# Medical Policy

the medication is available.



Name			ricultine	are services beparement
Service Category  Anesthesia   Medicine Services and Procedures   Surgery   Evaluation and Management Services   Pathology and Laboratory Procedures   DME/Prosthetics or Supplies   Pathology and Laboratory Procedures   DME/Prosthetics or Supplies   Pathology and Laboratory Procedures   Other IV Antibiotics   Service Description   Enclosed document contains but is not limited to the following information: classification of antibiotics, medical necessity guidelines, limitations and restrictions, the most common antibiotics within each class and references utilized.    Please note that all services described in this policy require prior authorization.   Please refer to the member's contract benefits in effect at the time of service to determine coverage or non-coverage of these services as it applies to an individual member.   Providers should report all services using the most up-to-date industry-standard procedure, revenue, and diagnosis codes, including modifiers where applicable.   Providers must submit all required and requested documentation for case evaluation and determination including but not limited to the following: Medical Order with all required documentation, History of previous and or failed treatment, History of allergies, General information related to diagnosis which include but is not limited to the following: CBS, Urinalysis, Urine Culture, Blood Culture, Studies that confirm diagnosis Ex: MRI, Bone Scan, Infectology and or Surgery Consults.   The plan may request additional documentation and information not received and or provided initially related to condition and diagnosis for case evaluation and determination.   Any additional documentation submitted specifying medical necessity criteria and considered important for case evaluation and determination will be reviewed by Clinical Team utilizing guidelines and regulation criteria.	Policy Name	Policy Number	Scope	
Anesthesia	IV Antibiotics	MP-IV-FP-01-24		
□ Anesthesia □ Medicine Services and Procedures □ Surgery □ Evaluation and Management Services □ DME/Prosthetics or Supplies			⊠ MMM MA	☐ MMM Multihealth
□ Surgery □ DME/Prosthetics or Supplies □ Pathology and Laboratory Procedures □ DME/Prosthetics or Supplies □ Pathology and Laboratory Procedures ☑ Other IV Antibiotics  Service Description  Enclosed document contains but is not limited to the following information: classification of antibiotics, medical necessity guidelines, limitations and restrictions, the most common antibiotics within each class and references utilized.  Please note that all services described in this policy require prior authorization.  ■ Please refer to the member's contract benefits in effect at the time of service to determine coverage or non-coverage of these services as it applies to an individual member. ■ Providers should report all services using the most up-to-date industry-standard procedure, revenue, and diagnosis codes, including modifiers where applicable. ■ Providers must submit all required and requested documentation for case evaluation and determination including but not limited to the following: Medical Order with all required documentation, History of previous and or failed treatment, History of allergies, General information related to diagnosis which include but is not limited to the following: CBS, Urinalysis, Urine Culture, Blood Culture, Studies that confirm diagnosis Ex: MRI, Bone Scan, Infectology and or Surgery Consults. ■ The plan may request additional documentation and information not received and or provided initially related to condition and diagnosis for case evaluation and determination. ■ Any additional documentation submitted specifying medical necessity criteria and considered important for case evaluation and determination will be reviewed by Clinical Team utilizing guidelines and regulation criteria.  ■ Sanford Guidelines are utilized for review of antibiotics criteria and recommended therapy.	Service Category	<u>i</u>	<u>i</u>	
□ Radiology Procedures □ DME/Prosthetics or Supplies □ Pathology and Laboratory Procedures ☑ Other IV Antibiotics  Service Description  Enclosed document contains but is not limited to the following information: classification of antibiotics, medical necessity guidelines, limitations and restrictions, the most common antibiotics within each class and references utilized.  Please note that all services described in this policy require prior authorization.  • Please refer to the member's contract benefits in effect at the time of service to determine coverage or non-coverage of these services as it applies to an individual member.  • Providers should report all services using the most up-to-date industry-standard procedure, revenue, and diagnosis codes, including modifiers where applicable.  • Providers must submit all required and requested documentation for case evaluation and determination including but not limited to the following: Medical Order with all required documentation, History of previous and or failed treatment, History of allergies, General information related to diagnosis which include but is not limited to the following: CBS, Urinalysis, Urine Culture, Blood Culture, Studies that confirm diagnosis Ex: MRI, Bone Scan, Infectology and or Surgery Consults.  • The plan may request additional documentation and information not received and or provided initially related to condition and diagnosis for case evaluation and determination.  • Any additional documentation submitted specifying medical necessity criteria and considered important for case evaluation and determination will be reviewed by Clinical Team utilizing guidelines and regulation criteria.	☐ Anesthesia	☐ Medicir	ne Services and Pro	ocedures
□ Pathology and Laboratory Procedures ☑ Other IV Antibiotics  Service Description  Enclosed document contains but is not limited to the following information: classification of antibiotics, medical necessity guidelines, limitations and restrictions, the most common antibiotics within each class and references utilized.  Please note that all services described in this policy require prior authorization.  • Please refer to the member's contract benefits in effect at the time of service to determine coverage or non-coverage of these services as it applies to an individual member.  • Providers should report all services using the most up-to-date industry-standard procedure, revenue, and diagnosis codes, including modifiers where applicable.  • Providers must submit all required and requested documentation for case evaluation and determination including but not limited to the following: Medical Order with all required documentation, History of previous and or failed treatment, History of allergies, General information related to diagnosis which include but is not limited to the following: CBS, Urinalysis, Urine Culture, Blood Culture, Studies that confirm diagnosis Ex: MRI, Bone Scan, Infectology and or Surgery Consults.  • The plan may request additional documentation and information nor received and or provided initially related to condition and diagnosis for case evaluation and determination.  • Any additional documentation submitted specifying medical necessity criteria and considered important for case evaluation and determination will be reviewed by Clinical Team utilizing guidelines and regulation criteria.  • Sanford Guidelines are utilized for review of antibiotics criteria and recommended therapy.	☐ Surgery	☐ Evaluat	ion and Manageme	ent Services
<ul> <li>Service Description</li> <li>Enclosed document contains but is not limited to the following information: classification of antibiotics, medical necessity guidelines, limitations and restrictions, the most common antibiotics within each class and references utilized.</li> <li>Please note that all services described in this policy require prior authorization.</li> <li>Please refer to the member's contract benefits in effect at the time of service to determine coverage or non-coverage of these services as it applies to an individual member.</li> <li>Providers should report all services using the most up-to-date industry-standard procedure, revenue, and diagnosis codes, including modifiers where applicable.</li> <li>Providers must submit all required and requested documentation for case evaluation and determination including but not limited to the following: Medical Order with all required documentation, History of previous and or failed treatment, History of allergies, General information related to diagnosis which include but is not limited to the following: CBS, Urinalysis, Urine Culture, Blood Culture, Studies that confirm diagnosis Ex: MRI, Bone Scan, Infectology and or Surgery Consults.</li> <li>The plan may request additional documentation and information not received and or provided initially related to condition and diagnosis for case evaluation and determination.</li> <li>Any additional documentation submitted specifying medical necessity criteria and considered important for case evaluation and determination will be reviewed by Clinical Team utilizing guidelines and regulation criteria.</li> <li>Sanford Guidelines are utilized for review of antibiotics criteria and recommended therapy.</li> </ul>	☐ Radiology Procedures	☐ DME/P	rosthetics or Suppli	ies
