

İNFEKTİF ENDOKARDİTTE YENİ KANITLAR, YENİ YAKLAŞIMLAR

ULUSAL KILAVUZ GÜNCELLEMESİNE DOĞRU

18 EKİM 2025

Ankara Üniversitesi İbn-i Sina Hastanesi
Hasan Ali Yücel Salonu, Ankara



KLİMİK DERNEĞİ İNFEKTİF ENDOKARDİT VE DİĞER
KARDİYOYASKÜLER İNFEKSİYONLAR ÇALIŞMA GRUBU



İnfektif Endokarditte Son İki Yılda Ne Oldu?



Dr. Özlem Kurt Azap

Başkent Üniversitesi Tıp Fakültesi

Enfeksiyon Hastalıkları ve Klinik Mikrobiyoloji AD

The screenshot shows a Gmail interface with a sidebar on the left containing folders like 'Compose', 'Inbox', 'Sent', 'Spam', 'Trash', 'Purchases', 'Social', 'Updates', 'Forums', 'Promotions', and 'Labels'. The main area displays an email from 'Sanford Guide' with the subject '100% Human Written Clinical Guidance'. The email content includes the Sanford Guide logo and a promotional banner for the Sanford Guide app, which is shown on a smartphone. The app interface displays a search bar, a list of actions, and a table of antibiotic susceptibility data for various bacteria.

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İNFİKTİF ENDOKARDİT 2023'TE NELER DEĞİŞTİ?

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İNFİKTİF ENDOKARDİT: GÜNCEL BİLGİLER, YEREL VERİLER

21-22 EKİM 2017

Sağlık Bilimleri Üniversitesi
Dr. Siyami Ersek Göğüs, Kalp ve Damar Cerrahisi
Eğitim ve Araştırma Hastanesi
İstanbul



ESCMID

Postgraduate course Programme

**Update on Endocarditis and Endovascular
Infections**

Milan, Italy
6 - 8 November 2025



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Son İki Yılda (2023 Nisan-2025 Ağustos) Yayımlanan Dikkat Çekici İnfektif Endokardit Makaleleri



Dr. Özlem Kurt Azap

Başkent Üniversitesi Tıp Fakültesi

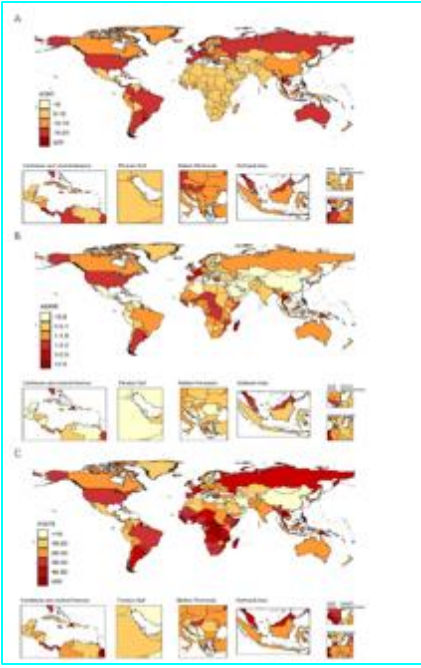
Enfeksiyon Hastalıkları ve Klinik Mikrobiyoloji AD

Başlıklar

- **Epidemiyolojik veriler:** Sıklık, mortalite, hastalık yükü
- **Tanı kriterleri/Tanı Yöntemleri:** Duke/ISCVİD tanı kriterleri
Mikrobiyolojik, Görüntüleme
- **Antibiyotik tedavisi:** Direnç Sorunu, Oral tedaviye geçiş
- **Cerrahi tedavi**
- **Profilaksi**
- **Endokardit ekibi**

Endokardit tanı kriterleri büyük bir değişim içinde 😊

- Epidemiyolojik verilerin değişmesi tanı kriterlerini etkiliyor
- Tanı kriterlerinin değişmesi epidemiyolojik verileri etkiliyor



Epidemiyolojik Veriler

İE Yüğü: 30 yıllık veri

Journal of Epidemiology and Global Health (2025) 15:69
https://doi.org/10.1007/s44197-025-00413-x

RESEARCH ARTICLE

Global Trends and Regional Differences in the Burden of Infective Endocarditis, 1990–2021: An Analysis of the Global Burden of Disease Study 2021

Huanhuan Miao¹ · Zhanyang Zhou¹ · Zheng Yin¹ · Xue Li¹ · Yuhui Zhang¹ · Yuqing Zhang¹ · Jian Zhang^{1,2}

Received: 11 February 2025 / Accepted: 23 April 2025
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Abstract

Background The study aimed to offer detailed insights into the global, regional, and national burden of IE in 2021, while also examining the temporal trends of IE from 1990 to 2021.

Methods Data on the absolute numbers and age-standardized rates (ASR) of incidence, deaths, and disability-adjusted life years (DALYs) related to IE were sourced from the Global Burden of Disease Study (GBD) 2021. The estimated annual percentage changes (EAPC) of ASR were calculated to quantify the temporal trends. Furthermore, joinpoint regression models were used to identify the temporal trends and the primary joinpoint year of ASR.

Results Globally, the age-standardized incidence rate (ASIR) for IE increased with an EAPC of 1.00 (95%CI: 0.93–1.08) from 9.35 per 100 000 population in 1990 to 12.61 per 100 000 population in 2021. Despite a rise in the absolute number of death cases and DALYs related to IE, the age-standardized mortality rate (ASMR) has remained stable (EAPC 0.06, 95%CI: -0.10–0.22), and the age-standardized DALYs rate (ASDR) has exhibited a decline (EAPC -0.34, 95%CI: -0.45–0.24) between 1990 and 2021. Males bore a higher burden of IE compared to females, with the peak burden gradually shifting towards older individuals. In 2021, the ASIR for IE exhibited an increase with the rise in socio-demographic index (SDI) quintiles, with the highest ASIR observed in the high SDI region (15.77 per 100 000 population). Moreover, the highest growth rates of ASIR, ASMR, and ASDR were also noted in the high SDI region. On the other hand, the ASMR (1.34 per 100 000 population) and ASDR (40.71 per 100 000 population) for IE were relatively high in the low SDI region. Joinpoint analysis demonstrated that the ASIR, ASMR, and ASDR did not experience any sudden surges either globally or across different SDI regions after 2007.

Conclusions The burden of IE remained relatively high, characterized by a rising ASIR and a stable ASMR on a global scale. This burden was notably prominent among males, the elderly, and in the high and low SDI regions. Region-specific prevention and management strategies might be warranted to reduce the burden of IE.



IHME = Institute for Health Metrics and Evaluation
(Sağlık Ölçümleri ve Değerlendirme Enstitüsü)

- 1990–2021 yılları arasında İE yükündeki değişimleri cinsiyet, yaş grubu, SDI bölgesi, coğrafi bölge ve ülke bazında incelemiştir
- Erkeklerde İE yükü, kadınlara göre daha yüksek bulunmuştur
- Zamanla, en yüksek hastalık yükü ileri yaş gruplarına kaymıştır



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Huanhuan Miao¹, Zhenyang Zhou², Zheng Yin², Xue Li², Yuhui Zhang², Yujing Zhang², Jian Zhang^{1,2}

Terimler önemli!

