#### Table 8. Adult parenteral antimicrobial dosage guidelines

Antibiotic	Usual Dosages <sup>a</sup>
ANTIBACTERIAL AGENTS	
Penicillins	
ampicillin	1-2 g q4-6h
cloxacillin	2 g q4-6h
penicillin G	2-4 million units q4-6h
piperacillin-tazobactam	3.375 g q6h
meropenem	500 mg q6h
Cephalosporins	
cefazolin	1-2 g q8h
cefoxitin	1-2 g q6-8h
cefuroxime	0.75-1.5 g q8h
ceftriaxone	1-2 g q24h
ceftazidime	1-2 g q8h
Fluoroquinolones	
ciprofloxacin	400 mg q12h
levofloxacin	500-750 mg q24h
moxifloxacin	400 mg q24h
Macrolides	
azithromycin	500 mg q24h
Aminoglycosides	
gentamicin or tobramycin	80 mg q8h
Others	
clindamycin	600 mg q8h
cotrimoxazole (TMP-SMX)	10-20 mg/kg/day trimethoprim in divided doses q6-8h
metronidazole	500 mg q8h
vancomycin	1 g q12h or 15 mg/kg q12h
ANTIFUNGAL AGENTS	
amphotericin B	0.5-1 mg/kg q24h
fluconazole	100-400 mg q24h
caspofungin	70 mg load then 50 mg q24h
ANTIVIRAL AGENTS	·
acyclovir	5-10 mg/kg/dose q8h
ganciclovir	5 mg/kg/dose q12h

\* Based on normal renal function in a 70 kg patient.

Table 9. Parenteral to oral conversion suggestions
--

Oral Therapy Options <sup>a</sup>
AGENTS
amoxicillin
cloxacillin or cephalexin
penicillin V
amoxicillin-clavulanate or cotrimoxazole (TMP-SMX) +/- metronidazole or ciprofloxacin +/- metronidazole
·
cephalexin or cloxacillin
cephalexin + metronidazole or cotrimoxazole + metronidazole or amoxicillin-clavulanate
cotrimoxazole or amoxicillin-clavulanate or azithromycin/clarithromycin
amoxicillin-clavulanate or cephalexin or ciprofloxacin/levofloxacin/moxifloxacin
ciprofloxacin
ciprofloxacin
levofloxacin
moxifloxacin
azithromycin
cloxacillin +/- metronidazole or cephalexin +/- metronidazole or clindamycin
ENTS
fluconazole
ITS
acyclovir or valacyclovir

<sup>a</sup> Patients should be clinically stable, demonstrate clinical improvement, and be able to tolerate oral feeding and medications. Selection of oral therapy should be based on cultures and sensitivities. In absence of useful cultures, oral therapy may be selected based on potential pathogens, community- versus hospital-acquired infection, pharmacokinetics, spectrum of activity, and cost of each oral agent. Oral agents listed above represent those currently on the WRHA Formulary and does not represent all commercially available oral agents.

