Bug Busters: A Clinician's Guide to Infectious Diseases

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This presenter has no current or potential conflicts to report.

Objectives



Understand the principles of infectious disease and summarize antibacterial drugs by class



Identify treatment and prophylaxis of specific infections



Review antifungals, antivirals, and associated disease states



Understand the importance of antimicrobial stewardship and how to apply it to daily practice

Introduction

Infectious diseases are caused by one or more pathogens (like bacteria, fungi, viruses, protozoa, parasites, or infectious proteins called prions

They are transmitted through various mechanisms such as physical contact or airborne transmission

The primary factors that impact treatment decisions include the bug (pathogen), the drug (the antimicrobial), and the patient (host). They should all be thought of together when learning about infectious disease

Mechanisms of Action





Time

Time-Dependent

- Goal: maintain drug level above MIC for most of the dosing interval
- Dosing Strategies: shorter dosing interval, extended or continuous infusion
- Antibiotics: Beta Lactams (penicillins, cephalosporins, carbapenems)

Beta Lactams



PENICILLINS	Antibiotics	Coverage
Natural Penicillins	Penicillin G Penicillin VK	Gram Positive Cocci - Streptococci - Enterococci Gram Positive anaerobes - mouth flora
Aminopenicillins	Amoxicillin Ampicillin	 Streptococci Enterococci Gram positive anaerobes Haemophilus Neisseria Proteus E. coli
Aminopenicillins combined with beta lactamase inhibitor (clavulanate, sulbactam, tazobactam)	Amoxicillin/Clavulanate Ampicillin/Sulbactam	- Adds activity against MSSA, some resistant strains of gram-negative bacteria, gram negative anaerobes (B. fragilis)
Extended Spectrum Penicillins with beta lactamase inhibitor	Piperacillin/tazobactam	Broad Spectrum - adds expanded coverage gram negative bacteria including pseudomonas
Antistaphylococcal penicillins	Dicloxacillin Nafcillin Oxacillin	 Streptococci enhanced MSSA coverage -NO activity against Enterococcus, Gram negative pathogens, and anaerobes

Key Features of Penicillins

Class Effects

• All penicillins increase the risk of seizures if accumulation occurs (i.e. in renal failure)

Oral		
Penicillin VK	First line treatment for strep throat and mild, nonpurulent skin infections (no abscess)	
Amoxicillin	 First line treatment for acute otitis media (80-90 mg/kg/day) Drug of choice for IE prophylaxis before dental procedures (2g x1 30-60 min prior to procedure) 	
Amoxicillin/Clavulanate (Augmentin)	First line treatment for acute otitis media (90 mg/kg/day) and for sinus infections (if antibiotics are indicated)	
Dicloxacillin, oxacillin	Cover MSSANo renal dose adjustment needed	

IE: Infective Endocarditis

Key Features of Penicillins

Parenteral		
Penicillin G Benzathine (Bicillin)	 Drug of choice for syphilis (2.4 million units IM x1) NEVER used IV 	
Nafcillin	Cover MSSANo renal dose adjustment needed	
Piperacillin/Tazobact am (Zosyn)	 Only penicillin active against pseudomonas Extended infusions (4 hours) can be used to maximize T > MIC 	

