

# Nyheter från EUCAST

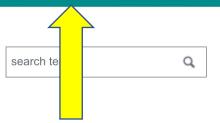
## **Gunnar Kahlmeter**

**EUCAST** 

The EUCAST Development Laboratory Clinical microbiology, Växjö, Sweden



European Society of Clinical Microbiology and Infectious Diseases



Organization

**Consultations** 

**EUCAST News** 

New definitions of S, I and R

Clinical breakpoints and dosing

Rapid AST in blood cultures

Expert rules and intrinsic resistance

Resistance mechanisms

**SOPs and Guidance documents** 

MIC and zone distributions and ECOFFs

**AST** of bacteria

**AST** of mycobacteria

**AST** of fungi

**AST** of veterinary pathogens

Frequently Asked Questions (FAQ)

Meetings

**Publications and documents** 

**Presentations and statistics** 



### The European Committee on Antimicrobial Susceptibility Testing - EUCAST

EUCAST is a standing committee jointly organized by ESCMID, ECDC and European national breakpoint committees. EUCAST was formed in 1997. It has been chaired by lan Phillips (1997 - 2001), Gunnar Kahlmeter (2001 - 2012), Rafael Canton 2012 - 2016) and Christian Giske (2016 - ). Its scientific secretary is Derek Brown (1997 - 2016) and John Turnidge (2016 - ). Its webmaster is Gunnar Kahlmeter (2001 - ). From 2016, Rafael Canton is the Clinical Data Co-ordinator and from 2012, Gunnar Kahlmeter is the Technical Data Co-ordinator and Head of the EUCAST Development Laboratory.

Since the very beginning, Johan Mouton, was important to EUCAST in that he was an ardent supporter of European harmonisation and of bringing PKPD to bear on breakpoint setting. Johan died in the evening of Tuesday 9 July, 2019 .The Steering Committee meeting 8 - 9 of July was the only meeting since 2002 to which Johan could not travel. We shall miss Johan for his courage and his passion for getting it right! Some of our memories of Johan are on the ESCMID website.

The EUCAST Development Laboratory for antibacterial agents is located in Sweden and breakpoints for Vibrio species can be addressed through gunnar.kahlmeter[at]eucast.org or erika.matuschek[at]eucast.org.

QUICK NAVIGATION





18 Oct 2021

**EUCAST** on "intrinsic resistance and unusual phenotypes" updated

06 Oct 2021

General consultation on colistin breakpoints

04 Oct 2021

Seminar on EUCAST Disk diffusion of anaerobic bacteria 9 December

04 Oct 2021

General consultation on breakpoints for anaerobic bacteria

30 Sep 2021

General consultation on clinical

**Links and Contacts** 

search term Q

<b>EUCAST News</b>					
Organization  Consultations  EUCAST News  New definitions of S, I and R	The European Committee on Antimicrobial Susceptibility Testing — EUCAST	EUCAST News	~		
Clinical breakpoints and dosing		Dec			
Rapid AST in blood cultures	FUGACTAL	RSS	<u> </u>		
Expert rules and intrinsic resistance	EUCAST News	RSS 2.0			
Resistance mechanisms	Here you can find the latest news and updates from EUCAST.	RDF			
SOPs and Guidance documents		<b>№</b> ATOM 0.3			
MIC and zone distributions and ECOFFs	18 Oct 2021				
AST of bacteria	EUCAST on "intrinsic resistance and unusual phenotypes" updated The EUCAST document Advice on intrinsic resistance and exceptional	EUCAST NEWSLETTE	EUCAST NEWSLETTER SIGN UP		
AST of mycobacteria		Email:			
AST of fungi	06 Oct 2021				
AST of veterinary pathogens	General consultation on colistin breakpoints	GN UP			
Frequently Asked Questions (FAQ)	EUCAST is proposing to change colistin breakpoints to breakpoints in				
Meetings		_			
Publications and documents	04 Oct 2021				
Presentations and statistics	Seminar on EUCAST Disk diffusion of anaerobic bacteria 9 December Disk diffusion of anaerobic bacteria in 16-20 hours (9 December, 2021; at				
Videos and online seminars					
Warnings!	04 Oct 2021				
Translations	General consultation on breakpoints for anaerobic bacteria				
Information for industry	EUCAST general and public consultation on breakpoints for anaerobic				