<ul> <li>Enclosed document contains but is not limited to the following information: classification of antibiotics, medical necessity guidelines, limitations and restrictions, the most common antibiotics within each class and references utilized.</li> <li>Please note that all services described in this policy require prior authorization.</li> <li>Please refer to the member's contract benefits in effect at the time of service to determine coverage or non-coverage of these services as it applies to an individual member.</li> <li>Providers should report all services using the most up-to-date industry-standard procedure, revenue, and diagnosis codes, including modifiers where applicable.</li> <li>Providers must submit all required and requested documentation for case evaluation and determination including but not limited to the following: Medical Order with all required documentation, History of previous and or failed treatment, History of allergies, General information related to diagnosis which include but is not limited to the following: CBS, Urinalysis, Urine Culture, Blood Culture, Studies that confirm diagnosis Ex: MRI, Bone Scan, Infectology and or Surgery Consults.</li> <li>The plan may request additional documentation and information not received and or provided initially related to condition and diagnosis for case evaluation and determination.</li> <li>Any additional documentation submitted specifying medical necessity criteria and considered important for case evaluation and determination will be reviewed by Clinical Team utilizing guidelines and regulation criteria.</li> <li>Sanford Guidelines are utilized for review of antibiotics criteria and recommended therapy.</li> </ul>	☐ Pathology and Laboratory Procedures	⊠ <u>Other I</u>	<u>V Antibiotics</u>	
<ul> <li>medical necessity guidelines, limitations and restrictions, the most common antibiotics within each class and references utilized.</li> <li>Please note that all services described in this policy require prior authorization.</li> <li>Please refer to the member's contract benefits in effect at the time of service to determine coverage or non-coverage of these services as it applies to an individual member.</li> <li>Providers should report all services using the most up-to-date industry-standard procedure, revenue, and diagnosis codes, including modifiers where applicable.</li> <li>Providers must submit all required and requested documentation for case evaluation and determination including but not limited to the following: Medical Order with all required documentation, History of previous and or failed treatment, History of allergies, General information related to diagnosis which include but is not limited to the following: CBS, Urinalysis, Urine Culture, Blood Culture, Studies that confirm diagnosis Ex: MRI, Bone Scan, Infectology and or Surgery Consults.</li> <li>The plan may request additional documentation and information not received and or provided initially related to condition and diagnosis for case evaluation and determination.</li> <li>Any additional documentation submitted specifying medical necessity criteria and considered important for case evaluation and determination will be reviewed by Clinical Team utilizing guidelines and regulation criteria.</li> <li>Sanford Guidelines are utilized for review of antibiotics criteria and recommended therapy.</li> </ul>	Service Description			
<ul> <li>Please refer to the member's contract benefits in effect at the time of service to determine coverage or non-coverage of these services as it applies to an individual member.</li> <li>Providers should report all services using the most up-to-date industry-standard procedure, revenue, and diagnosis codes, including modifiers where applicable.</li> <li>Providers must submit all required and requested documentation for case evaluation and determination including but not limited to the following: Medical Order with all required documentation, History of previous and or failed treatment, History of allergies, General information related to diagnosis which include but is not limited to the following: CBS, Urinalysis, Urine Culture, Blood Culture, Studies that confirm diagnosis Ex: MRI, Bone Scan, Infectology and or Surgery Consults.</li> <li>The plan may request additional documentation and information not received and or provided initially related to condition and diagnosis for case evaluation and determination.</li> <li>Any additional documentation submitted specifying medical necessity criteria and considered important for case evaluation and determination will be reviewed by Clinical Team utilizing guidelines and regulation criteria.</li> <li>Sanford Guidelines are utilized for review of antibiotics criteria and recommended therapy.</li> </ul>	medical necessity guidelines, limitations ar references utilized.	nd restrictions, the most	common antibiotic	
<ul> <li>or non-coverage of these services as it applies to an individual member.</li> <li>Providers should report all services using the most up-to-date industry-standard procedure, revenue, and diagnosis codes, including modifiers where applicable.</li> <li>Providers must submit all required and requested documentation for case evaluation and determination including but not limited to the following: Medical Order with all required documentation, History of previous and or failed treatment, History of allergies, General information related to diagnosis which include but is not limited to the following: CBS, Urinalysis, Urine Culture, Blood Culture, Studies that confirm diagnosis Ex: MRI, Bone Scan, Infectology and or Surgery Consults.</li> <li>The plan may request additional documentation and information not received and or provided initially related to condition and diagnosis for case evaluation and determination.</li> <li>Any additional documentation submitted specifying medical necessity criteria and considered important for case evaluation and determination will be reviewed by Clinical Team utilizing guidelines and regulation criteria.</li> <li>Sanford Guidelines are utilized for review of antibiotics criteria and recommended therapy.</li> </ul>	Please note that all services described in t	this policy require prior a	uthorization.	
<ul> <li>Guideline utilized for case determination will be furnished upon request.</li> <li>The approval of IV antibiotics is subject to step therapy in instances when an oral formulation of</li> </ul>	<ul> <li>Providers should report all services and diagnosis codes, including modes are determination including but not lir documentation, History of previous related to diagnosis which include Blood Culture, Studies that confirm</li> <li>The plan may request additional desinitially related to condition and disconding and disconding and regulation criteria.</li> <li>Sanford Guidelines are utilized for Guideline utilized for case determinations and services.</li> </ul>	as it applies to an individes using the most up-to-dated differs where applicable. It and requested documer mited to the following: Mais and or failed treatment but is not limited to the following is not limited to the following is for case evaluation and informations is for case evaluation with the following medical determination will be rever review of antibiotics critical in the following in the following is an individual of the following in the following is a subject to the following in the following is a subject to the following is a subject to the following in the following is a subject to the following i	ual member.  ate industry-standa  ntation for case eva  ledical Order with a  t, History of allergie  following: CBS, Urin  Scan, Infectology a  nation not received  on and determinati  al necessity criteria  viewed by Clinical T  iteria and recomme  I upon request.	rd procedure, revenue, fluation and fall required fes, General information finalysis, Urine Culture, fand or Surgery Consults. I and or provided fion. for and considered feam utilizing guidelines feeded therapy.