Abbreviations

APC	Annual percent change
AAPC	Average annual percentage change
ASDR	Age-standardized disability-adjusted life years rate
ASIR	Age-standardized incidence rate
ASMR	Age-standardized mortality rate
ASR	Age-standardized rates
CDRIE	Intracardiac device-related infective endocarditis
CI	Confidence interval
CODEm	Cause of Death Ensemble model
DALYs	Disability-adjusted life years
DisMod-MR	Disease Modelling Meta-Regression
EAPC	Estimated annual percentage change
GATHER	Guidelines for Accurate and Transparent Health Estimates Reporting
GBD	Global Burden of Diseases, Injuries, and Risk Factors Study
ICD	International Classification of Disease
IE	Infective endocarditis
NVE	Native valve endocarditis
PVE	Prosthetic valve endocarditis
SDI	Socio-demographic index
ST-GPR	Spatiotemporal Gaussian process regression
UI	Uncertainty interval

ASDR: yaşa göre standardize edilmiş engellilik yükü oranı

ASIR: yaşa göre standardize edilmiş insidans hızı

ASMR: yaşa göre standardize edilmiş mortalite hızı

DALY: engellilikle kaybedilen yaşam yılı

Dünyada İE- 2021

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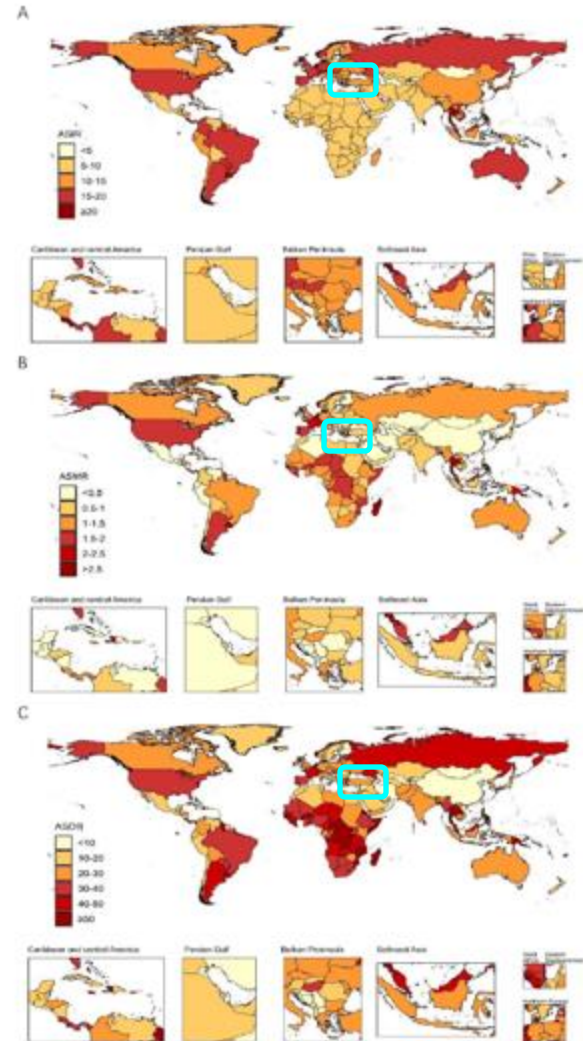
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Conclusions The burden of IE remained relatively high, characterized by a rising ASIR and a stable ASMR on a global scale. This burden was notably prominent among males, the elderly, and in the high and low SDI regions. Region-specific prevention and management strategies might be warranted to reduce the burden of IE.

Fig. 3 Age-standardized incidence (A), mortality (B), and DALYs (C) rates for IE in 204 countries and territories, for both sexes, 2021. ASDR, age-standardized disability-adjusted life year rate; ASIR, age-standardized incidence rate; ASMR, age-standardized mortality rate; DALYs, disability-adjusted life years



İE Yüğü: 30 yıllık veri

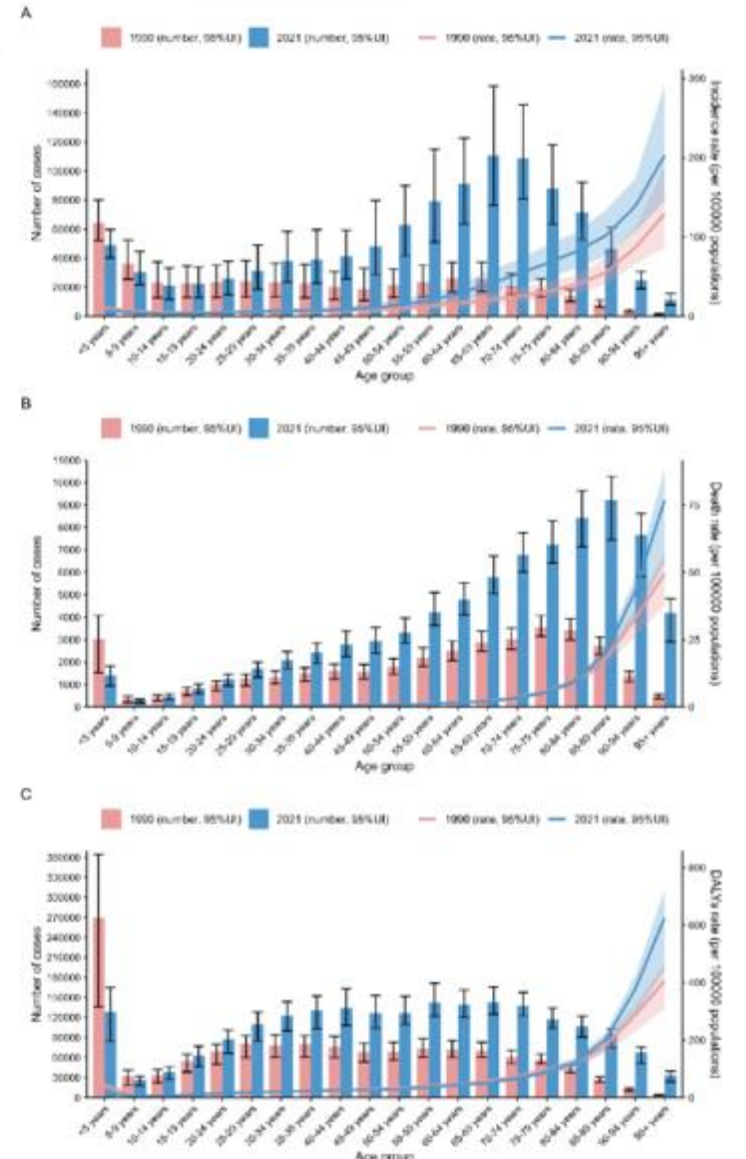
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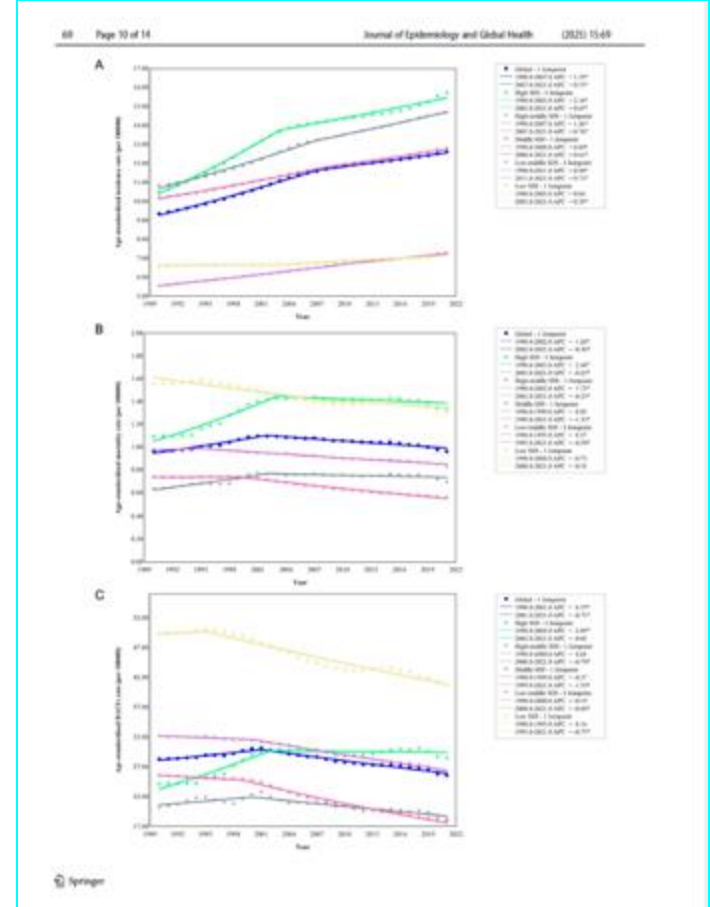
Fig. 1 Age-specific numbers and rates of incidence (A), deaths (B), and DALYs (C) for IE at the global level in 1990 and 2021. DALYs, disability-adjusted life years; UI, uncertainty interval



- Hem **yeni olgu sayısı** hem de yaşa göre standardize edilmiş insidans hızı (**ASIR**) **artış** göstermiştir
- Bu **artışın**;
 - nüfus artışı
 - yaşlanan toplum yapısı
 - invaziv girişimlerin artması ve tanı yöntemlerindeki ilerlemelerle ilişkili olduğu düşünülmektedir

Dünyada İE- 30 yıl

- Mutlak ölüm sayısı ve engellilikle kaybedilen yaşam yılı (DALY) artsa da yaşa göre standardize edilmiş mortalite hızı (ASMR) büyük ölçüde sabit kalmış, yaşa göre standardize edilmiş engellilik yükü oranı (ASDR) ise azalma göstermiştir
- Bu **olumlu eğilim**;
yeni nesil antibiyotiklerin kullanıma girmesi,
cerrahi tedavi endikasyonu olan olgularda cerrahiye yönelimin artması
multidisipliner “**Endokardit Ekipleri**”nin kurulmasıyla açıklanabilir





Son 30 yılda İE...