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#### Publications and documents

Organization

J Antimicrob Chemother 2015; 70: 2427-2439 doi:10.1093/jac/dkv145 Advance Access publication 18 June 2015 Journal of Antimicrobial Chemotherapy

... Consultations

Consultations

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Frequently Asked Questions (FAQ)

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#### **Publications and documents**

**Rationale Documents** 

**Standard Operation Procedures** 

Discussion documents

#### Consultations

Publications in journals

Technical notes

**Posters** 

**Other Documents** 

**External documents** 

#### **EUCAST Consultations**

November 14, 2020

#### **Current consultations (download form for comments)**

Address the form/responses to Prof John Turnidge of EUCAST.

 EUCAST has made available the manual, a reading guide and QC criteria related to disk diffusion of anaerobic bacteria.

The 2014 Garrod Lecture: EUCAST – are we heading towards international agreement?

Gunnar Kahlmeter\*

- Vibrio species genral consultation on EUCAST clinical breakpoints for five species paths (1 October - 15 November). Queries on the material can be addressed to G Kahlmeter (gunnar.kahlmeter@eucast.org)
- Breakpoints for anaerobic bacteria and supplementary MIC distributions (4 October 15 November, 2021). Queries on the material can be addressed to G Kahlmeter (gunnar.kahlmeter@eucast.org).

EUCAST has made available the manual, a reading guide and QC criteria related to disk diffusion of anaerobic bacteria.

There will be an online seminar on the new clinical breakpoints for anaerobic bacteria and the new disk diffusion method on the 9th of December, 2021 (8AM, 1PM and 4PM). More information.

- Colistin breakpoints (6 October 15 November, 2021). Queries can be addressed to J Turnidge (john.turnidge@eucast.org).
- Download form for comments.

#### Previous consultations with comments and EUCAST responses:

- Breakpoint table v. 11.0 (the 2021 table, valid from the 1 January, 2021). Consultation closed 18 December, 2020. Comments and EUCAST responses (19 December, 2020)
- Fosfomycin breakpoints revision. Consultation closed 30 November, 2020.
   Comments and EUCAST response (15 December, 2020).
- Fluoroquinolone breakpoint adjustments (Campylobacter, Corynebacteria, Bacillus).
   Consultation closed 31 October, 2020. There were no comments.

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#### Rapid AST in bloodcultures

Organization

Consultations

**EUCAST News** 

New definitions of S, I and R

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#### Rapid AST in blood cultures

Methods

Breakpoints for short incubation

Screening for resistance mechanisms

FAQ on RAST



Rapid AST in bloodcultures

#### Rapid AST directly from blood culture bottles

January 3, 2021

EUCAST has developed a method for rapid AST (reading at 4, 6 or 8h incubation) directly from positive blood culture bottles (RAST).

Following the initial development, published in 2019, a clinical trial in 55 laboratories was performed and published in 2020.

These are the essential steps in the RAST method:

### RAST – nyheter under 2022

- Brytpunkter för förlängd inkubation 16, 20, 24 timmar
- Tolkningskriterier för fler antibiotika
- Arter ingen utvidgning

AST of mycobacteria

**AST** of fungi

**AST** of veterinary pathogens

Frequently Asked Questions (FAQ)

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- identity of species must be known prior to interpretation of AST results.
- the method is validated for the following species.
  - Escherichia coli
  - Klebsiella pneumoniae
  - Pseudomonas aeruginosa
  - Staphylococcus aureus
  - Streptococcus pneumoniae
  - Enterococcus faecalis and Enterococcus faecium
  - Acinetobacter baumannii (added 2 May, 2019)

