ID Tips of the Week repository, updated quarterly. For questions, comments, or suggestions, contact SCardwell@peacehealth.org. Prepared by ID/AMS Columbia Network: Andy Root, MD, Mike Conte, DO, Jose Rivera Sarti, MD, and Sophia Cardwell, PharmD, MPH. Recommendations incorporate published research and guidelines, local epidemiology, and diagnostic and therapeutic tools available at PeaceHealth.

Keywords (to aid searching/Ctrl+f):

General ID, bacteremia, UTI, pneumonia, diarrhea, GNRs, oral tx, labs, osteomyelitis, SSTI, DFI, Flu/COVID

7/17/23: Recommended duration of antibiotic therapy for bacteremia

With a few notable exceptions, the duration of treatment for bacteremia is not determined by the presence of bacteremia but rather the <u>source or associated infectious syndrome</u>. For example, if the bacteremia is secondary to pyelonephritis then the preferred duration would be 7 days (or 10 depending on which agent is used) because that's how long we treat pyelonephritis.

Exceptions:

- Staph aureus. Staph aureus bacteremia should be treated with at least 14 days of therapy, often 4-6 weeks. ID consult is recommended/expected and associated with improved outcomes.
- Candida (not a bacterium, but we're including it here). Candidemia should also be treated with at least 14 days of therapy. Again, ID consult is recommended/expected and associated with improved outcomes.
- Bacteremia secondary to a source for which a short course (≤5 days) of antibiotic therapy might otherwise be recommended (e.g. CAP or intra-abdominal infection with source control). In this case the duration of therapy should be individualized based on the organism and clinical factors.
- Organisms that commonly cause endocarditis (such as Enterococcus faecalis or viridans streptococci). A short course may be appropriate, but exercise caution here, especially when a focal source is not evident.

Keywords: bacteremia

7/24/2023: When should surveillance (follow-up) blood cultures be drawn?

Surveillance blood cultures should be drawn within 1-3 days of starting antibiotic therapy in the following scenarios:

- Staph aureus bacteremia
- Candidemia
- Patients with established or possible endovascular infection including endocarditis, central line infection, cardiac device infection, vascular graft infection, etc (note that sustained bacteremia can be an important diagnostic clue that one of these is present)
- Patients with bacteremia who are not improving as expected on antibiotic therapy

In most other scenarios surveillance blood cultures are unnecessary, low yield, and – if contaminants are detected – could potentially lead to unnecessary antibiotics and/or delay in discharge.

Keywords: bacteremia

7/31/2023: What is the recommended duration of therapy for UTI?

Unlike most infections this depends on both a specific characterization of the syndrome AND the antibiotic chosen.

Re: syndrome, a number of classifications are used, but we encourage dividing UTI into 2 distinct categories:

- **Cystitis**: Symptoms include dysuria, frequency, and urgency. This includes both "uncomplicated" and "complicated" cases (the latter often used to indicate abnormal anatomy or particular host factors). Cystitis predominantly occurs in women but can occur in men.
- **Pyelonephritis -OR- UTI with systemic signs/symptoms:** Symptoms include fever/chills and flank pain/tenderness with or without the cystitis symptoms mentioned above.
- Note that **catheter-associated UTI** can fall into either of these categories.

Here are recommended durations of therapy:

Cystitis

- TMP-SMX 3 days
- nitrofurantoin 5 days
- O IV β-lactam 3 days
- O PO β-lactam 5 days
- o fluoroquinolones 3 days
- o PO fosfomycin single dose
- IV aminoglycoside single dose

• Pyelonephritis -OR- UTI with systemic sign/symptoms

- o fluoroquinolones 5-7 days
- TMP-SMX 7-10 days
- O IV β-lactam 7 days
- IV \rightarrow PO β -lactam 7-10 days

Not included above are other syndromes including renal/perinephric abscess, prostatitis, and urethritis.

Keywords: UTI

8/8/2023: Treatment recommendations for infectious diarrhea

Rule number 1: infectious diarrhea most often should not be treated with antibiotics

Rule number 2: in the comparatively rare scenario where treatment is indicated, fluoroquinolones are not the drug of choice.

For patients with diarrhea and severe sepsis, treating empirically for sepsis including activity against common GI pathogens (e.g. ceftriaxone +/- metronidazole) is appropriate. Definitive therapy against pathogens of infectious diarrhea can be started with identification and/or sensitivity data.

It's preferable for patients with diarrhea and without sepsis to receive supportive care without antibiotics, due to lack of efficacy, potential harm over and above that of unnecessary antibiotic use, and promotion of antibiotic resistance. Here is a short summary of pathogen-specific recommendations:

Antibiotics	Antibiotics	Antibiotics	Special considerations antibiotic use	TOC if
routinely	specifically	generally		indicated
recommended?	harmful?	ineffective?		

Campylobacter	No	No*	Yes	Bloodstream infection; age over 65; pregnant; HIV with AIDS; receiving chemotherapy	Azithromycin ¹ ; susceptible beta-lactam
Giardia	Yes	No	No	N/A	Metronidazole
E. coli (STEC)	No	Yes ²	Yes	Do not use	N/A
E. coli (ETEC/EHEC)	No	No*	Yes	N/A	Ceftriaxone
Norovirus	No	No*	Yes	Do not use	N/A
Salmonella	No	Yes ³		Bloodstream infection; age over 65; immune compromised	Ceftriaxone
Shigella	No			Bloodstream infection; immune compromised; transmission risk high	Azithromycin ¹ , ceftriaxone ⁴

^{*}unnecessary antibiotics are always harmful, but in this case they don't worsen the disease, they are just not helping an/or causing their regular toxicities and adverse effects.

- 1. Caution using macrolides for bloodstream infections, wait for susceptibility or use a likely active beta lactam if need be
- 2. Antibiotics are recommended against due to their association with potentially fatal cases of hemolytic uremic syndrome, in addition to lack of efficacy.
- 3. Antibiotics are recommended against due to their association with prolonged bacterial shedding and transmission, in addition to lack of efficacy.
- 4. If use indicated, recommended to wait for sensitivities before starting antibiotics. Special concern for using fluoroquinolones even if reported susceptible due to undetected resistance

Keywords: diarrhea

8/14/2023: Antipseudomonal agent of choice

Cefepime is the preferred anti-pseudomonal beta lactam here at PeaceHealth. It has reliable activity against *P. aeruginosa*, improved activity against AmpC producing enterobacterales (*E. cloaceae*, *K. aerogenes*, and *C. freundii*) and less nephrotoxicity compared with piperacillin/tazobactam, and a higher threshold for development of resistance compared with meropenem. All three are approximately equivalent for incidence of *C. difficile* infection, although cefepime is likely slightly lower in risk than meropenem or piperacillin/tazobactam. All three are also approximately equivalent in their activity against common pathogens like *E. coli* and *Streptococcus* spp., and no more effective than ceftriaxone – if anti-pseudomonal activity is not required, ceftriaxone is generally preferred over any of the 3.

For patients requiring empiric activity against *P. aeruginosa*, cefepime +/- metronidazole is preferred over piperacillin/tazobactam or meropenem, with a few exceptions:

Situation	Alternative

Activity against both <i>P. aeruginosa</i> and <i>Enterococcus</i> spp.	Piperacillin/tazobactam
Presumed or confirmed ESBL-producing <i>E. coli</i> , <i>Proteus</i> , <i>K. pneumoniae</i> , or <i>K. oxytoca</i> spp.	Meropenem
Genuine allergy or intolerance to 3 rd or 4 th generation cephalosporins	Piperacillin/tazobactam or meropenem

Standard antipseudomonal doses are:

Cefepime 1g IV Q8H or 2g Q12H (2g Q8H is aimed at intermediately susceptible isolates, central nervous system penetration, etc.)

Piperacillin/tazobactam 3.375 g IV q8H via 4 prolonged infusion (4.5 g prolonged infusion is aimed at intermediately susceptible isolates)

Meropenem 1 g IV Q8H (2 g Q8H is aimed at central nervous system penetration)

Keywords: GNR

8/21/2023: Which antibiotics have excellent oral bioavailability?

A number of commonly prescribed antibiotics are highly bioavailable, meaning that they have excellent oral absorption and are considered equally effective when administered orally vs. intravenously.

They <u>should be administered orally</u> whenever the patient is able to swallow, has a functional gut, and is not on vasopressors.

Potential benefits of this approach include decreased costs, earlier discharge, and reduced complications associated with IV access.

Highly bioavailable antibiotics:

- fluoroquinolones (levofloxacin, ciprofloxacin, moxifloxacin)
- trimethoprim-sulfamethoxazole
- metronidazole
- azithromycin
- doxycycline
- linezolid
- clindamycin (if you can find a use for it!)
- fluconazole

Keywords: general ID

8/28/2023: How do I know whether a positive blood culture is a contaminant?

First, a refresher on the basics ...

Number of sets:

Typically 2 sets are obtained. This is preferable to a single set as it increases sensitivity and allows for greater ability to distinguish bacteremia from contamination. Drawing 3 sets (typically over a period of several hours) further increases sensitivity.

Number of bottles:

A blood culture set has 2 bottles, 1 aerobic & 1 anaerobic (most aerobic pathogens can grow in anaerobic bottles). When a blood culture is positive the result will indicate whether the aerobic, anaerobic, or both bottles are positive. If the patient is a difficult draw there may be only a single/pediatric bottle.

Timing of positivity:

Most clinically relevant cultures turn positive within 24-48 hours. Some organisms are fastidious (e.g. anaerobes) and may take longer to grow.

Gram stain:

When growth of bacteria is detected a gram stain is reported.

DNA identification panel:

This is a rapid PCR with a number of targets for various species of bacteria & fungi (it is not exhaustive so a positive culture could have a negative DNA ID panel). Like a gram stain, it is performed after growth is detected and is resulted quickly.

Back to the original question ...

The following features, particularly in combination, suggest that a positive result is potentially/likely a contaminant:

- Isolation of coagulase-negative staphylococci (other than *lugdunensis*), diphtheroids, *Bacillus* spp, *Micrococcus* spp, *Cutibacterium acnes*
 - Other streptococci & Enterococcus spp can also be occasional contaminants but exercise caution
- Only 1-2 (out of 4) bottles are positive
- Growth occurs late (after 48 hrs of incubation)
- Pre-test likelihood of bacteremia with the isolated organism is low

A few other points:

- Staph aureus, gram negative rods, beta-hemolytic streptococci (groups A, B, and C/G), Strep pneumoniae, & Candida are virtually never considered contaminants
- Clinical context is very important for example, growth of Staph epidermidis in a patient presenting with a
 possible catheter infection or prosthetic valve endocarditis is very different than in a patient presenting with
 pneumonia
- When in doubt as to whether a positive result may be a contaminant, repeating cultures (ideally before antibiotics are administered) can be extraordinarily helpful

Keywords: bacteremia

9/5/23: How to handle positive blood cultures - part 1

In keeping with the blood culture theme, here is a breakdown on how to handle positive blood cultures with staphylococci. These are general guidelines and exceptions exist – when in doubt please call ID.

DNA ID Panel Result	Organism	Significance	Initial Management	Preferred Antibiotic
Staphylococcus spp	coagulase	Usually a	Likely contaminant?	vancomycin
detected S. aureus not detected	negative staph	contaminant	> Do nothing	(if indicated)
S. epidermidis not detected S. epidermidis not detected S. lugdunensis not detected mecA/C & MREJ detected or not Staphylococcus spp detected	(unspecified species) Staph epidermidis	Consider Number of bottles positive Clinical context	Possibly/probably real? Repeat cultures FIRST Consider starting antibiotics though can often wait for results of follow-up cultures to confirm infection	Change to cefazolin OR nafcillin if susceptible
S. aureus not detected S. epidermidis				
detected				
S. lugdunensis not detected				
mecA/C & MREJ detected <u>or</u> not				
Staphylococcus spp	Staph	Can be a	Likely contaminant?	cefazolin 2g IV q8
detected S. aureus not detected	lugdunensis (methicillin	contaminant but often a pathogen	Consider drawing repeat blood	- OR -
S. epidermidis not	susceptible)	and can behave similarly to Staph	cultures	nafcillin 2g IV q4
detected		aureus	Possibly/probably real?	(if indicated)
S. lugdunensis detected		Consider	Repeat blood cultures	
mecA/C & MREJ <u>not</u> detected		Number of bottles	Consider starting antibiotics	
Staphylococcus spp detected	Staph Iugdunensis	positive • Clinical	Consider ID consult	vancomycin
S. aureus not detected	(possibly	context		(if indicated)
S. epidermidis not detected	methicillin resistant)			Change to cefazolin OR nafcillin if susceptible

S. lugdunensis detected mecA/C & MREJ detected				
Staphylococcus spp detected S. aureus detected S. epidermidis not detected S. lugdunensis not detected	MSSA	Always a pathogen	 Start antibiotics Evaluate for source and pursue source control Echocardiogram Draw surveillance cultures in 1-2 days Consult ID 	cefazolin 2g IV q8 - OR - nafcillin 2g IV q4
mecA/C & MREJ <u>not</u> detected				
Staphylococcus spp detected	MRSA			vancomycin
S. aureus detected S. epidermidis not detected S. lugdunensis not detected mecA/C & MREJ detected				If allergy/contraindication, daptomycin

Keywords: bacteremia

9/11/23: How to handle positive blood cultures – part 2

Last week it was staph, this week it's strep.

Again, these are general guidelines and exceptions exist – when in doubt please call ID.