- ASIR (Yaşa göre standardize edilmiş insidans oranı) SDI düzeyi arttıkça yükselmiştir
Yüksek SDI bölgelerinde daha yüksek İE insidansı
- ASMR (Yaşa göre standardize edilmiş mortalite oranı) ve ASDR (Yaşa göre standardize edilmiş özürlülüğe göre ayarlanmış yaşam yılı kaybı)
düşük SDI bölgelerinde görece daha **yüksek** bulunmuştur



Son 30 yılda İE...

Bu veriler, küresel sağlık eşitsizliklerini ve bakım kalitesindeki farklılıkları göstermektedir

Politika yapıcılar:

- Bölgesel farklılıkları tanımlamalı
- Yerel epidemiyolojik özellikleri ve risk faktörlerini incelemeli
- Bölgeye özgü önleme ve yönetim stratejileri geliştirmelidir

Japonya: 2016–2021 Ulusal Kohort Verileri

Journal of the American Heart Association

ORIGINAL RESEARCH

Trends in the Clinical Characteristics and Outcome of Infective Endocarditis: A Nationwide Study From 2016 to 2021

Hiroshi Kamoto, MD, PhD; Takeshi Nishi, MD, PhD; Kyo Kamisaka, MD; Yoshitaka Sasahira, MD; Koshiro Kanacka, MD, PhD; Yoko Sumita, MD, PhD; Yoshitaka Iwanaga, MD, PhD; Chisato Izumi, MD, PhD; Shiro Uemura, MD, PhD

BACKGROUND: Infective endocarditis (IE) is still a fatal disease, and given its rarity, ongoing updates to patient characteristics and outcomes of IE are essential for providing precision diagnoses and effective treatments. This study sought to examine temporal trends in the clinical characteristics and in-hospital occurrence of adverse outcomes of IE.

METHODS AND RESULTS: Using the Japan nationwide administrative database, we identified patients with IE in Japan from 2016 to 2021. A total of 17 407 patients with IE (37.8% women; median age, 72 [interquartile range, 59–81] years) were identified. The incidence of IE increased from 2.02 per 100 000 population in 2016 to 2.59 per 100 000 population in 2021. The median age of the patients increased from 70 years in 2016 to 73 years in 2021, and patient backgrounds were becoming more complex. The in-hospital mortality rate was 14.5%, which significantly increased from 14.1% in 2016 to 15.4% in 2021. Higher age (odds ratio [OR], 1.34 [95% CI, 1.29–1.40]), heart failure (OR, 1.37 [95% CI, 1.18–1.59]), cerebrovascular complications (OR, 1.78 [95% CI, 1.59–1.99]), renal failure (OR, 2.36 [95% CI, 2.06–2.70]), and sepsis (OR, 2.79 [95% CI, 2.48–3.13]), were independently associated with an increased in-hospital mortality rate. In contrast, cardiac surgery (OR, 0.47 [95% CI, 0.41–0.55]), and a higher number of total cardiac surgeries (OR, 0.77 [95% CI, 0.65–0.92]) independently predicted a lower in-hospital mortality rate.

- **17 bin 407 İE olgusu**
- **insidans:**
2016'da 100.000'de **2,02** → 2021'de **2,59**
- **Ortanca yaş: 70'ten 73'e yükseldi; hastalar giderek yaşlı ve daha kompleks hale geldi**
- **Hastane içi mortalite: %14,1 → %15,4'e arttı**

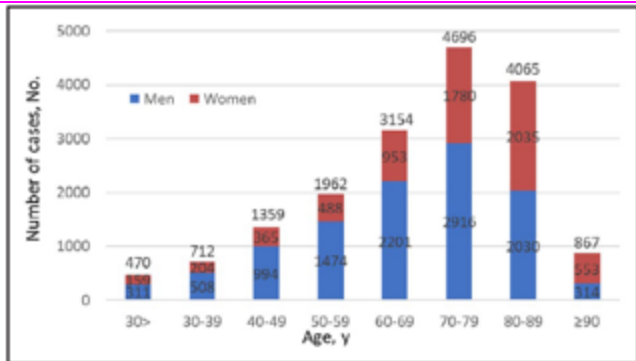


Figure 3. Age distribution of patients with IE. IE indicates infective endocarditis.

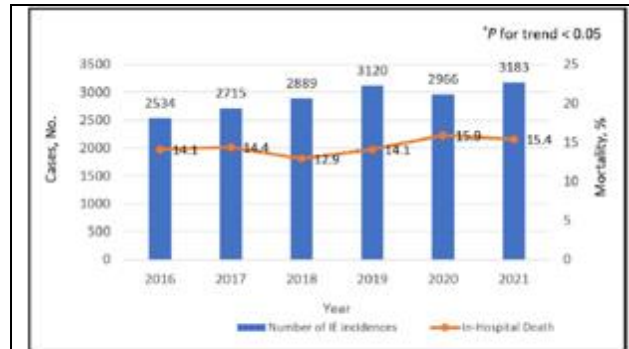


Figure 2. Number of IE cases and in-hospital death with IE from 2016 to 2021.

Japonya: 2016–2021 Ulusal Kohort Verileri

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Mortaliteyi Artıran Faktörler

- İleri yaş
- Kalp yetmezliği
- Serebrovasküler komplikasyonlar
- Böbrek yetmezliği
- Sepsis

Mortaliteyi Azaltan Faktörler

- Kardiyak cerrahi uygulanması
- Cerrahi hacmi yüksek merkezlerde tedavi

Türkiye’de neler değişti?

Tablo 1. Türkiye’de 1974-2017 Yılları Arasında İzlenmiş Enfektif Endokardit Olgularını İçeren Çalışmaların Özellikleri

Yazarlar	Dönem	Şehir	Olgu Sayısı	Yaş*	Erkek (%)	ARA (%)	Yapay Kapak (%)	NDU (%)	CIED (%)	Diyaliz (%)	Stafilokok (%)	Staphylococcus aureus (%)	MRSA (%)	Streptokok (%)	Enterokok (%)	Brucella spp. (%)	Kan Kültürü Negatif (%)	Ameliyat (%)	Nozokomiyal (%)	Kesin İET (%)	Mortalite (%)
Toplam	1974-2017	Türkiye	2712	47	61	37	27	2	7	9	31	21	5	19	9	7	37	50	25	90	24
Kaya et al. (47)	2005-2012	Şanlıurfa	24	41	50	13			4	8.3	21			17	4	33	21	37			8
İnanç et al. (49)	2006-2007	Kayseri	27	47	63	67	41				33	11		4	11	4	45	19			15
Çaylan et al. (48)	1997-2001	Trabzon	32	47	73	22					37	22		19	9		78	25			41
Zencir et al. (33)	2000-2015	Aydın	59	59	43	32						39						27		100	27
Tuğcu et al. (34)	2000-2007	İstanbul	68	51	59	40	56		3	12	41	28	4	25	1.5	0	21	59	28	84	25
Sucu et al. (35)	1997-2007	Gaziantep	72	45	57	36	29			3	27	10		23	4	4	36	28		100	15
Heper ve Yörükoglu (36)	1995-1999	Ankara	74	25	66	66	14				44	21		29	15	7	11				18
Cancan-Gursul et al. (32)	2006-2013	İzmir	80	51	70	12	0	22.5	7.5	49	27	2.5	20	17	12	51	56				15
Agca et al. (37)	2009-2014	Bursa	85	52	56	22	40		13		27	8	2	11	5	6	32	27		100	36
Taşbakan et al. (38)	2007-2017	İzmir	100	52	65	13	3	42.5				18		32	11		26				14
Erbay et al. (39)	2002-2008	Ankara	107	45	71		44										34				27
Akıl et al. (40)	2005-2012	Çok Merkezli	112	45	50	18	17	8		5.4	50	35		29	16		16	13		90	29
Şimşek-Yavuz et al. (50)	2015-2016	Çok Merkezli	112	55	70	12	33	4	8	13	31	15	3.5	22	8	1	29.5	56	35	77	28
Adademir et al. (41)	1997-2004	İstanbul	119	39	69	39	20		2		24	22	11	9	3	3	57	76		100	
Çay et al. (42)	2008-2011	Ankara	121	55	53	14	42		10	18	33	16		21	7	6	29	38		100	24
Turak et al. (43)	1997-2012	Ankara	157	59	55	15	42		9		30	13		22	7	6	29	39		100	26
Leblebicioğlu et al. (30)	2002-2012	Çok Merkezli	158	47	51	39	0				28		6	11	11	6	34	51			31
Özveren et al. (44)	1985-2011	İstanbul	174	39	79		15				15		2	29		6	47	100			16
Kocabaş et al. (45)	2009-2012	İzmir	194	48	66	22	12		3.1	6	30			16	6		43	52		100	20
Cetinkaya et al. (29)	1974-1999	Ankara	228	36	55	64	20	0	0.4		27	24		14	6	5.7	50	37	12	53	23
Elbey et al. (31)	2005-2012	Çok Merkezli	248	47	55	28	30		6		29	21	6	11	11	5	37.5	54		100	33
Şimşek-Yavuz et al. (4)	2004-2015	İstanbul	325	47	58	34	44	1	5.6	3	36	20	2	19	7	5	22	52	23	86	27
Toplam	1974-2017	Türkiye	2712	47	61	37	27	2	7	9	31	21	5	19	9	7	37	50	25	90	24