## **Medical Necessity Guidelines**

Service Description	Medical Necessity Guidelines	Limits or Restrictions	Most Common Antibiotics
1. Penicillins	Penicillin is one of the most commonly used antibiotics globally; it has a wide range of clinical indications. It is also considered one of the strongest. Interrupts proliferation of the bacteria.  Penicillin is effective against many different infections involving gram-positive cocci, gram-positive rods (e.g., Listeria), most anaerobes, and gram-negative cocci (e.g., Neisseria).  Importantly, certain bacterial species have obtained penicillin resistance, including enterococci. Enterococci infections now receive treatment with a combination of penicillin and streptomycin or gentamicin.  Certain gram-negative rods are also resistant to penicillin due to penicillin's poor ability to penetrate the porin channel.  However, later generations of broad-spectrum penicillins are effective against gram-negative rods.  Second-generation penicillins (ampicillin and amoxicillin) can also penetrate the porin channel, making these drugs effective against Proteus mirabilis, Shigella, H. influenzae, Salmonella, and E. coli.  Third-generation penicillin is also able	It's important to note that penicillins may interfere with the effectiveness of birth control pills.  Some individuals exhibit a severe allergic reaction to penicillin known as anaphylaxis. Anaphylaxis is a potentially lifethreatening condition that causes dysfunction in several body systems.  Penicillins and other betalactams do not penetrate well into phagocytes, thus limiting their ability to kill intracellular pathogens. In addition, penicillins only exert their bactericidal effect on bacteria that are actively replicating.	Pennicillin G Pennicillin VK Nafcillin Oxacillin Cloxacillin Flucoxacillin Ampicillin Amoxicillin Amox-Clav Amp-Sulb Pip-Tazo



	to penetrate gram-negative			
	bacterial porin channels.			
	Fourth-generation penicillins			
	such as piperacillin are			
	effective against the same			
	bacterial strains as third-			
	generation penicillins and			
	Klebsiella, enterococci,			
	Pseudomonas aeruginosa, and			
	Bacteroides fragilis.			
	Penicillins are commonly used			
	for the following conditions:			
	Pneumonia, Tonsillitis, Dental			
	Abscess, Strep Throat, Urinary			
	Tract.			
2. Carabalaan arina		Cida effects and similar to	C-f!:	
2. Cephalosporins	These types of antibiotics are	Side effects are similar to	Cefazolin	
	usually grouped into categories	those experienced with	Cefotetan	
	that are called generations.	penicillin. These include	Cefoxitin	
	There are five generations of	nausea, diarrhea, rash and	Cefuroxime	
	cephalosporins. The first	thrush. If someone is	Cefotaxime	
	generation of these antibiotics	allergic to penicillins it is	Ceftizoxime	
	is usually used for infections	likely they will be allergic	Cefuroxime	
	that are easier to treat. The	to cephalosporins since	Cefoperazone	
	latter generations are for more	they are similar in	Ceftriaxone	
	serious bacterial infections.	molecular structure.	Ceftazidime	
	Cephalosporins are often used	Depending on how severe	Cefepime	
		the allergy is, some	Ceftaz-Avibac	
	for strep throat, meningitis,			
	pneumonia, urinary tract	individuals may be able to	Ceftaroline	
	infections and ear infections.	still take third, fourth or	Ceftobiprole	
		fifth generation	Ceftobiprole	
	The fifth generation of	cephalosporins.	Cefto-Tazo	
	cephalosporins is called		Cefuderocol	
	Ceftaroline and is used for	Cephalosporins have the		
	antibiotic resistant infections	following limitations: Lack		
	such as MRSA.	of activity against		
		enterococci. Enterococcus		
	The cephalosporins that are	faecalis and E. faecium		
	primarily prescribed include	cause a variety of		
	1			
	cephalexin, cefaclor and	infections, including		
	ceftriaxone (as an injection).	endocarditis, urinary tract		
		infections.		
	Cefazolin, cefuroxime and			
	cefoxitin are not used as often			
	and normally prescribed for			
	individuals with cystic fibrosis			
	or those undergoing dialysis.			
3. Carbapenems	They are a class of antibiotics	Adverse effects include	Doripenem	
J. Carbapenenti	also known as beta lactam.	increased resistance to one	Ertapenem	
İ	also kilowii as beta lactaili.	moreusea resistance to one	Litapeneni	



