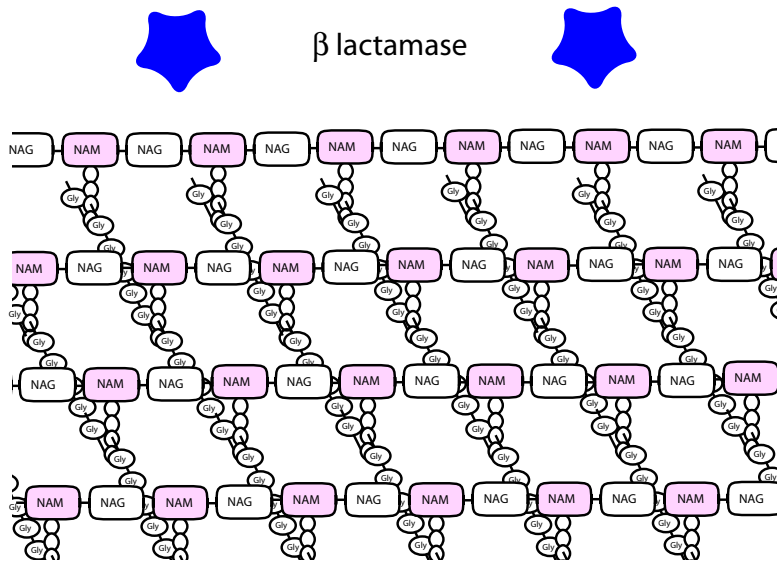
beta lactams



beta lactams



bacterial cell walls



Gram positive

cell wall

peptidoglycan

cell membrane

β lactamase