*Ortalama. *Modifiye Duke ölçütleriyle. ARA: akut romatizmal ateş. NDU: damar içi ilaç kullanıcısı. CIED: kardiyak implante edilebilir elektronik cihaz. MRSA: metisiline dirençli Staphylococcus aureus, IE: enfektif endokardit.

Klimik Derg. 2019; 32(Suppl. 1): 2-116



Results

- ✓ 745 patients (n=600 definite, n=145 possible IE).
- ✓ The median age of cohort was **56 years (18-91)**
- ✓ 460 (61.7%) patients were male.
- ✓ The most common predisposing heart conditions were native valve disease (40.8%)
 - Degenerative (20%); congenital heart disease (12.2%), rheumatic valve disease (8.6%),
- ✓ Previous valve surgery (30.9%), previous IE episode (5.1%).
- ✓ The median duration of illness prior to admission was 15(1-365) days.
- ✓ Most of the patients (65.8%) had native valve IE
- ✓ Most episodes were left-sided
 - Mitral 272 (36.5%), aortic 244 (32.8%).
- ✓ The causative microorganisms were identified by blood cultures in 549(73.7%) cases.
- ✓ The most common causative microorganisms;
 - Staphylococcus aureus (23.6%),
 - Streptococcus spp (13.4%)
 - Coagulase negative Staphylococcus spp. (13.3%)
 - Enterococcus spp (12.5%).
- ✓ The changing epidemiology of causative agents according to years were shown in Figure 1.
- ✓ During this period, 286(39%) patients underwent valve surgery.
- ✓ The in hospital mortality of IE was 27.4% .

Türkiye: 2013–2023 Ulusal Kohort Verileri

European Journal of Clinical Microbiology & Infectious Diseases (2025) 44:1325–1333
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RESEARCH

Epidemiological, clinical and microbiological aspects of infective endocarditis in Türkiye

Elif M. Saricaoglu¹ · Seniha Başaran² · Derya Seyman³ · Merve Arslan⁴ · Serpil Özkan-Oztürk⁵ · Yasemin Tezer-Takca⁶ · Yeşim Uygun-Kırmaz⁷ · Nuran Sarı⁸ · Danel Berzag-Dantzi⁹ · Alpay Azap¹ · Serap Simsek-Yavuz² · Özlem Kurt-Azap⁹ · Türkiye Endocarditis Group (TEG) Investigators

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Abstract

Purpose Infective endocarditis (IE) is an evolving disease with a shifting epidemiology and disease burden over time. This study aimed to compare the epidemiological and clinical aspects of IE over three time periods across eleven years.

Methods This was a retrospective cohort, multicenter study conducted in Türkiye, comparing three periods: 2013–2016, 2017–2020, and 2021–2023. Epidemiological and microbiological characteristics, as well as patient outcomes, were analyzed and compared across these periods.

Results A total of 1,044 patients diagnosed with IE were included. The median (IQR) age was 57 (44–68) years, with an increasing pattern ($p < 0.001$). Throughout the study period, the prevalence of intracardiac devices increased, whereas the prevalence of degenerative and congenital heart diseases declined. Among all patients, the most frequently identified pathogens were staphylococci (36.4%), followed by streptococci (14.0%) and enterococci (11.9%). Throughout the three periods, there was a significant increase in staphylococci, with *S. aureus* emerging as the predominant pathogen in all type IE. The in-hospital mortality rate among all patients was 22.5%. Independent risk factors for in-hospital mortality included ≥ 65 age (OR=1.9), chronic kidney disease (OR=1.9), nosocomial acquisition (OR=2.1), *Candida* spp. infection (OR=2.9), prosthetic valve IE (OR=1.9), vegetation size > 15 mm (OR=1.6), and central nervous system emboli (OR=2).

Conclusion The epidemiology of IE is undergoing significant changes, leading to shifts in microbiological profiles and clinical presentations. Effective management of IE should be guided by established clinical guidelines while integrating up-to-date epidemiological data to ensure comprehensive and evidence-based patient care.

Keywords Epidemiology · Infective endocarditis · Mortality · Türkiye

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Table 1 Epidemiological and clinical characteristics for infective endocarditis cohort stratified by three time periods

	Total (n=1044)	2013–2016 (n=205)	2017–2020 (n=491)	2021–2023 (n=348)	p value	Post-hoc analysis
Demographics						
Age in years	57(44.68)	54 (40.65)	55(43.66)	60(47.70)	<0.001	1=2, 2<3
18–39 y	202(19.3)	48(23.4)	98(20)	56(16.1)	<0.001	1,2<3
40–64 y	517(49.6)	104(50.7)	256(52.1)	157(45.1)		
≥65 y	325(31.1)	53(25.9)	137(27.9)	135(38.8)		
Male sex	666(63.8)	133(64.9)	310(63.1)	223(64.1)	0.901	
Comorbidities						
Hypertension	382(36.6)	66(32.2)	157(32)	159(45.7)	<0.001	1=2, 2<3
Diabetes mellitus	282(27)	39(19)	114(23.2)	129(37.1)	<0.001	1,2<3
Chronic kidney disease	220(21.1)	40(19.5)	97(19.8)	83(23.9)	0.297	
Malignancy	57(5.5)	9(4.4)	19(3.9)	29(8.3)	0.015	1=2, 2<3
Autoimmune disease	44(4.2)	3(1.5)	23(4.7)	18(5.2)	0.086	
Predisposition conditions						
Prosthetic valve	323(30.9)	59(28.8)	166(33.8)	98(28.2)	0.166	
Degenerative heart disease	183(17.5)	48(23.4)	89(18.1)	46(13.2)	0.009	1,2>3
Hemodialysis	169(16.2)	27(13.2)	84(17.1)	58(16.7)	0.419	
Intracardiac device	102(9.8)	15(7.3)	39(7.9)	48(13.8)	<0.001	1=2<3
Rheumatic fever	87(8.3)	23(11.2)	39(7.9)	25(7.2)	0.060	
Bicuspid aortic valve	64(6.1)	16(7.8)	35(7.1)	13(3.7)	0.070	
Congenital heart disease	60(5.7)	17(8.3)	32(6.5)	11(3.2)	0.026	1,2>3
Previous IE episode	52(5)	14(6.8)	22(4.5)	16(4.6)	0.397	
Intravenous drug user	47(4.5)	6(2.9)	28(5.7)	13(3.7)	0.192	
Type of acquisition						
Community acquired	619(59.3)	133(64.9)	269(54.8)	217(62.4)	0.709	
Nosocomial	157(15)	29(14.1)	72(14.7)	56(16.1)	0.010	1=2, 2<3
Non-nosocomial healthcare-associated infection	268(25.7)	43(21)	150(30.5)	75(21.6)	0.484	
Endocarditis site						
Mitral	469(44.9)	90(43.9)	217(44.2)	162(46.6)	0.510	
Aortic	402(38.5)	102(49.8)	203(41.3)	97(27.9)	<0.001	1,2>3
Tricuspid	111(10.6)	15(7.3)	51(10.4)	45(12.9)	0.035	1,2<3
Pulmonary	13(1.3)	3(1.5)	7(1.4)	3(0.1)	0.561	
Lead	70(6.7)	8(3.9)	28(5.7)	34(9.8)	0.011	1,2<3
Endocard	21(2)	4(2)	8(1.6)	9(2.6)	0.189	
Type of IE						
Native-valve IE	659(63.1)	136(66.4)	303(61.7)	221(63.2)	0.482	
Prosthetic-valve IE	307(29.4)	59(28.7)	156(31.8)	93(26.7)	0.266	
CIED-IE	78(7.5)	10(4.9)	32(6.5)	35(10.1)	0.048	1=2, 2<3
Complications						
Cranial embolism	203(19.4)	37(18)	106(21.6)	60(17.2)	0.250	
Splenic infarct/abscess	23(2.2)	2(1)	19(3.9)	2(0.6)	0.353	
Glomerulonephritis	29(2.8)	1(0.5)	21(4.3)	7(2)	0.593	
Pulmonary embolism	77(7.4)	2(1)	32(6.5)	43(12.4)	<0.001	1,2<3
Spondylodiscitis	24(2.3)	1(0.5)	7(1.4)	16(4.6)	0.002	1=2, 2<3
Heart failure	120(11.5)	10(4.9)	52(10.6)	58(16.7)	<0.001	1,2<3
Cardiac surgery for IE	406(38.8)	102(49.8)	196(39.9)	108(31.1)	<0.001	1,2>3
In-hospital mortality	233(22.3)	38(18.5)	112(22.8)	83(24.4)	0.271	