DNA ID Panel Result	Organism	Significance	Management/Antibiotics
Streptococcus spp detected S. agalactiae not detected S. pneumoniae not detected	Unspecified streptococcus Likely S. dysgalactiae (group C or G) OR viridans strep	S. dysgalactiae is virtually always a pathogen viridans streptococci are usually pathogens but occasionally contaminants, consider	 If contaminant suspected → repeat cultures FIRST, consider starting antibiotics Antibiotic choice depends upon clinical context and syndrome Consider using ceftriaxone while awaiting species ID

S. pyogenes not detected		Number of bottles positiveClinical context	S. dysgalactiae is uniformly susceptible to penicillin and other beta-lactams
Streptococcus spp detected S. agalactiae not detected S. pneumoniae not detected S. pyogenes detected Streptococcus spp detected S. agalactiae detected S. pneumoniae not detected S. pneumoniae not detected S. pyogenes not detected	S. pyogenes (group A strep) S. agalactiae (group B strep)	Always a pathogen	 Start (or change to) active antibiotics Uniformly susceptible to penicillin and other betalactams cefazolin, penicillin, or ampicillin are preferred group A Strep only: clindamycin should be added for shock but not used as monotherapy
Streptococcus spp detected S. agalactiae not detected S. pneumoniae detected S. pyogenes not detected	S. pneumoniae (pneumococcus)		 Start (or change to) active antibiotics ceftriaxone 1g IV q24 preferred If meningitis, ceftriaxone 2g IV q12 PLUS vancomycin

Keywords: bacteremia

9/19/2023: Preferred empiric antibiotic therapy for diabetic foot infection

Remember:

- Most patients do NOT need empiric (up front) MRSA or *Pseudomonas* coverage
- Excessively broad coverage is often harmful, not just risk-neutral
- Do NOT obtain or feel compelled to act upon results of superficial/swab cultures
- Final antibiotic selection should be based on deep/tissue/surgical cultures
- Chronic infections may exhibit necrosis/gangrene on exam and gas on x-ray but these are distinct from necrotizing infection or gas gangrene which are acute and often rapidly progressive and accompanied by severe sepsis or shock

Scenario	Preferred empiric therapy	Comments
Absence of ischemia, necrosis, devitalized tissue, or sepsis	cefazolin 2g IV q8	Main pathogens are MSSA, streptococci, and coag-neg staph
Presence of ischemia, necrosis, devitalized tissue, and/or sepsis	ampicillin-sulbactam 3g IV q6 - OR - ceftriaxone 1g IV q24 +/- metronidazole 500mg PO/IV q8	Pathogens include above organisms plus enteric gram-negatives, anaerobes, & enterococci
Increased risk for Pseudomonas	cefepime 2g IV q12 +/- metronidazole 500mg PO/IV - OR - piperacillin-tazobactam 3.375g IV q8	 Indications for <i>Pseudomonas</i> activity: Necrotizing infection Recent positive culture from relevant site
Increased risk for MRSA	Above - PLUS - vancomycin IV per pharmacy	 Indications for MRSA activity: Requiring vasopressors Necrotizing infection Recent positive culture from relevant site

Keywords: DFI

9/25/23: Interpretation of C diff test results

C. difficile testing at PeaceHealth employs a strategy of initial NAAT (DNA) testing. If negative, no further testing is done. If positive, this reflexes to toxin EIA testing. NAAT testing is highly sensitive though lacks specificity (positive result indicates presence of toxigenic strain but not toxin itself). Toxin EIA testing is highly specific to infection but only about 75% sensitive, so false negatives can occur.

Reminder that C diff testing should only be ordered when ALL of the following criteria are met

- Acute onset diarrhea
- 3+ liquid/loose stools within 24 hours
- No evident alternative explanation (another illness, medication, laxatives, etc)

Here's a breakdown of how to interpret results:

RESULT	INTERPRETATION	ACTION
C diff DNA negative	C diff infection ruled out	Do not treat for C diff infectionDo not repeat testing
C diff DNA positive Toxin EIA negative	C diff infection is possible (vs. colonization)	➤ If clinical suspicion low → favor observation
TOXIII LIA Hegative	(vs. colonization)	

		AA	If clinical suspicion high \rightarrow consider treatment for C diff infection If treatment does not lead to clinical improvement, reconsider diagnosis
C diff DNA positive	C diff infection confirmed	\wedge	Treat for C diff infection
Toxin EIA positive			

Keywords: diarrhea

10/5/23: FAQ on pneumonia, Part 1

Guidance re: pneumonia therapy is generally broken down into **CAP** (community acquired pneumonia) and **HAP** (hospital acquired pneumonia). There are other syndromes that might require modified approaches, including necrotizing pneumonia, empyema, lung abscess, and aspiration pneumonitis.

Many of you may also be familiar with the designation <u>HCAP</u> (healthcare-associated pneumonia), which was included in prior versions of guidelines and applied to some patients who were thought to be at elevated risk for pneumonia due to resistant organisms (including MRSA and *Pseudomonas*). <u>This designation has been abandoned</u> as years of data demonstrated that this approach was both ineffective and harmful. Anyone who develops pneumonia while not admitted to a hospital is considered to have CAP.

What is the preferred empiric therapy for CAP?

ceftriaxone 1g IV q24

+/- azithromycin 500mg PO/IV q24

What are alternatives in the setting of allergy/intolerance/contraindication?

For ceftriaxone: ampicillin-sulbactam 3g IV q6

For azithromycin: doxycycline 100mg PO/IV q12

What about levofloxacin?

Quinolones have a much less favorable safety profile as compared to beta-lactams. Levofloxacin should only be used when a patient truly cannot tolerate the above agents OR in the rare instance that anti-Pseudomonal therapy is warranted (we'll address this in an upcoming tip).

What is the recommended duration of therapy for CAP?

5 days

When should therapy potentially be extended beyond 5 days?

- Lack of clinical improvement after 72 hrs (e.g. still febrile)
- Parapneumonic effusion (maybe)
- Empyema
- Lung abscess
- Necrotizing pneumonia

What should be used for stepdown oral therapy? (assumes that a specific culprit organism was not identified)

- amoxicillin 500mg PO TID (preferred)
- amoxicillin-clavulanate 875mg PO BID
- cefpodoxime 200mg PO BID (non-preferred; reserve for penicillin allergy)
- doxycycline 100mg PO BID

Keywords: pneumonia

10/11/23: FAQ on pneumonia, Part 2

MRSA & *Pseudomonas* (and other resistant gram negatives) are rare pathogens in community acquired pneumonia (CAP). Additionally, use of drugs active against MRSA & *Pseudomonas* is associated with risk of drug resistance, AKI, *C. difficile* infection, secondary infection, prolonged length of stay, and death. These drugs should only be used when there is reasonable likelihood of benefit.

So ...

Which patients with CAP should receive empiric MRSA coverage?

- Prior isolation of MRSA from respiratory culture (not including MRSA nasal screen)
- Necrotizing or cavitary pneumonia
- Empyema, if severely ill (e.g. ICU)

Which patients with CAP should receive empiric Pseudomonas coverage?

- Prior isolation of Pseudomonas from respiratory culture
- Underlying structural lung disease, if severely ill (e.g. ICU)

In all cases, therapy should be tailored to results of microbiologic results (sputum gram stain & culture, respiratory virus testing, etc). If no resistant organisms are found, revert to standard therapy (e.g. ceftriaxone alone).

MRSA nasal screening should only be utilized in patients who are *appropriately* started on empiric MRSA therapy based on the criteria above. In those cases a negative result should lead to de-escalation of therapy. While MRSA nasal screening has good negative predictive value in the proper setting, there is no setting where positive predictive value has been demonstrated.

Keywords: pneumonia

10/17/2023: FAQ on pneumonia, Part 3

What is the preferred empiric therapy for hospital acquired pneumonia (HAP) & ventilator associated pneumonia (VAP)?

cefepime 2g IV q12 for HAP or q8 for VAP

+/- vancomycin IV per pharmacy

What are alternatives in the setting of allergy/intolerance/contraindication?

For cefepime: piperacillin-tazobactam 3.375g IV q6

For vancomycin: linezolid 600mg PO/IV q12

When is it most important to include empiric MRSA coverage (with vancomycin or linezolid)?

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- Prior isolation of MRSA from respiratory culture
- Necrotizing or cavitary pneumonia
- Empyema

Keywords: pneumonia

What is the recommended duration of therapy for HAP/VAP?

7 days

When should we double-cover for resistant gram negatives?

Almost never. Fortunately our local rates of resistance are low enough that double coverage is only indicated in the rare severely ill patient with a history of highly-resistant *Pseudomonas* or other gram negatives. ID or ID pharmacy guidance should be requested in these scenarios.

Keywords: pneumonia

10/23/23: The MRSA nares PCR

The MRSA nares PCR has poor positive predictive value and should not be used to start or maintain therapy that is not otherwise indicated. It has good negative predictive value for pneumonia in low-prevalence settings, meaning its utility is in *stopping* recommended empirical therapy.

Predictive values are closely related to disease prevalence, so no test will have good positive predictive value in a low-prevalence setting. MRSA pneumonia rates range from <1 to 10%, depending on the study. Assuming the highest, a 10% population rate of MRSA pneumonia, positive and negative predictive values of the MRSA nasal PCR are:

	PPV	NPV
All pneumonia	44.8%	96.5%
VAP	35.7%	94.8%

When ordering a test with good negative predictive value in a low prevalence setting, it's important to consider whether the test is needed at all/what other tools are available to answer the question the test is purporting to answer. To illustrate: a pregnancy test for a male has excellent negative predictive value, but a positive result doesn't indicate you need to start prenatal vitamins, and you have other ways of knowing the patient is not pregnant.

So when should you use the MRSA PCR?

If you have a patient with pneumonia for whom empiric vancomycin is recommended (prior isolation of MRSA from a respiratory culture [not prior nasal PCR], necrotizing or cavitary pneumonia, empyema), the MRSA PCR has sufficient (>90%) negative predictive value to stop vancomycin.

When should you NOT use the MRSA PCR?

- To continue vancomycin that was started for a patient without relevant risk factors for MRSA. Stopping vancomycin regardless of the nasal PCR is appropriate in this setting.
- To aid in your decision to start MRSA therapy for any indication
- To aid in your decision to start or stop MRSA therapy for non-pulmonary infections

The decision to include anti-MRSA treatment should be made based on clinical status and relevant risk factors, for which there are clearly defined and evidence-based recommendations, regardless of the MRSA nasal PCR.

Keywords: labs, general ID, pneumonia, SSTI

10/21/23: Here are some SCARY <spooky ghost sound!> stats about antibiotic (mis)use and resistance:

- 33-50% of all antibiotic prescriptions are unnecessary or inappropriate (this includes inpatient and has remained stable for more than a decade)
- Inappropriate prescribing is the leading factor driving antibiotic resistance (overly broad therapy, unnecessary combo therapy, excessive duration, wrong dose)
- ~1 in 5 infections in wealthy countries are caused by resistant bacteria
- Antibiotic resistance killed nearly 5 million people in 2019 (the last year this analysis was done)
- In the US, 2.8 million antibiotic resistant infections occur annually, killing more than 35,000 people (data from 2012, current numbers likely substantially higher) and costing nearly \$5 billion

On a positive note: since inappropriate prescribing is the main driver of these problems, using antibiotics wisely (right drug at the right dose for the right duration and only when necessary) goes a long way towards solving them.

Credit to our ID pharmacist, Sophia Cardwell, for the stats.

Keywords: general ID

11/9/2023: Aspiration pneumonitis v. pneumonia:

- Approximately 25% of patients with aspiration pneumonitis will progress to pneumonia, regardless of administration of prophylactic (within the first 2 days of aspiration) antibiotics
- For aspiration pneumonitis, observing patients off antibiotics is (and should be) the standard of care, as antibiotics do not affect progression to pneumonia
- Patients with aspiration pneumonitis who are given prophylactic antibiotics are more likely to require escalation
 of antibiotics should pneumonia develop, due to selection of resistant pathogens

Aspiration		Aspiration pneumonitis		Aspiration pneumonia		Pulmonary abscess
Aspiration events (inhalation of gastric or oropharyngeal contents) are common, and not infectious.	→ Based on volume, pH, etc.	An acute inflammatory response in the first 48 hours after aspiration event. May have \triangle WBC, fever, SOB. Not infectious.	→ ~25% develop pneumonia, whether you give antibiotics or not	Pneumonia (standard dx criteria) 2-7 days post aspiration. Infectious. Treat as CAP (e.g. no metronidazole)	→ Takes weeks to months	Chronic, rather than acute, may need to include metronidazole, may require surgery and prolonged treatment.
Observe off antibiotics – giving prophylactic antibiotics here does not				Ceftriaxone m pathogens	onotherapy are oral ana	

change the rate of progression to pneumonia, and causes specific harm.

| Bacteroides spp. May add metronidazole for abscesses with Fusobacterium spp.

| Infection | Infectio

Keywords: pneumonia

11/9/23: Gram negative resistance: AmpC beta-lactamase

<u>Cefepime</u> is the preferred agent for treatment of serious infection with *Citrobacter freundii, Enterobacter cloacae,* & *Klebsiella aerogenes*.

With the exception of cystitis and non-severe soft tissue infection, ceftriaxone, piperacillin/tazobactam, and oral cephalosporins & penicillins should be avoided <u>regardless of susceptibility testing results</u>. With susceptibility ciprofloxacin, levofloxacin, trimethoprim/sulfamethoxazole, and if cystitis, nitrofurantoin can be considered as alternatives to cefepime or if transitioning to oral therapy.

Background/explanation:

Citrobacter freundii, Enterobacter cloacae, and Klebsiella aerogenes all reliably carry genes for inducible resistance on treatment with 3rd generation cephalosporins and piperacillin/tazobactam via production of the <u>AmpC beta lactamase</u>. Current technology is unable to detect the AmpC beta lactamase prior to emergence of resistance, making susceptibility data from initial cultures misleading. Susceptibility reports for systemic infections to piperacillin/tazobactam and ceftriaxone should NOT be used to make treatment decisions. Cefepime is reliably active against AmpC producing enterobacterales, and an MIC of ≤2 affirms susceptibility to cefepime.

More info here: https://www.peacehealth.org/sites/default/files/2023-03/PeaceHealth-AmpC.pdf

Keywords: GNR

11/28/23: Gram negative resistance: extended-spectrum beta-lactamase (ESBL)

ESBL-producing *E. coli, Klebsiella pneumoniae/oxytoca*, or *Proteus* can be identified by resistance to ceftriaxone or ceftazidime OR by detection of CTX-M gene in blood isolates.

Serious or systemic infection

- Meropenem is preferred
- Oral stepdown therapy with SMX-TMP, ciprofloxacin, or levofloxacin may be considered (if susceptible)

Uncomplicated cystitis

- Nitrofurantoin or SMX-TMP are preferred
- Alternatives
 - Single dose aminoglycoside
 - o Ciprofloxacin or levofloxacin for 1-3 days
 - Single dose fosfomycin (E. coli only)
 - Meropenem (if the only susceptible agent)
- Patients initially started on cefepime or piperacillin/tazobactam who are clinically improving may complete treatment without change or extension
- Doxycycline & amoxicillin/clavulanate should NOT be used (regardless of susceptibility results)

ID consult is encouraged for bacteremia and other serious systemic infections.

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Keyword: GNR

12/28/23: Influenza & oseltamivir

For inpatients with influenza, antiviral treatment with <u>oseltamivir should be started as early as possible</u>.