#### Table 10. Adult dosing recommendations in renal impairment<sup>a</sup>

Drug	g Creatinine Clearance (CrCl) in mL/min <sup>b</sup> (suggested dosage adjustment based on normal dose)						
Penicillins							
ampicillin	> 30 (q6h)	10-30 (q6-12h)	< 10 (q12h)				
cloxacillin		NO CHANGE	NECESSARY				
penicillin	> 50 (q4-6h)	10-50 (q6-8h)	< 10 (20-50% of usual dose) <sup>a</sup>				
piperacillin- tazobactam	> 40 (q6h)	20-40 (q8h)	< 20 (q12h)				
Carbapenems							
meropenem	> 50 (q6h)	30-49 (q8h)	10-29 (q12h)	< 10 (q24h)			
Cephalosporins							
cefazolin	> 50 (q8h)	10-50 (q12h)	< 10 (q24h)				
cefoxitin	> 30 (q6-8h)	10-30 (q12-24h)	< 10 (q24h)				
cefuroxime	> 20 (q8h)	10-20 (q12h)	< 10 (q24h)				
ceftriaxone		NO CHANGE	NECESSARY				
ceftazidime	> 50 (q8h)	30-50 (q12h)	10-30 (q24h)	< 10 (50% q24-48h)			
Aminoglycosides							
gentamicin/ tobramycin/ amikacin Fluoroquinolones	Contac	ct the Pharmacist at yo	ur facility for dosing assis	stance			
ciprofloxacin	> 30	< 30					
оргопохаонт	(q12h)	(q24h)					
levofloxacin (e.g. CAP)	> 50 (q24h)	20-49 (500 mg load, then 50% q24h)	10-19 (500 mg load, then 50% q48h)				
moxifloxacin			NECESSARY				
Macrolides	<u> </u>						
azithromycin	1	NO CHANGE	NECESSARY				
Antifungal Agents							
fluconazole	. 50						
	> 50 (q24h)	20-50 (50% q24h)	< 20 (25% of usual dose q24h)				
caspofungin		(50% q24h)	(25% of usual				
caspofungin Antiviral Agents		(50% q24h)	(25% of usual dose q24h)				
		(50% q24h) NO CHANGE 25-50 (q12h)	(25% of usual dose q24h)	< 10 (50% q24h)			
Antiviral Agents	(q24h)	(50% q24h) NO CHANGE 25-50	(25% of usual dose q24h) NECESSARY 10-25	(50% q24h) < 10			
Antiviral Agents acyclovir ganciclovir	(q24h) > 50 (q8h) 50-69	(50% q24h) NO CHANGE 25-50 (q12h) 25-49 2.5 mg/kg q24h	(25% of usual dose q24h) NECESSARY 10-25 (q24h) 10-25 1.25 mg/kg q24h	(50% q24h) < 10			
Antiviral Agents acyclovir ganciclovir (induction doses) Miscellaneous clindamycin	(q24h) > 50 (q8h) 50-69	(50% q24h) NO CHANGE 25-50 (q12h) 2.5-49 2.5 mg/kg q24h NO CHANGE	(25% of usual dose q24h) NECESSARY 10-25 (q24h) 1.0-25 1.25 mg/kg q24h NECESSARY	(50% q24h) < 10			
Antiviral Agents acyclovir ganciclovir (induction doses) Miscellaneous clindamycin metronidazole	(q24h) > 50 (q8h) 50-69 2.5 mg/kg q12h	(50% q24h) NO CHANGE 25-50 (q12h) 25-49 2.5 mg/kg q24h NO CHANGE NO CHANGE	(25% of usual dose q24h) NECESSARY 10-25 (q24h) 10-25 1.25 mg/kg q24h NECESSARY NECESSARY	(50% q24h) < 10 1.25 mg/kg 3x/wk			
Antiviral Agents acyclovir ganciclovir (induction doses) Miscellaneous clindamycin	(q24h) > 50 (q8h) 50-69 2.5 mg/kg q12h > 25 (q6-8h)	(50% q24h) NO CHANGE 25-50 (q12h) 25-49 2.5 mg/kg q24h NO CHANGE 15-25 (50% q6-8h)	(25% of usual dose q24h) NECESSARY 10-25 (q24h) 1.0-25 1.25 mg/kg q24h NECESSARY	(50% q24h) < 10 1.25 mg/kg 3x/wk 15 y not recommended)			

<sup>a</sup> Suggested dosages-for individualized dosage modifications or more information contact the Pharmacy Department at your facility.