#### Antimicrobial wild type distributions of microorganisms

Mic distributions include collated data from multiple sources, geographical areas and time periods an

Programvaran för MIC- och Zondistributioner helt omprogrammerad Tillgänglig från EUCASTs hemsida Samtliga distributioner kurerade 2021 och 2022. Nya distributioner för ett flertal antibiotika och flertal arter tillförda





#### MIC and Inhibition zone diameter distributions of microorganisms without and with phenotypically evident resistance mechanisms

#### MIC and inhibition zone diameter distributions

Distributions are shown as "aggregated distributions" and as "aggregated weighted distributions". For aggregated weighted distributions (as defined in SOP 10) were added to form one common distribution. For aggregated weighted distributions each individual distribution was converted to contribute equally to the common aggregated distributions. In this way large distributions are prevented from drowning out smaller distributions.

#### 1. MIC distributions

The website gives MIC distributions for individual micro-organisms (bacteria and fungi) and antimicrobial agents in tables and histograms. The distributions are based on collated data from an increasing total of more than 30 000 MIC distributions from worldwide sources. Unless otherwise specifically stated, the data are representative of results obtained with MIC methods performed by or calibrated to reference broth microdilution using ISO-20776-2. Different methods do not give exactly the same results, but the results rarely vary by more than one doubling dilution step. In this way the aggregated MIC distributions encompass the variation between different investigators, laboratories, geographic locations and time periods.

#### 2. Inhibition zone diameter distributions

The website gives inhibition zone diameter distributions for individual organisms and antimicrobial agents in tables and histograms. The distributions are based on collated data from an increasing number of sources worldwide. The data are representative of results obtained with the EUCAST disk diffusion method (launched in 2009 - see www.eucast.org).

#### Clinical MIC and Zone diameter breakpoints

These are not shown on this website - please consult the EUCAST breakpoint tables for bacteria and fungi on www.eucast.org.

#### Epidemiological cut-off values (ECOFF) and tentative epidemiological cut-off values (TECOFF)

ECOFFs (and TECOFFs) distinguish microorganisms without (wild type) and with phenotypically detectable acquired resistance mechanisms (non-wild type) to the agent in question. The epidemiological cut-off value is shown in the tables and the bottom left-hand corner of each MIC and zone diameter graph. TECOFFs (ECOFFs in parentheses) are based on 3 or 4 distributions and ECOFFs on at least 5 and up to 100 or more distributions.

#### Limitations

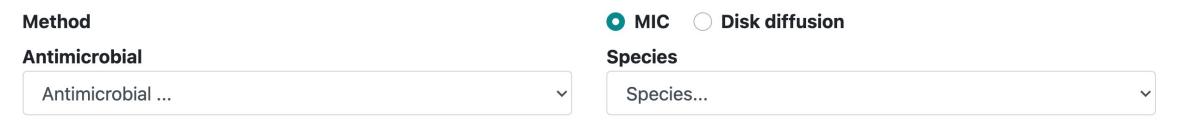
The database is being curated in accordance with EUCAST SOP 10.

## Antimicrobial wild type distributions of microorganisms

Mic distributions include collated data from multiple sources, geographical areas and time periods and can never be used to infer rates of resistance

### Search for ANTIMICROBIAL or SPECIES

### Search database



EUCAST 2021

#### Search database

Method	MIC  Disk diffusion					
Antimicrobial	Species					
Gentamicin	Species					

Elements per page 50 v

#### MIC distributions for Gentamicin, 2021-10-22

Antimicrobial: Gentamicin (Method: MIC)