This recommendation applies <u>regardless of timing of symptom onset</u>. There is good evidence supporting oseltamivir for any hospitalized patient with influenza to reduce death and duration of hospitalization, <u>especially when started within 48 hours from admission</u>. Withholding oseltamivir for inpatients based on symptom duration is NOT recommended and may be harmful. Even if delayed until hospital day 3 or later, it is still officially recommended wherever possible.

1/2/24: Influenza & bacterial pneumonia

How common does secondary bacterial pneumonia complicate influenza?

Uncommonly! About 1-3% of cases (in a non-pandemic year), skewing a bit higher in hospitalized patients.

When should I suspect secondary bacterial pneumonia?

- Biphasic illness, e.g. relapsed fever after prior improvement
- Persistent fever after 3-5 days
- Purulent sputum production
- Lobar consolidation

What are the most common pathogens?

- Strep pneumoniae
- Staph aureus (MSSA & MRSA)

How should I evaluate patients with suspected secondary bacterial pneumonia?

- Repeat chest imaging
- Sputum gram stain & culture (tracheal aspirate or BAL if intubated)
- Nasal MRSA PCR if starting vancomycin

What empiric antibiotics should be used if I suspect secondary bacterial pneumonia?

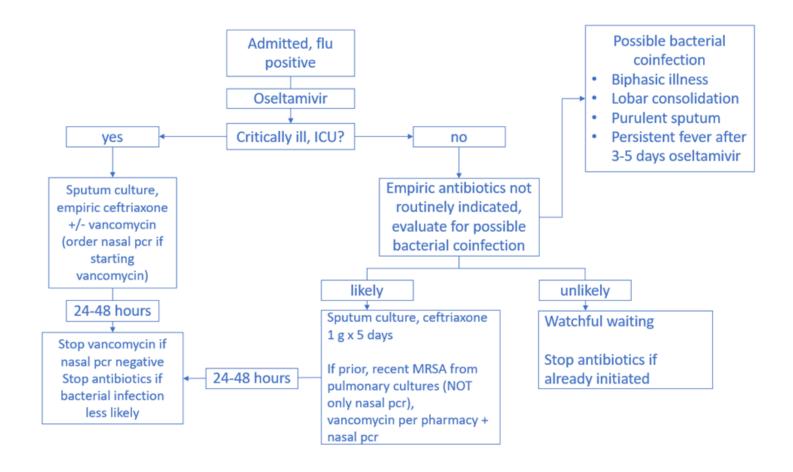
- ceftriaxone 1g IV q24
- +/- vancomycin IV per pharmacy *IF* critically ill or MRSA recently isolated from respiratory culture (not just nasal PCR)

What should I NOT do in this scenario?

- Routinely start antibiotics in patients with influenza
- Prescribe atypical agents (e.g. azithromycin)
- Prescribe anti-Pseudomonal agents (e.g. cefepime)

Keywords: Flu/COVID, pneumonia

Prefer a flow-chart? Here you go!



1/26/24: Antibiotic management of hematogenous osteomyelitis

<u>Hematogenous</u> osteomyelitis (most commonly vertebral osteomyelitis-diskitis with/without epidural abscess) is a distinct syndrome from <u>contiguous</u> osteomyelitis (e.g. diabetic foot infection). Hematogenous ostemyelitis is most commonly mono-microbial, and *Staphylococcus* spp. (MSSA, MRSA, CoNS) are the most frequently identified pathogens with gram negative organisms are less common. It's essential that the culprit pathogen be isolated in order to ensure successful management. Accordingly, <u>antibiotics should NOT be started reflexively</u>.

Here is a recommended approach to initial management:

Stable patients →

- HOLD ANTIBIOTICS until cultures can be obtained surgically or percutaneously (in addition to blood cultures)
- Once cultures obtained, start vancomycin IV per pharmacy +/- ceftriaxone 1-2g IV q24

Stable patients with positive blood cultures ->

• Assuming that a contaminant is not suspected, start antibiotics targeting the isolated organism

Patients with hemodynamic instability OR neurologic deficits >

Start vancomycin IV per pharmacy +/- ceftriaxone 1-2g IV q24

ID consult is strongly encouraged for these cases.

Keywords: osteomyelitis

2/7/2024: Antibiotic management of contiguous osteomyelitis

As mentioned with our last tip, this is a distinct syndrome from hematogenous osteomyelitis.

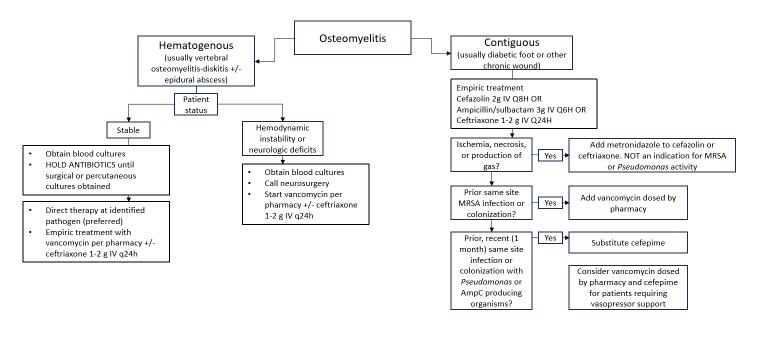
We most commonly see contiguous osteomyelitis in the setting of diabetic foot infection, though the same general principles apply to other scenarios.

- Bone cultures (e.g. with surgical debridement) are STRONGLY preferred over soft tissue or superficial cultures (as the latter will often pick up irrelevant superficial colonizers)
- Streptococci (group A/pyogenes and B/agalactiae) and MSSA are the most common pathogens
 - o Enteric pathogens like E. coli are less common
 - MRSA, Pseudomonas, & anaerobes are unlikely in absence of specific risk factors
- Most patients DO NOT require MRSA or Pseudomonas coverage up front
 - o Antibiotics targeting these organisms are no more effective for the most common pathogens
 - o Starting these antibiotics unnecessarily is more likely to be harmful than beneficial
 - You have plenty of time to start these later if indicated based on cultures

Scenario	Preferred empiric therapy				
Absence of ischemia, necrosis, gas, devitalized tissue, or sepsis	cefazolin 2g IV q8				
Presence of ischemia, necrosis, devitalized tissue, gas, and/or sepsis	ampicillin-sulbactam 3g IV q6 - OR -				
	ceftriaxone 1g IV q24				
	+/- metronidazole 500mg PO/IV q8				
Increased risk for <i>Pseudomonas</i> or other resistant	cefepime 2g IV q12				
gram negatives	+/- metronidazole 500mg PO/IV				
 Recent (~1 month) same site culture with these organisms 	- OR -				
Critically ill patients (e.g. requiring pressors)	piperacillin-tazobactam 3.375g IV q8				
Increased risk for MRSA	Above				
Recent (~1 month) same site culture with MRSA	- PLUS -				
Critically ill patients (e.g. requiring pressors)	vancomycin IV per pharmacy				

Keywords: osteomyelitis, DFI

2/16/24: Differential antibiotic management of hematogenous v. contiguous osteomyelitis



Hematogenous osteomyelitis is a distinct syndrome from contiguous osteomyelitis. It is likely monomicrobial; *Staphylococcus* spp. most common. Gram negative pathogens less likely. Antibiotics should be held before appropriate cultures obtained wherever possible.

Vancomycin is a mainstay of empiric treatment.

Contiguous osteomyelitis is a distinct syndrome from hematogenous osteomyelitis. Beta hemolytic strep (group A and B, or *S. pyogenes* and *S. agalactiae*) and MSSA are most common. Enterics like *E. coli* are less common, and MRSA, *Pseudomonas*, and strict anaerobes are unlikely without specific risk factors. Vancomycin, cefepime, and piperacillin/tazobactam should be avoided for routine empiric treatment.

Keywords: osteomyelitis, DFI

2/26/24: Management of sacral decubitus ulcers

(Sticking with the recent theme of osteomyelitis ...)

The mainstay of therapy for sacral decubitus ulcers is local wound care and off-loading.

If there is evidence of local soft tissue infection, empiric antibiotics targeting skin & enteric flora are appropriate. Reasonable options include ampicillin-sulbactam OR ceftriaxone +/- metronidazole. Anti-MRSA & anti-pseudomonal agents should be avoided for routine empiric therapy.

Imaging studies, including MRI, often over-estimate the presence of osteomyelitis and cannot reliably distinguish osteomyelitis from bone remodeling.

There is no evidence of benefit of antibiotic therapy for sacral osteomyelitis associated with sacral decubitus ulcers unless concomitant surgical debridement and wound coverage is performed. In that scenario, a course of culture-directed antibiotic therapy is appropriate following closure.

Unfortunately, to our knowledge, there are no surgeons in SW Washington that perform these surgeries. As such, patients are generally referred to an academic center and can be maintained with wound care, off-loading, nutritional optimization and – when appropriate – a short course of antibiotics for soft tissue infection until evaluated by a surgeon.

Keywords: osteomyelitis

3/15/24: How to use this year's antibiogram, gram negative edition! See here

https://www.peacehealth.org/pages/providers-and-medical-professionals/resources/infectious-diseases or instructions for Epic link in any micro result below (using SW rather than SJ given it has more unique isolates):

Species identification on blood pcr/preliminary culture should be used with the antibiogram to focus antibiotics for responding patients

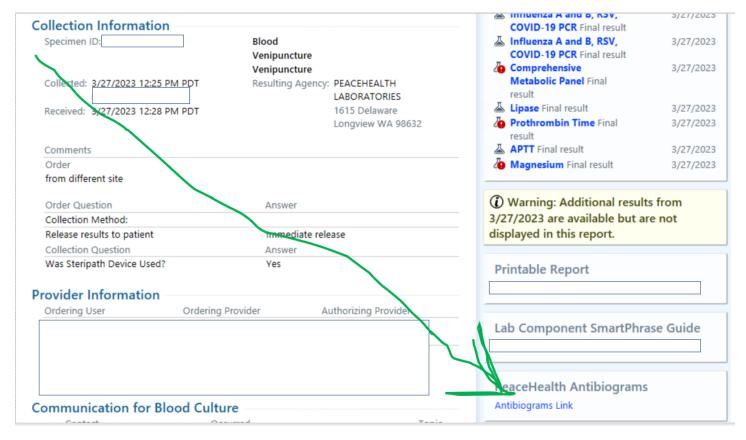
- 1. If GNRs identified, stop vancomycin wherever possible
- 2. If cefepime, piperacillin/tazobactam, or meropenem is ordered, change to ceftriaxone for E. coli, C. koseri, K. oxytoca, K. pneumoniae with no CTX-M gene identified
 - a. CTX-M is an accurate predictor of ceftriaxone non-susceptibility, if positive, meropenem should be used, if negative, ceftriaxone is appropriate
- 3. If ceftriaxone or piperacillin/tazobactam is ordered, change to cefepime for AmpC producers C. freundii, E. cloacae, K. aerogenes

A note on piperacillin/tazobactam: it should not routinely be used for empiric therapy in our population. It is not reliably active against AmpC producers, which are more common than P. aeruginosa. It is no better than ceftriaxone for E. coli and other non AmpC producing enterobacterales; ceftriaxone resistance precludes the use of piperacillin/tazobactam for these, regardless of final susceptibility. It is less reliable for P. aeruginosa than cefepime, and it has a much higher risk of nephrotoxicity than any cephalosporin.

Further definitive treatment with susceptibility data (narrow spectrum, PO, stop date) should still happen 😊

						Beta-lactams							oro-	Amir	oglycos	idos	Miscellaneous			
		P	enicillir	\$			Cephal	osporin	s	Carbap	enems	quino	lones	Amir	logiyeo:	sides	IVIE	cenane	ous	
GRAM NEGATIVE ORGANISM PERCENT SUSCEPTIBLE	# Isolates Tested	Ampicillin	Amoxicillin/clavulanic acid	Piperacillin/tazobactam	Cefazolin - urine isolates only	Cefoxitin	Ceftriaxone	Ceftazidime	Cefepime	Ertapenem	Meropenem	Ciprofloxacin	Levofloxacin	Amikacin	Gentamicin	Tobramycin	Nitrofurantoin - urine only	Tetracycline	Trimethoprim/sulfamethoxazole	
Citrobacter freundii	63	R	R	-	-	R	-	-	100	98	100	97	95	100	97	98	94	98	97	
Citrobacter koseri	30	R	100	100	-	87	100	100	100	100	100	100	97	100	100	100	-	97	100	
Enterobacter cloacae	138	R	R	-	-	R	-	-	96	95	100	94	93	100	99	98	52	87	93	
Escherichia coli	1529	61	88	98	88	94	92	97	98	100	100	84	79	100	94	95	98	81	81	
Klebsiella aerogenes	51	R	R	-	-	R	-	-	100	98	100	96	98	100	100	98	17	94	98	
Klebsiella oxytoca	75	R	92	97	-	97	91	96	100	100	100	96	95	99	95	95	82	90	93	
Klebsiella pneumoniae	333	R	92	97	87	97	89	93	97	99	99	88	88	100	95	92	35	84	89	
Morganella morganii	35	R	R	97	-	51	89	74	97	100	100	71	71	100	91	100	-	50	74	
Proteus mirabilis	201	74	85	100	93	91	95	99	100	100	100	82	82	100	94	96	R	R	79	
Pseudomonas aeruginosa	295	R	R	89	-	-	R	91	91	R	99	91	84	99	-	100	-	-	R	
Serratia marcescens	45	R	R	98	-	R	93	100	100	100	100	93	91	100	98	87	R	41	100	
Stenotrophomonas maltophilia	56	R	R	R	-	-	R	-	-	R	R	-	95	R	R	R	-	-	100	

NOTE: Third generation cephalosporins and piperacillin/tazobactam are unreliable for systemic infections with AmpC producing organisms (E. cloacae, K. aerogenes, or C. freundii), regardless of susceptibility data, due to the development of induced resistance on treatment.