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Table 3 Microbiological differences between three time periods of overall cohorts

	Total (n = 1044)	2013–2016 (n = 205)	2017–2020 (n = 491)	2021–2023 (n = 348)	p value
Culture negative IE	281(26.9)	67(32.7)	136(27.7)	78(22.4)	0.027
<i>Staphylococcus</i> spp.	380(36.4)	57(27.8)	180(36.7)	143(41.1)	0.007
<i>S. aureus</i>	245(23.5)	32(15.6)	116(23.6)	97(27.9)	0.004
Methicillin resistance	72(29.4)	14(43.8)	29(25)	29(29.9)	0.185
CoNS	135(12.9)	25(12.2)	64(13.1)	46(13.2)	0.962
Methicillin resistance	97(71.9)	24(96)	45(70.3)	28(60.9)	0.052
<i>Streptococcus</i> spp.	146(14)	35(17.1)	60(12.2)	51(14.7)	0.220
<i>Enterococcus</i> spp.	124(11.9)	21(10.2)	68(13.8)	35(10)	0.129
Enterobacterales	24(2.3)	8(3.9)	14(2.9)	12(3.4)	0.659
Non-fermentatives	18(1.7)	4(2)	11(2.2)	3(0.9)	0.252
<i>Candida</i> spp.	27(2.6)	5(2.4)	8(1.6)	14(4)	0.134
<i>Granulicatella</i> and <i>Abiotrophia</i> spp.	11(1.1)	2(1)	6(1.2)	3(0.9)	0.574
<i>Corynebacterium</i> spp.	10(1)	3(1.5)	5(1)	2(0.6)	0.252
<i>Coxiella</i> spp.	9(0.9)	5(2.4)	3(0.6)	2(0.6)	0.688
<i>Brucella</i> spp.	7(0.7)	2(1)	1(0.2)	4(1.1)	0.665
<i>Bartonella</i> spp.	1(0.1)	-	-	1(0.3)	0.782
HACEK	2(0.2)	1(0.5)	1(0.2)	0(0)	0.594



Tanı Kriterleri / Tanı Yöntemleri



Tanı kriterlerinin «öyküsü»

Year	Diagnostic Criteria	Improvement
1981	Von Reyn	1. Definite, Probable, Possible, Rejected
1994	Duke Criteria	1. Major & Minor Criteria 2. Clinical Definite Criteria 3. Add Echocardiography
2000	Modified Duke Criteria	1. Remove "Possible" Echo 2. Expand <i>S. aureus</i> inclusion
2015	ESC Criteria	1. Add PET-CT, Cardiac CT
2023	Duke ISCVI Criteria	1. Expand Microbiology - <i>E. faecalis</i> , most <i>Streptococci</i> - Add "Typical" PVE Pathogens 2. Add Molecular Diagnostics - 16S rRNA PCR, FISHseq from valve - next generation sequencing from blood 3. Add Surgical Major Criteria 4. Remove Blood culture Restrictions - Timing & Separate Venipunctures



Vance Fowler
United States

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Vance Fowler
United States

25 Ağustos 2023- Avrupa Kardioloji Derneği Rehberi

2023 Duke-ISCVID Endokardit Kriterleri



2023

Duke ISCVID Criteria

1. Expand Microbiology

- *E. faecalis*, most *Streptococci*
- Add "Typical" PVE Pathogens

2. Add Molecular Diagnostics

- 16S rRNA PCR, FISHseq from valve
- next generation sequencing from blood

3. Add Surgical Major Criteria

- Remove Blood Culture Restrictions
- Timing & Separate Venipunctures

Clinical Infectious Diseases

VIEWPOINTS

The 2023 Duke-International Society for Cardiovascular Infectious Diseases Criteria for Infective Endocarditis: Updating the Modified Duke Criteria

Vance G. Fowler Jr.,^{1,2,3} David T. Durack,⁴ Christine Seifried-Sady,⁵ Eugene Athan,⁶ Arnold S. Bayer,^{7,8} Anna Lisa Chanen,⁹ Anders Dahl,¹⁰ Louis DiLandro,¹¹ Emanuele Durante-Mangoni,¹² Xavier Dvorak,¹³ Claudia Garcia-Forbes,¹⁴ Emil Fouad,¹⁵ Margaret M. Hansen,¹⁶ Barbara Kozak,¹⁷ Bruno Rios,¹⁸ Adolf W. Karchhous,¹⁹ Carlos A. Morones,²⁰ Gaby A. Pelt,²¹ Maria Nazzareno Picot,²² Stephen D. Prokter,²³ Albert Rojas,²⁴ Francesco Vendemmiere,²⁵ Jan T. M. van der Meer,²⁶ Thomas W. van de Ven,²⁷ and Jose M. Muso²⁸

OBJECTIVE: Infective endocarditis (IE) is a life-threatening condition. The 2023 Duke-International Society for Cardiovascular Infectious Diseases (ISCVID) criteria for IE were updated to reflect advances in microbiology, epidemiology, and treatment. The 2023 Duke-ISCVID IE criteria propose significant changes, including new microbiology diagnostics (enzyme immunoassay for *Bartonella* species, polymerase chain reaction, amplicon/metagenomic sequencing, in situ hybridization), imaging (positron emission computed tomography with 18F-fluorodeoxyglucose, cardiac computed tomography), and inclusion of intraoperative inspection as a new Major Clinical Criterion. The list of "typical" microorganisms causing IE was expanded and includes pathogens to be considered as typical only in the presence of intracardiac prostheses. The requirements for timing and separate venipunctures for blood cultures were removed. Last, additional predisposing conditions (transcatheter valve implants, endovascular cardiac implantable electronic devices, prior IE) were clarified. These diagnostic criteria should be updated periodically by making the Duke-ISCVID Criteria available online as a "Living Document."

KEYWORDS: endocarditis; Duke Criteria; PET/CT; echocardiography; ISCVID.

The Duke Criteria for diagnosis of infective endocarditis (IE) were originally published in 1994 [1] and modified in 2000 [2]. Their primary purpose was to serve as a research tool to standardize the definition of a clinically proven condition. Their presence paved the way for a steady stream of institutional investigations [3–7] that transformed our understanding

of the disease. However, the microbiology, diagnostics, epidemiology, and treatment of IE have changed significantly since the debut of these criteria. For example, endovascular cardiac implantable electronic devices (CIEDs), including permanent pacemakers and cardioverter-defibrillators, are now present in at least 10% of contemporary IE case series [8], and constitute a significant risk factor for infection [8, 9]. Transcatheter-implanted valves are infected at rates comparable to surgically implanted valves, and are an increasing component of prosthetic valve endocarditis (PVE). In 2015, the European Society of Cardiology [10] proposed changes to the Modified Duke Criteria; however, recent advances require further modifications of the formal diagnostic criteria for IE.