They work by inhibiting synthesis of the bacterial cell wall. Carbapenems are often used for serious urinary infections, abdominal infections, blood infections and pneumonia.

Carbapenems possess the broadest spectrum of activity and greatest potency against Gram-positive and Gramnegative bacteria. As a result, they are often used as "last-line agents" or "antibiotics of last resort" when patients with infections become gravely ill or are suspected of harboring resistant bacteria.

The carbapenem antibiotics and their role in our antimicrobial armamentarium. Among the β-lactams currently available, carbapenems are unique because they are relatively resistant to hydrolysis by most βlactamases, in some cases act as "slow substrates" or inhibitors of β-lactamases, and still target penicillin binding proteins. This "value-added feature" of inhibiting βlactamases serves as a major rationale for expansion of this class of βlactams. Interferes with membrane proteins.

The most common are: Mero-Meropenem, IMP-cila -rele, Imp-cilastatin, Ertapenem, Doripenem

Doripenem, ertapenem, imipenem, and meropenem are each drugs in the Carbapenem class that are usually

of the drugs used in the combination, as well as a lack of synergy or additivity and strain dependence.

Carbapenems have low oral bioavailability and thus do not cross gastrointestinal membranes readily and must be administered intravenously.

Are eliminated predominately by renal excretion. Carbapenems exhibit unique pharmacological properties and are typically used to treat complicated bacterial infections. A carbapenem is often combined with an antibiotic that targets Gram-positive bacteria when used for the empirical treatment of patients with serious nosocomial infections of unidentified origin.

Safety and tolerability.
Nephrotoxicity,
neurotoxicity, and
immunomodulation have
been reported with the use
of carbapenems, and thus
predisposing factors should
be considered when
administering any
carbapenem, they alter the
intestinal microflora and
select for carbapenemresistant isolates.

Imp-cilastatin Imp-cila-rele Meropenem Mero-Vabor Aztreonam



,					
		administered intravenously or			
		injected into a muscle. These			
		drugs are often prescribed for			
		infections that aren't easily			
		treated with other antibiotics.			
		treated with other antibiotics.			
		Combonomono ono sincilor to			
		Carbapenems are similar to			
		penicillins. These types of			
		antibiotics, however, so far			
		seem unaffected by the			
		increasing problem of antibiotic			
		resistance.			
	4. Fluoroquinolone	The fluoroquinolones are a	It is generally	Ciprofloxacin	
	·	family of broad spectrum,	recommended to use these	Delafloxacin	
		systemic antibacterial agents	antibiotics only after other	Gemifloxacin	
		that have been used widely as	courses of treatment have	Ofloxacin	
		therapy of respiratory and	failed.	Levofloxacin	
			ianeu.	Moxifloxacin	
		urinary tract infections.	Elimenticales I		
		Interferes with bacteria DNA	Fluroquinolones have also	Norfloxacin	
		replication and transcription.	been linked in recent years	Prulifloxacin	
			to mental health problems,	Gemifloxacin	
		Fluoroquinolones are active	disturbances with blood	Gatifloxacin	
		against a wide range of aerobic	sugar and specifically aortic		
		gram-positive and gram-	aneurysms.		
		negative organisms.			
		Gram-positive	Within the last year the		
		coverage includes	FDA has required labeling		
		penicillinase- and non-	changes to strengthen the		
		•	warnings. There may be		
		penicillinase			
		producing	some cases, however, such		
		Staphylococci,	as when treating bacterial		
		Streptococcus	pneumonia, that the		
		pneumoniae and	potential benefits		
		viridans, Enterococcus	outweigh the risks. Serious		
		faecalis, Listeria	cases of pneumonia and		
		monocytogenes, and	abdominal infections may		
		Nocardia species.	require the use of		
		Gram negative	fluoroquinolones.		
		coverage includes			
		Neisseria meningitides			
		and gonorrhoeae,			
		Haemophilus			
		influenzae, and most			
		clinically important			
		Enterobacteriaceae			
		species, Pseudomonas			
		aeruginosa and Vibrio			
		species.			
	5. Aminoglycosides	The aminoglycosides are	The aminoglycosides all	Gentamicin	1
		natural products and	have serious toxicities	Tobramycin	
		I natarai products and	Have Serious toxicities	Tobramyoni	1



semisynthetic derivatives from a variety of actinomycetes and have potent activity against many gram-negative bacteria. The first aminoglycoside used in clinical practice was streptomycin which was derived from Streptomyces griseus and was the first effective agent against mycobacterium tuberculosis. The aminoglycosides are believed to act by binding to ribosomes of bacteria and blocking protein synthesis.