Keywords: general ID, labs

3/25/24: How to use this year's antibiogram, gram positive edition! See here

https://www.peacehealth.org/pages/providers-and-medical-professionals/resources/infectious-diseases or instructions for Epic link in any micro result below (using SW rather than SJ given it has more unique isolates):

Species identification on blood pcr/preliminary culture should be used with the antibiogram to focus antibiotics for responding patients

- 1. If common pathogens other than MRSA identified, stop vancomycin wherever possible (GNRs, MSSA, strep, most coagulase negative staph and enterococcus)
- 2. If MSSA identified, change to cefazolin or nafcillin wherever possible, there is a mortality/morbidity benefit for these agents for MSSA compared with vancomycin or ceftriaxone
- 3. If beta hemolytic strep identified (*S. pyogenes, S. agalactiae*, *S. dysgalactiae*), change to cefazolin these organisms are uniformly susceptible
 - a. Clindamycin is more harmful than beneficial except for infection with *S. pyogenes* in patients requiring vasopressors
- 4. If *Enterococcus faecalis* identified, change to an ampicillin containing regimen wherever possible; cephalosporins do not have enterococcal activity

A note on vancomycin: it is inferior to beta lactams for treatment of common pathogens other than MRSA/MRSE (MSSA, strep, and enterococcus), from both an efficacy and safety standpoint. It's used for MRSA largely because the alternatives are even worse than vancomycin, but it is neither safe nor particularly effective when used for treating infections where beta lactam antibiotics are active. Because of its unfavorable safety profile, vancomycin should only be

used empirically for infectious disease states where MRSA (or more rarely coagulase negative staphylococcus) are likely pathogens, unless there is a genuine contraindication to preferred therapy.

Further definitive treatment with susceptibility data (narrow spectrum, PO, stop date) should still happen 😊



			Be	ta lactar	ns		Flu	oro-	Am	ino-				N. Alice	scellane	OLIE			
		P	enicillins	/Cepha	losporin	5	quino	lones	glyco	sides				IVIII	scenarie	ous			
GRAM POSITIVE ORGANISM PERCENT SUSCEPTIBLE		Penicillin	Oxacillin	Ampicillin	Cefazolin	Ceftriaxone	Ciprofloxacin	Levofloxacin	Gentamicin Synergy	Streptomycin Synergy	Clindamycin	Daptomycin	Erythromycin	Minocycline	Nitrofurantoin - urine only	Rifampin	Tetracycline	Trimethoprim/sulfamethoxazole	Vancomycin
Enterococcus faecalis	300	-	-	100	-	-	-	-	83	96	-	-	-	-	99	-	-	-	100
Enterococcus faecium	34	-	-	29	-	-	-	-	-	-	-	-	-	-	-	-	-	-	64
Staphylococcus aureus , MRSA	318	-	0	-	0	-	-	-	-	-	79	100	-	100	-	100	74	99	100
Staphylococcus aureus , MSSA	696	-	100	-	100	-	-	-	-	-	87	-	-	100	-	-	92	100	-
Staphylococcus epidermidis	89	-	42	-	42	-	-	-	-	-	62	-	-	100	-	-	74	63	100
Staphylococcus lugdunensis	47	-	89	-	89	-	-	-	-	-	84	-	-	100	-	-	100	98	100
Streptococcus agalactiae (Group B)	89	100	-	-	100	100	-	-	-	-	65	-	49	-	-	-	-	-	100
Streptococcus anginosus	92	98	-	-	-	99	-	-	-	-	-	-	-	-	-	-	-	-	100
Streptococcus constellatus	43	91	-	-	-	93	-	-	-	-	-	-	-	-	-	-	-	-	100
Streptococcus intermedius	38	100	-	-	-	100	-	-	-	-	-	-	-	-	-	-	-	-	100
Streptococcus mitis/oralis	57	75	-	-	-	100	-	98	-	-	-	-	-	-	-	-	88	-	100
Streptococcus pneumoniae	70	100*	-	-	-	99*	-	100	-	-	-	-	74	-	-	-	81	-	-
Streptococcus pyogenes (Group A)	117	100	-	-	100	100	-	-	-	-	71	-	71	-	-	-	-	-	100

Keywords: general ID, labs, bacteremia

4/8/24: Ceftriaxone remains our antibiotic of choice for empiric treatment of most community acquired infections.

This includes (but is not limited to) community-acquired pneumonia, pyelonephritis, intra-abdominal infection, and diabetic foot infection.

Local rates of resistant organisms (e.g. MRSA, Pseudomonas, other resistant gram negatives) remain low and predictable based on history and site of infection. They are unlikely pathogens in the above scenarios.

Anti-pseudomonal agents (including cefepime, piperacillin-tazobactam, quinolones) and vancomycin are more toxic, less effective against more likely pathogens, or both.

Keywords: general ID

5/15/2024: Individual species of coagulase negative staphylococci (CoNS) are now being reported in microbiology results.

SC 10_2025

This is a change from our previous process of identifying them only as 'Coagulase negative *Staphylococcus* species, not *Staphylococcus luqdunensis*'.

For example, what would have previously been reported as:

Isolated from the anaerobic bottle Coagulase negative Staphylococcus species, not Staphylococcus lugdunensis!! (Critical)

May now appear as:

Isolated from the anaerobic bottle Staphylococcus hominis!! (Critical)

Here's what you should know/remember about CoNS:

- They make up ~25% of all blood isolates; species include *Staph epidermidis, lugdunensis, haemolyticus, simulans, auricularis, capitis, hominis*
- They usually represent blood culture contamination
- <u>Staph lugdunensis</u> is the one exception while it can also be a contaminant, it is more likely to cause disease and behaves similarly to Staph aureus
- Presence or absence of the MecA gene is irrelevant with respect to significance of a positive culture

If you think a patient may have a "real" CoNS bacteremia, the best course of action is to <u>collect more blood cultures</u> <u>before starting antibiotics</u>.

A true bacteremia is more likely present with one or more of the following:

- Multiple (typically > 2) bottles positive with the same organism (species and susceptibility pattern)
- Risk factors, especially a central line or other endovascular device/hardware
- Fever and other signs/symptoms of infection without a more likely explanation

Keywords: bacteremia, labs

6/13/2024

Management of diabetic foot infection, Part 1

Management of diabetic foot infection includes a decision regarding empiric (up-front) antibiotics as well as evaluation to determine pathogens involved, depth of infection, concurrent vascular disease and need for revascularization, and surgical debridement (or amputation). Subsequent management decisions include interpretation of culture results, adjustment of antibiotic therapy, and decision re: intensity (e.g. IV vs PO) and duration of antibiotic therapy.

For empiric antibiotic therapy, remember:

- Most patients do NOT benefit from empiric (up front) MRSA or Pseudomonas coverage
- The presence or absence of osteomyelitis should not alter antibiotic selection (for example, osteomyelitis does not necessitate broader therapy)
- Excessively broad coverage is harmful
- Do NOT obtain or feel compelled to act upon results of superficial/swab cultures
- Chronic infections may exhibit necrosis/gangrene on exam and gas on x-ray these changes are distinct from necrotizing infection or gas gangrene which are acute and often rapidly progressive and accompanied by severe sepsis or shock

Scenario	Preferred empiric therapy	Comments
Absence of ischemia, necrosis, devitalized tissue, or sepsis	cefazolin 2g IV q8	Likely pathogens are MSSA, streptococci, and coagulase-neg staph
Presence of ischemia, necrosis, devitalized tissue, and/or sepsis	ampicillin-sulbactam 3g IV q6 - OR - ceftriaxone 1g IV q24 +/- metronidazole 500mg PO/IV q8	Pathogens include above organisms plus enteric gram-negatives, anaerobes, & enterococci
Increased risk for Pseudomonas	cefepime 2g IV q12 +/- metronidazole 500mg PO/IV - OR - piperacillin-tazobactam 3.375g IV q8	 Indications for <i>Pseudomonas</i> activity: Necrotizing infection Recent positive culture from relevant site
Increased risk for MRSA	Above - PLUS - vancomycin IV per pharmacy	 Indications for MRSA activity: Requiring vasopressors Necrotizing infection Recent positive culture from relevant site

Keywords: DFI

7/3/2024

Management of diabetic foot infection, Part 2

Aside from selection of empiric (up-front) antibiotic therapy there are a number of other management considerations and principals to follow.

Initiation of antibiotic therapy

- For <u>patients with osteomyelitis who are stable</u> (afebrile, without significant cellulitis, abscess, or septic arthritis) it is best to <u>hold antibiotics</u> until deep/bone cultures are obtained (e.g. surgically)
- For all other patients, it is reasonable/appropriate to start antibiotics while also obtaining cultures as quickly as possible so as to maximize yield
- Remember that most patients should NOT receive empiric anti-MRSA or anti-Pseudomonas therapy (see table below)

Cultures

- Do NOT obtain specimens for culture by swabbing a superficial wound (studies show superficial wound swabs do not correlate with deeper/tissue/bone samples)
- Cleanse and debride wounds before obtaining any specimen for culture
- For suspected osteomyelitis of the foot, collect a sample of bone (percutaneously or surgically) for culture and histopathology

Additional evaluation & management

- Most patients, including any with evidence of ischemia, necrosis, devitalized tissue, and/or sepsis, should be evaluated by podiatry/orthopedics
- Consider vascular studies +/- vascular surgery evaluation
- Consider wound care team and diabetes educator evaluations

Keywords: DFI

7/17/2024

Management of diabetic foot infection, Part 3

Determining definitive antibiotic therapy

- Final antibiotic selection can be made based on clinical improvement, relevant culture results, and upon completion of any surgical procedures
- If started initially, drugs targeting MRSA, *Pseudomonas*, or other resistant gram negatives should be discontinued if these organisms are not isolated from relevant cultures
- Patients without osteomyelitis can be treated with oral antibiotics (unless cultures dictate otherwise, e.g. quinolone-resistant *Pseudomonas*)
- Patients with osteomyelitis may require IV antibiotics for the duration of treatment and should typically be managed with ID assistance

Scenario	Typical Duration of Therapy
Curative amputation without residual soft tissue infection	24-48 hours post-op
Soft tissue infection	10-14 days
Residual osteomyelitis following debridement/amputation	3-6 weeks
Osteomyelitis managed non-operatively	6 weeks

Keywords: DFI

8/2/2024

Hi everyone,

Along the recent tip of the week theme of Diabetic Foot Infections, wanted to make you aware of the recent ID related State of the Art Reviews (StAR) being written and published in the Clinical Infectious Disease (CID) journal.

There is a great review on Diabetic Foot Infections here:

<u>Evaluation and Management of Diabetes-related Foot Infections | Clinical Infectious Diseases | Oxford Academic (oup.com)</u>

Other topics published thus far include Staph aureus bacteremia, Enterococcal bacteremia, Prosthetic Joint Infection, Neurosyphilis, and Encephalitis. The homepage can be found here:

State-of-the-Art Reviews | Clinical Infectious Diseases | Oxford Academic (oup.com)

These are intended for ID, Non-ID providers, and trainees alike. Maybe the easiest way to digest this is likely in its Podcast form, via the highly regarded Febrile podcast (should be a free podcast in all app stores):

febrile - A Cultured Podcast About All Things Infectious Disease (febrilepodcast.com)

Keywords: DFI, general ID

8/19/2024

Hello,



For this week, please find the attached document for Diabetic Foot Infection management I created with the help of the ID team and input from Dr. Fish with podiatry.

This hopes to serve as a guideline to streamline management of patients with foot infection and concern for osteomyelitis. There is an algorithm at the end for quick reference.

Keywords: DFI, osteomyelitis

8/29/2024

Metronidazole dosing

Key points:

- Dosage interval of metronidazole at PeaceHealth has changed from Q8 to Q12 for most infections (and for both PO & IV routes)
- Pharmacy will adjust dosing as appropriate based on an interchange policy
- Please note your patient's current dosage interval if continuing metronidazole on discharge
- While the above dosing is different from commonly used references (e.g. UpToDate, Lexicomp, Sanford) it is supported by PK data and several clinical studies and is employed at other reputable institutions

Background:

In response to a nationwide shortage of IV metronidazole, a review of alternatives yielded good information about Q12H dosing, which is different from recommendations from tertiary databases and national guidelines. In light of this review, the dosage interval of metronidazole at PH has recently changed from a standard of Q8H for most infections to Q12H.

This change is supported by the pharmacokinetics of metronidazole (oral and IV) and several clinical studies (here, here, and here) examining outcomes with Q8 versus Q12 hour dosing. Metronidazole has an approximately 8-12 hour half-life, achieves 98-100% oral bioavailability, and plasma levels are unaffected by infection, acute or chronic renal disease, renal replacement, etc. It also has a low volume of distribution (0.5-1L), meaning it does *not* require weight based dose adjustments (please ask a pharmacist colleague for more about volume of distribution (3). 500 mg Q12H dosing provides serum levels well in excess of susceptible MICs for *Bacteroides fragilis* and other *Bacteroides spp.*, which are the primary anaerobic pathogens is it used for in our institutions.

Based on these data, a few institutions have examined clinical outcomes comparing Q8 versus Q12 hour dosing. The first, above Soule at al. at Novant Health, compared outcomes for primarily intra-abdominal infections, and found no difference in duration of treatment, length of stay, escalation of treatment, or death. Cure rates were ~80% in both groups. The second, above by Shah et al. at Yale New Haven, compared outcomes specifically for anaerobic bacteremia, of which the majority were *Bacteroides spp.*, and found no difference in escalation of treatment, length of stay, or death. Lastly, Béïque et al. compared outcomes for diverticulitis/appendicitis and found no difference in outcomes. These excluded parasitic or amoebic infections, *C. difficile*, and CNS infections. Studies suggesting superiority of Q8 over Q12 hour dosing are not available.

It's important to note that the cure rates in the first paper (80%) are similar to pooled cure rates for intra-abdominal infections with sepsis overall. That's not to suggest that patients on Q12H metronidazole have no risk of escalation of treatment, but that both pharmacokinetic data and clinical evidence demonstrate that escalation of treatment is most likely related to factors *other* than metronidazole dosing.