<sup>b</sup> To estimate creatinine clearance (CL<sub>CR</sub>) (mL/min) use the following calculation normalized for a 72 kilogram person.

```
CL<sub>CR</sub> male = (140-age) x 88.4
                                                   CL<sub>CR</sub> female = 0.85 x CL<sub>CR</sub> male
```

S<sub>CR</sub> (µmoles/L) <sup>c</sup> Monitor serum concentrations. Table 11. Antimicrobial Restrictions at Health Sciences Centre (revised 2017)

Antimicrobial (alphabetical order)	Status <sup>a</sup>	Exception Criteria	Criteria For Use⁵
Acyclovir IV	Consultation		
Amphotericin B Lipid Complex	Consultation		Yes
Caspofungin	Consultation		Yes
Ceftazidime	Consultation	Hemodialysis (HSC or SBH) or Peritoneal (SBH)	
Ertapenem	Consultation (CIVP <sup>c</sup> only)		
Fluconazole IV	Consultation		
Ganciclovir IV	Consultation	under protocol	
Linezolid	Consultation		Yes
Meropenem	Consultation		
Piperacillin- Tazobactam	Consultation		
Daptomycin	Consultation		Yes
Voriconazole	Consultation		Yes

<sup>a</sup> Consultation or verbal approval should be obtained from the Infectious Diseases consult service.

<sup>b</sup> Reviewed by the WRHA Antimicrobial Pharmacotherapy Subcommittee November 2014. Criteria for use may be obtained from a WRHA Pharmacy Department.

<sup>c</sup> CIVP = Community IV Program



# Health Sciences Centre Antibiogram for 2022

## (Adult patients only - Based on data from 2021)

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#### DISCLAIMERS

This guide is provided as an educational resource for physicians and other healthcare professionals caring for adult patients at the Health Sciences Centre. The authors of the guide have made every effort to ensure that the information contained in it was accurate at the time of publication. Users of the guide are encouraged to consult other references to confirm the information presented in it. The authors are not responsible for errors, omissions, inaccuracies, or the continued completeness of the information contained in the guide. The information in the guide should not be used or relied upon to replace the skill and professional judgment required to determine appropriate patient care and treatment. Also, the quide is not intended to replace or to be used as a substitute for the complete prescribing information prepared by each pharmaceutical manufacturer for their anti-infective agents. Because of possible changes in anti-infective indications, changes in dosage information, differences in patients' responses to therapy, newly described toxicities, drug-drug interactions, and other items of importance, reference to complete prescribing information is recommended before any of the anti-infective agents described in the guide are used.

### HOW TO USE THE ANTIBIOGRAM PORTION OF THE GUIDE (Tables 1-6)

- The information presented in the antibiogram is intended only to guide initial empiric anti-infective agent therapy at the Health Sciences Centre.
- Initial broad-spectrum empiric therapy should be focused to the most appropriate narrow-spectrum agent(s) based on the laboratory identification of pathogen(s) and known susceptibility patterns/results, if the situation permits.
- Consideration should be given to equally efficacious but less expensive anti-infective agents for empiric therapy or when streamlining of therapy is desired, if the situation permits.

#### SUGGESTED CRITERIA FOR IV TO ORAL ANTIBIOTIC CONVERSION IN ADULTS

- Clinical improvement of infectious signs and symptoms (e.g., temperature defervescence, decreased white blood cell count).
- Patient is clinically stable (excludes patients in the intensive care unit, patients with febrile neutropenia, or patients with life threatening infections).
- · Patient can tolerate oral feeding and medications (bowel sounds, no diarrhea/nausea/ vomitina).
- For rapid step-down, choose agents with high bioavailability (e.g., clindamycin, cotrimoxazole (TMP-SMX), fluoroguinolones).
- If anti-infective agent susceptibilities are known, anti-infective therapy should be tailored based on available data.