CEPHALOSPORINS

Generation	Agents	Pearls
First	Cefazolin (Ancef, IV, IM) Cephalexin (Keflex, PO)	Cefazolin is commonly used for MSSA, has good CNS penetration, and commonly used for surgical prophylaxis
	Cefadroxil	Cephalexin is commonly used for skin infections and strep throat
Second	Cefuroxime (PO, IV, IM) Cefotetan (IV, IM)	 Cefuroxime is often used for acute otitis media, community acquired pneumonia (CAP), sinus infections (if antibiotics are indicated)
	Cefaclor (PO) Cefoxitin (IV, IM)	 Cefotetan and Cefoxitin offer anaerobic coverage (B. fragilis) and are commonly used for colorectal surgical prophylaxis Cefotetan can cause a disulfiram-like reaction with alcohol ingestion
Third (group 1)	Cefdinir (PO) Ceftriaxone (IV, IM) Cefotaxime (IV, IM)	 Cefdinir is commonly used for CAP and sinus infections (if antibiotics are indicated)
Cefixime (PO) Cefpodoxime (PO)	Cefixime (PO) Cefpodoxime (PO)	 Ceftriaxone and Cefotaxime are commonly used for CAP, meningitis, spontaneous bacterial peritonitis (SBP), pyelonephritis Ceftriaxone does NOT require renal dose adjustments and should NOT be used in neonates (can cause biliary sludging, kernicterus)
Third (group 2)	Ceftazidime Ceftazidime/Avibactam (Avvcaz, IV)	Ceftazidime is active against Pseudomonas
Ceftolozar	Ceftolozane/Tazobactam (Zerbaxa, IV)	Ceftazidime/Avibactam and Ceftolozane/Tazobactam are used for MDR gram- negative organisms (like Pseudomonas)
Fourth	Cefepime (Maxipime, IV, IM)	Cefepime is active against Pseudomonas
Fifth	Ceftaroline fosamil (Teflaro, IV)	Only beta lactam active against MRSA
Siderophore Cephalosporin	Cefiderocol	Broad spectrum against gram-negative bacteria

CARBAPENEMS

- Broad spectrum, reserved for MDR Gram negative infections
- Active against most Gram positive, Gram negative (including ESBL producers), and anaerobes
- NO activity against atypical pathogens, MRSA, VRE, C. difficle, Stenotrophomonas

Agents	Key Featu	ures	
Doripenem (IV)	NOT used for pneumonia		
Imipenem (IV) Imipenem/Cilastatin (IV) Imipenem/Cilastatin/Relebactam (Recarbio, IV)	 Imipenem/Cilastatin/Relebactam is approved for complicated UTI/Pyelonephritis and intraabdominal infections 	CNS adverse effects	
Meropenem (Merrem, IV)		 Major drug interaction with valproic acid and derivatives 	
Meropenem/Vaborbactam (Vabomere, IV)	Approved for complicated UTI/Pyelonephritis		
Ertapenem (Invanz, IV)	 Ertapenem No coverage for Pseudomonas, Acinetobacter, Enterococcus Commonly used for diabetic foot infections 		

Aztreonam

- Monobactam
- Its monobactam structure makes cross-reactivity with beta-lactam allergy unlikely
- Spectrum
 - Gram negative organisms (including Pseudomonas)
 - NO gram positive activity
 - NO anaerobic activity

Aminoglycosides



Aminoglycosides			
Agents	Pearls	Class Features	
Gentamicin (IV, IM, ophthalmic, topical)	Can by synergistic with beta- lactams or vancomycin for certain gram positive organisms	 Coverage: gram negative (including Pseudomonas – EXCEPT Gentamicin) 	
Tobramycin (IV, IM, ophthalmic, inhaled)	Often used in cystic fibrosis	 ADRs: renal toxicity, ototoxicity 2 Dosing Strategies Traditional: uses lower doses more 	
Amikacin (IV, IM)	Second-line treatment for Mycobacterial infections	 frequently <u>Extended Interval:</u> uses higher doses 	
Plazomicin (IV)	For complicated UTI; use only when there are no alternative options	less frequently leading to less accumulation and less toxicity	

Fluorouinolones



Fluoroquinolones

Broad spectrum \rightarrow Variety of gram negative, gram positive, and atypical pathogens

• Notably, resistance to fluoroquinolones is increasing rapidly

AGENT		PEARLS	CLASS EFFECTS
	Levofloxacin	Antipseudomonal	Black Box Warning:
Respiratory Fluoroquinolones Used for pneumonia, reliable activity against S. pneumoniae	Moxifloxacin	 Does not reach adequate concentrations in the urine so should not be used for UTIs No renal dose adjustments needed 	 Tendon inflammation/rupture Peripheral Neuropathy CNS effects Avoid in patients with myasthenia gravis
	Gemifloxacin	Rarely used	QTc prolongation
	Ciprofloxacin	 Antipseudomonal Cannot put cipro suspension through NG tube CAN crush immediate release tablets – hold tube feeds at least 1 hour before and 2 hours after the dose 	 Hypo- and hyperglycemia Photosensitivity Counseling pearls: Avoid sun exposure Separate from cations (to optimize absorption) Monitor blood glucose