SC 10_2025

Keywords: general ID, oral tx

9/13/2024

Management of septic arthritis

(native/non-prosthetic joints)

Diagnosis

- For stable patients, aspirate suspected septic joints BEFORE administration of antibiotics
- Evaluate synovial fluid for: GS & culture, cell count with differential, crystals
- Collect blood cultures for febrile patients or in those with suspected concomitant bacteremia (also before antibiotics)
- CRP is of some value (if normal, argues against septic arthritis; if elevated, support diagnosis)

Microbiology

- MSSA is most common
- Strep, gram negatives, MRSA are less common
- Neisseria gonorrhoeae is uncommon but worth noting as requires some pre-test suspicion to ensure appropriate testing

Preferred empiric therapy

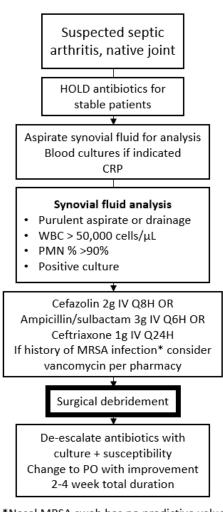
- Cefazolin or ampicillin/sulbactam are preferred for most patients
- Ceftriaxone is a reasonable alternative if GNRs are suspected
- Consider adding vancomycin for hemodynamic instability, injection drug use, or history of MRSA infection (note: nasal PCR has no predictive value for joint infections)

Management

- Consult orthopedics source control with surgical debridement is recommended wherever possible
- De-escalate therapy to the narrowest spectrum effective agent ASAP.
- IV antibiotics can be transitioned to PO in most cases
- Total duration of antibiotics with source control is 2-4 weeks
- ID consult is generally appropriate

Keywords: general ID, osteomyelitis

For those of you who prefer flow charts:



^{*}Nasal MRSA swab has no predictive value

9/19/2024:

Management of C. difficile diarrhea (for interpretation of test results, please see the tip from 9/25/23)

Step 1: Stop or de-escalate the inciting antibiotic wherever possible

This is step is extremely important, continuation of broad spectrum antibiotics is directly associated with severe disease and recurrence. Non-infections conditions for which antibiotics are commonly prescribed inpatient include hypoxia or peripheral edema due to volume overload, asymptomatic bacteriuria, aspiration pneumonitis, viral respiratory tract infections, vascular insufficiency, or diverticulosis without diverticulitis. Fluoroquinolones, ceftriaxone, antipseudomonal beta lactams, and clindamycin are the highest-risk antibiotics. Non-fluoroquinolone oral antibiotics, tetracyclines, metronidazole, and cefazolin are lower risk. Probiotics do not mitigate the effects of antibiotics in the setting of *C. difficile* (or any other setting, for that matter).

Step 2: Treat with oral vancomycin at 125 mg four times a day for 10 days

Colonic concentrations of vancomycin 125 mg average 500-1000 μ/g ; elevated MICs to vancomycin are rare, and 125 mg dosing yields concentrations 30 to 250 times above even intermediately susceptible MICs. Studies comparing doses ranging from 125 to 500 mg have identified no difference in time to resolution of symptoms, clinical cure, or other outcomes. There is little evidence to support the 500 mg dose, even in critically ill patients, however it is recommended as a last line option to prevent colectomy, despite the lack of data. Systemic concentrations of vancomycin have been measured with the 500 mg dose, especially in patients with renal failure – monitoring is recommended in this

circumstance. Fidaxomicin is equally effective when compared with PO vancomycin at treating CDI, but cost prohibitive (\$5977 v. \$57 per standard treatment), so it remains non formulary.

Other management considerations

The most important modifiable risk factor for *C. difficile* is exposure to antibiotics. The overwhelming majority of cases are directly preceded by exposure to antibiotics. 30-50% of antibiotics prescribed in the US are unnecessary or ineffective as written, for both the inpatient and outpatient setting. Reducing unnecessary use of antibiotics, in number, duration, by class, or overall has repeatedly demonstrated drastic decreases in *C. difficile* incidence, in a variety of settings, without adverse outcomes on the management of infectious diseases.

There is an emerging body of literature related to chemoprophylaxis in high-risk patients, alternative therapies/doses, and management of recurrence and severe disease. There are no clear recommendations in these populations. Often, these interventions are demonstrated to either have no effect or worsen outcomes, depending on the scenario. ID consultation is recommended for recurrent or severe disease. Your pharmacy colleagues are always available to assist with antibiotic de-escalation. Wherever possible, observing patients with questionable bacterial infection v. non-infectious syndromes *off* of antibiotics is safer than administration of antibiotics, with or without probiotics, chemoprophylaxis, etc.

Keywords: diarrhea

10/10/2024:

Transmission precautions

Below is an incomplete list of common infections for which transmission precautions (above & beyond standard precautions) may be required.

Bear in mind that patients in isolation generally receive less (and therefore less good) care, and for this reason precautions should not be put in place unless they are indicated.

Note that infection/colonization with MRSA is ***not*** an indication for contact precautions (this was a historical practice but now we know better).

For any questions regarding transmission precautions, including if they're indicated or can be discontinued, please call Infection Prevention (IP) at 360-514-2210.

Infection	Precautions
MRSA	None
MRSA with uncontained drainage	Contact
Meningococcal, H. influenzae, or	Droplet
undetermined bacterial meningitis	(discontinue after 24 hours antibiotics)
Influenza	Droplet
COVID-19	Special Contact Droplet with Eye

Pulmonary TB (including rule out)	Airborne (without contact)
C. difficile	Contact Enteric
Diarrhea, noninfectious	None
Diarrhea, infectious (confirmed/suspected)	Varies by organism - check with IP
Shingles	None (cover lesions until dry/crusted)
Shingles, immunocompromised or disseminated	Airborne & Contact

Keywords: general ID, labs

10/18/2024

IV to PO conversion

We're going to focus on IV to PO as we continue to address the ongoing shortages of the fluids needed to administer IV meds. See ID facts from 7/17, 7/31, and 8/21 of 2023 for more information of bioavailability and duration of therapy Θ .

Changing antibiotics from IV to PO once patients are stable and tolerating oral medications is a best practice, demonstrated to <u>facilitate earlier hospital discharge</u> and <u>reduce complications</u> including readmission. For highly bioavailable medications, we have good information about switching directly between IV and PO forms of the same drug (your pharmacy colleagues can facilitate this with minimal fuss). Recommendations for IV beta lactams and vancomycin are a little trickier, as the menu of options differs by disease state, but nonetheless a sizeable body of evidence recommends we do this <u>earlier</u> and <u>more often</u> than we think. General tips:

- Amoxicillin, amox/clav, and cephalexin have excellent oral bioavailability when dosed at the upper end of their ranges
 - With susceptibility data, these are good choices for step down therapy for CAP, UTI, gram negative and some gram positive bacteremia
- Fluoroquinolones and SMX-TMP are good choices for gram negative bacteremia with sensitivity data
 - More toxicities are associated with their use compared with beta lactams
- Cefpodoxime has reliable activity against respiratory pathogens and most quinolone and SMX-TMP resistant GNRs
 - Its poorer bioavailability relegates it to when use of other oral options is precluded by allergy, resistance, etc.

Keywords: general ID, oral tx

See attached guide for more detailed recommendations.



Clinical decision support IV PO.pdf

12/11/2024 Ceftriaxone dosing

Ceftriaxone dosed at 1g Q24H is appropriate for most infections.

Below is a breakdown of recommendations, including select disease states for which higher dosing is recommended based on available evidence.

As ceftriaxone is generally safe and well-tolerated, it may be tempting to think *more is better*. But keep in mind that giving double the dose of any drug without need is likely to increase adverse events without corresponding benefit.

Dose	Disease state
1g IV Q24H	Pneumonia, UTI (including pyelonephritis), skin & soft tissue infection, intra-abdominal infection (including SBP*), streptococcal bacteremia, enterobacterales bacteremia
2g IV Q24H	Infective endocarditis, some osteomyelitis/large abscesses if surgical source control is not feasible
2g IV Q12H	Bacterial meningitis, adjunctive therapy for enterococcal infective endocarditis

^{*}citations for the recommendation in UpToDate for 2g for SBP are 2 papers demonstrating no difference between 1 and 2 gram dosing for this indication

1g Q24H of ceftriaxone achieves optimal kinetics (100% time/MIC), and studies comparing 1 v 2-gram dosing (non CSF/IE indications) in critical illness, obesity, hypoalbuminemia, etc. have demonstrated no difference in outcomes. Tertiary references contain mixed messaging, e.g. the pages for bacteremic pyelonephritis and pneumonia recommend 1g, where the pages for pneumococcal bacteremia and enterobacterales bacteremia recommend either ranges or 2 g in UpToDate, but without citation.

Keywords: general ID, bacteremia

1/8/2025

Happy 2025! We're going to recycle one from the fall since we're seeing so much flu and there's been *another* publication.

Oseltamivir and influenza

The official (CDC) recommendation for influenza is: oseltamivir should be started **as early as possible** for influenza patients who are:

- Hospitalized
- Suffering from severe, complicated, or progressive illness
- At high risk for complications (this is mostly for outpatient folks, as symptomatic inpatients should technically all be treated anyway)

Note this recommendation applies <u>regardless</u> of <u>timing</u> of <u>symptom onset</u>. There is good evidence that oseltamivir on admission for any hospitalized patient can reduce length of stay, readmission, ICU admission, AKI, and death (<u>here, here, here, here, and here</u>) – research looking at timing anywhere from '**on admission**' to using a cutoff of 48 hours from admission notes maximal benefit when treatment is started in this window. Denying oseltamivir for inpatients based on total symptom duration is harmful and **NOT** recommended. If for some reason oseltamivir was delayed to hospital day 3 or later, it should still be started if possible.

Keywords: influenza, oseltamivir

SC 10_2025

1/31/2025: TB evaluation & isolation

What precautions are recommended for patient with possible pulmonary tuberculosis?

Patients who *might* have pulmonary TB (i.e. It's on the differential diagnosis, whether high or low) should be placed in "Airborne Respirator" precautions (this is the name in CareConnect). Contact precautions are NOT recommended for TB.

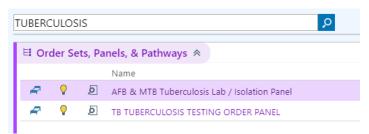
What tests should be ordered to evaluate for pulmonary tuberculosis?

Sputum testing for AFB smear & culture x3 and MTB PCR x2.

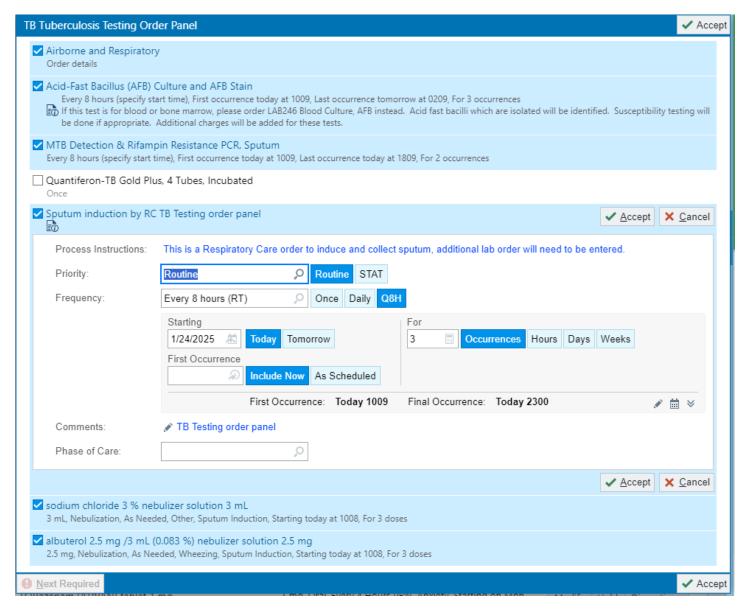
<u>PLEASE use the order set available in CareConnect</u> as this will ensure that appropriate tests, RT sputum induction (if needed), and appropriate precautions will be ordered. Note that AFB smear & culture and MTB PCR should be performed on the same samples (which might not happen if you order them separately, leading to extra sputum being collected and delaying the process). The samples can be collected as frequently as 8 hours apart, so if everything goes according to plan this can all be done pretty quickly.

If you type "tuberculosis" in the order field, you should see something like this:

Order and Order Set Search



Selecting one of these will pull up the whole panel, which is pre-selected with the option to add a Quantiferon:



Should I order a Quantiferon?

It depends on who you ask. It is reasonable, understanding that IGRAs (and TB skin tests) are NOT good tests for active TB. If positive, they don't distinguish between active and latent infection. If negative, they are not sensitive enough to rule out active infection. They may influence your clinical suspicion for TB disease but are insufficient on their own and arguably don't change management.

The smears and PCRs returned negative - can I discontinue airborne precautions?

Probably, but this needs to be approved by Infection Prevention (who will discuss with ID if needed). In cases where there is still some clinical suspicion for TB and no alternative diagnosis, we might continue precautions out of an abundance of caution.

Keywords: tuberculosis, isolation

2/7/2025: Avoid prescribing antibiotics for incidental sinusitis and/or bronchitis noted on imaging, with or without mild, short term, or resolving symptoms.

Sinusitis and bronchitis are primarily viral infections (98% for sinusitis and 95% for bronchitis), yet antibiotics are prescribed in 50-75% of diagnoses (including incidental inpatient diagnoses). Research into this phenomenon indicates sinusitis in particular is a diagnosis associated with the most unnecessary antibiotic use, see the NEJM commentary below on the linked research. In any case, bacterial sinusitis and bronchitis are rare and the vast majority of cases, inpatient and outpatient, should not receive antibiotics.

Officially, bacterial etiology of sinusitis should be considered if

- NO improvement in symptoms (fever, purulent nasal discharge, facial pain) for 10 days or more
- Fever greater than 39 C (102 F) for more than 3 consecutive days along with purulent nasal discharge or facial pain
- Incidence of fever, headache, or purulent nasal discharge after a 5-6 day URI with initial improvement

Treatment, where indicated, is amox/clav for no more than 5 days, doxycycline or cefpodoxime are acceptable substitutions in the setting of a true penicillin allergy; macrolides and fluoroquinolones should be avoided.

NEJM Journal Watch



SUMMARY AND COMMENT | GENERAL MEDICINE, INFECTIOUS DISEASES, HOSPITAL MEDICINE, EMERGENCY MEDICINE

INFORMING PRACTICE

June 13, 2019

"I'm Calling It Sinusitis. Here's Your Antibiotic!"

Abigail Zuger, MD, reviewing Martinez KA et al. J Gen Intern Med 2019 Jun

Doctors who lavish antibiotics on patients with upper respiratory infections were far more likely to diagnose "sinusitis" than were others.

Almost all upper respiratory tract infections are viral. The one exception to this generalization is sinusitis, which sometimes does have a bacterial component and, in severe cases, can benefit from empirical antibiotics. Researchers examined the records of a U.S. nationwide direct-to-consumer telemedicine service. During a 40-month period, 105 physicians treated more than 13,000 patients (mean age, 39) with upper respiratory complaints, coded as sinusitis, pharyngitis, bronchitis, influenza, or unspecified respiratory tract infection.

Physicians ranged widely in their likelihood of prescribing antibiotics for these patients: Mean antibiotic prescribing rates were 40% in the lowest quartile of prescribers and 87% in the highest. Physicians in the highest prescribing quartile were significantly more likely to code encounters as "sinusitis" than were those in the lowest quartile (59% vs. 35%), and the highest-quartile prescribing rate for patients with diagnosed sinusitis was 95%. These physicians also prescribed antibiotics at a higher rate than their peers for other diagnoses.