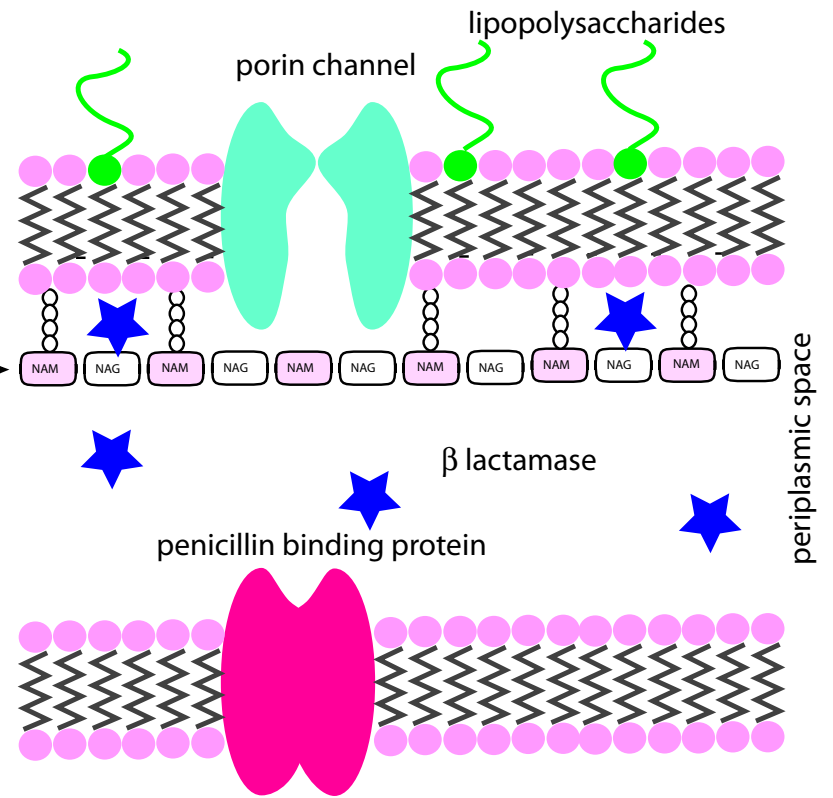
porin channel

lipopolysaccharides

β lactamase

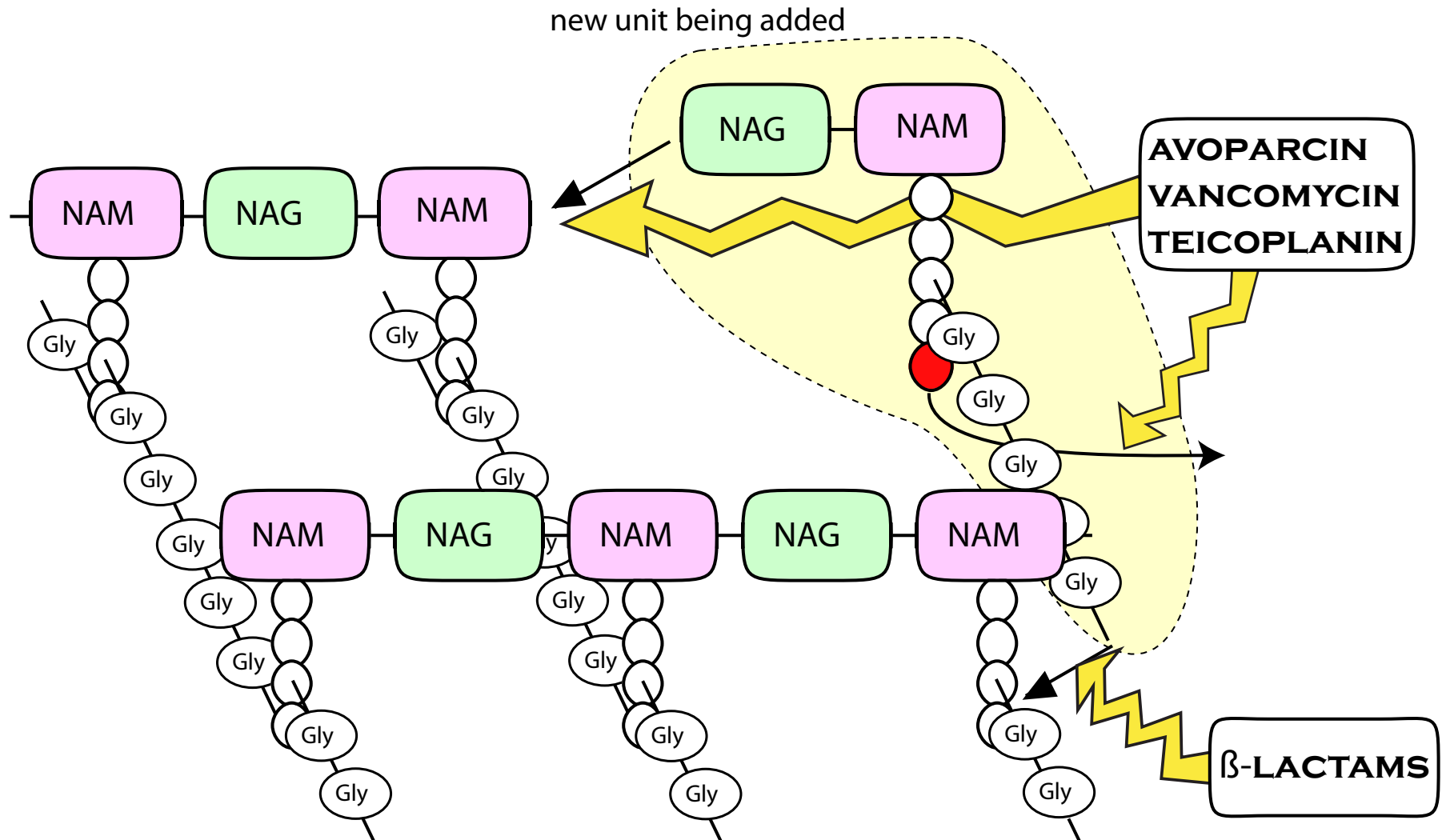
penicillin binding protein