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reliable activ S. pneumon	AVO	D IN ELDE	RLY IF
		POSSIBLE	
	Ciprofloxacin	 through NG tube CAN crush immediate release tablets – hold tube feeds at least 1 hour before and 2 hours after the dose 	 Counseling pearls: Avoid sun exposure Separate from cations (to optimize absorption) Monitor blood glucose

Sulfonamaides



Sulfamethoxazole/Trimethoprim (Bactrim)

Spectrum	 Staphylococci (including MRSA) Some broad gram-negative coverage: Haemophilus Proteus E. coli Klebsiella Enterobacter, Shigella Salmonella Stenotrophomonas 	 Some opportunist infections (OI): Nocardia Pneumocystis Toxoplasmosis NO coverage for: Pseudomonas Enterococci Atypicals Anaerobes
Formulations	PO, IV Single Strength (SS) tablets: 400 mg SMX/80 mg TMP Double Strength (DS) tablets: 800mg SMX/160 mg TMP	
Pearls	 Dosed based on TMP component ADRs: Skin reactions (SJS/TEN), thrombotic thrombocytopenic purpura (TTP), photosensitivity, hyperkalemia, AKI 	

Select Gram Positive agents



Agent	Route	Coverage	Pearls
Vancomycin	PO (C. diff only), IV	 Only gram-positive bacteria Staphylococci (MRSA) Streptococci Enterococci (not VRE) C. difficile (PO route only) 	 Monitor for AKI, infusion reaction Requires therapeutic drug monitoring For severe MRSA infections: AUC/MIC target is 400-600, trough target is 15-20 mcg/mL
Daptomycin	IV	Only gram-positive bacteriaStaphylococci (MRSA)Enterococci	 Deactivated by lung surfactant so not to be used in pneumonias Can elevate CK (monitor at baseline then weekly) Can falsely elevate PT/INR but does not increase bleed risk
Linezolid	PO, IV	Only gram-positive bacteriaStaphylococci (MRSA)Enterococci	ADRs:Serotonin syndromeMyelosuppression

Disease States

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Upper Respiratory Tract Infections

Disease State	Common Pathogens	Treatment
Acute Otitis Media (AOM)	S. Pneumoniae H. Influenzae	 Amoxicillin (80-90 mg/kg/day in 2 divided doses) Amoxicillin/Clavulanate (90 mg/kg/day in 2 divided doses) Ceftriaxone (50 mg/kg IM or IV for 1 or 3 days)
Influenza	H. Influenzae	Oseltamivir for 5 days
Pharyngitis	S. pyogenes	 Penicillin, Amoxicillin, or 1st or 2nd generation cephalosporin for 10 days Azithromycin x5 days
Sinusitis	Respiratory Viruses S. Pneumoniae H. Influenzae M. catarrhalis	 No antibiotics needed for respiratory viral infections Amoxicillin/Clavulanate – treatment duration varies



Lower Respiratory Tract Infections

DISEASE STATE	COMMON PATHOGENS	TREATMENT
Community Acquired Pneumonia (CAP)	S. Pneumoniae H. Influenzae M. Catarrhalis	 Empiric coverage should include: S. pneumoniae and atypical pathogens Outpatient No co-morbidities: Amoxicillin (high dose, 1 g 3x daily) OR doxycycline OR azithromycin Co-morbidities: Beta-lactam (Amoxicillin/Clavulanate or cephalosporin) PLUS azithromycin OR monotherapy with respiratory fluoroquinolone Inpatient Beta Lactam (ceftriaxone) + macrolide (azithromycin) OR doxycycline OR doxycycline OR Monotherapy with respiratory fluoroquinolone (levofloxacin, moxifloxacin – NOT ciprofloxacin) If risk factors for MRSA: add vancomycin or linezolid
		antipseudomonal agent