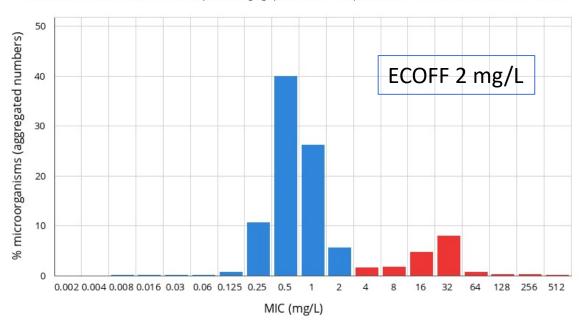
	0.002	0.004	0.008	0.016	0.03	0.06	0.125	0.25	0.5	1	2	4	8	16	32	64	128	256	512	Distributions	Observations	(T)ECOFF	Confidence interval
Achromobacter xylosoxidans	0	0	0	0	0	0	0	0	0	0	0	0	2	17	73	0	0	0	0	1	92	-	
Acinetobacter anitratus	0	0	0	0	0	0	0	6	29	37	25	7	5	7	0	0	0	0	0	1	116	ID	
Acinetobacter baumannii	0	0	0	0	0	6	21	226	607	451	150	79	48	80	38	73	36	107	7	22	1929	2	1 - 4
Acinetobacter calcoaceticus	0	0	0	0	0	0	0	6	21	30	18	15	13	7	0	0	0	0	0	1	110	ID	
Actinobaculum schaalii	0	0	0	0	0	0	1	6	9	22	10	0	0	0	0	0	0	0	0	1	48	ID	
Alcaligenes faecalis	0	0	0	0	0	0	0	0	0	1	20	2	0	0	0	0	0	0	0	1	23	ID	
Bifidobacterium longum	0	0	0	0	0	0	0	0	0	0	0	2	8	15	21	9	5	1	1	1	62	ID	
Campylobacter coli	0	0	0	0	4	38	160	1550	4160	1911	175	4	2	1	5	99	0	0	0	50	8109	1	0.5 - 2
Campylobacter jejuni	0	0	0	2	33	141	1688	6829	14541	2346	108	12	5	0	5	62	0	0	1	41	25773	1	0.5 - 2
Citrobacter freundii	0	0	0	1	0	0	17	269	798	363	56	36	37	93	15	55	1	0	32	7	1773	2	1 - 4
Citrobacter koseri	0	0	0	0	0	2	8	42	145	58	12	2	3	1	1	0	1	0	0	3	275	(2)	0.5 - 4
Enterobacter agglomerans	0	0	0	0	0	1	5	43	5	0	0	0	0	0	0	0	0	0	0	1	54	ID	
Enterobacter cloacae	1	0	1	3	26	125	324	1202	3647	1384	160	87	124	92	19	103	50	52	60	38	7460	2	0.25 - 2
Enterococcus faecalis	0	0	0	0	0	2	2	7	18	37	73	156	880	1761	704	61	57	69	686	25	4513	64	32 - 128
Enterococcus faecium	0	0	0	0	0	0	0	1	5	14	83	555	1147	575	93	13	11	46	320	24	2863	32	16 - 64
	0.002	0.004	0.008	0.016	0.03	0.06	0.125	0.25	0.5	1	2	4	8	16	32	64	128	256	512	Distributions	Observations	(T)ECOFF	Confidence interval
Enterococcus hirae	0	0	0	0	0	0	0	0	1	4	50	98	234	151	28	0	0	0	2	8	568	32	4 - 32
Escherichia coli	0	0	4	6	26	49	525	8521	32063	21068	4423	1213	1399	3728	6362	512	138	167	70	93	80274	2	1 - 2

EUCAST 2021

## The EUCAST MIC and Zone distribution database

# Gentamicin / Escherichia coli International MIC distribution - Reference database 2021-08-27 Based on aggregated distributions

MIC distributions include collated data from multiple sources, geographical areas and time periods and can never be used to infer rates of resistance

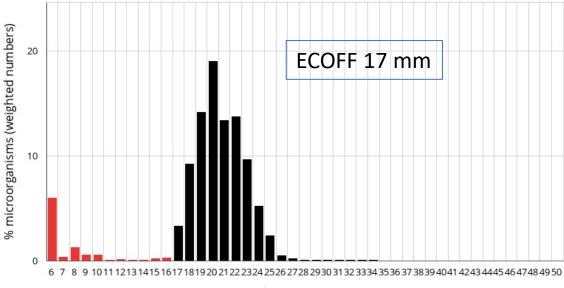


MIC Epidemiological cut-off (ECOFF): 2 mg/L Wildtype (WT) organisms: ≤ 2 mg/L

Confidence interval: 1 - 2 80274 observations (93 data sources) Gentamicin / Escherichia coli International zone diameter distribution - Reference database 2021-08-27 EUCAST disk diffusion method