#### Table 1. In vitro activity of selected anti-infective agents tested against Gram-negative bacillia

							Percer	nt Susc	eptible						
Organism (number tested): January through December 2021 = Not tested, not routinely reported, or not recommended	Ampicillin	Amoxicillin- Clavulanate	Piperacillin- Tazobactam	Cefazolin	Cephalexin <sup>b</sup>	Cefuroxime	Ceftriaxone	Ceftazidime	Ertapenem	Meropenem	Gentamicin	Tobramycin	Ciprofloxacin	Trimethoprim- Sulfamethoxazole	Nitrofurantoin°
Acinetobacter baumannii complex (35)			89				57	77		n.d.	100	97	94	100	
Citrobacter spp. (65)			75				68	66	98	98	85	92	72	82	94
Enterobacter cloacae complex (218)			80				70	72	94	97	94	93	87	81	34
Escherichia coli (348) systemic	41	79	93	55			80	85	100	100	91	90	61	67	
Escherichia coli (828) urine	45	81	96	56	n.d.		82	89	100	100	92	91	59	64	96
Haemophilus influenzae (75) <sup>d</sup>	65	n.d.				97								64	
Klebsiella aerogenes (35)			71				71	74	100	100	100	100	97	100	3
Klebsiella pneumoniae (296)		87	92	79	n.d.		88	87	99	99	93	92	82	83	25
Klebsiella/Raoultella spp. (125) <sup>e</sup>		89	92	24			92	99	100	100	98	98	90	90	79
Morganella morganii (51)			100				94	84	100	100	92	96	75	82	
Proteus mirabilis (85)	81	96	100	n.d.	n.d.		100	100	100	100	94	96	92	85	
Pseudomonas aeruginosa (419)			90					87		89	96	99	82		
Serratia marcescens (80)			98				93	99	99	99	98	96	81	98	
Stenotrophomonas maltophilia (118)														98	

a Isolates tested and reported are from all sources combined, Jan 1 to Dec 31, 2021 with the exception of Escherichia coli (subdivided into systemic isolates and urine isolates); data compiled according to the recommendations of the Clinical and Laboratory Standards Institute (CLSI) in their document M39-A4 (2014).

- Cephalexin is only indicated for the treatment of uncomplicated lower urinary tract infections.
- Nitrofurantoin is only indicated for acute cystitis.
- <sup>d</sup> H. influenzae data obtained from isolates tested at Health Sciences Centre, Jan 1 to Dec 31, 2017, Sixty-one isolates were tested for Cefuroxime and 69 isolates were tested for Trimethoprim-Sulfamethoxazole. Data from adult and pediatric patients.
- The current laboratory identification system is unable to differentiate Klebsiella oxytoca from Raoultella spp.

n.d. = no data - absence of data for certain drug-organism combinations reflects limitations of the testing method currently used by Shared Health Clinical Microbiology Jaboratories

#### Table 2. In vitro activity of selected anti-infective agents tested against Gram-positive coccia

						Pe	rcent S	uscepti	ble					
Organism (number tested): January through December 2021 Not tested, not routinely reported, or not recommended	Penicillin	Ampicillin	Oxacillin <sup>b</sup>	Vancomycin	Daptomycin	High-Level Gentamicin <sup>o</sup>	High-Level Streptomycin <sup>c</sup>	Erythromycin <sup>d</sup>	Clindamycin	Trimethoprim- Sulfamethoxazole	Rifampin <sup>®</sup>	Linezolid	Tetracycline	Nitrofurantoin <sup>6</sup>
Enterococcus faecalis (235)		100		98	100	85	91					98		99
Enterococcus faecium (119)		11		30	n.d.	85	89					87		n.d.
Staphylococcus aureus (1790)			64	100	100			53	73	99	99	100	96	96
Staphylococcus epidermidis (304)			31	100	100			33	56	65	99	100	87	99
Staphylococcus haemolyticus (39)			13	100	100			13	28	33	59	100	87	100
Staphylococcus hominis (39)			54	100	100			26	54	72	100	100	87	98
Staphylococcus lugdunensis (166)			89	100	100			83	85	99	100	100	93	99
Coagulase-negative staphylococci (35)			70	100	100			40	62	80	100	100	97	100
Streptococcus pyogenes (n.a.) <sup>9</sup> (Group A Streptococcus)	100													
Streptococcus agalactiae (162) <sup>h</sup> (Group B Streptococcus)	100			100					60					

a Isolates tested and reported are from all sources (surveillance isolates excluded) with the exception of Enterococcus faecalis and Enterococcus faecium (systemic isolates only), Jan to Dec, 2021; data compiled according to the recommendations of the Clinical and Laboratory Standards Institute (CLSI) in their document M39-A4 (2014).

