

M100

Performance Standards for Antimicrobial Susceptibility Testing

This document includes updated tables for the Clinical and Laboratory Standards Institute antimicrobial susceptibility testing standards M02, M07, and M11.

A CLSI supplement for global application.

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Abstract

The data in the tables are valid only if the methodologies in CLSI documents M02,¹ M07,² and M11³ are followed. These standards contain information about disk diffusion (M02¹) and dilution (M07² and M11³) test procedures for aerobic and anaerobic bacteria. Clinicians depend heavily on information from the microbiology laboratory for treating their seriously ill patients. The clinical importance of antimicrobial susceptibility test results demands that these tests be performed under optimal conditions and that laboratories have the capability to provide results for the newest antimicrobial agents. The tables presented in M100 represent the most current information for drug selection, interpretation, and quality control using the procedures standardized in M02,¹ M07,² and M11.³ Users should replace previously published tables with these new tables. Changes in the tables since the previous edition appear in boldface type.

Clinical and Laboratory Standards Institute (CLSI). *Performance Standards for Antimicrobial Susceptibility Testing*. 32nd ed. CLSI supplement M100 (ISBN 978-1-68440-134-5 [Print]; ISBN 978-1-68440-135-2 [Electronic]). Clinical and Laboratory Standards Institute, USA, 2022.

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Suggested Citation

CLSI. *Performance Standards for Antimicrobial Susceptibility Testing*. 32nd ed. CLSI supplement M100. Clinical and Laboratory Standards Institute; 2022.

Previous Editions:

December 1986, December 1987, December 1991, December 1992, December 1994, December 1995, January 1997, January 1998, January 1999, January 2000, January 2001, January 2002, January 2003, January 2004, January 2005, January 2006, January 2007, January 2008, January 2009, January 2010, June 2010, January 2011, January 2012, January 2013, January 2014, January 2015, January 2016, January 2017, January 2018, January 2019, January 2020, March 2021

M100-Ed32
ISBN 978-1-68440-134-5 (Print)
ISBN 978-1-68440-135-2 (Electronic)
ISSN 1558-6502 (Print)
ISSN 2162-2914 (Electronic)

Volume 42, Number 2

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Overview of Changes

M100-Ed32 replaces the previous edition of the supplement, M100-Ed31, published in 2021. The major changes in M100-Ed32 are listed below. Other minor or editorial changes were made to the general formatting and to some of the table footnotes and comments. Changes to the tables since the previous edition appear in boldface type. The following are additions or changes unless otherwise noted as a “*deletion.*”

Users of M100-Ed32 should note recent formatting changes to Tables 2, including:

- **An intermediate (I) with a ^ in Tables 2 indicates agents that have the potential to concentrate in the urine. The I^ is for informational use only. The decision to report I^ is best made by each laboratory based on institution-specific guidelines and in consultation with appropriate medical personnel.**

M100 is updated and reviewed annually as new data and new agents become available. Use of outdated documents is strongly discouraged.

Section/Table	Changes
General	
CLSI Breakpoint Additions/Revisions Since 2010	<p>Revised:</p> <ul style="list-style-type: none"> • Cefiderocol <ul style="list-style-type: none"> – Disk diffusion breakpoints for Enterobacterales (p. xxiii) and <i>Acinetobacter</i> spp. (p. xxv) – Disk diffusion and MIC breakpoints for <i>Stenotrophomonas maltophilia</i> (p. xxvi) • Ceftolozane-tazobactam disk diffusion breakpoints for Enterobacterales (p. xxiii) • Piperacillin MIC breakpoints for Enterobacterales (p. xxiv) • Piperacillin-tazobactam disk diffusion and MIC breakpoints for Enterobacterales (p. xxiv) • Amoxicillin-clavulanate MIC breakpoints for <i>Haemophilus influenzae</i> and <i>Haemophilus parainfluenzae</i> (p. xxvii) • Lefamulin disk diffusion breakpoints for <i>H. influenzae</i> only (p. xxvii) and <i>Streptococcus pneumoniae</i> (p. xxviii) <p>Deleted:</p> <ul style="list-style-type: none"> • Piperacillin disk diffusion breakpoints for Enterobacterales • Amoxicillin-clavulanate disk diffusion breakpoints for <i>H. influenzae</i>

Overview of Changes (Continued)

Section/Table	Changes
Tables 1. Suggested Groupings of Antimicrobial Agents Approved by the US Food and Drug Administration for Clinical Use That Should Be Considered for Testing and Reporting by Microbiology Laboratories in the United States	
Table 1A. Nonfastidious Organisms	<p>Added:</p> <p>Cefiderocol to Group B for Enterobacterales, <i>Pseudomonas aeruginosa</i>, <i>Acinetobacter</i> spp., and <i>S. maltophilia</i> (pp. 20 and 22)</p>
Tables 2. Zone Diameter and/or MIC Breakpoints	
Table 2A. Zone Diameter and MIC Breakpoints for Enterobacterales	<p>Added:</p> <ul style="list-style-type: none"> • Ampicillin dosage regimen comments for parenteral and oral administration (p. 36) • Comment clarifying removal of piperacillin disk diffusion breakpoints (p. 36) • Comment for β-lactam combination agents regarding susceptibility of combination agents when the primary single agent is susceptible (replaced previous imipenem-relebactam surrogate testing comment) (p. 37) • Amoxicillin-clavulanate dosage regimen comments for parenteral and oral administration (p. 37) • Ampicillin-sulbactam dosage regimen comment (p. 37) • Piperacillin-tazobactam dosage regimen comment (p. 38) <p>Revised:</p> <ul style="list-style-type: none"> • General comment regarding reporting of results of amoxicillin testing with ampicillin (p. 34) • General comment regarding I[^] (p. 35) • General comment and reference to associated tables regarding direct blood culture susceptibility testing of Enterobacterales with select antimicrobial agents (p. 35) • Piperacillin MIC breakpoints (p. 36) • Ceftolozane-tazobactam disk diffusion breakpoints and associated dosage regimen comment (p. 37) • Piperacillin-tazobactam disk diffusion and MIC breakpoints (p. 38) • Cefiderocol test group and disk diffusion breakpoints (p. 40) <p>Deleted:</p> <ul style="list-style-type: none"> • Piperacillin disk diffusion breakpoints • Imipenem-relebactam surrogate testing comment