#### Based on aggregated distributions where each distribution has equal weight \*

Distributions include collated data from multiple sources, geographical areas and time periods and can never be used to infer rates of resistance



Zone diameter (mm)

Disk content: 10 Epidemiological cut-off (ECOFF): 17 mm Wildtype (WT) organisms: ≥ 17 mm

Confidence interval: -29120 observations (15 data sources)

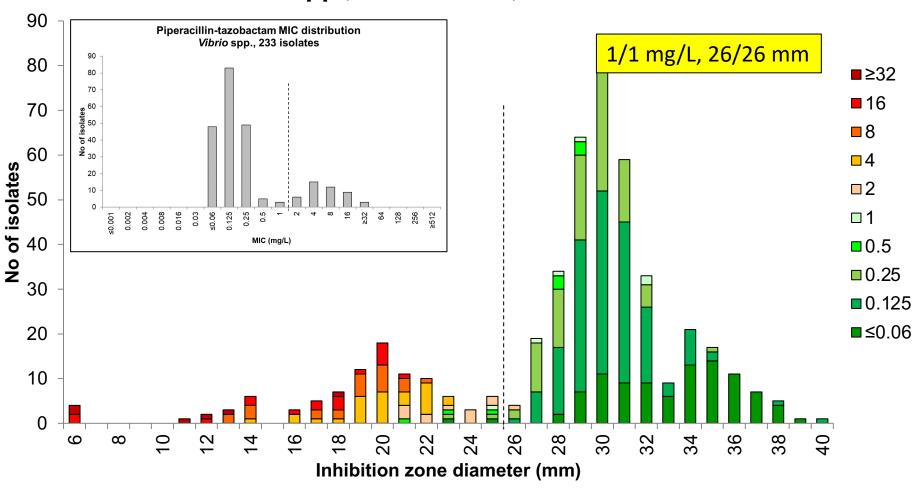
Available at www.eucast.org

<sup>\*</sup> individual distributions were converted to percentages of their individual total and then aggregated

## **EUCAST** projekt

- New agents
  - Cefiderocol, tedizolid, delafloxacin
  - Betalactam betalactamase inhibitor agents and methods (meropenem-vaborbactam, imipenem-relebactam, ....)
    - Disk diffusion f\u00f6r meropenem-vaborbactam i brytpunktstabell version 12.0 (2022)
- Species, methods and breakpoints
  - Achromobacter xylosoxidans
  - Vibrio cholerae, V. parahemoyticus, V. vulnificus, V. alginolyticus, V. fluvialis.
  - Revised breakpoints and new disk diffusion methodology for Anaerobic bacteria
    - FAA, 16-20h, 5 fast-growing anaerobic bacteria (*Bacteroides spp, Prevotella, F.necrophorum, Cl.perfringens, Cut.acnes*).
  - Nocardia spp (in collaboration with France and Australia) breakpoints and methods during 2022/23

# Piperacillin-tazobactam 30-6 μg vs. MIC *Vibrio* spp., 233 isolates, 466 correlates



For species not listed below, see EUCAST Guidance document on how to test and interpret results when there are no breakpoints Expert Rules and Intrinsic Resistance Tables

MIC determination (agar dilution)

Medium: Fastidious Anaerobe Agar (FAA)

Inoculum: 105 CFU/spot

Incubation: Anaerobic environment, 35-37°C, 48h

Reading: Unless otherwise stated, read MICs at the lowest concentration of the agent where a noticeable

difference is seen in visible growth between the test and control plate.

Quality control: Bacteroides fragilis ATCC 25285 and Clostridium perfringens ATCC 13124.