Duration: 5 days



Lower Respiratory Tract Infections

DISEASE STATE	COMMON PATHOGENS	TREATMENT
 S. Pneumoniae H. Influenzae M. Catarrhalis MRSA Gram negative rods P. aeruginosa Acinetobacter spp. Enterobacter spp. E. coli Klebsiella spp 	S. Pneumoniae H. Influenzae M. Catarrhalis MRSA Gram negative rods • P. aeruginosa	Choose 1 antibiotic to cover pseudomonas and MSSA if low risk for MRSA or MDR pathogens <i>Examples:</i> Cefepime or piperacillin/tazobactam
		 Choose 2 antibiotics, one for MRSA and one for Pseudomonas if risk for MRSA but low risk for MDR pathogens <i>Examples:</i> Cefepime + Vancomycin OR linezolid
	 Choose 3 antibiotics, one for MRSA and two for pseudomonas if risk for both MRSA and MDR pathogens <i>Examples:</i> Piperacillin/tazobactam + ciprofloxacin + vancomycin Cefepime + gentamicin + linezolid 	

Duration: 7 days (may vary depending on clinical response)

Skin and Soft Tissue Infections (SSTI) - Cellulitis

CAUSES

Previous trauma to the area Underlying lesions PATHOGENS

S. pyogenes S. areus Gram negative bacilli Anaerobes

LOCATION

Anywhere, common on the limbs and face

Skin and Soft Tissue Infections (SSTI) - Cellulitis

Non-purulent			
Mild	PO RouteCephalexinPenicillin VKAmoxicillinDicloxacillin		
Moderate	IV Route Cefazolin	Duration 5-10 days	
Severe	Debridement with cultures IV Route Vancomycin + piperacillin/tazobactam		

Penicillin allergy: can use clindamycin

Skin and Soft Tissue Infections (SSTI) - Cellulitis

Purulent			
Mild	Incision and drainage (I & D) only		
Moderate	Incision and drainage (I & D) and cultures PO Route Sulfamethoxazole/trimethoprim (Bactrim)* Doxycycline* Clindamycin* Dicloxacillin Cephalexin	Duration	
Severe	Incision and drainage (I & D) and cultures IV Route Vancomycin* Daptomycin* Linezolid* Dalbavancin Oritavancin	Variable	

*If community acquired MRSA suspected

Urinary Tract Infection (UTI)



DIAGNOSIS	COMMON PATHOGENS	TREATMENT
Asymptomatic Bacteriuria	Any of the below	NONE! Exceptions: pregnancy, endourological procedures
Acute Uncomplicated Cystitis	Proteus, E. coli (majority), S. saprophyticus, Enterococci	Nitrofurantoin 100 mg by mouth twice daily x5 days OR SMX/TMP 1 DS tablet twice daily x3 days OR Fosfomycin 3g x1 dose
Acute uncomplicated pyelonephritis	E. coli, Enterococci, Proteus, Klebsiella, Pseudomonas	Ceftriaxone 1-2g IV once daily OR
Complicated UTI	E. coli, Klebsiella, Enterobacter, Serretia, Pseudomonas, Enterococci, Staphylococci	Ciprofloxacin (500mg by mouth twice daily) OR Levofloxacin (750mg by mouth twice daily) OR Alternative beta-lactam
Candida in the urine	Treatment is NOT recommended Exceptions: symptomatic, high-risk	patients include neutropenic patients, very low-birth-

weight infants (<1500 g), and patients who will undergo urologic manipulation

Sexually Transmitted Infections (STI)

INFECTION	DRUG OF CHOICE	ALTERNATIVE
Syphilis (primary, secondary, early latent)	Penicillin G benzathine (2.4 million units IM x1)	Doxycycline 100mg by mouth twice daily for 14 days
Syphilis (Late latent)	Penicillin G benzathine (2.4 million units IM weekly x3 weeks)	Doxycycline 100mg by mouth twice daily for 28 days
Neurosyphilis	Penicillin G aqueous crystalline (1-24 million units daily, divided into 6 doses or by continuous infusion for x10-14 days)	Penicillin G procaine
Gonorrhea	Ceftriaxone (250mg IM x1 dose) + Azithromycin (1g PO x1 dose)	Ceftriaxone (250mg IM x1) + doxycycline (100mg by mouth twice daily x7 day) Monotherapy is NOT recommended due to resistance
Chlamydia	Azithromycin (1g by mouth x1 dose)	Doxycycline (100mg by mouth twice daily x7 days)
Bacterial Vaginosis	Metronidazole (500mg by mouth twice daily x7 days) OR metronidazole 0.75% gel (1 applicator daily or twice daily x5 day)	Clindamycin 300mg by mouth twice daily x7 days
Trichomoniasis	Metronidazole (2g by mouth x1 dose)	Metronidazole 500mg by mouth twice daily x7 days