The aminoglycosides are poorly absorbed orally and typically are given parenterally, either by intravenous or intramuscular injection. Gentamicin, tobramycin and amikacin are given parenterally and are used for severe gram negative bacterial infections usually in combination with penicillins or cephalosporins. Streptomycin is now rarely used and largely as adjunctive therapy of multi-drug resistant tuberculosis. Plazomicin is a recently introduced agent and is given intravenously as monotherapy for complicated urinary tract infections or acute pyelonephritis. Plazomicin is a semi-synthetic aminoglycoside which has been modified to evade conventional forms of aminoglycoside resistance. Neomycin is used orally to treat hepatic encephalopathy. Because it is poorly absorbed orally, neomycin causes a decrease in intestinal bacteria, thereby decreasing ammonia production and absorption from the colon.

which often limit their applicability and the dose and duration of therapy. The common serious adverse effects of the aminoglycosides are ototoxicity, neuropathy, and nephrotoxicity.

Liver injury from the aminoglycosides is rare, perhaps because the other side effects of aminoglycosides limit the amount that can be given. Isolated case reports of idiosyncratic hepatotoxicity have been published for most, but not all of the aminoglycosides.

Amikacin Plazomicin



6	. Macrolides	Aminoglycosides are broadspectrum bactericidal antibiotics used mainly to treat aerobic Gram-negative bacteria and selected Grampositive bacteria often in combination with other antibiotics.  Aminoglycosides entered widespread clinical use to combat infections caused by members of the Enterobacterales order of Gram-negatives including Escherichia coli and Klebsiella pneumonia (Krause et al. 2016), and they have also been used effectively against Pseudomonas aeruginosa (Karlowsky et al. 2003) and Staphylococcus aureus (Lee and Lee 2016).  They are usually given as oral medication. Macrolides are often used to treat very basic bacterial infections.  Inhibits synthesis of proteins by bacteria, occasionally leading to cell death.  These antibiotics are often used for specific types of pneumonia, chlamydia, and urethritis. Macrolides are sometimes prescribed to prevent a bacterial infection.  If an individual has had their spleen removed or suffers from sickle-cell disease the person may need to use one of these antibiotics on a regular basis to prevent an infection.  Specific drugs in this class include roxithromycin, clarithromycin, azithromycin, clarithromycin, azithromycin, and erythromycin.	Minor side effects can include nausea, diarrhea and ringing in the ears.  Macrolides are often a good alternative for individuals that are allergic to penicillins or cephalosporins. However, potential complications regarding these antibiotics are that they do have some drug interaction concerns that could lead to serious heart complications.	Eriythromycin Azithromycin Clarithromycin Telithromycin	
---	--------------	---	---	--	--



7. Tetracyclines	Tetracyclines (tetracycline,	The most common side	Doxycicline
•	doxycycline, minocycline,	effects may include	Eravacycline
	tigecycline) are a class of	nausea, diarrhea, swollen	Minocycline
	medication used to manage	tongue, troubling	Omadacycline
	and treat various bacterial	swallowing and soreness or	Tetracycline
	infections.	swelling in the genital area.	Tigecycline
	Tetracyclines classify as protein		0 ,
	synthesis inhibitor antibiotics	A rare but potential serious	
	and are considered to be	side effect is possible	
	broad-spectrum.	blindness due to	
	· ·	intracranial hypertension.	
	Tetracyclines activity against a	,,	
	wide range of microorganisms	Tetracycline should be	
	including gram-positive and	taken on an empty	
	gram-negative bacteria,	stomach, at least 1 hour	
	chlamydiae, mycoplasmas,	before or 2 hours after	
	rickettsiae, and protozoan	meals or snacks. Drink a	
	parasites.	full glass of water with	
	·	each dose of tetracycline.	
	Tetracycline resistance now	Do not take tetracycline	
	occurs in an increasing number	with food, especially dairy	
	of pathogenic, opportunistic,	products such as milk,	
	and commensal bacteria. The	yogurt, cheese, and ice	
	presence of tetracycline-	cream.	
	resistant pathogens limits the		
	use of these agents in	Tetracyclines are	
	treatment of disease.	contraindicated in	
		pregnancy because of the	
	Tetracycline resistance is often	risk of hepatotoxicity in the	
	due to the acquisition of new	mother, the potential for	
	genes, which code for energy-	permanent discoloration of	
	dependent efflux of	teeth in the fetus (yellow	
	tetracyclines or for a protein	or brown in appearance),	
	that protects bacterial	as well as impairment of	
	ribosomes from the action of	fetal long bone growth.	
	tetracyclines. Many of these	Tetracycline usage is also	
	genes are associated with	associated with teeth	
	mobile plasmids or	discoloration in children	
	transposons and can be	under the age of eight.	
	distinguished from each other	Thus, it should be avoided	
	using molecular methods	in pediatric patients under	
	including DNA-DNA	that age.	
	hybridization with		
	oligonucleotide probes and	Clinicians should also avoid	
	DNA sequencing.	tetracyclines in patients	
		with renal failure due to	
	A limited number of bacteria	the excretion of the drug	
	acquire resistance by	being primarily by the	
	mutations, which alter the	kidneys. If tetracyclines	
	permeability of the outer	must be used in this group	