Overview of Changes (Continued)

Section/Table	Changes
Tables 2. (Continued)	
Table 2B-1. Zone Diameter and MIC Breakpoints for <i>Pseudomonas aeruginosa</i>	<p>Added:</p> <ul style="list-style-type: none"> Positive blood culture broth as an inoculum to the testing conditions box (p. 48) General comment regarding direct blood culture susceptibility testing of <i>P. aeruginosa</i> with select antimicrobial agents (p. 49) Comment for β-lactam combination agents regarding susceptibility of combination agents when the primary single agent is susceptible (replaced previous imipenem-relebactam surrogate testing comment) (p. 50) <p>Revised:</p> <ul style="list-style-type: none"> General comment regarding I[^] (p. 48) Ceftolozane-tazobactam dosage regimen comment (p. 50) Cefiderocol test group (p. 50) <p>Deleted:</p> <ul style="list-style-type: none"> Imipenem-relebactam surrogate testing comment
Table 2B-2. Zone Diameter and MIC Breakpoints for <i>Acinetobacter</i> spp.	<p>Added:</p> <ul style="list-style-type: none"> Comment for β-lactam combination agents regarding susceptibility of combination agents when the primary single agent is susceptible (p. 55) Cefiderocol testing and reporting comment (p. 55) <p>Revised:</p> <ul style="list-style-type: none"> Cefiderocol test group, disk diffusion breakpoints, and associated dosage regimen comment (p. 55)
Table 2B-4. Zone Diameter and MIC Breakpoints for <i>Stenotrophomonas maltophilia</i>	<p>Revised:</p> <ul style="list-style-type: none"> Cefiderocol test group, disk diffusion and MIC breakpoints, and reporting comment (p. 61)
Table 2B-5. MIC Breakpoints for Other Non-Enterobacterales	<p>Added:</p> <ul style="list-style-type: none"> Comment for β-lactam combination agents regarding susceptibility of combination agents when the primary single agent is susceptible (p. 63)
Table 2C. Zone Diameter and MIC Breakpoints for <i>Staphylococcus</i> spp.	<p>Added:</p> <ul style="list-style-type: none"> Dalbavancin, oritavancin, and telavancin dosage regimen comments (p. 72) Tedizolid dosage regimen comment (p. 75) <p>Revised:</p> <ul style="list-style-type: none"> <i>Staphylococcus</i> spp. indications for vancomycin (p. 72) and lefamulin (p. 75) to include methicillin-resistant <i>S. aureus</i>

Overview of Changes (Continued)

Section/Table	Changes
Tables 2. (Continued)	
Table 2D. Zone Diameter and MIC Breakpoints for <i>Enterococcus</i> spp.	<p>Added:</p> <ul style="list-style-type: none"> • Penicillin and ampicillin dosage regimen comments for parenteral and oral administration (p. 79) • Dalbavancin, oritavancin, and telavancin dosage regimen comments (p. 81) • Tedizolid dosage regimen comment (p. 83) <p>Revised:</p> <ul style="list-style-type: none"> • General comment regarding I[^] (p. 78) • Rx combination therapy comment (p. 79)
Table 2E. Zone Diameter and MIC Breakpoints for <i>Haemophilus influenzae</i> and <i>Haemophilus parainfluenzae</i>	<p>Added:</p> <ul style="list-style-type: none"> • Ampicillin dosage regimen comment (p. 86) • Comment for β-lactam combination agents regarding susceptibility of combination agents when the primary single agent is susceptible (p. 87) • Ampicillin-sulbactam, amoxicillin-clavulanate, and ceftolozane-tazobactam dosage regimen comments (p. 87) <p>Revised:</p> <ul style="list-style-type: none"> • Amoxicillin-clavulanate MIC breakpoints for susceptible and intermediate (p. 87) • Lefamulin disk diffusion breakpoint (for <i>H. influenzae</i> only) (p. 89) <p>Deleted:</p> <ul style="list-style-type: none"> • Amoxicillin-clavulanate disk diffusion breakpoints
Table 2F. Zone Diameter and MIC Breakpoints for <i>Neisseria gonorrhoeae</i>	<p>Revised:</p> <ul style="list-style-type: none"> • Tetracycline dosage regimen comment (p. 92)
Table 2G. Zone Diameter and MIC Breakpoints for <i>Streptococcus pneumoniae</i>	<p>Added:</p> <ul style="list-style-type: none"> • Amoxicillin (nonmeningitis) and amoxicillin-clavulanate (nonmeningitis) dosage regimen comment (p. 96) <p>Revised:</p> <ul style="list-style-type: none"> • Lefamulin disk diffusion breakpoint (p. 98)
Table 2H-1. Zone Diameter and MIC Breakpoints for <i>Streptococcus</i> spp. β-Hemolytic Group	<p>Added:</p> <ul style="list-style-type: none"> • Dalbavancin, oritavancin, and telavancin dosage regimen comments (p. 102) • Tedizolid dosage regimen comment (p. 104)
Table 2H-2. Zone Diameter and MIC Breakpoints for <i>Streptococcus</i> spp. Viridans Group	<p>Added:</p> <ul style="list-style-type: none"> • Dalbavancin, oritavancin, and telavancin dosage regimen comments (pp. 107-108) • Tedizolid dosage regimen comment (p. 109)