- Oxacillin accurately predicts the activity of all semi-synthetic penicillins, including cloxacillin, beta-lactam/beta-lactamase inhibitor combinations, cephalosporins, and carbapenems for Staphylococcus aureus and coagulase-negative staphylococci.
- Susceptibility to high-level gentamicin or high-level streptomycin indicates that these agents can be used in combination with a cell wall active agent (e.g., ampicillin or vancomycin) for synergy. Gentamicin and streptomycin should never be used alone as treatment for Enterococcus faecalis or Enterococcus faecium
- d Erythromycin activity predicts the activity of azithromycin and clarithromycin for staphylococci and streptococci.
- Rifampin should NOT be used alone as treatment for infection.
- Nitrofurantoin is indicated for acute cystitis only.
- In.a. = not applicable Susceptibility testing of Streptococcus pyogenes is not routinely performed as 100% are susceptible to penicillin. If treating infection in a penicillin allergic patient, contact the lab for testing of second line agents.
- Streptococcus agalactiae isolates were obtained from vaginal/rectal swabs submitted for Group B Streptococcus detection to the Health Sciences Centre, St. Boniface Hospital, and Westman Laboratory in 2012.

n.d. = no data - absence of data for certain drug-organism combinations reflects limitations of the testing method currently used by Shared Health Clinical Microbiology laboratories.

#### Table 3. In vitro activity of selected anti-infective agents tested against Streptococcus nneumoniae

Infection Type (number tested)		Percent Susceptible						
= Not tested, not routinely reported, or not recommended	Penicillin (oral)	Penicillin (intravenous)	Ceftriaxone	Vancomycin	Levofloxacin	Clarithromycin	Doxycycline	Trimethoprim- Sulfamethoxazole
Systemic Isolates (Blood + CSF) <sup>b</sup>								
Meningitis (180)		83	96	100				86
Non-Meningitis infection (180)	83	98	99	100	99	68	92	86
Respiratory Isolates <sup>c</sup>								
Non-Meningitis infection (40)	78	95	98	100	100	68	83	80

a For Streptococcus pneumoniae, different susceptibility breakpoints for penicillin and ceftriaxone exist depending on whether meningitis or a non-meningitis infection is being treated ICLSI, M100, 31ª edition). For penicillin, when treating a non-meningitis infection different breakpoints exist for oral and intravenous dosing. For non-meningitis infections, susceptibility to oral penicillin predicts susceptibility to amoxicillin. Oral agents are not appropriate for the treatment of bacterial meningitis.

<sup>b</sup> Systemic isolates were obtained from patients across Manitoba as part of the SAVE Study between January and December. 2017 CSE = cerebrospinal fluid

Respiratory isolates were obtained from patients at the Health Sciences Centre (HSC) and St. Boniface Hospital (SBH) between January and December, 2018.

#### Table 4. In vitro activity of selected anti-infective agents tested against Methicillin-Susceptible and Methicillin-Resistant Staphylococcus aureus isolates<sup>a</sup>

Organism (number tested)				Percent S	usceptible			
= Not tested, not routinely reported, or not recommended	Oxacillin <sup>b</sup>	Vancomycin	Trimethoprim- Sulfamethoxazole	Erythromycin	Clindamycin	Tetracycline	Linezolid	Daptomycin
Methicillin-Susceptible Staphylococcus aureus (1121)	100		99	69	72	96		
Methicillin-Resistant Staphylococcus aureus (657)	0	100	99	23	75	96	100	100

a Isolates tested and reported are from all sources (surveillance isolates excluded). Jan to Dec. 2021; data compiled according to the recommendations of the Clinical and Laboratory Standards Institute (CLSI) in their document M39-A4 (2014).