Meningitis



- Common pathogens: Streptococcus pneumoniae, Neisseria meningitidis, and Haemophilus influenzae
 - Add coverage for Listeria monocytogenes in neonates, age >50 years and immunocompromised patients

MENINGITIS: EMPIRIC TREATMENT			
Age <1 month (neonates)	Age 1 month to 50 years	Age >50 years or immunocompromised	
Ampicillin +	Vancomycin +	Ampicillin (for Listeria coverage) +	
Cefotaxime OR gentamicin	Ceftriaxone OR cefotaxime	Vancomycin + Ceftriaxone OR cefotaxime	

Do not used Ceftriaxone in neonates

• Can cause biliary sludging and kernicterus in neonates

Antifungals

Fungal Classifications

YEASTS	MOLDS	DIMORPHIC FUNGI	
 Candida spp C. albicans C. glabrata C. tropicalis C. parapsilosis C. krusei 	Aspergillus spp Zygomycetes (Mucor spp)	Histoplasma capsulatum Blastomyces dermatitidis Coccidioides immitis	
Cryptococcus neoformans			

Antifungals

CLASS	AGENTS	CLASS EFFECTS
Amphotericin	Amphotericin B deoxycholate (conventional) Amphotericin B Lipid Complex Liposomal Amphotericin B	ADRs: infusion reactions, hypokalemia, hypomagnesia, nephrotoxicity Monitoring: renal function, electrolytes, LFTs
Azole Antifungals	Fluconazole (PO, IV) Itraconazole (PO) Ketoconazole (PO, topical) Voriconazole (PO, IV) Posaconazole (PO, IV) Isavuconazole (PO, IV)	 All can cause increased LFTs All have a risk for QTc prolongation (except Isavuconazole) All azoles are moderate-strong CYP3A4 inhibitors Fluconazole is the only azole that requires renal dose adjustments Itraconazole can cause heart failure Voriconazole can cause visual changes and phototoxicity
Echinocandins	Caspofungin Micafungin Anidulafungin	ADRs: increased LFTs, severe skin reactions with caspofungin)
Flucytosine		ADRs: dose related myelosuppression, AKI

Fungal Infection – Empiric Treatment

PATHOGEN	PREFERRED REGIMEN	ALTERNATIVE REGIMEN
Candida albicans Oropharyngeal infection (thrush)	Fluconazole	Nystatin
Candida albicans Esophageal infection	Fluconazole	Echinocandin
Candida krusei and glabrata Bloodstream infection	Echinocandin	Amphotericin B
Aspergillus (invasive)	Voriconazole, Posaconazole	Amphotericin B, isavuconazole
Cryptococcus neoformans meningitis	Amphotericin B + flucytosine	



ANTIVIRAL	COMMON INDICATION	PEARLS	
Acyclovir		WarningsNeeds renal dose adjustment	
Valacyclovir	Herpes Simplex Virus (HSV) Varicella Zoster Virus (VZV)	 Acyclovir requires adequate hydration to reduce risk of renal tubular damage Side Effects Increased LFTs Neutropenia Increased Scr/BUN with crystal neuropathy Monitoring Renal function, LFTs, CBC 	
Ganciclovir		Boxed WarningsMyelosuppression, carcinogenic, teratogenic	
Valganciclovir	Cytomegalovirus (CMV)	 Side Effects Thrombocytopenia Neutropenia Leukopenia Monitoring CBC with differential Scr Retinal exam (valganciclovir) 	

Opportunist Infections (OI)