Overview of Changes (Continued)

Section/Table	Changes
Tables 2. (Continued)	
Table 2I. Zone Diameter and MIC Breakpoints for <i>Neisseria meningitidis</i>	Added: <ul style="list-style-type: none"> Ampicillin dosage regimen comment (p. 111)
Table 2J. MIC Breakpoints for Anaerobes	Added: <ul style="list-style-type: none"> Comment for β-lactam combination agents regarding susceptibility of combination agents when the primary single agent is susceptible (replaced previous imipenem-relebactam surrogate testing comment) (p. 115) Revised: <ul style="list-style-type: none"> Imipenem-relebactam test group (p. 115) Deleted: <ul style="list-style-type: none"> Imipenem-relebactam surrogate testing comment
Tables 3. Specialized Resistance Testing	
Table 3D. Tests for Colistin Resistance for Enterobacterales and <i>Pseudomonas aeruginosa</i>	Revised: <ul style="list-style-type: none"> Nomenclature for <i>Escherichia coli</i> ATCC^{®a} BAA-3170[™] (formerly <i>E. coli</i> AR Bank #0349 <i>mcr-1</i>) (p. 148) QC range for <i>E. coli</i> ATCC[®] BAA-3170[™] (p. 148) QC range for <i>P. aeruginosa</i> ATCC[®] 27853 (p. 148)
Table 3E-1. Test for Performing Disk Diffusion Directly From Positive Blood Culture Broth	Revised: <ul style="list-style-type: none"> Applicable organism groups (p. 152) Antimicrobial concentration information (refer to new Tables 3E-2 and 3E-3) (p. 152) Applicable incubation length (p. 152) Results reporting procedure (p. 152) QC recommendations (p. 153)
Table 3E-2. Zone Diameter Disk Diffusion Breakpoints for Enterobacterales Direct From Blood Culture (new table)	Added: <ul style="list-style-type: none"> Enterobacterales disk diffusion breakpoints for antimicrobial agents approved for susceptibility testing directly from blood culture bottles and associated comments and references (pp. 154-155)

Overview of Changes (Continued)

Section/Table	Changes
Tables 3. (Continued)	
Table 3E-3. Zone Diameter Disk Diffusion Breakpoints for <i>Pseudomonas aeruginosa</i> Direct From Blood Culture (new table)	Added: <ul style="list-style-type: none"> <i>P. aeruginosa</i> disk diffusion breakpoints for antimicrobial agents approved for susceptibility testing directly from blood culture bottles and associated comments and references (p. 156)
Table 3K. Test for Detecting High-Level Aminoglycoside Resistance in <i>Enterococcus</i> spp.	Revised: <ul style="list-style-type: none"> Additional testing and reporting comments (p. 175)
Tables 5. MIC QC Ranges and Associated Tables	
Table 5A-1. MIC QC Ranges for Nonfastidious Organisms and Antimicrobial Agents Excluding B-Lactam Combination Agents	Added: <ul style="list-style-type: none"> Gepotidacin QC range for <i>Enterococcus faecalis</i> ATCC® 29212 Colistin QC ranges <ul style="list-style-type: none"> <i>E. coli</i> NCTC 13486 <i>E. coli</i> ATCC® BAA-3170™ (formerly AR Bank #0349 <i>mcr-1</i>) Revised: <ul style="list-style-type: none"> Imipenem QC range for <i>E. coli</i> ATCC® 25922 Ozenoxacin QC range for <i>E. faecalis</i> ATCC® 29212 Tebipenem QC range for <i>S. aureus</i> ATCC® 29213
Table 5A-2. MIC QC Ranges for Nonfastidious Organisms and B-Lactam Combination Agents	Added: <ul style="list-style-type: none"> Ceftibuten QC ranges: <ul style="list-style-type: none"> <i>E. coli</i> ATCC® 25922 <i>E. coli</i> NCTC 13353 <i>Klebsiella pneumoniae</i> ATCC® BAA-1705™ <i>K. pneumoniae</i> ATCC® BAA-2814™ Meropenem QC ranges: <ul style="list-style-type: none"> <i>E. coli</i> NCTC 13353 <i>Acinetobacter baumannii</i> NCTC 13304 Revised: <ul style="list-style-type: none"> Imipenem QC ranges: <ul style="list-style-type: none"> <i>E. coli</i> ATCC® 25922 <i>K. pneumoniae</i> ATCC® 700603 Imipenem-relebactam QC ranges: <ul style="list-style-type: none"> <i>E. coli</i> ATCC® 25922 <i>K. pneumoniae</i> ATCC® 700603 Meropenem-nacubactam QC range for <i>E. coli</i> ATCC® 25922