membrane porins and/or lipopolysaccharides in the outer membrane, change the regulation of innate efflux systems, or alter the 16S rRNA.  These drugs can treat rickettsial infections, ehrlichiosis, anaplasmosis, leptospirosis, amebiasis, actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
outer membrane, change the regulation of innate efflux systems, or alter the 16S rRNA.  These drugs can treat rickettsial infections, ehrlichiosis, anaplasmosis, leptospirosis, amebiasis, actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
regulation of innate efflux systems, or alter the 16S rRNA.  These drugs can treat rickettsial infections, ehrlichiosis, anaplasmosis, leptospirosis, amebiasis, actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
systems, or alter the 16S rRNA.  These drugs can treat rickettsial infections, ehrlichiosis, anaplasmosis, leptospirosis, amebiasis, actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
These drugs can treat rickettsial infections, ehrlichiosis, anaplasmosis, leptospirosis, amebiasis, actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
rickettsial infections, ehrlichiosis, anaplasmosis, leptospirosis, amebiasis, actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
rickettsial infections, ehrlichiosis, anaplasmosis, leptospirosis, amebiasis, actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
ehrlichiosis, anaplasmosis, leptospirosis, amebiasis, actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
leptospirosis, amebiasis, actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
actinomycosis, nocardiosis, brucellosis, melioidosis, tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
tularemia, chlamydial infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
infections, pelvic inflammatory disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
disease, syphilis, traveler's diarrhea, early Lyme disease, acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
acne, legionnaire's disease, and Whipple disease. They cover Borrelia recurrentis,
and Whipple disease. They cover Borrelia recurrentis,
cover Borrelia recurrentis,
March actorium marrinum
Mycobacterium marinum,
Mycoplasma pneumoniae,
Staphylococcus aureus
(including methicillin-resistant
S. aureus [MRSA]), Vibrio
vulnificus, and vancomycin-
resistant enterococcus (VRE)
(susceptible strains).
Meningococcal prophylaxis is
also achievable.
Other indications of
tetracyclines include rosacea,
bullous dermatoses,
sarcoidosis, Kaposi sarcoma,
pyoderma gangrenosum,
hidradenitis suppurativa,
Sweet syndrome, a1-
antitrypsin deficiency,
panniculitis, pityriasis
lichenoides chronica,
rheumatoid arthritis,
scleroderma, cancer, and
cardiovascular diseases
(abdominal aortic aneurysm
and acute myocardial
infarction).
8. Glico-Lipo The term glycopeptide refers to Daptomycin
a group of antimicrobial agents Vancomycin
a proup of antimicropial agents   Valicomyem
that includes vancomycin and Teicoplanin



	VISA isolates in the United States were also resistant to teicoplanin, the term glycopeptide-intermediate S. aureus (GISA) was used to indicate this broader resistance profile.	Oritavancin Dalbavancin	
	While GISA may be a more specific term for strains intermediate to both vancomycin and teicoplanin, not all VISA strains are intermediate to teicoplanin; therefore, VISA is a more accurate and widely used term.		
 9. Ox-Lid Oxazolidinones)	Oxazolidinones are a new class of antibiotics used to treat serious skin and bacterial infections, often after other antibiotics have been ineffective.	Linezolide Tedizoline	
	Target protein synthesis in a wide spectrum of grampositive and anaerobic bacteria. Inhibits synthesis of proteins by bacteria, preventing growth.		
	Oxazolidinones are a recent class of synthetic antibiotics with a chemical structure characterized by a basic nucleus of 2-oxazolidone active against a wide spectrum of multidrug-resistant Grampositive bacteria (GPB), namely vancomycin-resistant Enterococcus (VRE), MRSA and Mycobacterium tuberculosis (Mtb).		
	Oxazolidinones bind to the 50S ribosomal subunit, inhibiting the biosynthesis of bacterial proteins. The first oxazolidinone clinically available was Linezolid (LNZ), discovered in 1996 and		



	approved in 2000 for clinical			
	use by the FDA (U.S. Food and			
	Drug Administration). LNZ is			
	widely employed for GPB			
	infections, and it is considered			
	an efficient drug for surgical			
	infections and in the treatment			
	of drug-resistant pulmonary			
	infections and MDR-TB			
	infections.			
	infections.			
	Among oxazolidinones, only			
	LNZ and Tedizolid are clinically			
	approved for MDR-TB			
	infections. Tedizolid (TZD)			
	belongs to the second			
	generation of oxazolidinones			
	_			
	and is also indicated for the			
	treatment of skin infections.			
	Radezolid (RZD), belonging to			
	the biaryl oxazolidinone family,			
	is effective against resistant			
	LNZ strains. Although clinical			
	trials into community-acquired			
	pneumonia and into skin and			
	soft tissue infections have			
	concluded, studies on its			
	acceptability are not yet			
	finished.			
	In the field of treating MDR-TB			
	infections, many efforts have			
	been made to discover the			
	next generation of			
	oxazolidinones having better			
	antibacterial efficacy and fewer			
	adverse effects. Recently,			
	several oxazolidinone analogs			
	_			
	have been developed at well-			
	known pharmaceutical			
	companies, some of which			
	have been found to be suitable			
	for treating MDR-TB.			
10. Poly	Polymyxins comprise a class of	Hypersensitivity to	Polymyxin B	
	antibiotics targeting gram-	polymyxin B, colistin	Colistin	
			Lefamulin	
	negative bacterial infections.	methanesulfonate, colistin,	Leidiliulli	
		or any formulation		
	Polymyxin B and Polymyxin E	component.		
	(colistin) are the two drugs			
 				i



within this antibiotic class used primarily in clinical practice. They are FDA approved for serious infections with multidrug-resistant gramnegative bacteria, especially those caused by Enterobacteriaceae, Pseudomonas aeruginosa and Acinetobacter baumannii.