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OI - Primary Prophylaxis			
Infection	Indication in HIV	Regimen	Discontinuation
Pneumocystis pneumonia (PCP or PJP)	CD4+ count <200 cells/mm³	Preferred: SMX/TMP DS or SS by mouth daily Alternatives: SMX/TMP DS by mouth 3x weekly OR Dapsone OR Atovaquone OR Inhaled pentamidine	CD4+ count ≥200 cells/mm ³ for ≥ 3 months on ART
Toxoplasma gondii encephalitis (commonly called Toxo)	Toxoplasma IgG positive with CD4+ count <100 cells/mm ³	Preferred: SMX/TMP DS by mouth daily Alternatives: SMX/TMP DS by mouth 3x weekly or 1 SS by mouth daily OR Dapsone + pyrimethamine + leucovorin OR Atovaquone	CD4+ count >200 cells/mm ³ for ≥ 3 months on ART
Mycobacterium avium complex (MAC) infection	Not recommended if ART is started immediately Initiate if NOT taking ART and CD4+ count <50 cells/mm ³ Must rule out active disseminated MAC disease	Preferred: Azithromycin 1,200 mg by mouth weekly Alternatives: Azithromycin 600 mg by mouth twice weekly OR Clarithromycin 500 mg by mouth twice daily	Taking fully suppressive ART

OI - Treatment				
Infection	Preferred Regimen	Alternative	Secondary Prophylaxis	
Candidiasis (thrush)	Fluconazole	Itraconazole, posaconazole	None	
Cryptococcal meningitis	Induction therapy: Amphotericin B (deoxycholate or liposomal) + flucytosine	Fluconazole +/- flucytosine	Fluconazole (low dose)	
Cytomegalovirus (CMV)	Valgancyclovir OR Ganciclovir	If toxicities to ganciclovir or resistant strains: letermovir, foscarnet, cidofovir	N/A	
Mycobacterium avium complex (MAC) infection	Clarithromycin OR azithromycin + ethambutol	Add a 3 rd or 4 th agent using rifabutin, amikacin, or streptomycin, moxifloxacin, or levofloxacin	Same as treatment regimen	
Pneumocystis pneumonia (PCP or PJP)	SMX/TMP +/- prednisone or methylprednisolone Duration: 21 days	Atovaquone OR Clindamycin + primaquine OR Pentamidine IV OR Dapsone + trimethoprim	Same as primary prophylaxis	
Toxoplasma gondii encephalitis (commonly called Toxo)	Pyrimethamine + leucovorin + sulfadiazine	SMX/TMP	Same as treatment; reduced doses	

Antimicrobial Stewardship

(Not so) Shocking Facts



Antimicrobial Stewardship

- Antimicrobial stewardship: efforts to coordinate interventions designed to optimize antimicrobial use and ensure the best clinical outcomes while minimizing unintended consequences
 - Combat rising antimicrobial resistance
 - Reduce healthcare costs
 - Improving patient outcomes

New National Estimate*

Each year, antibiotic-resistant bacteria and fungi cause at least an estimated: *Clostridioides difficile*** is related to antibiotic use and antibiotic resistance:









AND INCREASES IN INFECTIONS CAUSED BY:

▲ 315%

Erythromycin-resistant invasive group A strep Drug-resistant *Neisseria gono<u>rrhoeae</u>*

124%

ESBL-producing Enterobacteriaceae

4 50%

Why?

Inconsistency in adoption of CDC recommended strategies to minimize infection spread

Challenges in preventing spread of germs in non-hospital settings

Emerging threats from outside of the United States

Spread of resistant threats in the food supply, animals

Limited outpatient stewardship efforts

The Threats

Urgent Threats

These germs are public health threats that require urgent and aggressive action:







CARBAPENEM-RESISTANT ACINETOBACTER

CANDIDA AURIS

CLOSTRIDIOIDES DIFFICILE





CARBAPENEM-RESISTANT ENTEROBACTERIACEAE

DRUG-RESISTANT NEISSERIA GONORRHOEAE

The Threats

Serious Threats

- Drug-resistant Campylobacter
- Drug-resistant Candida
- Extended-spectrum beta-lactamase (ESBL)-producing Enterobacteriaceae
- Vancomycin-resistant Enterococci (VRE)
- Multidrug-resistant Pseudomonas aeruginosa (P. aeruginosa)
- Drug-resistant nontyphoidal Salmonella
- Drug-resistant Salmonella serotype Typhi
- Drug-resistant Shigella
- Methicillin-resistant Staphylococcus aureus (MRSA)
- Drug-resistant Streptococcus pneumoniae (S. pneumoniae)
- Drug-resistant Tuberculosis (TB)

Concerning Threats

- Erythromycin-resistant group A Streptococcus
- Clindamycin-resistant group B Streptococcus

Watch List

- Azole-resistant Aspergillus fumigatus (A. fumigatus)
- Drug-resistant Mycoplasma genitalium (M. genitalium)
- Drug-resistant Bordetella pertussis (B. pertussis)

Available data show an alarming increase in resistant infections starting during hospitalization, growing at least 15% from 2019 to 2020.