periplasmic space



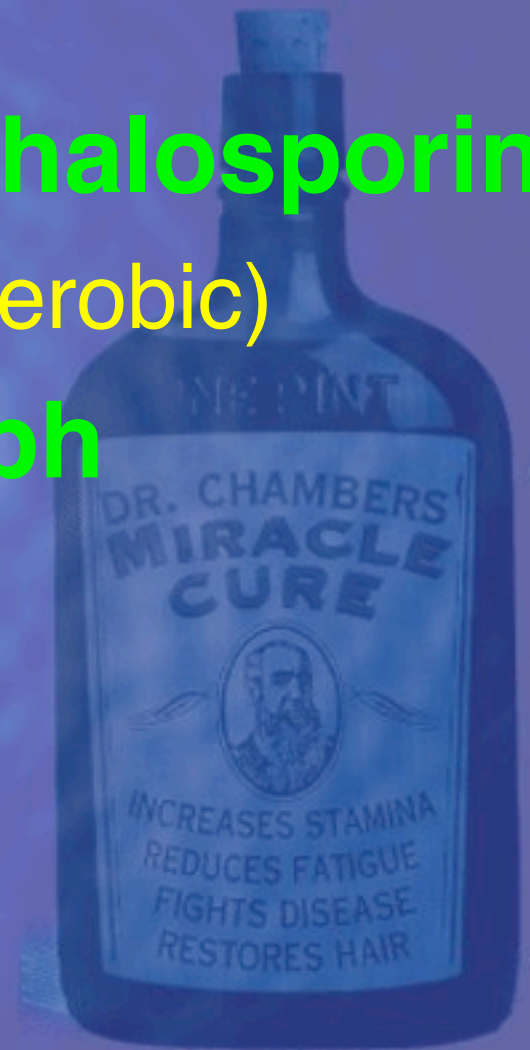
Gram negative

mechanism

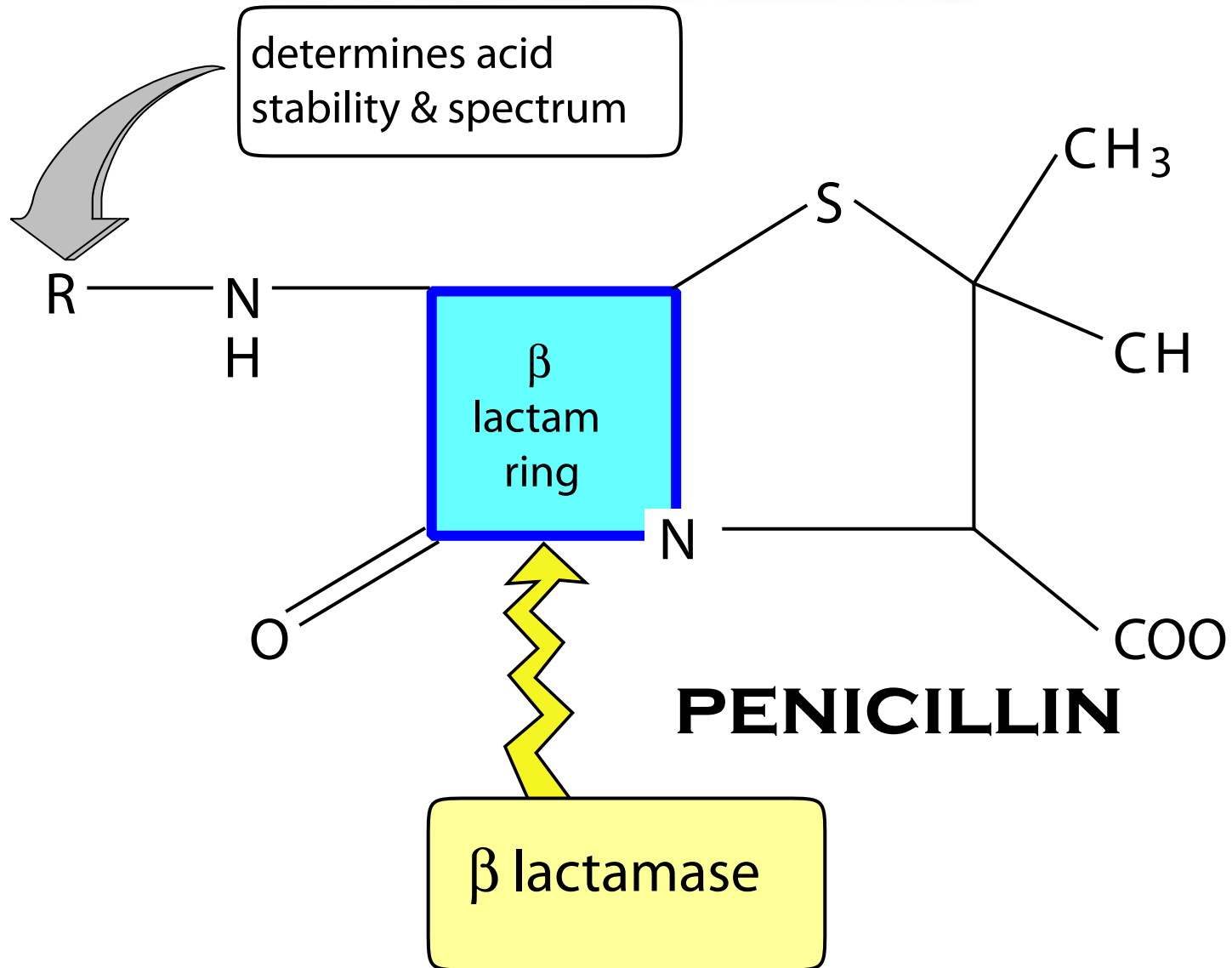


spectrum

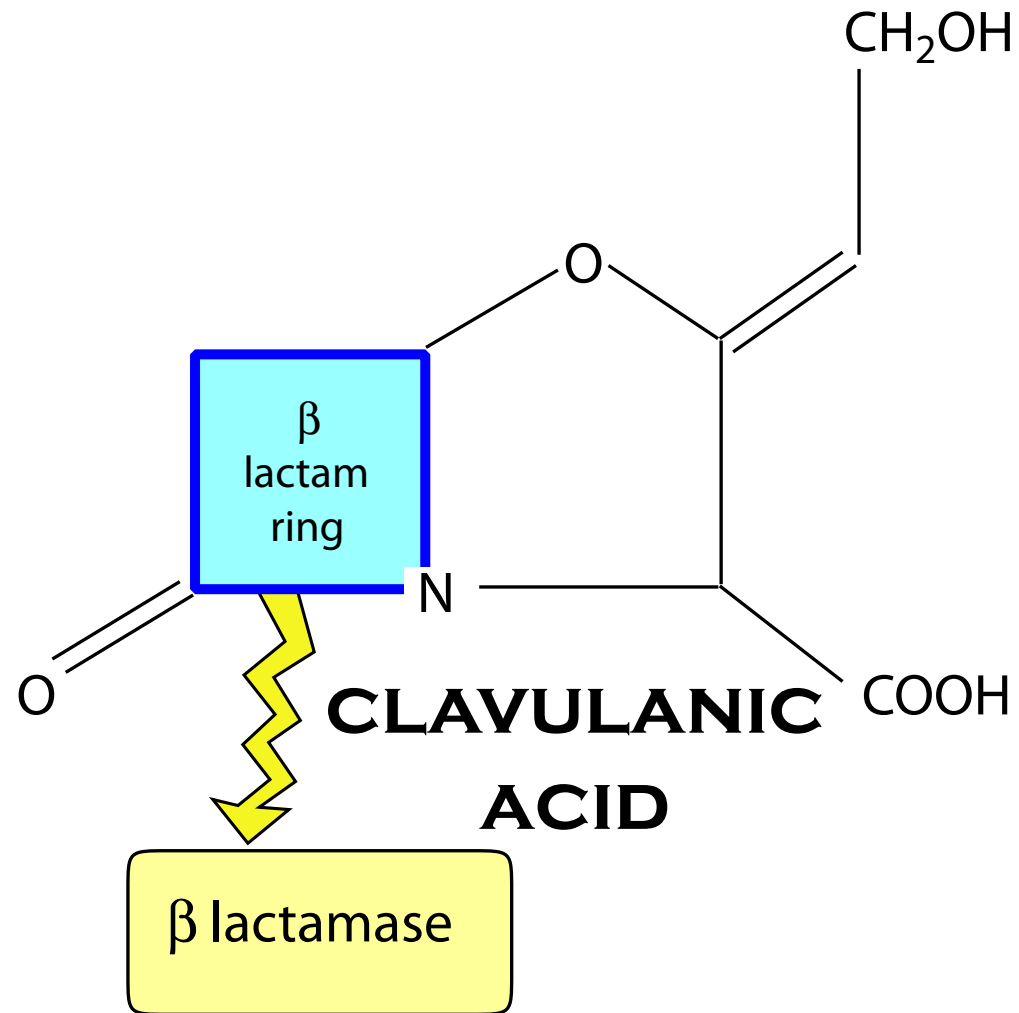
- **simple penicillins & cephalosporins**
 - mainly G+ (aerobic and anaerobic)
- **semisynthetic pen & ceph**
 - BL producing G+
 - G-
 - *Pseudomonas*