<sup>b</sup> Oxacillin accurately predicts the activity of all semi-synthetic penicillins, including cloxacillin, beta-lactam/beta-lactamase inhibitor combinations, cephalosporins, and carbapenems for Staphylococcus aureus,

#### Table 5. In vitro activity of selected anti-infective agents tested against anaerobic isolates collected from hospitals in Winnipega

			Percent S	usceptible		
Organism (number tested) = Not tested, not routinely reported, or not recommended	Penicillin	Amoxicillin- Clavulanate	Piperacillin- Tazobactam	Clindamycin	Meropenem	Metronidazole
Bacteroides fragilis (108)		93	n.d.	44	93	100
Bacteroides thetaiotaomicron (37)		94	n.d.	14	97	100
Prevotella bivia (54)	7	100	n.d.	32	100	96
Prevotella disiens (34)	32	97	n.d.	18	100	100

<sup>a</sup> Isolates were obtained from WRHA hospitals between Jan 2019 and Dec 2020; data compiled according to the recommendations of the Clinical and Laboratory Standards Institute (CLSI) in their document M39-A4 (2014).

n.d. = no data – absence of data for certain drug-organism combinations reflects limitations of the testing method currently used by Shared Health Clinical Microbiology laboratories.

#### Table 6. In vitro activity of selected anti-fungal agents tested against Candida species collected from hospitals in Winnipega

ANTIE Penic amo amo clox peni Cepha cept Macro azith clari Fluoro cipro levof moxi Other clind cotri doxy nitro metr ANTIF flucon itracor

ANTI acyclo

valacy

	Pe	ercent Susceptib	le
Organism (number tested)	Fluconazole <sup>b</sup>	Voriconazole	Micafungin
Candida albicans (33)	100	100	100
Candida glabrata (34)	91	n.d.	97

<sup>a</sup> Data obtained by testing a random sample of C. albicans and C. glabrata isolates from Health Sciences Centre and St. Boniface Hospital, collected between Jan 2017 and Dec 2018, Susceptibility interpretations are based on updated CLSI breakpoints (M60, 2<sup>nd</sup> Edition). Isolates tested and reported are from blood only.

<sup>b</sup> For fluconazole, there is only a susceptible-dose dependent (SDD) breakpoint for C. glabrata. The percentage of C. glabrata isolates that tested SDD to fluconazole was 91%. Susceptibility of SDD isolates to fluconazole is dependent on achieving the maximum blood level possible (i.e., should use the maximum dosage regimen). Consultation with infectious diseases is recommended for further guidance.

n.d. = breakpoints have not been defined for voriconazole versus C. alabrata.

#### Table 7. Adult oral antimicrobial dosage guidelines

Antibiotic	Usual Dosages	Cost (\$) per day <sup>a</sup>
BACTERIAL AGENTS		
cillins		
oxicillin	500 mg tid	1.10
oxicillin-clavulanate	500 mg tid or 875 mg bid	2.75-3.00
kacillin	500 mg qid	1.50
icillin V	300 mg qid	0.30
alosporins	· ·	
halexin	500 mg qid	1.80
olides		
hromycin	250-500 mg daily	1.25-2.50
ithromycin	250-500 mg bid	2.25-3.25
roquinolones	· · ·	
ofloxacin	250-750 mg bid	1.40-2.50
ofloxacin	500-750 mg daily	3.50-6.50
xifloxacin	400 mg daily	1.50
rs		
damycin	450-600 mg tid	1.50-3.00
imoxazole (TMP-SMX)	1 DS (double strength) tab bid	0.25
ycycline	100 mg bid	1.30
ofurantoin (Macrobid®)	100 mg bid	1.50
tronidazole	500 mg tid	0.35
FUNGAL AGENTS		
nazole	100-400 mg daily	5.55-22.20
onazole	200-400 mg daily	8.00-16.00
VIRAL AGENTS		
ovir	200-800 mg 5x/day	5.00-16.00
yclovir	1 g tid	5.25

<sup>a</sup> Approximate cost per inpatient day excluding dispensing costs as of February 2017 based on the Manitoba Drug Interchangeability Formulary and Manufacturer's List Prices. Prices have been rounded.