Polymyxins are often the only effective antibiotic agent against multidrug-resistant organisms, particularly carbapenem-resistant Enterobacteriaceae. They have become the last line of treatment for infections that are resistant to other antibiotics. They are useful in treating infections of the urinary tract, meninges, and bloodstream by susceptible strains of pseudomonas aeruginosa, Enterobacteriaceae, and Acinetobacter baumannii.

Drugs act on the outer membrane of gram-negative bacteria by destabilizing the phospholipids and lipopolysaccharides (LPS) present. There is an electrostatic interaction between the positively charged polymyxin and the phosphate groups of the negatively charged lipid A membrane, which causes displacement of divalent cations such as calcium and magnesium from the phosphate groups within these membrane lipids. This activity leads to increased permeability, a disrupted outer cell membrane, and intracellular contents begin to

Renal function requires close monitored during the administration of intravenous polymyxins as a result of the high frequency of nephrotoxicity and potential severity.

Therapeutic drug monitoring of polymyxins is also a recommendation due to a narrow therapeutic window for efficacy and toxicity. However, therapeutic drug monitoring for the polymyxins is not universally available. Decreasing urine output, increasing BUN, and creatinine may require discontinuation of systemic therapy with polymyxins.

The recommended target serum concentration level is 2 mg/mL for susceptible strains.



	leak out, resulting in cellular		
	bacterial death.		
11. Anti fungals	Fungi are unicellular or multi-	All formulations of	Amphotericin B
	cellular eukaryotic organisms	amphotericin B (AMB-d, L-	Micafungin
	that exist in all environments	AMB, ABLC, ABCD) are	Casposfungin
	worldwide. While most fungi	contraindicated in patients	Anidulafungin
	do not play a significant role in	with a known or likely	Isavuconazonium
	human disease, there are	hypersensitivity to	Sulfate
	several hundred fungi that do,	amphotericin B or any	Posaconazole
	resulting in fungal infection or	components of the L-AMB,	Voriconazole
	disease. Fungal infections	ABLC, or ABCD	Itraconazole
	(mycoses) range from common	formulations.	Fluconazole
	benign infections like 'jock itch'		
	to serious, life-threatening	Nystatin is contraindicated	
	infections such as cryptococcal	in patients with	
	meningitis. Antifungal	hypersensitivity to the drug	
	antimicrobials are one drug	or any additional	
	class that can combat these	components in the dosage	
	mycoses.	formulation.	
		Torritalation.	
	Clinically, fungal infections are	All azoles should be	
	best categorized first according		
	to the site and extent of the	avoided in patients with	
	infection, then the route of	hypersensitivities to azole	
	acquisition, and finally, the	drugs or dosage form	
	virulence of the causative	components and used with	
	organism. These classifications	caution in patients with	
	are essential when	renal impairment/failure	
	determining the most effective	and or hepatic	
	treatment regimen for a	impairment/failure.	
	particular mycosis. Mycoses		
	classify as local (superficial,	Fluconazole requires	
	cutaneous, subcutaneous) or	cautious administration in	
	systemic (deep, bloodborne).	patients with electrolyte	
	The acquisition of the fungal	abnormalities, torsades de	
	infection is either an	pointes, and or medical	
	exogenous	history, family history, and	
	(airborne/inhalation,	or current QTc	
	cutaneous exposure,	prolongation.	
	percutaneous inoculation) or	Itraconazole has an FDA	
	an endogenous process	boxed warning against the	
	(normal flora or reactivated	use in treating	
	infection). The virulence of the organism is classified as either	onychomycosis in patients with CHF. Itraconazole is	
		contraindicated in	
	a primary infection (disease	pregnancy, left ventricular	
	arising in a healthy host) or	dysfunction, and current or	
	opportunistic infection	active congestive heart	
	(disease arising in human hosts that have a compromised	failure. This drug should be	
	-	used cautiously in patients	
 <u> </u>	immune system or other	asca cautiously iii patietits	



defenses).

Aspergillosis - Aspergillus fumigatus, A. flavus Blastomycosis - Blastomyces dermatitidis Candidiasis - Candida albicans, C. glabrata, C. krusei, C. parasilosis, C. tropicalis Chromoblastomycosis (Chromomycosis) -Cladosporium carrionii, Phialophora verrucosa, Fonsecaea pedrosoi Coccidioidomycosis -Coccidioides imitis, C. posadasii Cryptococcosis - Cryptococcus neoformans, C. gattii Dermatophytosis (Tinea) -Microsporum spp., Epidermophytum spp., Trichophyton spp. Fusariosis - Fusarium oxysporum, F. proliferatum, F. verticillioides Histoplasmosis - Histoplasma capsulatum Mucormycosis (Zygomycosis) -Mucor spp., Rhizopus spp. Paracoccidioidomycosis -Paracoccidioides brasiliensis Pneumocystis pneumonia -Pneumocystis jirovecii (formerly called P. carinii)\* \*While this is an essential and prevalent fungal disease, it is not treated with typical antifungal agents. Sporotrichosis - Sporothrix schenckii Tinea (Pityriasis) Versicolor -Malassezia furfur (also called Pityrosporum orbiculare), M. globosa

with cystic fibrosis, cardiovascular disease, pulmonary disease, and the elderly. Ketoconazole carries several FDA boxed warnings:

- This agent should be used only when another effective antifungal, including azoles, cannot be tolerated or is not available
- This agent carries a significant risk of hepatotoxicity, even in patients without predisposing factors, and thus any treatment with ketoconazole should include close liver function monitoring.
- Ketoconazole has several contraindicated drug interactions that may cause QTc prolongation by increasing concentrations of cisapride, disopyramide, dofetilide, dronedarone, methadone, quinidine, or ranolazine. Ketoconazole is a cytochrome P450 inhibitor.