beta lactams

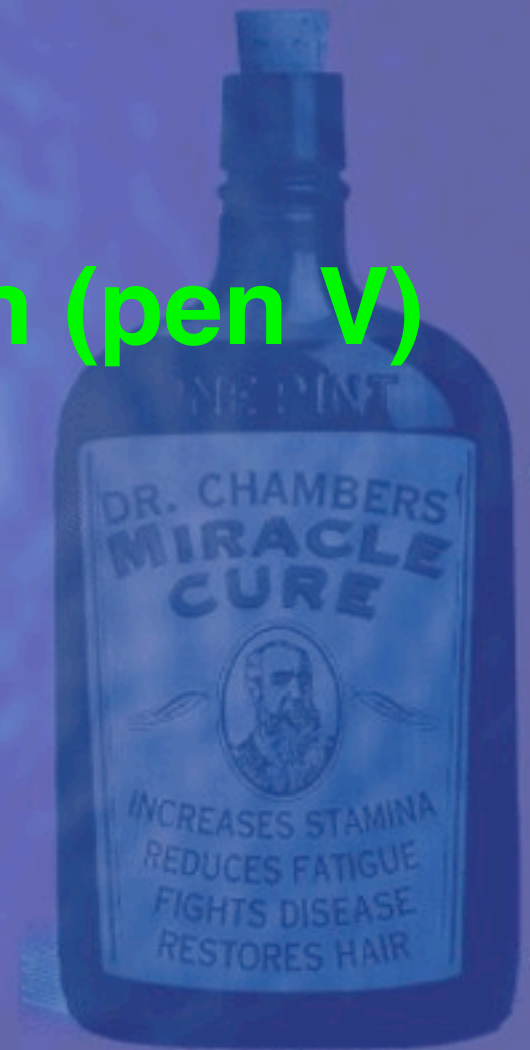


clavulanate



narrow spec penicillins

- benzylpenicillin (pen G)
- phenoxymethylpenicillin (pen V)
 - non BL G+
- cloxacillin
- flucloxacillin
 - BL G+



broad spec penicillins

- amoxycillin
- ampicillin
 - non BL G+ & G-
- co-amoxiclav
 - BL G+ & G-



anti-Pseudomonas

- ticarcillin (\pm clavulanate)
- piperacillin (\pm tazobactam)



use

- **benzylpenicillin**
 - most horse infections
 - most mastitis
- **co-amoxiclav**
 - general purpose broad spectrum drug



penicillin resistance

- rare (apart from *Staph aureus*)
- develops slowly
- usually not clinically significant
 - animals
 - owner