Overview of Changes (Continued)

Section/Table	Changes
Tables 5. (Continued)	
Table 5B. MIC QC Ranges for Fastidious Organisms (Broth Dilution Methods)	Revised: <ul style="list-style-type: none"> • Grepafloxacin QC range for <i>H. influenzae</i> ATCC® 49247
Table 5C. MIC QC Ranges for <i>Neisseria gonorrhoeae</i> (Agar Dilution Method)	Added: <ul style="list-style-type: none"> • Gentamicin QC range for <i>Neisseria gonorrhoeae</i> ATCC® 49226
Table 5D. MIC QC Ranges for Anaerobes (Agar Dilution Method)	Added: <ul style="list-style-type: none"> • Tebipenem QC ranges: <ul style="list-style-type: none"> – <i>Bacteroides fragilis</i> ATCC® 25285 – <i>Bacteroides thetaiotaomicron</i> ATCC® 29741 – <i>Clostridioides difficile</i> ATCC® 700057 – <i>Eggerthella lenta</i> ATCC® 43055 Revised: <ul style="list-style-type: none"> • Fidaxomicin QC range for <i>C. difficile</i> ATCC® 700057
Appendixes	
Appendix B. Intrinsic Resistance; B1. Enterobacterales	Added: <ul style="list-style-type: none"> • Polymyxin B and colistin for <i>Hafnia alvei</i> (p. 249) • Footnote regarding intrinsic resistance for <i>Hafnia paralvei</i> (p. 249)
Appendix C. QC Strains for Antimicrobial Susceptibility Tests	Added: <ul style="list-style-type: none"> • QC strain <i>E. coli</i> NCTC 13486 (p. 257) Revised: <ul style="list-style-type: none"> • Nomenclature for <i>E. coli</i> ATCC® BAA-3170™ (formerly <i>E. coli</i> AR Bank #0349 <i>mcr-1</i>) (p. 257)

Overview of Changes (Continued)

Section/Table	Changes
Appendixes (Continued)	
Appendix E. Dosage Regimens Used to Establish Susceptible or Susceptible-Dose Dependent Breakpoints	<p>Added:</p> <ul style="list-style-type: none"> • Enterobacteriales <ul style="list-style-type: none"> – Ampicillin (parenteral and oral) (p. 268) – Amoxicillin-clavulanate (parenteral and oral) (p. 268) – Ampicillin-sulbactam (p. 268) – Indications for cefazolin (uncomplicated UTIs and infections other than uncomplicated UTIs) (p. 268) – Imipenem-relebactam exclusion for the family <i>Morganellaceae</i> (p. 269) – Piperacillin-tazobactam (p. 270) • <i>Staphylococcus</i> spp. <ul style="list-style-type: none"> – Dalbavancin, oritavancin, tedizolid, and televancin as applicable to <i>S. aureus</i> only (p. 271) • <i>Enterococcus</i> spp. <ul style="list-style-type: none"> – Ampicillin (parenteral and oral) (p. 271) – Dalbavancin as applicable to vancomycin-susceptible <i>E. faecalis</i> only (p. 271) • <i>H. influenzae</i> and <i>H. parainfluenzae</i> <ul style="list-style-type: none"> – Ampicillin (p. 272) – Ampicillin-sulbactam (p. 272) • <i>S. pneumoniae</i> <ul style="list-style-type: none"> – Amoxicillin (p. 272) – Amoxicillin-clavulanate (p. 272) • <i>Streptococcus</i> spp. β-hemolytic group <ul style="list-style-type: none"> – Dalbavancin as applicable to <i>S. pyogenes</i>, <i>S. agalactiae</i>, and <i>S. dysgalactiae</i> only (p. 273) – Tedizolid as applicable to <i>S. pyogenes</i> and <i>S. agalactiae</i> only (p. 273) • <i>Streptococcus</i> spp. viridans group <ul style="list-style-type: none"> – Dalbavancin and tedizolid as applicable to <i>S. anginosus</i> group only (p. 273) • <i>Neisseria meningitis</i> <ul style="list-style-type: none"> – Ampicillin (p. 273) <p>Deleted:</p> <ul style="list-style-type: none"> • <i>P. aeruginosa</i> <ul style="list-style-type: none"> – Ticarcillin

Overview of Changes (Continued)

Section/Table	Changes
Glossaries	
Glossary II. Antimicrobial Agent Abbreviation(s), Route(s) of Administration, and Drug Class	<p>Added:</p> <ul style="list-style-type: none"> • Rifapentine <p>Corrected:</p> <ul style="list-style-type: none"> • Cefepime-nacubactam abbreviation • Rifaximin abbreviation • Sulbactam-durlobactam route of administration
Glossary III. List of Identical Abbreviations Used for More Than One Antimicrobial Agent in US Diagnostic Products	<p>Added:</p> <ul style="list-style-type: none"> • Cefdinir • Cefditoren • Clinafloxacin • Cloxacillin • Cefpirome • Cefprozil • Ceftolozane-tazobactam • Colistin • Tobramycin • Trimethoprim

Abbreviations: ATCC®, American Type Culture Collection; MIC, minimal inhibitory concentration; UTI, urinary tract infection.