Voriconazole is contraindicated in galactose malabsorption/intolerance, Lapp lactase deficiency, glucose malabsorption, uncorrected electrolyte abnormalities, and



pregnancy. Clinicians should use this agent with caution in patients with a medical or family history of QTc prolongation, history of torsades de pointes, and or hematologic malignancy. Isavuconazole is contraindicated in patients with familial short QTc syndrome and should be used with caution in patients with hematologic malignancies. Posaconazole is contraindicated in pregnancy. Caution is advisable in patients with electrolyte abnormalities, renal insufficiency, cardiomyopathy, torsades de pointes, or medical history/family history/congenital prolonged QTc interval. Terbinafine should be utilized with caution or avoided in patients with hypersensitivity reactions, depression, gastrointestinal issues, liver failure, and immune suppression secondary to hematologic effects. All echinocandins are contraindicated in patients with hypersensitivities to any of the echinocandin drugs or dosage form components. Caspofungin should be used with caution in hepatic impairment. Treatment with griseofulvin should include considerations for potential adverse events in



	susceptible patients and	
	those with existing disease	
	states; particularly patients	
	with a hypersensitivity to	
	griseofulvin, a	
	hypersensitivity to	
	penicillins (there is a	
	possible cross-reaction	
	between penicillins and	
	griseofulvin), hepatic	
	failure, patients with	
	known porphyrias, and	
	patients that are pregnant	
	or nursing.	
	Flucytosine carries an FDA	
	boxed warning that this	
	agent should be used with	
	extreme caution in renal	
	impairment and that	
	hematologic, hepatic, and	
	renal function should have	
	close monitoring. This	
	agent is contraindicated in	
	patients with	
	hypersensitivity to this	
	drug or its components,	
	first trimester pregnancies, and breastfeeding women.	
	Caution is advisable with	
	this agent in patients with	
	-	
	renal impairment, hepatic	
	impairment, bone marrow	
	depression, and pregnant	
	patients in their second or third trimester.	
	The quinolines iodoquinol	
	and clioquinol are	
	contraindicated in patients	
	with hypersensitivities to	
	the drugs or their	
	components.	
	Antifungals, which are	
	utilized only as topical	
	agents, including	
	ciclopirox, potassium	
	iodide, and zinc pyrithione,	
	should be avoided in	
	patients with	
	hypersensitivities to these	
	agents.	

## **Medical Policy**



		Health	care Services Department
			_

## **Medical Policy**



### **Healthcare Services Department**

#### **Reference Information**

Gilbert, David N., M.D., Chambers, Henry F., M.D., Saag, Michael S., M.D., Pavia, Andrew T., M.D., Boucher, Helen W., M.d., **The Sanford Guide to Antimicrobial Therapy 2022**, 22<sup>nd</sup>. Edition.

Links:

Infectious Diseases Society of America <a href="https://www.idsociety.org/practice-guideline/alphabetical-guidelines/">https://www.idsociety.org/practice-guideline/alphabetical-guidelines/</a>

National Library of Medicine https://www.nlm.nih.gov/



## **Policy History**

Date	Version	Comments
12/07/20	Draft	New Medical Policy
23	Diait	New Medical Policy
12/15/20 23	Final	<ul> <li>Approved by Medical Policy Committee with inclusion of the following wording:         The approval of IV antibiotics is subject to step therapy in instances when an oral formulation of the medication is available.     </li> </ul>
04/08/20	Revision/Fi	Removed the following wording from Policy Name/Title:
24	nal	Hydration Administration (IV)/External Infusion Pumps
		Removed from service description the following:
		The general information related nursing services and general antibiotic use.
		Removed wording from the General Description of services specifically in the following section: Please note that all services described in this policy require prior authorization.  • Removed: LCD, Articles and other evidence-based guidelines such as MCG (Milliman) are utilized to determine hydrations, supplies and equipment as per standard regulation and as applicable.
		Removed wording from the following paragraph: Enclosed document contains but is not limited to the following information:  Specifically removed the following wording:  • brief service description
		Removed from Medical Necessity Guidelines the following numbers:  12. Hydration Administration 13. External Infusion Pumps
		Removed the following links from references:  • <a href="https://www.hopkinsguides.com/hopkins/index/Johns_Hopkins_ABX_G">https://www.hopkinsguides.com/hopkins/index/Johns_Hopkins_ABX_G</a> <a href="mailto:uide/Antibiotics">uide/Antibiotics</a>
		A52732 Billing and Coding: Hydration Services <a href="https://www.cms.gov/medicare-coverage-database/view/ncd.aspx">https://www.cms.gov/medicare-coverage-database/view/ncd.aspx</a>
		Guidance on the Treatment of Antimicrobial Resistant Gram-Negative     Infections     https://www.idsociety.org/practice-guideline/amr-guidance/#null



- L33794 External Infusion Pumps
   Medicare Coverage Database
   https://www.cms.gov/medicare-coverage-database/view/ncd.aspx
- National Library of Medicine
   Fluoroquinolones
   https://www.ncbi.nlm.nih.gov/books/NBK547840/
- National Library of Medicine
   Article: Oxazolidinone Antibiotics: Chemical, Biological and Analytical Aspects
   https://www.ncbi.nlm.nih.gov/pmc/articles/PMC8305375/
- US Food and Drug Administration https://www.fda.gov/