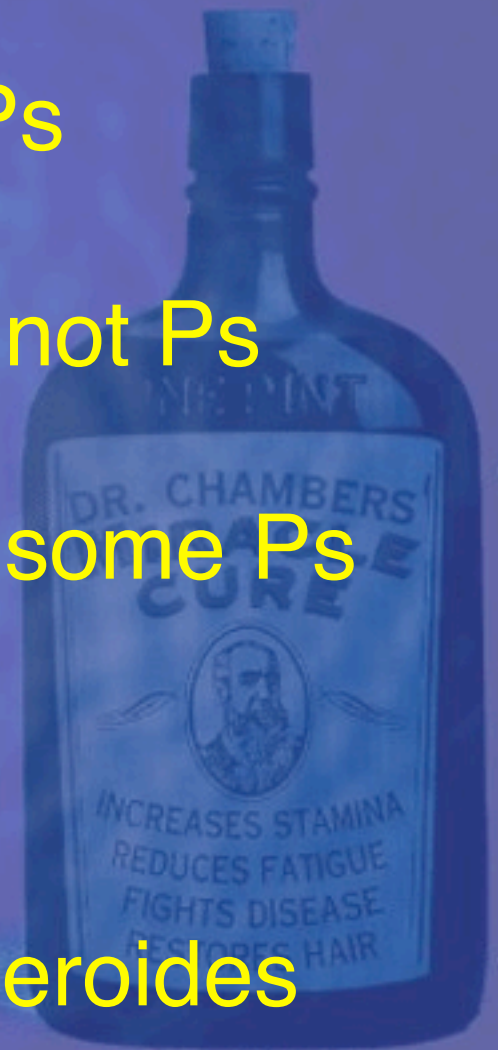
human use

- **penicillin**
 - URT infections
- **flucloxacillin**
 - Staph aureus skin infections
- **amoxycillin**
 - general purpose broad spectrum drug
- **co-amoxiclav**
 - 2nd line broad spectrum drug



cephalosporins

- **G1**
 - good G+, moderate G-, not Ps
- **G2**
 - very good G+, moderate G-, not Ps
- **G3**
 - moderate G+, very good G-, some Ps
- **G4**
 - good all round
- **cephamycins**
 - mainly G-, not Ps, good Bacteroides



cephalosporins

- G1
 - skin disease dogs
- G2
 - mastitis cows
- G3
 - foot rot cows!
- G4
 - mastitis cows
- cephamycins
 - not used



cephalosporin resistance

- rare
- develops slowly
- can be clinically significant
 - animals
- very significant
 - people



human use

- **G1 / 2**
 - second / third line drugs
- **G3 / 4**
 - life-threatening G- infections



pharmacokinetics

absorption

- **iv**
 - Na or K salts (duration 4-6h)
- **im / sc**
 - procaine, benzathine penicillin (duration 12-24h)
 - amoxycillin trihydrate
- **po**
 - phenoxymethyl penicillin
 - flucloxacillin
 - many cephalosporins



pharmacokinetics

distribution

- weak acids
- generally low V_d
 - poor penetration into tissues
 - cephs mainly better than pens



pharmacokinetics

elimination

- eliminated by kidneys
- anion pumps in PCT
 - probenecid
 - aspirin
- most modern cephs rapid elimination in animals



other cell wall inhibitors

- bacitracin
- fosfomycin
- carbapenems
- monobactams
- glycopeptides
 - vancomycin
 - teicoplanin



What would you do?

- calved 10 days ago
- all quarters painful and swollen
- clots & blood in milk



problems?



problems?

- mastitis



decision process

- Does the drug kill the bacteria?
- Does the drug get to where the bacteria are?
- Is clinically significant resistance likely?
 - in the cow?
 - in the herd?
 - in the farmer?



bacteria?

- Strep uberis?
- Staph aureus?
- other Streps?
- other Staphs?
- E. coli?



treatment?

- sample for C & S?
- intramammary drugs?
- parenteral drugs?
- withholding times?



resistance?

- **narrow spectrum G+**
 - penicillin
 - cloxacillin
 - G1 cephalosporins
- **broad spectrum**
 - ampicillin
 - co-amoxiclav
 - cefquinome



withholding times

- at peak lactation
- cure vs value of discarded milk?



penicillins & cephalosporins

- block formation of cell walls
- early drugs G+, modern drugs broad spectrum
- relatively polar - poor penetration
- safe and widely used
- resistance to cephalosporins potentially serious in people