Footnote

- a. ATCC® is a registered trademark of the American Type Culture Collection.

CLSI Breakpoint Additions/Revisions Since 2010

Previous breakpoints can be found in the edition of M100 that precedes the document listed in the column labeled “Date of Addition/Revision (M100 edition).” For example, previous breakpoints for aztreonam are listed in M100-S19 (January 2009).

Antimicrobial Agent	Date of Addition/Revision (M100 edition)	Disk Diffusion Breakpoints		MIC Breakpoints		Comments
		New ^a	Revised ^b	New ^a	Revised ^b	
Enterobacterales						
Azithromycin	January 2015 (M100-S25)	X		X		<i>S. enterica</i> ser. Typhi only
	March 2021 (M100-Ed31)	X		X		<i>Shigella</i> spp. Previously assigned an ECV
Aztreonam	January 2010 (M100-S20)		X		X	
Cefazolin (parenteral)	January 2010 (M100-S20)				X	Removed disk diffusion breakpoints January 2010 (M100-S20)
	January 2011 (M100-S21)	X			X	
	January 2016 (M100-S26)	X		X		For uncomplicated UTIs
Cefazolin (oral)	January 2014 (M100-S24)	X		X		Surrogate test for oral cephalosporins and uncomplicated UTIs
Cefepime	January 2014 (M100-S24)		X		X	Revised breakpoints include SDD
Cefiderocol	January 2019 (M100, 29th ed.)			X		
	January 2020 (M100, 30th ed.)	X				
	February 2022 (M100-Ed32)		X			
Cefotaxime	January 2010 (M100-S20)		X		X	
Ceftaroline	January 2013 (M100-S23)	X		X		
Ceftazidime	January 2010 (M100-S20)		X		X	
Ceftazidime-avibactam	January 2018 (M100, 28th ed.)	X		X		
Ceftizoxime	January 2010 (M100-S20)		X		X	
Ceftolozane-tazobactam	January 2016 (M100-S26)			X		
	January 2018 (M100, 28th ed.)	X				
	February 2022 (M100-Ed32)		X			
Ceftriaxone	January 2010 (M100-S20)		X		X	

CLSI Breakpoint Additions/Revisions Since 2010 (Continued)

Antimicrobial Agent	Date of Addition/Revision (M100 edition)	Disk Diffusion Breakpoints		MIC Breakpoints		Comments
		New ^a	Revised ^b	New ^a	Revised ^b	
Enterobacterales (Continued)						
Ciprofloxacin	January 2012 (M100-S22)		X		X	<i>Salmonella</i> spp. (including <i>S. enterica</i> ser. Typhi)
	January 2019 (M100, 29th ed.)		X		X	Non- <i>Salmonella</i> spp.
Colistin	January 2020 (M100, 30th ed.)			X		Previously assigned an ECV
Doripenem	June 2010 (M100-S20-U)	X		X		
Ertapenem	June 2010 (M100-S20-U)		X		X	
	January 2012 (M100-S22)		X		X	
Imipenem	June 2010 (M100-S20-U)		X		X	
Imipenem-relebactam	March 2021 (M100-Ed31)	X		X		
Levofloxacin	January 2013 (M100-S23)		X		X	<i>Salmonella</i> spp. (including <i>S. enterica</i> ser. Typhi)
	January 2019 (M100, 29th ed.)		X		X	Non- <i>Salmonella</i> spp.
Meropenem	June 2010 (M100-S20-U)		X		X	
Meropenem-vaborbactam	January 2019 (M100, 29th ed.)	X		X		
Norfloxacin	January 2020 (M100, 30th ed.)	X		X		Reinstated breakpoints deleted from M100, 29th ed.
Ofloxacin	January 2013 (M100-S23)			X		<i>Salmonella</i> spp. (including <i>S. enterica</i> ser. Typhi)
Pefloxacin	January 2015 (M100-S25)	X				<i>Salmonella</i> spp. (including <i>S. enterica</i> ser. Typhi) Surrogate test for ciprofloxacin
Piperacillin	February 2022 (M100-Ed32)				X	Removed disk diffusion breakpoints due to reassessment of disk correlates for revised MIC breakpoints
Piperacillin-tazobactam	February 2022 (M100-Ed32)		X		X	
Polymyxin B	January 2020 (M100, 30th ed.)			X		

CLSI Breakpoint Additions/Revisions Since 2010 (Continued)