Clostridium perfringens DSM 25589 with a metronidazole 5 µg disk to monitor the anaerobic atmosphere.

Disk diffusion (EUCAST standardised disk diffusion method)

**Medium:** Fastidious Anaerobe Agar (FAA). The plates should be dried prior to inoculation (at 20-25°C overnight or at

35°C, with the lid removed, for 15 min).

Inoculum: McFarland 1.0

Incubation: Anaerobic environment, 35-37°C, 18±2h

Reading: Unless otherwise stated, read zone edges as the point showing no growth viewed from the front of the plate with the lid removed and with reflected light. See pictures below and the EUCAST Reading Guide for disk duffsion of

anaerobic bacteria for further information.

Quality control: Bacteroides fragilis ATCC 25285 and Clostridium perfringens ATCC 13124.

Clostridium perfringens DSM 25589 with a metronidazole 5 µg disk to monitor the anaerobic atmosphere.

#### Bacteroides spp.

Antimicrobial agent	MIC	breakpo	ints	Disk	Zone diameter		ter	Notes						
		(mg/L)		content	breakpoints (mm)		breakpoints (mm)		breakpoints (mm)		breakpoints (mm)		mm)	Numbered notes relate to general comments and/or MIC breakpoints.
	S≤	R>	ATU	(µg)	S≥	R <	ATU	Lettered notes relate to the disk diffusion method.						
Piperacillin-tazobactam	8 <sup>1</sup>	8 <sup>1</sup>		30-6	20	20		1. For susceptibility testing purposes, the concentration of tazobactam is fixed at 4 mg/L.						
Piperacillin-tazobactam, B. thetaiotaomicron	IE	ΙE			IE	IE		2/A. The meropenem zone diameter breakpoint will detect all cfiA gene mediated carbapenem resistance in Bacteroides						
Meropenem	1 <sup>2</sup>	1 <sup>2</sup>		10	28 <sup>A</sup>	28 <sup>A</sup>		fragilis. Some isolates with an MIC of 1 mg/L may harbour the cfiA gene.						
Clindamycin	(4) <sup>3</sup>	(4) <sup>3</sup>		2	(10) <sup>B,C</sup>	(10) <sup>B,C</sup>		3/B. For explanation of breakpoints in brackets, see the Notes sheet at the beginning of the table.						
Metronidazole	4	4		5	25	25		C. Examine zones carefully for colonies within zones. Colonies should be taken into account when reading.						

#### Prevotella spp.

Antimicrobial agent	MIC breakpoints		Disk	Zone diameter			Notes					
		(mg/L)		content	breakpoints (mm)		(mm)	Numbered notes relate to general comments and/or MIC breakpoints.				
	S≤	R>	ATU	(µg)	S≥	R<	ATU	Lettered notes relate to the disk diffusion method.				
Benzylpenicillin	0.5	0.5		1 unit	20	20		1. For susceptibility testing purposes, the concentration of tazobactam is fixed at 4 mg/L.				
Piperacillin-tazobactam	0.5 <sup>1</sup>	0.5 <sup>1</sup>		30-6	26	26						
Meropenem	0.25	0.25		10	34	34		A. Examine zones carefully for colonies within zones. Colonies should be taken into account when reading.				
Clindamycin	0.25	0.25		2	31 <sup>A</sup>	31 <sup>A</sup>						
Metronidazole	4	4		5	22	22						

## ATU 2022

- Inga större förändringar 2022, men sannolikt utvecklas ATU ytterligare 2023.
- Vi arbetar på att göra ciprofloxacin/Enterobacterales mer robust
- Piperacillin-tazobaktam har nu bara en smal ATU på 16 mg/L (19 mm) vi får frågor på varför?

## CEFIDEROCOL ....as an example

You can report S and I results outside the ATU with confidence!

You can decide to perform an MIC on results inside the ATU and gain confidence before reporting a category S, I or R. You can decide to NOT report results inside the ATU.

You can decide to trust your method and ignore the ATU to report as tested.