In response to this need, in 2023, the International Society for Cardiovascular Infectious Diseases (ISCVID) convened a Working Group of 25 subject matter experts from 5 continents

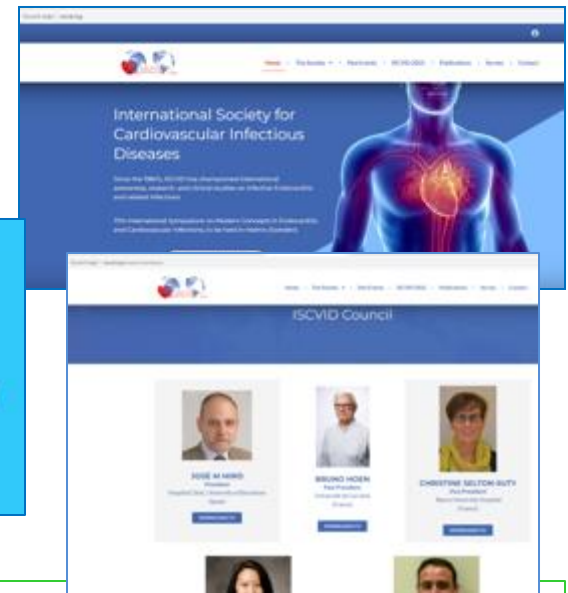


Table 3. Updates to Modified Duke Criteria Proposed by 2023 Duke-International Society for Cardiovascular Infectious Diseases Infective Endocarditis (IE) Criteria

CRITERIA	Change
PATHOLOGIC CRITERIA	
Microorganism identification	Microorganisms identified in appropriate sample by PCR, amplicon or metagenomic sequencing, or in situ hybridization
MAJOR CLINICAL CRITERIA	
Microbiology	
Blood cultures	Removed requirements for timing and separate venipunctures for blood cultures.
Definition of typical organisms	Added typical pathogens: 1) <i>S. lugdunensis</i> ; <i>E. faecalis</i> ; all streptococci except <i>S. pneumoniae</i> and <i>S. pyogenes</i> ; <i>Granulicatella</i> spp.; <i>Abiotrophia</i> spp.; and <i>Gemella</i> spp. 2) Organisms to be considered "typical" IE pathogens in the setting of intracardiac prosthetic material: coagulase negative staphylococci, <i>Corynebacterium striatum</i> , <i>C. jeikeium</i> , <i>Serratia marcescens</i> , <i>Pseudomonas aeruginosa</i> , <i>Cutibacterium acnes</i> , nontuberculous mycobacteria, and <i>Candida</i> spp.
Other microbiologic tests	Added new Major Criteria for fastidious pathogens: 1) PCR or amplicon/metagenomic sequencing identifies <i>C. burnetii</i> , <i>Bartonella</i> spp., or <i>T. whipplei</i> from blood; or 2) IFA ≥1:800 for IgG antibodies identifies <i>B. henselae</i> or <i>B. quintana</i> .
Imaging	
Echocardiography	Similar to earlier versions. Cornerstone of imaging criterion.
Cardiac computed tomography	Added new Major Criterion.
[18F]FDG PET/CT	Findings equivalent to echocardiography.
[18F]FDG PET/CT	Added new Major Criterion.
Findings for native valve, cardiac device, or prosthetic valve >3 mo after cardiac surgery are equivalent to echocardiography.	
Surgical	
Intraoperative inspection	Added new Major Criterion.
Intraoperative inspection constitutes Major Criterion in absence of Major Criterion by cardiac imaging or histopathology.	
MINOR CLINICAL CRITERIA	
Predisposition	Added transcatheter valve implant/repair, endovascular CIED, and prior diagnosis of IE.
Fever	Unchanged.
Vascular phenomena	Added splenic and cerebral abscess.
Immunologic phenomena	Added definition for immune complex mediated glomerulonephritis.
Microbiological	Added PCR or amplicon/metagenomic sequencing evidence of typical pathogen.
Imaging	Added PET/CT evidence <3 mo of cardiac surgery.
Physical examination	New auscultation of regurgitant murmur when echocardiography is unavailable.

Abbreviations: [18F]FDG PET CT, positron emission computed tomography with 18F-fluorodeoxyglucose; CIED, cardiac implantable electronic device; IFA, immunofluorescence assay; PCR, polymerase chain reaction.

«Cerrahi» kriter



NEW MAJOR CRITERION – SURGICAL EVIDENCE

The intraoperative inspection of cardiac pathology by cardiovascular surgeons is invaluable in a case of suspected IE, particularly if further pathologic or microbiologic confirmation is not available. As a result, the ISCVID Working Group has added intraoperative evidence of IE (eg, vegetations, abscess, valvular destruction, dehiscence or loosening of prosthetic valve, other direct evidence of IE) as a new Major Criterion in the 2023 Duke-ISCVID IE Criteria when other definitive criteria (eg, cardiac imaging, histology, microbiology) IE are unavailable (Table 2).

Cerrahin ameliyatta endokardit olduğunu düşünmesi
–başka hiçbir bulgu olmaksızın-
majör kriter olarak değerlendiriliyor

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Mikrobiyolojik kriterlerde neler deđiřti?



Table 3. Updates to Modified Duke Criteria Proposed by 2023 Duke-International Society for Cardiovascular Infectious Diseases Infective Endocarditis (IE) Criteria

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- Kan kltr «halen» ok nemli
- Kan kltrn alma zamanına iliřkin nerilerde deđiřiklik var
 - nceki nerilere gre 1 saat iinde 3 set kan kltr almak gerekiyordu
 - řimdiki nerilerde sre belirtilmiyor

Mikrobiyolojik kriterlerde neler değişti?



Microbiology

Blood cultures	Removed requirements for timing and separate venipunctures for blood cultures.
Definition of typical organisms	Added typical pathogens: 1) <i>S. lugdunensis</i> ; <i>E. faecalis</i> ; all streptococci except <i>S. pneumoniae</i> and <i>S. pyogenes</i> ; <i>Granulicatella</i> spp.; <i>Abiotrophia</i> spp.; and <i>Gemella</i> spp.

- Endokardit açısından «tipik» kabul edilecek etkenlere yeni eklenenler:
- S.lugdunensis*
 - E.faecalis*
 - S.pneumoniae* ve *S.pyogenes* dışındaki tüm streptokoklar
 - Granulicatella spp.*
 - Abiotrophia spp.*
 - Gemella spp.*
- | Majör Ölçütler |
|---|
| 1. Infektif endokarditle uyumlu pozitif kan kültürü |

Tablo 11. Avrupa Kardiyoloji Derneği 2015 Modifikasyonlarına Göre Infektif Endokardit Tanı Ölçütleri (65)

Majör Ölçütler

1. **İnfektif endokarditle uyumlu pozitif kan kültürü**
 - a. İki ayrı kan kültüründe İE ile uyumlu tipik mikroorganizmaların üremesi (viridans streptokoklar, *Streptococcus bovis*, HACEK grubu, *Staphylococcus aureus*; ya da başka bir odak odak olmaması koşuluyla, toplumdaki edinilmiş enterokoklar
 - ya da
 - b. İE ile uyumlu mikroorganizmaların kan kültürlerinde sürekli üremesi >12 saat arayla alınmış en az iki kan kültüründe pozitif sonuç alınması; ya da üç ayrı kan kültürünün hepsinde ya da 4 ayrı kan kültürünün çoğunda (birinci ve son örnekler arasında en az 1 saat olması koşuluyla) pozitif sonuç alınması

Mikrobiyolojik kriterlerde neler deđiřti?



Microbiology

Blood cultures

Removed requirements for timing and separate venipunctures for blood cultures.

Definition of typical organisms

Added typical pathogens:

- 1) *S. lugdunensis*; *E. faecalis*; all streptococci except *S. pneumoniae* and *S. pyogenes*; *Granulicatella* spp.; *Abiotrophia* spp.; and *Gemella* spp.
- 2) Organisms to be considered "typical" IE pathogens in the setting of intracardiac prosthetic material: coagulase negative staphylococci, *Corynebacterium striatum*; *C. jeikeium*, *Serratia marcescens*, *Pseudomonas aeruginosa*, *Cutibacterium acnes*, nontuberculous mycobacteria, and *Candida* spp.

➤ **İntra-kardiyak yabancı cisim** olduđuunda endokardit açısından «tipik» kabul edilecek etkenler:

Koagülaz negatif stafilokok

Corynebacterium striatum

Corynebacterium jeikeium

Serratia marcescens

Pseudomonas aeruginosa

Cutibacterium acnes

Non-tuberculosis mikobakteriler

Candida spp.