Antimicrobial Agent	Date of Addition/Revision (M100 edition)	Disk Diffusion Breakpoints		MIC Breakpoints		Comments
		New ^a	Revised ^b	New ^a	Revised ^b	
<i>Pseudomonas aeruginosa</i>						
Cefiderocol	January 2019 (M100, 29th ed.)			X		
	January 2020 (M100, 30th ed.)	X				
Ceftazidime-avibactam	January 2018 (M100, 28th ed.)	X		X		
Ciprofloxacin	January 2019 (M100, 29th ed.)		X		X	
Colistin	January 2017 (M100, 27th ed.)				X	
	January 2020 (M100, 30th ed.)				X	
Doripenem	January 2012 (M100-S22)	X		X		
Imipenem	January 2012 (M100-S22)		X		X	
Imipenem-relebactam	March 2021 (M100-Ed31)	X		X		
Levofloxacin	January 2019 (M100, 29th ed.)		X		X	
Meropenem	January 2012 (M100-S22)		X		X	
Norfloxacin	January 2020 (M100, 30th ed.)	X		X		Reinstated breakpoints deleted from M100, 29th ed.
Piperacillin	January 2012 (M100-S22)		X		X	
Piperacillin-tazobactam	January 2012 (M100-S22)		X		X	
Polymyxin B	January 2020 (M100, 30th ed.)				X	
Ticarcillin	January 2012 (M100-S22)		X		X	
Ticarcillin-clavulanate	January 2012 (M100-S22)		X		X	
<i>Acinetobacter spp.</i>						
Cefiderocol	January 2019 (M100, 29th ed.)			X		
	January 2020 (M100, 30th ed.)	X				
	February 2022 (M100-Ed32)		X			
Colistin	January 2020 (M100, 30th ed.)				X	
Doripenem	January 2014 (M100-S24)	X		X		
Imipenem	January 2014 (M100-S24)		X		X	
Meropenem	January 2014 (M100-S24)		X		X	
Polymyxin B	January 2020 (M100, 30th ed.)				X	

CLSI Breakpoint Additions/Revisions Since 2010 (Continued)

Antimicrobial Agent	Date of Addition/Revision (M100 edition)	Disk Diffusion Breakpoints		MIC Breakpoints		Comments
		New ^a	Revised ^b	New ^a	Revised ^b	
<i>Stenotrophomonas maltophilia</i>						
Cefiderocol	January 2019 (M100, 29th ed.)			X		
	January 2020 (M100, 30th ed.)	X				
	February 2022 (M100-Ed32)		X		X	
Other Non-Enterobacterales						
Norfloxacin	January 2020 (M100, 30th ed.)	X		X		Reinstated breakpoints deleted from M100, 29th ed.
<i>Staphylococcus</i> spp.						
Cefoxitin	January 2019 (M100, 29th ed.)		X			<i>S. epidermidis</i> Surrogate test for oxacillin
Ceftaroline	January 2013 (M100-S23)	X		X		
	January 2019 (M100, 29th ed.)		X		X	Revised breakpoints include SDD
Dalbavancin	January 2018 (M100, 28th ed.)			X		
Lefamulin	March 2021 (M100-Ed31)	X		X		
Norfloxacin	January 2020 (M100, 30th ed.)	X		X		Reinstated breakpoints deleted from M100, 29th ed.
Oritavancin	January 2016 (M100-S26)			X		
Oxacillin	January 2016 (M100-S26)		X		X	<i>S. pseudintermedius</i>
	January 2018 (M100, 28th ed.)		X		X	<i>S. schleiferi</i>
	January 2019 (M100, 29th ed.)		X			<i>S. epidermidis</i>
	March 2021 (M100-Ed31)				X	<i>Staphylococcus</i> spp. except <i>S. aureus</i> and <i>S. lugdunensis</i>
Tedizolid	January 2016 (M100-S26)			X		
Telavancin	January 2016 (M100-S26)	X		X		
	January 2017 (M100, 27th ed.)					Removed disk diffusion breakpoints January 2017 (M100, 27th ed.)

CLSI Breakpoint Additions/Revisions Since 2010 (Continued)

Antimicrobial Agent	Date of Addition/Revision (M100 edition)	Disk Diffusion Breakpoints		MIC Breakpoints		Comments
		New ^a	Revised ^b	New ^a	Revised ^b	
<i>Enterococcus</i> spp.						
Dalbavancin	January 2018 (M100, 28th ed.)			X		
Daptomycin	January 2019 (M100, 29th ed.)				X	
	January 2020 (M100, 30th ed.)				X	Separated into two sets of breakpoints: <ul style="list-style-type: none"> • <i>Enterococcus</i> spp other than <i>Enterococcus faecium</i> • <i>E. faecium</i> (includes SDD)
Norfloxacin	January 2020 (M100, 30th ed.)	X		X		Reinstated breakpoints deleted from M100, 29th ed.
Oritavancin	January 2016 (M100-S26)			X		
Tedizolid	January 2016 (M100-S26)			X		
Telavancin	January 2016 (M100-S26)	X		X		
	January 2017 (M100, 27th ed.)					Removed disk diffusion breakpoints January 2017 (M100, 27th ed.)
<i>Haemophilus influenzae</i> and <i>Haemophilus parainfluenzae</i>						
Amoxicillin-clavulanate	February 2022 (M100-Ed32)				X	Removed disk diffusion breakpoints February 2022 (M100-Ed32)
Ceftaroline	January 2013 (M100-S23)	X		X		
Ceftolozane-tazobactam	March 2021 (M100-Ed31)			X		
Doripenem	January 2012 (M100-S22)	X		X		
Lefamulin	March 2021 (M100-Ed31)	X		X		
	February 2022 (M100-Ed32)		X			For <i>H. influenzae</i> only

CLSI Breakpoint Additions/Revisions Since 2010 (Continued)