COMMENT

This study provides an interesting glimpse into the psychology of antibiotic overprescription. Doctors who overprescribe might know perfectly well that antibiotics are not indicated for most respiratory infections, but, whether subconsciously or not, use coding habits to justify their inability to say "no." In theory, medical coding summarizes the clinical encounter, but in this study, it appeared to dictate the outcome of many encounters — a depressing (if unsurprising) observation, and one, I suspect, that is not confined to the telemedicine sphere.

CITATIONS

Martinez KA et al. Coding bias in respiratory tract infections may obscure inappropriate antibiotic use. *J Gen Intern Med* 2019 Jun; 34:806. (https://doi.org/10.1007/s11606-018-4823-x)

Keywords: sinusitis

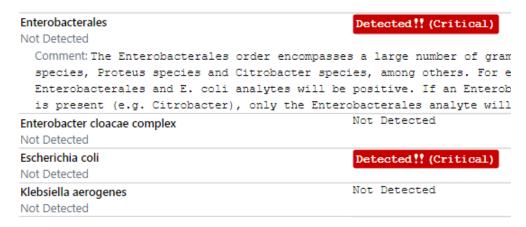
4/4/2025: As a reminder, when interpreting the rapid blood culture molecular identification PCR panel, "Enterobacterales" and "Enterobacter" are NOT equivalent:

-"Enterobacterales" = More generic term for an **Order** of bacteria (as in Kingdom, Phylum, Class, **Order**, Family, Genus, Species...and the associated mnemonic involving King Phillip...) that ultimately includes common gram negative rod bacteria Genera such as Escherichia (E. Coli), Klebsiella, Salmonella, Proteus, Shigella, Citrobacter, *AND* Enterobacter.

-So Enterobacter cloacae is a specific organism belonging to the order of Enterobacterales and will show up like this:

Enterobacterales	Detected!! (Critical)
Not Detected	
Comment: The Enterobacterales order encompasses a la	arge number of gram-negativ
species and Citrobacter species, among others. For positive. If an Enterobacterales order member that be positive.	
Enterobacter cloacae complex	Detected!! (Critical)
Not Detected	
Escherichia coli	Not Detected
Not Detected	
Klebsiella aerogenes	Not Detected
Not Detected	

-However, you can have other members of Enterobacterales such as E coli seen here. Again, this does NOT mean that Enterobacter is also present (Enterobacter cloacae is NOT detected):



-The reason for even having Enterobacterales as a target on the blood culture rapid PCR panel is nuanced, but it is also possible to receive an Enterobacterales result alone without a more specific genus/species name like this:

Enterobacterales Not Detected	Detected!! (Critical)
Comment The Enterobacterales order enco species and Citrobacter species, among positive. If an Enterobacterales order : be positive.	others. For example: when E. coli is
Enterobacter cloacae complex Not Detected	Not Detected
Escherichia coli Not Detected	Not Detected
Klebsiella aerogenes Not Detected	Not Detected
Klebsiella oxytoca Not Detected	Not Detected
Klebsiella pneumoniae group Not Detected	Not Detected
	57 . 70

This patient ended up having Shigella bacteremia. See a more complete list of Enterobacterales positive and no other targets detected in Dr. Root's previous ID tip of the week.

This is a key distinction because empiric treatment for Enterobacter cloacae, an AMP-C producer (like Citrobacter freundii and Enterobacter/Klebsiella aerogenes), is IV Cefepime. However, for bactermia due to other Enterobacterales (E coli, Klebsiella pneumoniae, etc.), IV Ceftriaxone is the empiric antibiotic of choice for our patients.

Keywords: Enterobacter, enterobacterales, PCR

4/24/2025:

Inpatient HIV testing on patients known to have HIV

Consider the following points when considering ordering HIV Viral Load and CD4 testing inpatient for patients living with HIV:

- -If a patient is known to have well controlled HIV and good adherence, inpatient HIV Viral Load and CD4 testing is not likely to change management and opportunistic infection risk is very low
- -CD4 count can often be falsely low in the setting of acute illness
- -If a patient is not taking HIV medication for >6 months, we know that their HIV Viral Load is going to be high and CD4 low, so the exact numbers are unlikely to change management and opportunistic infection should be considered in the differential diagnosis
- -A person who screens positive for HIV as a first time diagnosis will certainly need HIV Viral Load and CD4 testing. Several other tests are indicated for new HIV diagnosis, which can be guided by ID consultation.
- -On average, a CD4 test cost ~\$40 and HIV viral load test ~\$76

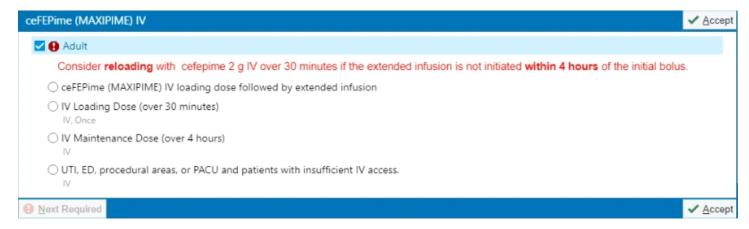
Just like any test, keep in mind how the HIV Viral Load and CD4 tests will change your management inpatient.

Of course, this is a separate topic from HIV *screening* inpatient in persons without HIV. HIV screening with the 4th generation ab/ag test is still recommended and an underutilized tool to screen those at risk of HIV acquisition (persons who use drugs, houseless individuals, sexual practice risk factors, etc.) as well as for those individuals that have never had screening.

Keywords: HIV, testing

5/1/2025: Extended infusion beta lactams

In our next Epic update, the antipseudomonal beta lactams will include an extended infusion panel with a loading dose over 30 minutes, followed by q8h infusions over 3 hours (cefepime or meropenem) or 4 hours (piperacillin/tazobactam). Here's what cefepime is going to look like, for example:



This is designed to maximize pharmacokinetic parameters (time spent with concentration over that of the MIC) for intermediately susceptible gram negative rods. Data for extended infusion beta lactams are mixed, with any potential benefit more likely in those with intermediately susceptible isolates or those with critical illness, see here. Increased ADEs (catheter or toxicity associated) are higher with extended infusions; it is recommended to use this dosing strategy primarily for those who are critically ill or with MDR GNRs.

UTIs are unlikely to benefit given the already favorable kinetics; standard infusions remain sufficient, as well as for circumstances where extended infusion is not feasible. If you would like to keep 30 minute infusions, select the bottom option for UTI/procedural areas/insufficient IV access. Recommended dosing by indication is:

Drug	Indication	Dose
Cefepime	Most indications	1g Q8H over 3 hours
	Meningitis, febrile neutropenia, <i>P. aeruginosa</i> , or AmpC producing enterobacterales with MIC 4-8	2g Q8H over 3 hours
Piperacillin tazobactam	Most indications	3.375g Q8H over 4 hours
	Febrile neutropenia, weight greater than 120 kg, cystic fibrosis	4.5g Q8H over 4 hours
Meropenem	Most indications	1g Q8H over 3 hours
	Meningitis, or intermediate susceptibility	2g Q8H over 3 hours

Fortunately for us, the vast majority of our bloodstream infections are not MDROs! In 2024, between SW and SJ, less than 5% of BSIs were *P. aeruginosa* or AmpC producing GNRs, so we should continue to use these agents only where indicated.

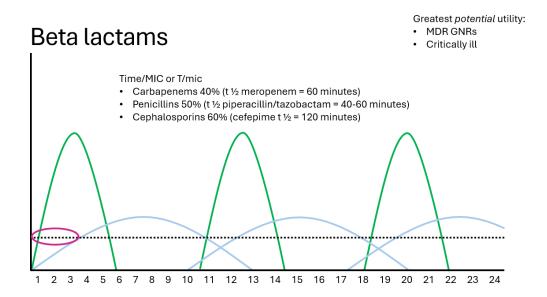
A reiteration of 3 key points from last week's tip:

- 1. Ceftriaxone remains the empirical treatment of choice for most infections, with reliable activity against *Streptococcus* spp. and non-AmpC, non-ESBL producing enterobacterales.
- 2. Cefepime is preferred over piperacillin/tazobactam for empiric treatment where AmpC producers or *P. aeruginosa* are likely, from both an efficacy and safety standpoint.
- 3. Cefepime, piperacillin/tazobactam, or meropenem offer no advantage over ceftriaxone for non-ESBL producing *E. coli, Proteus, K. pneumo*, or *K. oxytoca spp.*, but are associated with more toxicities and complications with use.

Keywords: extended infusion

Pharmacokinetics refresher, if you want refreshment

For beta lactams, the time spent at concentrations over the MIC is the pharmacokinetic parameter predicting success (e.g. no utility of high concentrations, AUC, etc.) Continuous or extended infusions are sometimes demonstrated to be more effective, particularly when the pathogen MIC is elevated, theoretically due to less time in the dosage interval spent at concentrations below the MIC. When we do this, it takes a little longer to get above the MIC (the pink oval in the diagram) because of the slower rate of infusion. For most patients, this doesn't matter, but for those with severe sepsis or septic shock, a loading dose is recommended when extended infusion doses are being used.



5/10/2025: Which oral beta lactam should I prescribe?

Cephalexin (or cefadroxil) is preferred over cefdinir or cefpodoxime wherever possible, and particularly for UTI and skin/soft tissue infections, as it has much greater oral bioavailability/potency and an improved toxicity profile. All 4 drugs are similarly active against enteric or urinary pathogens. Cephalexin or cefadroxil have greater activity against skin flora than cefdinir or cefpodoxime.

Use of cefpodoxime or cefdinir is most appropriate when allergies preclude the use of preferred agents.

	Bioavailability	Dose	Toxicity
Amoxicillin	70-90%	2-3x daily	Less diarrhea
Cephalexin	90%	2-4x daily	Less diarrhea
Cefadroxil	80-90%	1-2x daily	Less diarrhea
Amox/clav	70-90%	2x daily	More diarrhea
Cefpodoxime	50%	2x daily	More diarrhea
Cefdinir	20-25%	2x daily	More diarrhea

Spectrum of activity:

- E. coli are approximately 90% susceptible to each of cephalexin/cefadroxil, amox/clav, cefpodoxime, and cefdinir between SW and SJ, with a few percentage points of variability between hospitals.
- All beta strep (Group A, B, and C/G or S. pyogenes, S. agalactiae, and S. dysgalactiae) are uniformly sensitive to beta lactams, however cephalexin, cefadroxil, and amoxicillin have the most potent activity.
- Cephalexin, cefadroxil, and amox/clav have superior activity against MSSA than do cefpodoxime or cefdinir.
- S. pneumoniae remains 100% penicillin susceptible in our institutions, and amoxicillin or amoxicillin/clavulanate are preferred over cefdinir or cefpodoxime due to greater potency and bioavailability.
- H. influenzae produces a beta lactamase in 16-20% of isolates, amoxicillin is the treatment of choice for nonbeta lactamase producers, and either amox/clav (preferred) or cefpodoxime are appropriate for beta lactamase producing strains. Amoxicillin or amox/clav have favorable bioavailability/potency and less toxicity than 3rd generation cephalosporins.

Keywords: oral cephalosporin

5/16/2025: Antibiograms are posted! See here and also in the lower right of every micro

PeaceHealth Antibiograms

result:

Antibiograms Link

No major changes from last year, but a few things to call out

- 1. Ceftriaxone remains the empirical treatment of choice for most infections, with reliable activity against non-AmpC producing enterobacterales and Streptococcus spp.
- 2. Cefepime is preferred over piperacillin/tazobactam for empiric treatment where AmpC producers or P. aeruginosa are likely, from both an efficacy and safety standpoint.
- 3. Clindamycin, ciprofloxacin, and levofloxacin remain unreliable (50s-70s percent susceptible) for the common pathogens they used to have activity against (beta strep and anaerobes for clindamycin, E. coli and other enterobacterales for quinolones)
 - a. As an aside, these drugs are all much more toxic than their beta lactam equivalents, use should be avoided if penicillins or cephalosporins are an option
- 4. Cefepime, piperacillin/tazobactam, or meropenem offer no advantage over ceftriaxone for non-ESBL producing E. coli, Proteus, K. pneumo, or K. oxytoca spp., but are associated with more toxicities and complications with use.



INPATIENT DATA

PeaceHealth Southwest Medical Center

January 1 - December 31, 2024

Dash (-) = not tested, inappropriate, or < 30 strains tested. R = intrinsically resistant.

						Be	ta-lacta	ms					Flu	oro-	Amir	noglycos	idos	Miscellaneous					
			Penicillin	8			Cepi	hems			Carbap	enems	quint	lones	Ami	юдусия	ilites	MI	cenamen	Jus			
GRAM NEGATIVE ORGANISM PERCENT SUSCEPTIBLE	# Isolates Tested	Ampicillin	Amoxicillin/clavulanic acid	Piperacillin/tazobactam	Cefazolin	Ceftazidime	Ceftriaxone	Cefepime	Cefoxitin	Cefpodoxime	Ertapenem	Meropenem	Ciprofloxacin	Levofloxacin	Amikacin	Gentamicin	Tobramycin	Nitrofurantoin - urine only	Tetracycline	Trimethoprim/sulfamethoxazole			
Citrobacter freundii complex	55	R	R	-	R	-	-	100	R	-	98	100	87	84	97	96	96	89	81	87			
Enterobacter cloacae complex	121	R	R	-	R	-	-	91	R	61	90	100	95	95	100	99	98	36	88	93			
Escherichia coli	1618	59	88	96	85	96	91	96	94	88	100	100	79	75	100	92	93	96	80	81			
Klebsiella aerogenes	61	R	R	-	R	-	-	100	R	-	95	100	97	95	100	100	100	5	98	100			
Klebsiella oxytoca	90	R	96	97	-	98	96	97	99	91	100	100	96	99	100	96	94	100	92	96			
Klebsiella pneumoniae	402	R	93	93	85	91	88	94	96	88	100	100	89	86	100	95	94	29	85	89			
Morganella morganii	34	R	R	100	R	-	-	-	45	-	100	100	74	74	-	-	94	R	-	87			
Proteus mirabilis	237	73	100	100	98	99	98	98	93	98	100	100	77	78	100	92	95	R	R	74			
Pseudomonas aeruginosa	290	R	R	90	-	94	R	97	-	-	R	99	89	82	99	-	100	-	R	R			
Serratia marcescens	65	R	R	92	R	100	92	100	R	-	100	100	86	86	100	100	94	R	55	100			
Stenotrophomonas maltophilia	42	R	R	R	-	-	R	-	-	-	-	R	-	93	R	R	R	-	R	95			

NOTE: Third generation cephalosporins and piperacillin/tazobactam are unreliable for systemic infections with AmpC producing organisms (E. cloacae, K. aerogenes, or C. freundii), regardless of susceptibility data, due to the development of induced resistance on treatment.