- Carbapenem-resistant Acinetobacter (†78%)
- Antifungal-resistant Candida auris (+60%)*
- Carbapenem-resistant Enterobacterales (+35%)
- Antifungal-resistant Candida (†26%)

- ESBL-producing Enterobacterales (+32%)
- Vancomycin-resistant Enterococcus (+14%)
- Multidrug-resistant P. aeruginosa (†32%)
- Methicillin-resistant Staphylococcus aureus (+13%)

The COVID-19 pandemic did not help



	Hospital Core Elements	Priorities for Hospital Core Element Implementation
Hospita	I Leadership Commitment	
	Dedicate necessary human, financial, and information technology resources.	Antibiotic stewardship physician and/or pharmacist leader(s) have antibiotic stewardship responsibilities in their contract, job description, or performance review.
Accoun	tability	
	Appoint a leader or co-leaders, such as a physician and pharmacist, responsible for program management and outcomes.	Antibiotic stewardship program is co-led by a physiciar and pharmacist.*
Pharma	cy/Stewardship Expertise	
	Appoint a pharmacist, ideally as the co-leader of the stewardship program, to help lead implementation efforts to improve antibiotic use.	Antibiotic stewardship physician and/or pharmacist leader(s) have completed infectious diseases specialty training, a certificate program, or other training on antibiotic stewardship.
Action		
	Implement interventions, such as prospective audit and feedback or preauthorization, to improve antibiotic use.	Antibiotic stewardship program has facility-specific treatment recommendations for common clinical condition(s) and performs prospective audit/feedback or preauthorization.
Trackin	g	
	Monitor antibiotic prescribing, impact of interventions, and other important outcomes, like <i>C. difficile</i> infections and resistance patterns.	Hospital submits antibiotic use data to the NHSN Antimicrobial Use Option.
Reporti	ng	
*	Regularly report information on antibiotic use and resistance to prescribers, pharmacists, nurses, and hospital leadership.	Antibiotic use reports are provided at least annually to target feedback to prescribers. In addition, the antibiot stewardship program monitors adherence to facility- specific treatment recommendations for at least one common clinical condition.
Educati	on	
	Educate prescribers, pharmacists, nurses, and patients about adverse reactions from antibiotics, antibiotic resistance, and optimal prescribing.	No implementation priority identified.

Updated Joint Commission requirements for accreditation

New accreditation requirements → Implemented January 2023

> 12 **new and revised** requirements to decrease unnecessary antimicrobial utilization in the hospital setting

• Including critical access hospitals



Stewardship Tactics

Prospective Audit and Feedback (PAF)

Preauthorization protocols

Institutional Guidelines, Protocols, Ordersets

Prospective Audit and Feedback (PAF)

Process most commonly involves an ID physician or a clinical pharmacist with ID training reviewing active antimicrobial prescriptions to identify opportunities to enhance the safety and/or effectiveness of therapy

Widely recognized as one of the most effective antibiotic stewardship practices

Viewed as a core component of many hospital ASPs

Outcomes of Successful PAF

8-year, pharmacist-led PAF program:

- Decreased rates of MRSA
- Increased Pseudomonas aeruginosa susceptibility to carbapenems
- Decreased antimicrobial therapy duration
- Decreased number of days on IV therapy
- Increased cost savings

Preauthorization



Enhance the effectiveness of both PAF and preauthorization

Can optimize antibiotic selection and duration for common indications

Can include guidance on diagnostic approaches (especially empirically)

Based on national guidelines, optimized based on local antibiograms

Institutional Guidelines

Members of the Stewardship Team







Infectious Diseases trained Physicians, Pharmacists, APPs Infection Prevention and Microbiology colleagues

Nurses



We're all stewards!







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