Antimicrobial Agent	Date of Addition/Revision (M100 edition)	Disk Diffusion Breakpoints		MIC Breakpoints		Comments
		New ^a	Revised ^b	New ^a	Revised ^b	
<i>Neisseria gonorrhoeae</i>						
Azithromycin	January 2019 (M100, 29th ed.)			X		Previously assigned as ECV
	March 2021 (M100-Ed31)	X				
<i>Streptococcus pneumoniae</i>						
Ceftaroline	January 2013 (M100-S23)	X		X		
Doripenem	January 2012 (M100-S22)			X		
Doxycycline	January 2013 (M100-S23)	X		X		
Lefamulin	March 2021 (M100-Ed31)	X		X		
	February 2022 (M100-Ed32)		X			
Tetracycline	January 2013 (M100-S23)		X		X	
<i>Streptococcus</i> spp. β-Hemolytic Group						
Ceftaroline	January 2013 (M100-S23)	X		X		
Dalbavancin	January 2018 (M100, 28th ed.)			X		
Doripenem	January 2012 (M100-S22)			X		
Oritavancin	January 2016 (M100-S26)			X		
Tedizolid	January 2016 (M100-S26)			X		
Telavancin	January 2016 (M100-S26)	X		X		
	January 2017 (M100, 27th ed.)					Removed disk diffusion breakpoints January 2017 (M100, 27th ed.)
<i>Streptococcus</i> spp. Viridans Group						
Ceftolozane-tazobactam	January 2016 (M100-S26)			X		
Dalbavancin	January 2018 (M100, 28th ed.)			X		
Doripenem	January 2012 (M100-S22)			X		
Oritavancin	January 2016 (M100-S26)			X		
Tedizolid	January 2016 (M100-S26)			X		
Telavancin	January 2016 (M100-S26)	X		X		
	January 2017 (M100, 27th ed.)					Removed disk diffusion breakpoints January 2017 (M100, 27th ed.)

CLSI Breakpoint Additions/Revisions Since 2010 (Continued)

Antimicrobial Agent	Date of Addition/Revision (M100 edition)	Disk Diffusion Breakpoints		MIC Breakpoints		Comments
		New ^a	Revised ^b	New ^a	Revised ^b	
Anaerobes						
Doripenem	January 2012 (M100-S22)			X		
Imipenem-relebactam	March 2021 (M100-Ed31)			X		
Piperacillin-tazobactam	January 2017 (M100, 27th ed.)			X		
	January 2018 (M100, 28th ed.)			X		

Abbreviations: ECV, epidemiological cutoff value; SDD, susceptible-dose-dependent; UTI, urinary tract infection.

Footnotes

- “New” indicates the breakpoints are listed for the first time for a specific organism or organism group in the respective Table 2.
- “Revised” indicates previously established breakpoints for a specific organism or organism group in the respective Table 2 have changed. In some cases, unique breakpoints were added for a specific genus or species previously included within the organism or organism group breakpoints (eg, “*Salmonella* spp. [including *S. enterica* ser. Typhi]” was previously grouped with Enterobacterales).

CLSI Archived Resources

Resource	Web Address for Archived Table
Breakpoints that have been eliminated from M100 since 2010 have been relocated to the CLSI website.	https://clsi.org/media/pqlom3b5/_m100_archived_drugs_table.pdf
Methods that have been eliminated from M100 have been relocated to the CLSI website.	https://clsi.org/media/nszl4tbc/_m100_archived_methods_table.pdf
QC ranges that have been eliminated from M100 since 2010 have been relocated to the CLSI website.	https://clsi.org/media/r31oari2/_m100_archived_qc_table.pdf
ECVs that have been replaced by breakpoints have been relocated to the CLSI website.	https://clsi.org/media/3mekwxft/_m100_archived_ecvs_table.pdf

Abbreviations: ECV, epidemiological cutoff value; QC, quality control.

NOTE: The content of this document is supported by the CLSI consensus process and does not necessarily reflect the views of any single individual or organization.

Summary of CLSI Processes for Establishing Breakpoints and Quality Control Ranges

The Clinical and Laboratory Standards Institute (CLSI) is an international, voluntary, not-for-profit, interdisciplinary, standards-developing, and educational organization accredited by the American National Standards Institute that develops and promotes the use of consensus-developed standards and guidelines within the health care community. These consensus standards and guidelines are developed in an open and consensus-seeking forum to cover critical areas of diagnostic testing and patient health care. CLSI is open to anyone or any organization that has an interest in diagnostic testing and patient care. Information about CLSI can be found at www.clsi.org.

The CLSI Subcommittee on Antimicrobial Susceptibility Testing reviews data from a variety of sources and studies (eg, *in vitro*, pharmacokinetics-pharmacodynamics, and clinical studies) to establish antimicrobial susceptibility test methods, breakpoints, and QC parameters. The details of the data necessary to establish breakpoints, QC parameters, and how the data are presented for evaluation are described in CLSI document M23.⁴

Over time, a microorganism's susceptibility to an antimicrobial agent may decrease, resulting in a lack of clinical efficacy and/or safety. In addition, microbiological methods and QC parameters may be refined to ensure more accurate and better performance of susceptibility test methods. Because of these types of changes, CLSI continually monitors and updates information in its documents. Although CLSI standards and guidelines are developed using the most current information available at the time, the field of science and medicine is always changing; therefore, standards and guidelines should be used in conjunction with clinical judgment, current knowledge, and clinically relevant laboratory test results to guide patient treatment.

Additional information, updates, and changes in this document are found in the meeting summary minutes of the Subcommittee on Antimicrobial Susceptibility Testing at <https://clsi.org/meetings/ast-file-resources/>.