Haemophilus influenzae beta lactamase positive: 16.4% (n=79)



INPATIENT DATA

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	ľ		Be	ta lactar	ns		Flu	10ro-	Am	ino-											
		F	enicilina	s/Cepha	losporins	s	quine	olones	glyco	sides	Miscellaneous										
GRAM POSITIVE ORGANISM PERCENT SUSCEPTIBLE	# Isolates Tested	Penicillin	Oxacillin	Ampicillin	Cefazolin	Ceftriaxone	Ciprofloxacin	Levofloxacin	Gentamicin Synergy	Streptomycin Synergy	Clindamycin	Daptomycin	Erythromycin	Minocycline	Nitrofurantoin - urine only	Rifampin	Tetracycline	Trimethoprim/sulfamethoxazole	Vancomycin		
Enterococcus faecalis	349	-	=	100	R	R			84	94	R	-	-	*	100	*	100	R	99		
Enterococcus faecium	37		-2	32	R	R		(*)	93	48	R	194	- 67	12	100	-		R	78		
Staphylococcus aureus , MRSA	359		0	4.	0	(60		(a)	*	14	82	100	¥0.	100	1	100	65	100	100		
Staphylococcus aureus , MSSA	718		100	-	100	-	-	12.5	1.0		86		20	100	100	4	93	100	-		
Staphylococcus epidermidis	107		39	-	39	1	2	40	2	-	67	92	¥	100		2	73	64	100		
Staphylococcus lugdunensis	64	1021	91	-	91	100	2	130	0	12	88	182	20	100	72:	9	98	100	100		
Streptococcus agalactiae (Group B)	100	100	15	353	100	100			0.0	=	60	15	55	8	-	8	153		100		
Streptococcus anginosus	66	98	-	-		100	-	(etc)	42	-	-	5.7	*	-	-		· t		100		
Streptococcus constellatus	38	100		-		100	-	92	- 51	12	*				*		30		100		
Streptococcus mitis/oralis	32	75	- 12		-	97	-	150	2	127		10.7	+	->-	-		**	-	100		
Streptococcus pneumoniae	126	100*	-	10	*	100*	*	100	*	3	×	-	78	×		*	91	*	-		
Streptococcus pyogenes (Group A)	125	100	3		100	100	9		æ	14	71	12	70	-	1	14	(.)	2	100		

*Non-meningitis breakpoint



INPATIENT DATA

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						B4	ta-lactar	ns					Flu	0f0-	Amir	noglycos	ides	Miscellaneous				
		ı	Penicillin	8			Cept	ems			Carbag	enems	quino	lones	~****	7		mie	rcenaire.	,,,,,		
GRAM NEGATIVE ORGANISM PERCENT SUSCEPTIBLE	# Isolates Tested	Ampicillin	Amoxicillin/clavulanic acid	Piperacillin/tazobactam	Cefazolin	Ceftazidime	Ceftriaxone	Cefepime	Cefoxitin	Cefpodoxime	Ertapenem	Meropenem	Ciprofloxacin	Levofloxacin	Amikacin	Gentamicin	Tobramycin	Nitrofurantoin - urine only	Tetracycline	Trimethoprim/sulfamethoxazole		
Enterobacter cloacae complex	56	R	R	-	R	-	-	95	R	-	93	99	95	95	100	100	100	39	96	93		
Escherichia coli	979	63	90	98	90	97	94	98	95	91	100	100	83	79	100	95	95	98	82	84		
Klebsiella oxytoca	46	R	93	91	-	-	93	100	98	-	100	100	96	98	-	98	98	-	95	96		
Klebsiella pneumoniae	153	R	96	97	96	98	95	99	97	95	100	100	94	92	100	99	97	26	90	92		
Proteus mirabilis	70	83	100	100	100	100	100	100	94	100	100	100	83	83	100	96	97	R	R	83		
Pseudomonas aeruginosa	90	R	R	88	-	83	R	89	-	-	R	99	86	76	100	-	100	-	R	R		

NOTE: Third generation cephalosporins and piperacillin/tazobactam are unreliable for systemic infections with AmpC producing organisms (E. cloacae, K. aerogenes, or C. freundii), regardless of susceptibility data, due to the development of induced resistance on treatment.

Haemophilus influenzae beta lactamase positive: 20.9% (n=43)



INPATIENT DATA

PeaceHealth St. John Medical Center

January 1 - December 31, 2024

Dash (-) = not tested, inappropriate, or < 30 strains tested. R = intrinsically resistant.

				ta lacta				oro-		ino-				Mis	cellaneo	ous			
GRAM POSITIVE ORGANISM PERCENT SUSCEPTIBLE	# Isolates Tested	Penicillin	Oxacillin	Ampicillin	Cefazolin	Ceftriaxone	Ciprofloxacin	Levofloxacin	Gentamicin Synergy	Streptomycin Synergy	Clindamycin	Daptomycin	Erythromycin	Minocycline	Nitrofurantoin - unine only	Rifampin	Tetracycline	Trimethoprim/sulfamethoxazole	Vancomycin
Enterococcus faecalis	174	-	-	99	R	R	-	-	81	92	R	-	-	-	100		-	R	99
Staphylococcus aureus , MRSA	215	-	0	-	0	-	-	-	-	-	90	-	-	100	-	-	60	100	100
Staphylococcus aureus , MSSA	223	-	100		100	-	-	-	-	-	89	-	-	100	-	-	87	100	-
Staphylococcus epidermidis	31	-	32		32	-	-	-	-	-	-	-	-	-	-	-	-	-	100
Streptococcus agalactiae (Group B)	44	100	-	-	100	100	-	-	-	-	52	-	44	-	-	-	-	-	100
Streptococcus anginosus	41	100	-	-	-	100	-	-	-	-	-	-	-	-	-	-	-	-	100
Streptococcus pneumoniae	74	100*	-	-	-	100*	-	100	-	-	-	-	-	-	-	-	-	-	-
Streptococcus pyogenes (Group A)	73	100	-	-	100	100	-	-	-	-	72	-	72	-	-	-	-	-	100

*Non-meningitis breakpoint

To reiterate last week's point, *E. coli* susceptibilities for cephalexin or cefadroxil v. cefpodoxime or cefdinir are virtually identical, and cephalexin and cefadroxil have 2x the potency (bioavailability) of cefpodoxime, as well as lower rates of ADEs. A win for everybody!

Keywords: antibiogram

5/20/2025: Should Patients Hospitalized for Community Acquired Pneumonia be Treated with Additional Anaerobic Coverage?

Building off last week's ID tip of the week, Ceftriaxone is our empiric antibiotic of choice for community acquired pneumonia (CAP). Is there a role for anaerobic coverage for pneumonia? A recent article reviewing multiple studies found there was NO benefit to adding anaerobic coverage for treatment of CAP (such as adding Metronidazole or Clindamycin to Ceftriaxone). This includes aspiration pneumonia:

- Aspiration pneumonia, seen as lung infiltrate on imaging, is due to oropharyngeal and gastric organisms, predominantly covered by Ceftriaxone
- Aspiration pneumonitis is a temporary and self-limiting inflammatory response after lung injury from a witnessed aspiration event that should not be treated with antibiotics
- Anaerobes have been found to only be cultured in 0.5% of aspiration pneumonia cases
- Oral cavity anaerobes may play a protective role in those at risk for developing pneumonia
- Adding anaerobic coverage for CAP has been linked to an increase of *Clostridium difficile* infection, disruption to the natural gut microbiome, and longer hospital stays
- Adding anaerobic coverage for CAP does not provide mortality benefit or change outcomes
- · Empiric anaerobic coverage is warranted in the setting of lung abscess or empyema

We would advise against the empiric addition of Metronidazole to Ceftriaxone in the treatment of CAP including aspiration pneumonia. Interestingly, in another recent <u>study</u>, Ceftriaxone was shown to have slight mortality benefit (absolute difference of 1.5% benefit) when compared to Ampicillin/sulbactam as initial choice for CAP in older adults. This was true for aspiration pneumonia cases as well.

American Journal of Therapeutics



<u>Should Patients Hospitalized for Community-Acquired...:</u>
<u>American Journal of Therapeutics</u>

journals.lww.com

Keywords: CAP, anaerobes

5/27/2025: Antibiotics for CAP in Patients Admitted with COVID:

In a recent JAMA article entitled, *Antibiotic Treatment in Patients Hospitalized for Nonsevere COVID-19* (Pulia et al., 2025), investigators looked at clinical outcomes of patients hospitalized for COVID-19 receiving antibiotics for community acquired pneumonia (CAP).

Key points from article:

• Superimposed bacterial pneumonia in patients with COVID is rare – estimated at 5% of cases, yet patients admitted with COVID receive antibiotics >30% of the time!

- · Over 500,000 encounters of patients admitted with non-severe COVID were evaluated and split into two groups, those that received antibiotics on day 1 and those that did not.
- The composite outcome was clinical deterioration during the hospital visit based on need for ICU transfer, high-flow or mechanical ventilation, vasopressors, or in-hospital mortality.
- · Conclusions showed that antibiotics did not provide better outcomes. In fact, in propensity-matched scoring, patients who received CAP antibiotics had higher odds (1.33 odds ratio) of the composite deterioration and in-hospital mortality.

This study provides further data that antibiotics covering CAP in patients with COVID-19 are not beneficial and can be potentially harmful. We know that unnecessary antibiotics carry risk of adverse events, gut dysbiosis, and resistance. Patients admitted with COVID that develop a hospital-acquired infection are more likely to have this infection be resistant if they received antibiotics earlier on in the hospital stay.

This study is also relevant to our hospital as they found patients admitted for COVID were more likely to be started on antibiotics at non-academic centers. And for patients that received antibiotics on day one, 67% of these patients received additional antibiotics compared to only 14% that did not receive antibiotics on day 1. Underscoring the point, that just because the first provider started CAP treatment, does NOT mean it needs to be continued.

Keywords: CAP, COVID

6/26/2025: Inpatient management of dental infections

- Ampicillin/sulbactam is preferred empirically for dental infections (even for those who have 'failed' amoxicillin/clavulanate).
- Ceftriaxone, with or without metronidazole, is a suitable alternative in the setting of true penicillin allergy.
- Clindamycin, vancomycin, and antipseudomonal agents should not routinely be used, as they are less effective, more toxic, or both.

Streptococcus spp. are the primary cause of dental infections, followed by oral anaerobes. Infection progression on oral antibiotics is **expected** without dental treatment. Progression is due to a lack of source control/inability of oral antibiotics to achieve effective concentrations at the site of infection.

Oral anaerobes are typically susceptible to ceftriaxone monotherapy, the addition of metronidazole adds activity against *Bacteroides* spp., which are uncommon oral pathogens. Ampicillin/sulbactam has excellent activity against both oral and GI anaerobes. Resistance to clindamycin has increased sharply among both oral and GI anaerobes, and is approaching 50% for *Streptococcus* spp. Clindamycin does not provide suitable activity against likely pathogens. Additionally, it is the highest risk antibiotic for *C. difficile*; use should be avoided wherever possible.

Anti-MRSA agents are less effective for *Streptococcus* spp. than recommended antibiotics, and have significantly greater toxicities. For stable patients, these agents may be added later in the unlikely event MRSA is identified, but empiric use is far more likely to cause harm than improve outcomes. *S. aureus* is a possible pathogen of sialadenitis; MSSA is much more common than MRSA – empiric treatment against MRSA is not recommended for patients who are hemodynamically stable.

P. aeruginosa and enterobacterales spp. (*E. coli, Klebsiella,* AmpC producers, etc.) are exceedingly uncommon pathogens in dental infections – antipseudomonal agents are no better or more potent against likely pathogens compared with ampicillin/sulbactam or ceftriaxone, but are associated with greater toxicities.

Keywords: dental

7/23/2025: Complicated UTI Guidelines:

The much anticipated update to the IDSA UTI treatment guidelines has arrived!

The most recent publication centers on complicated UTI (cUTI) management. A future guideline will be made for uncomplicated (uUTI), however the presence of local bladder signs and symptoms (dysuria, urgency, frequency, and suprapubic pain) are included in the treatment definition (non-specific symptoms like altered mental status remain NOT diagnostic, even in the presence of a 'positive' UA)

Key summary points:

*The main change is a clear definition of complicated v. uncomplicated

- Uncomplicated
 - Infection confined to the bladder in afebrile men or women (e.g. cystitis, regardless of age, comorbidities, etc.)
- Complicated
 - o Infection beyond the bladder in men or women
 - Pyelonephritis
 - Fever or bacteremia
 - Presence of catheter with systemic symptoms (catheterized patients without systemic symptoms fall into uncomplicated UTI)

- *Antibiogram use is encouraged for empiric therapy! The recommendations are
 - Sepsis: at least 80% activity against likely pathogens recommended (we continue to be above 90% for *E. coli* with ceftriaxone, this is our preferred agent, 1 g dosing is recommended)
 - Septic shock (ICU/vasopressor support): at least 90% activity recommended; cefepime is preferred for its activity against AmpC producers, which piperacillin/tazobactam does not have
 - o In reiteration of the fluoroquinolone point, our *E. coli* % susceptibility remains in the 70s, empirical use is not appropriate
 - Initial de-escalation using the antibiogram with pathogen identification is encouraged, e.g. if started on cefepime, non-esbl producing E. *coli, K. pneumo/oxytoca,* or *Proteus* spp. can/should go to ceftriaxone even before final susceptibility.

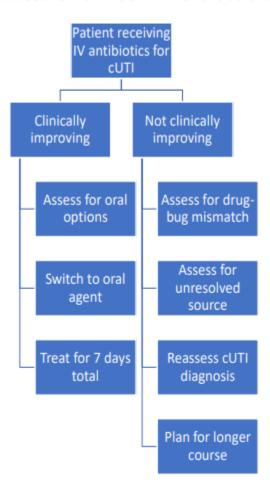
^{*}Prostatitis, perinephric abscess, or epididymitis are not included in the guidelines and treated as separate entities.

^{*}Fluoroquinolones should be avoided with fluoroquinolone exposure in the preceding 12 months - this is a class specific recommendation, it does not apply to beta lactams like penicillins or cephalosporins (another tip for another day).