Tablo 11. Avrupa Kardiyoloji Derneđi 2015 Modifikasyonlarının Göre İnfektif Endokardit Tanı Ölçütleri (65)

Majör Ölçütler

1. İnfektif endokarditle uyumlu pozitif kan kültürü

- a. İki ayrı kan kültüründe İE ile uyumlu tipik mikroorganizmaların üremesi (viridans streptokoklar, *Streptococcus bovis*, HACEK grubu, *Staphylococcus aureus*; ya da başka bir odak odak olmaması koşuluyla, toplumdan edinilmiş enterokoklar

ya da

- b. İE ile uyumlu mikroorganizmaların kan kültürlerinde sürekli üremesi >12 saat arayla alınmış en az iki kan kültüründe pozitif sonuç alınması; ya da üç ayrı kan kültürünün hepsinde ya da 4 ayrı kan kültürünün çoğunda (birinci ve son örnekler arasında en az 1 saat olması koşuluyla) pozitif sonuç alınması

Mikrobiyolojik kriterlerde neler değişti?



Other microbiologic tests

Added new Major Criteria for fastidious pathogens:

- 1) PCR or amplicon/metagenomic sequencing identifies *C. burnetii*, *Bartonella* spp., or *T. whipplei* from blood; or
- 2) IFA $\geq 1:800$ for IgG antibodies identifies *B. henselae* or *B. quintana*.

Zor üreyen bakteriler için «diğer» mikrobiyolojik testler eklendi:

- *Coxiella burnetii*, *Bartonella* spp veya *Tropheryma whipplei* için kanda PCR veya amolikon/metagenomik sekanslama
- *Bartonella henselae* ve *Bartonella quintana* Ig G antikorlarının IFA yöntemi ile 1/800 ve üzerinde olması

c. *Coxiella burnetii* için tek şişe pozitif kan kültürü ya da faz I antijenlerine karşı IgG antikor titresinin $>1:800$ olması

Görüntüleme kriterlerinde neler değişti?



Imaging

Echocardiography	Similar to earlier versions. Cornerstone of imaging criterion.
Cardiac computed tomography	Added new Major Criterion. Findings equivalent to echocardiography.
[18F]FDG PET/CT	Added new Major Criterion. Findings for native valve, cardiac device, or prosthetic valve >3 mo after cardiac surgery are equivalent to echocardiography.

Kardiyak BT ve PET-BT bulguları

- EKO bulguları ile benzer şekilde-
majör kriter olarak değerlendiriliyor

Clinical Infectious Diseases
VIEWPOINTS

Yves S. Fassin ^{1,2}, David T. Denek ¹, Christine Sellen-Sale ¹, Suzanne Miles ¹, David S. Bann ^{1,2}, Anne-Lise Chénier ¹, Andrew Bell ¹, Louis D'Amato ¹,
Isabelle Derode-Wang ¹, Karine Duvet ¹, Claudia Guerin-Fortin ¹, Emil Fiedler ¹, Wenguo W. Han ¹, Barbara Koon ¹, Brian Koon ¹,
André W. Kucharski ¹, Carlos A. Mestas ¹, Gaby A. Pett ^{1,2}, Maria Nieves-Perez ¹, Stephen S. Proctor ¹, Albert Suga ¹, Francis Verheul ^{1,2}

[illegible]

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Değişmeyenler



Table 1. Definitions of Infective Endocarditis According to the 2023 Duke-International Society for Cardiovascular Infectious Diseases Infective Endocarditis (IE) Criteria, With Proposed Changes in Bold Type

I. DEFINITE ENDOCARDITIS

A. Pathologic Criteria

(1) **Microorganisms identified^a in the context of clinical signs of active endocarditis in a vegetation; from cardiac tissue; from an explanted prosthetic valve or sewing ring; from an ascending aortic graft (with concomitant evidence of valve involvement); from an endovascular intracardiac implantable electronic device (CIED); or from an arterial embolus**

or

(2) **Active endocarditis^b (may be acute^c or subacute/chronic^d) identified in or on a vegetation; from cardiac tissue; from an explanted prosthetic valve or sewing ring; from an ascending aortic graft (with concomitant evidence of valve involvement); from a CIED; or from an arterial embolus**

B. Clinical Criteria

(1) 2 Major Criteria

or

(2) 1 Major Criterion and 3 Minor Criteria

or

(3) 5 Minor Criteria

II. POSSIBLE ENDOCARDITIS

A. 1 Major Criterion And 1 Minor Criterion

or

B. 3 Minor Criteria

III. REJECTED ENDOCARDITIS

A. Firm alternate diagnosis explaining signs/symptoms^e

or

B. **Lack of recurrence despite antibiotic therapy for less than 4 d.**

or

C. No pathologic or macroscopic evidence of IE at surgery or autopsy, with antibiotic therapy for less than 4 d

or

D. Does not meet criteria for possible IE, as above

- Kesin endokardit
- Olası endokardit
- Dışlanmış endokardit

- Majör kriterler
- Minör kriterler

Minör kriterlerde neler değişti?



Vaküler fenomenlere eklenenler:

- Splenik abse
- Serebral abse

Minör Ölçütler*

1. **Yatkınlık:** IE ye yatkınlık oluşturan kalp hastalığı, IVDU olma
2. **Ateş:** vücut sıcaklığının $>38^{\circ}\text{C}$ olması
3. **Vasküler olaylar** (sadece görüntülemeyle saptananlar dahil): majör arteriyel embolizm, septik pulmoner infarktler, mikotik anevrizma, intrakraniyal kanama; konjunktival kanamalar ve Janeway lezyonları
4. **İmmünolojik olaylar:** glomerülonefrit, Osler nodülleri, Roth lekeleri, romatoid faktör pozitifliği
5. **Mikrobiyolojik kanıtlar:** majör ölçütleri karşılamayan kan kültürü pozitiflikleri ya da IE ile uyumlu bir mikroorganizmayla aktif infeksiyonu gösteren serolojik kanıtlar

Eğilim yaratan durumlara eklenenler:

- TAVİ gibi işlemler
- Kalp pili gibi elektronik cihazlar
- Daha önce İE geçirmiş olma

NEW MINOR CLINICAL CRITERIA

Clinical features added to the list of possible Minor Criteria by the ISCVID Working Group as predisposing conditions included additional types of cardiac prosthetic material (eg, transcatheter valve implant/repair, endovascular leads of CIEDs), an updated list of congenital heart conditions [43, 44], and a prior diagnosis of IE [45]. The ISCVID Working group recognized additional vascular phenomenon, including cerebral abscess and splenic abscess. Last, the ISCVID Working group developed a practical definition of immune complex mediated glomerulonephritis within the immunologic phenomena category.

Yeni kriterlere ilişkin değerlendirme



- Önerilen yöntemlerin tamamının kaynakları kısıtlı ülkelerde yapılması mümkün olmayabilir
- Validasyon çalışmaları önemli
- Değişen bilgiler ve veriler ışığında sürekli güncellenmeli

2023 Duke ISCVID kriterleri vs Modifiye Duke Kriterleri

Clinical Infectious Diseases
MAJOR ARTICLE

External Validation of the 2023 Duke–International Society for Cardiovascular Infectious Diseases Diagnostic Criteria for Infective Endocarditis

Thomas W. van der Vaart,^{1,2,3,4} Patrick M. M. Bessnot,⁵ David T. Durack,⁶ Larry M. Baddour,^{7,8} Arnold S. Bayer,^{9,10} Emanuele Durante-Mangoni,¹¹ Thomas L. Hilland,¹² Adolf W. Karchner,¹³ Jesse M. Mice,^{14,15} Philippe Mercillon,¹⁶ Magnus Rasmussen,¹⁷ Christine Sehn-Sety,^{18,19} Vance G. Fowler Jr.,²⁰ and Jan T. M. van der Meer²¹

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(See the Invited Commentary by Chambers et al. on pages 964–7.)

Background. The 2023 Duke–International Society of Cardiovascular Infectious Diseases (ISCVID) criteria for infective endocarditis (IE) were introduced to improve classification of IE for research and clinical purposes. External validation studies are required.

Methods. We studied consecutive patients with suspected IE referred to the IE team of Amsterdam University Medical Center (from October 2016 to March 2021). An international expert panel independently reviewed case summaries and assigned a final diagnosis of “IE” or “not IE,” which served as the reference standard, to which the “definite” Duke ISCVID classifications were compared. We also evaluated accuracy when excluding cardiac surgical and pathologic data (“clinical” criteria). Finally, we compared the 2023 Duke ISCVID with the 2000 modified Duke criteria and the 2015 and 2023 European Society of Cardiology (ESC) criteria.