CLSI Reference Methods vs Commercial Methods and CLSI vs US Food and Drug Administration Breakpoints

It is important for users of M02,¹ M07,² and M100 to recognize that the standard methods described in CLSI documents are reference methods. These methods may be used for routine antimicrobial susceptibility testing of patient isolates, for evaluating commercial devices that will be used in medical laboratories, or by drug or device manufacturers for testing new agents or systems. Results generated by reference methods, such as those included in CLSI documents, may be used by regulatory authorities to evaluate the performance of commercial susceptibility testing devices as part of the approval process. Clearance by a regulatory authority indicates the commercial susceptibility testing device provides susceptibility results that are substantially equivalent to results generated using reference methods for the organisms and antimicrobial agents described in the device manufacturer's approved package insert.

CLSI breakpoints may differ from those approved by various regulatory authorities for many reasons, including use of different databases, differences in data interpretation, differences in doses used in different parts of the world, and public health policies. Differences also exist because CLSI proactively evaluates the need for changing breakpoints. The reasons why breakpoints may change and the manner in which CLSI evaluates data and determines breakpoints are outlined in CLSI document M23.⁴

Following a decision by CLSI to change an existing breakpoint, regulatory authorities may also review data to determine how changing breakpoints may affect the safety and effectiveness of the antimicrobial agent for the approved indications. If the regulatory authority changes breakpoints, commercial device manufacturers may have to conduct a clinical trial, submit the data to the regulatory authority, and await review and approval. For these reasons, a delay of one or more years may be needed if a breakpoint and interpretive category change is to be implemented by a device manufacturer. In the United States, it is acceptable for laboratories that use US Food and Drug Administration (FDA)-cleared susceptibility testing devices to use existing FDA breakpoints. Either FDA or CLSI susceptibility breakpoints are acceptable to laboratory accrediting organizations in the United States. Policies in other countries may vary. Each laboratory should check with the manufacturer of its antimicrobial susceptibility test system for additional information on the breakpoints and interpretive categories used in its system's software.

Following discussions with appropriate stakeholders (eg, infectious diseases and pharmacy practitioners, the pharmacy and therapeutics and infection prevention committees of the medical staff, and the antimicrobial stewardship team), newly approved or revised breakpoints may be implemented by laboratories. Following verification, CLSI disk diffusion test breakpoints may be implemented as soon as they are published in M100. If a device includes antimicrobial test concentrations sufficient to allow interpretation of susceptibility and resistance to an agent using the CLSI breakpoints, a laboratory could choose to, after appropriate verification, interpret and report results using CLSI breakpoints.

Subcommittee on Antimicrobial Susceptibility Testing Mission Statement

The Subcommittee on Antimicrobial Susceptibility Testing is composed of representatives from the professions, government, and industry, including microbiology laboratories, government agencies, health care providers and educators, and pharmaceutical and diagnostic microbiology industries. Using the CLSI voluntary consensus process, the subcommittee develops standards that promote accurate antimicrobial susceptibility testing and appropriate reporting. The mission of the Subcommittee on Antimicrobial Susceptibility Testing is to:

- Develop standard reference methods for antimicrobial susceptibility tests.
- Provide quality control parameters for standard test methods.
- Establish breakpoints and interpretive categories for the results of standard antimicrobial susceptibility tests and provide epidemiological cutoff values when breakpoints are not available.
- Provide suggestions for testing and reporting strategies that are clinically relevant and cost-effective.
- Continually refine standards and optimize detection of emerging resistance mechanisms through development of new or revised methods, breakpoints, and quality control parameters.
- Educate users through multimedia communication of standards and guidelines.
- Foster a dialogue with users of these methods and those who apply them.

The ultimate purpose of the subcommittee's mission is to provide useful information to enable laboratories to assist the clinician in the selection of appropriate antimicrobial therapy for patient care. The standards and guidelines are meant to be comprehensive and to include all antimicrobial agents for which the data meet established CLSI guidelines. The values that guide this mission are quality, accuracy, fairness, timeliness, teamwork, consensus, and trust.

Instructions for Use of Tables

These instructions apply to:

- **Tables 1A and 1B:** suggested groupings of antimicrobial agents that should be considered for testing and reporting by microbiology laboratories. These guidelines are based on antimicrobial agents approved by the US Food and Drug Administration (FDA) for clinical use in the United States. In other countries, placement of antimicrobial agents in Tables 1A and 1B should be based on available drugs approved for clinical use by relevant regulatory organizations.
- **Tables 2A through 2I:** tables for each organism group that contain:
 - Recommended testing conditions
 - Routine QC recommendations (also see Chapter 4 in M02¹ and M07²)
 - General comments for testing the organism group and specific comments for testing particular agent/organism combinations
 - Suggested agents that should be considered for routine testing and reporting by medical microbiology laboratories, as specified in Tables 1A and 1B (test/report groups A, B, C, U)
 - Additional drugs that are appropriate for the respective organism group but would generally not warrant routine testing by a medical microbiology laboratory in the United States (test/report group O for “other”; test/report group Inv. for “investigational” [not yet FDA approved])
 - Zone diameter and minimal inhibitory concentration (MIC) breakpoints
- **Tables 1C and 2J:** tables containing specific recommendations for testing and reporting results on anaerobes and some of the information listed in the bullets above
- **Tables 3A to 3K:** tables describing tests to detect particular resistance types in specific organisms or organism groups