^{*}Transition to oral therapy for cUTI should happen as soon as patient is clinically improving, can take oral therapy, and has an oral option available based on sensitivities.

* Treatment is 7 days! This includes concurrent bacteremia. No more 10-14 day vagueness.

Figure 1.2: Stepwise assessment of IV to oral switch and duration of antibiotic therapy



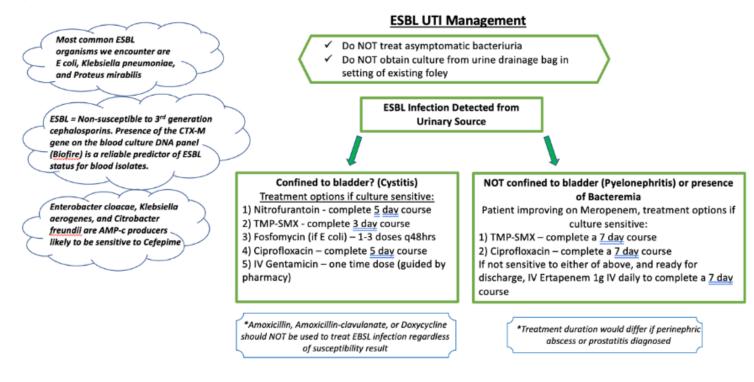
Abbreviations: IV=intravenous, cUTI=complicated UTI. Drug-bug mismatch means that the causative organism is not susceptible to the antibiotic prescribed.

Find the executive summary attached and here

Link to the full guideline webpage: <u>Complicated Urinary Tract Infections (cUTI)</u>: <u>Clinical Guidelines for Treatment and Management</u>

Keywords: complicated UTI

8/7/2025:



Keywords: esbl

9/17/2025: Antibiotics for sepsis refresher/FAQ

Ceftriaxone remains preferred for most community onset sepsis.

FAQ

Aren't we supposed to give broad-spectrum antibiotics?

'Broad-spectrum' in the context of the CMS sepsis measure means activity against both gram-negative and gram-positive organisms, rather than activity against MDROs like MRSA or *P. aeruginosa*. The federal list of antibiotics suitable for monotherapy includes these from our formulary: ampicillin/sulbactam, ceftriaxone, cefepime, levofloxacin, piperacillin/tazobactam, and meropenem, with a recommendation to base empiric treatment on local epidemiology and disease severity.

What are our local recommendations, then?

Here are ours, with the rationale (antibiograms are still here, if needed):

Local recommendations for preferred monotherapy for sepsis										
Ampicillin/sulbactam	Below 90% activity for <i>E. coli</i>									
Ceftriaxone	Preferred monotherapy for most community onset sepsis, given its local activity and safety profile									
Levofloxacin	(far) below 90% activity for <i>E. coli</i>									

Piperacillin/tazobactam*	Not active against AmpC producers, increased risk of nephrotoxicity compared with cefepime or meropenem
Cefepime*	Preferred monotherapy for septic shock (requiring vasopressors) given its local activity and safety profile
Meropenem*	Lower threshold for development of resistance compared with cefepime, associated with higher rates of <i>C. Difficile</i> . Preferred if relevant history of ESBL enterobacterales.

^{*}Including activity against AmpC producers, ESBL, and/or *P. aeruginosa* is recommended based on disease severity (septic shock requiring vasopressors) and validated risks for MDROs

Why not use cefepime and vancomycin for everyone?

The mortality risk of including unnecessarily broad-spectrum empiric activity is <u>equal</u> to that of providing inadequate empirical activity! Numerous recent studies (<u>here</u>, <u>here</u>, and <u>here</u>) also note sustained increases in use of anti-*Pseudomonal* and anti-MRSA antibiotics in US hospitals for sepsis, despite low rates of MRDOs.

So when should we use something other than ceftriaxone?

Activity against 90% or greater of likely pathogens is appropriate for those with community onset sepsis without shock, given the risks with broader activity and lack of association with excess mortality in the population in the unlikely event the causative pathogen is not susceptible. For those with shock, expanding the spectrum of empirical activity to include our more common MDR gram negative organisms (AmpC producers and *P. aeruginosa*) with cefepime is reasonable. Similarly, for those with prior, relevant history of an MDRO, including activity against the previous isolate is recommended. We'll talk more about MRSA next week.

Keywords: sepsis

9/26/2025: Role of anti-MRSA treatment (vancomycin and daptomycin) in sepsis and FAQ

Most empiric MRSA treatment for sepsis is both unnecessary and dangerous. Recent <u>data</u> demonstrate MRSA activity is initiated in over 40% of community onset sepsis diagnoses, despite rates of MRSA culture positivity of <u>3.4</u> to <u>3.7%</u> in community onset sepsis. This indication accounts for half of overall anti-MRSA antibiotic use in US hospitals! US MRSA prevalence is decreasing, with the rate of inpatient stays with MRSA present on admission <u>declining</u> from 0.097% to 0.084% between 2016 and 2021.

When compared with receipt non-MRSA treatment, anti-MRSA treatment is associated with significantly *increased* rates of mortality, as well as ICU transfer, readmission, and prolonged LOS. This relationship is significant and persists across multiple large US studies (see here, here, and here), with similar populations to those we treat here at PeaceHealth.

When should I include MRSA activity empirically?

Prior MRSA cultured from a relevant site is the only locally validated risk for MRSA infection. Of note, MRSA nares pcr has no positive predictive value in any setting, and should never be used to start empiric anti-MRSA treatment.

S. aureus generally is a common pathogen in purulent SSTI, *hematogenous* osteomyelitis (also MRSE here), right sided infective endocarditis, and post-influenza bacterial pneumonia - including MRSA activity empirically for those with hemodynamic instability is recommended.

What are our biggest opportunities to improve?

^{*}Including activity against AmpC producers, ESBL, and/or *P. aeruginosa* is recommended based on disease severity (septic shock requiring vasopressors) and validated risks for MDROs

Empiric treatment for MRSA is recommended against for

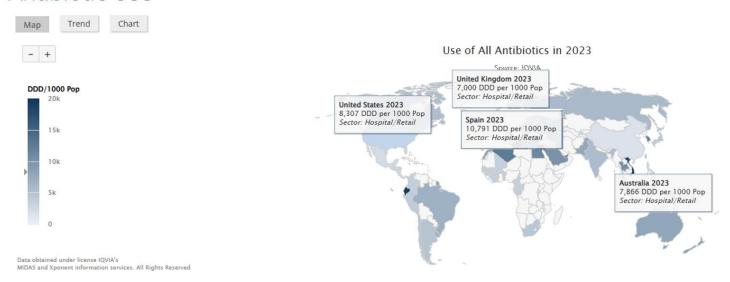
- Most UTI and intra-abdominal infection
- Diabetic foot infections without prior cultures for MRSA at the site (with or without possible osteomyelitis)
- Most community acquired pneumonia without prior respiratory culture (NOT pcr) positive for MRSA
- Most cases of febrile neutropenia
- Stable patients in whom the likelihood bacterial infection is unclear

Paradoxically, some empiric anti-MRSA treatment at PeaceHealth is started, 'just to be safe.' While well intentioned, this practice offers no benefit and is demonstrably harmful.

For fun, here are some data about global use of anti MRSA antibiotics:)

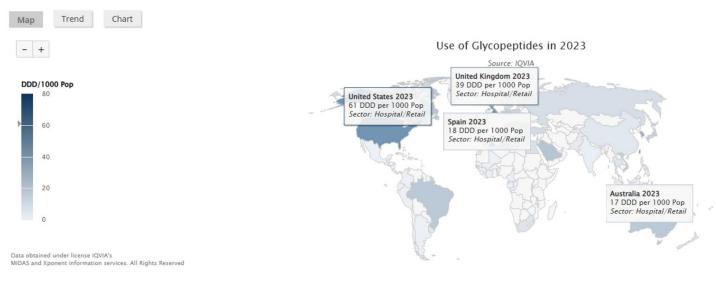
The US is the primary consumer of anti-MRSA antibiotics globally, a practice that is inconsistent with rates of MRSA. Here is the US and some of its nearest neighbors, overall antibiotics prescribing wise (Scandinavian nations prescribe antibiotics at approximately half our rates, if you're curious):

Antibiotic Use



And here is the same comparison for anti-MRSA antibiotics (glycopeptides = vancomycin here, but the data for daptomycin (a lipopeptide) are similar:) - we use more than anyone! Here's a study on the <u>disparate</u> use globally, for CAP:

Antibiotic Use



Keywords: sepsis, MRSA, vancomycin

10/3/2025: Role of anti-*Pseudomonas* treatment (cefepime, secondarily piperacillin/tazobactam or meropenem) in sepsis and FAQ

Most empiric *P. aeruginosa* treatment for sepsis is both unnecessary and potentially dangerous.

Recent <u>data</u> demonstrate anti-*Pseudomonas* activity is initiated in almost 60% of community onset sepsis diagnoses, despite rates of *P. aeruginosa* culture positivity of <u>2.3%</u> to <u>2.9%</u> for that indication. Rates of AmpC producing organisms (*E. cloacae, K. aerogenes, and C. freundii*) are not reported as commonly, but are approximately equivalent to those of *P. aeruginosa* at PH; the vast majority of our community onset sepsis pathogens are *E. coli* and strep. A common misconception is that anti-*Pseudomonal* antibiotics are 'stronger' or more potent than those not targeting resistant organisms. For susceptible *E. coli* and similar pathogens, there is no efficacy difference between any of the beta lactams, but expanded spectrum of activity *is* associated with harm. Every additional day of anti-pseudomonal treatment is <u>associated</u> with new development of resistance, and here's <u>one</u> of so many demonstrating the clear association between expanded spectrum and *C. difficile*.

When compared with non-anti *Pseudomonal* beta lactams, treatment targeting *Pseudomonas* is also associated with significantly *increased* rates of mortality, as well as ICU transfer, readmission, and prolonged LOS (see those here and here and here). Additionally, this study demonstrates that delaying anti-pseudomonal antibiotics (e.g. starting people on ceftriaxone and switching to cefepime if needed based on culture) is safe/possibly better overall, and certainly reasonable for most (not needing vasopressors/without prior relevant MDROs) patients with community onset sepsis here.

When should I include P. aeruginosa activity empirically?

Cefepime is recommended for those with sepsis and prior isolation of *P. aeruginosa* or AmpC producing enterobacterales from a relevant site.

Meropenem is recommended for those with sepsis and prior ESBL from a relevant site.

Piperacillin/tazobactam should not be routinely used empirically, its spectrum of activity and safety profile make it a poorer choice than cefepime or meropenem.

For febrile neutropenia, and those sepsis from a presumed urine or intra-abdominal source with hemodynamic instability, cefepime is recommended empirically, with rapid de-escalation where possible.

What are our biggest opportunities to improve?

Empiric treatment for P. aeruginosa is recommended against for

- Most skin and soft tissue infections
- Diabetic foot infections without prior cultures for P. aeruginosa or AmpC producers at the site (with or without possible osteomyelitis)
- Most community acquired pneumonia without prior respiratory culture positive for P. aeruginosa or AmpC producers
- Stable patients in whom the likelihood bacterial infection is unclear

Use of anti-pseudomonal agents where not needed is clearly associated with increased resistance, toxicities are drug specific but are generally worse (except piperacillin/tazobactam, which is pretty clearly worse) than ceftriaxone.

10/9/2025: For this week's ID Tip of the Week we're highlighting something mentioned a couple weeks back.

It is common for well-intended clinicians to view selection of excessively broad-spectrum empiric antibiotics as a strategy that is in their patient's best short term interest.

However, while it may feel that one is "erring on the side of caution" in taking this approach, the reality is that selection of excessively broad antibiotics poses similar mortality risk to selection of antibiotics that are too **narrow-spectrum**, as evidenced by this study cited in a recent Tip.

Interestingly, in a recent study looking at gram-negative rod infections (non-ICU, not in septic shock), patients in a matched cohort that received initial narrower empiric coverage (e.g. ceftriaxone or ampicillin-sulbactam) but then required escalation to broader coverage (e.g. cefepime or piperacillin-tazobactam) had similar outcomes to those given broader coverage up front. This provides reassurance that, in the rare instance that empiric coverage is not sufficiently broad, patients will not have poor outcomes.

Aside from the impact on individual patients, we know that use of excessively broad-spectrum antibiotics has a negative effect at a community level, encouraging the emergence of resistant flora.

It is good practice to keep this in mind when approaching the selection of empiric antibiotics for our patients and avoid prescribing anti-MRSA or anti-Pseudomonal antibiotics (for example) when not indicated or supported by local & national guidelines.

Keywords: sepsis

10/15/2025: Penicillin Allergy Delabeling Part 1

Antibiotics are a critical tool for treating infection. Unfortunately, inappropriate prescribing of antibiotics will lead to resistance and we will simply lose these tools. New antimicrobials in the pipeline are limited, therefore it is vital that we preserve the effectiveness of the antibiotics we have!

One strategy is antibiotic allergy delabeling. Most commonly seen with penicillin/amoxicillin, this is the satisfying action of removing an unnecessary antibiotic allergy from a patient's chart. Here are some facts about penicillin allergy:

-About 10-20% of patients have a reported penicillin allergy in their chart, although it is estimated that <1% of the US population has a true type I hypersensitivity reaction to penicillin

- -80% of persons with an IgE mediated response will lose their allergy altogether within 10 years of that previous reaction to penicillin. Almost all will be non-allergic after 20 years.
- -Giving second-line antibiotics to somebody without a true penicillin allergy can be less effective, more toxic, and increase risk of developing future resistance
- -Patients with penicillin allergy have higher rates of C Diff, MRSA, and VRE compared with matched controls
- -Documented penicillin allergy in the chart is associated with longer hospital stays, higher drug costs, and more frequent health care complications including surgical site infections compared to those without
- -More than 94% of individuals with a penicillin allergy label can tolerate penicillin

Link to paper with Data

Many patients can be delabeled based on a few questions or simple chart review alone. We encourage you to remove a penicillin allergy designation from a patient's chart if there is record of tolerating a penicillin-based antibiotic previously or if the patient-reported reaction is an intolerance rather than allergy (GI upset, headache, yeast infection, etc). These scenarios do not require direct oral graded challenge and taking a brief moment to remove a false penicillin allergy from the chart can go a long way in combatting antibiotic resistance at the personal and community level.

More about oral allergy challenge next week, stay tuned!

Keywords: allergy, delabel, penicillin, oral challenge