Results. A total of 595 consecutive patients with suspected IE were included; 399 (67%) were adjudicated as having IE; 111 (19%) had prosthetic valve IE, and 48 (8%) had a cardiac implantable electronic device IE. The 2023 Duke ISCVID criteria were more sensitive than either the modified Duke or 2015 ESC criteria (84.2% vs 74.9% and 80%, respectively; $P < .001$) without significant loss of specificity. The 2023 Duke ISCVID criteria were similarly sensitive but more specific than the 2023 ESC criteria (94% vs 82%; $P < .001$). The same pattern was seen for the clinical criteria (excluding surgical/pathologic results). New modifications in the 2023 Duke ISCVID criteria related to “major microbiological” and “imaging” criteria had the most impact.

Conclusions. The 2023 Duke ISCVID criteria represent a significant advance in the diagnostic classification of patients with suspected IE.

Keywords. infective endocarditis; diagnosis; Duke criteria; validation.

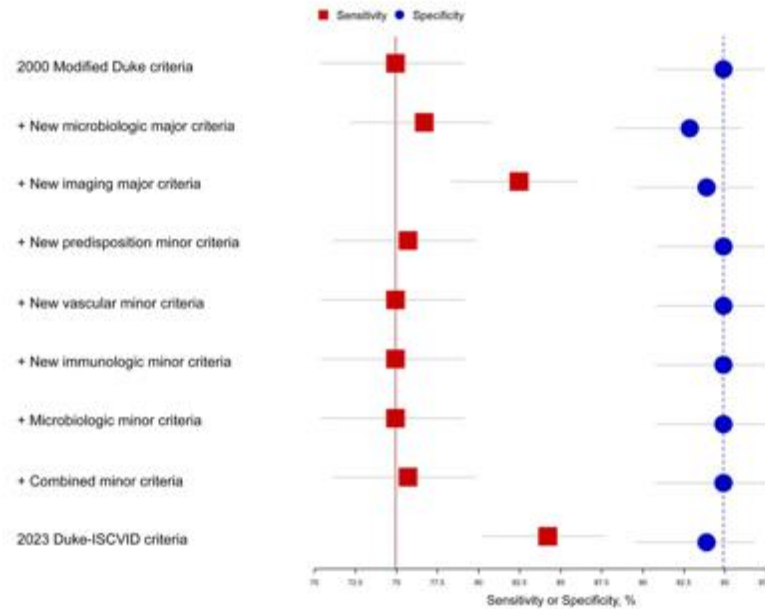


Figure 1. Added value of each change in the Duke–International Society of Cardiovascular Infectious Diseases (ISCVID) criteria compared with the modified Duke criteria. Each point (square for sensitivity, circle for specificity) provides the diagnostic accuracy measure for the 2000 modified Duke criteria with the addition of a specific change from the Duke-ISCVID criteria. Horizontal bars represent 95% confidence intervals for the point estimate of the complete Duke-ISCVID criteria.

2023 Duke-ISCVID kriterleri, Modifiye Duke kriterlerine göre daha yüksek duyarlılığa (sensitivite) ve benzer özgüllüğe (spesifisite) sahiptir

Our investigation provides evidence that the 2023 Duke-ISCVID criteria should replace previous versions of the Duke criteria because they have superior sensitivity and comparable specificity. While the 2023 Duke-ISCVID criteria can provide a useful framework to assist in the diagnosis of IE, clinicians should be aware that the sensitivity of the criteria is not perfect. This is apparent in the sensitivity analysis of the 2023 Duke-ISCVID clinical criteria in surgically treated patients. The sensitivity of the clinical criteria in this group was 75%, indicating that 25% of patients who will have surgically confirmed IE did not meet the clinical criteria for definite IE before surgery. Using a combination of both the definite and the possible classifications to define IE would be of limited use, as the specificity of this combination would be unacceptably low (21%). Clinicians should always consider pretest prob-

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European Society
of Cardiology

European Heart Journal (2023) 00, 1–95
<https://doi.org/10.1093/eurheartj/ehad193>

ESC GUIDELINES

2023 ESC Guidelines for the management of endocarditis

**Developed by the task force on the management of endocarditis
of the European Society of Cardiology (ESC)**

***Endorsed by the European Association for Cardio-Thoracic Surgery
(EACTS) and the European Association of Nuclear Medicine (EANM)***

Authors/Task Force Members: Victoria Delgado  ^{*,†}, (Chairperson) (Spain),
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Suzanne de Waha [‡], (Task Force Co-ordinator) (Germany), Nikolaos Bonaros 
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Stefano Caselli  (Switzerland), Torsten Doenst  (Germany),
Stephane Ederhy  (France), Paola Anna Erba  ¹ (Italy), Dan Foldager (Denmark),
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Eduard Quintana  ³ (Spain), Trine Bernholdt Rasmussen  (Denmark),
Arsen D. Ristić  (Serbia), Josep Rodés-Cabau (Canada), Alessandro Sionis 
(Spain), Liesl Joanna Zühlke  (South Africa), Michael A. Borger  ^{*,†},
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2023 Avrupa Endokardit Rehberi'nde neler değişti?

Review article

OPEN

The 2023 new European guidelines on infective endocarditis: main novelties and implications for clinical practice

Massimo Imazio^{a,b}

The 2023 European Society of Cardiology (ESC) guidelines for the management of infective endocarditis update the previous 2015 guidelines with main novelties in five areas: (1) antibiotic prevention for high-risk patients, and prevention measures for intermediate-risk and high-risk patients; (2) diagnosis with emphasis on multimodality imaging to assess cardiac lesions of infective endocarditis; (3) antibiotic therapy allowing an outpatient antibiotic treatment for stabilized, uncomplicated cases; (4) cardiac surgery with an emphasis on early intervention without delay for complicated cases; and (5) shared management decision by the endocarditis team. Most evidence came from observational studies and expert opinions. The guidelines strongly support a patient-centred approach with a shared decision process by a multidisciplinary team that should be implemented either in tertiary referral centres, becoming heart valve centres, and referral

centres. A continuous sharing of data is warranted in the hospitals' network between heart valve centres, which are used for referrals for complicated cases of infective endocarditis, and referral centres, which should be able to manage uncomplicated cases of infective endocarditis.

J Cardiovasc Med. 2024; 25:718–726

Keywords: antibiotic therapy, cardiac surgery, heart team, infective endocarditis, prevention

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Introduction

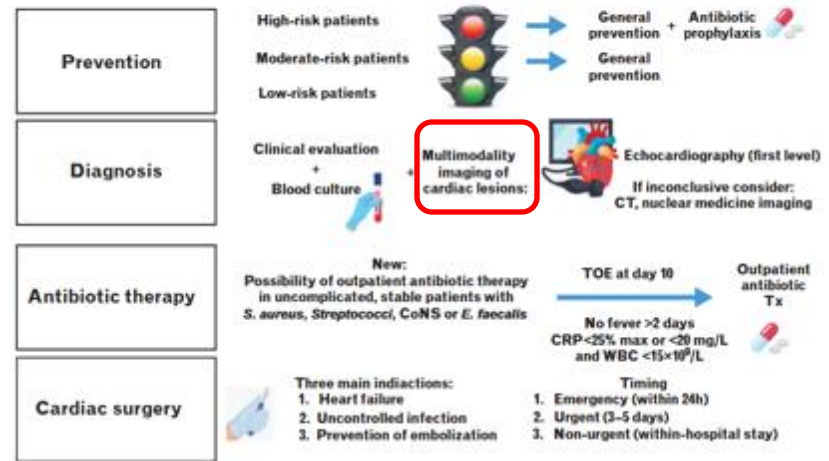
The 2023 European guidelines for the management of infective endocarditis¹ update the old 2015 guidelines² providing new important novelties in five main areas: prevention of infective endocarditis; diagnosis with multimodality imaging; antibiotic therapy and possibility of outpatient antibiotic therapy for stabilized, uncomplicated cases; timing of cardiac surgery; and role of the endocarditis team with a network of heart centres and referral centres (Fig. 1).

on antibiotic prophylaxis based on previous guidelines, have also contributed to this increase in the incidence of infective endocarditis.^{1,3,4}

The development of infective endocarditis requires a predisposing condition, pathogens entering the bloodstream (from oral cavity, skin, gastrointestinal or genitourinary tract), and host's immune response with a reduced capacity for clearance of the infectious agent. The role of pre-

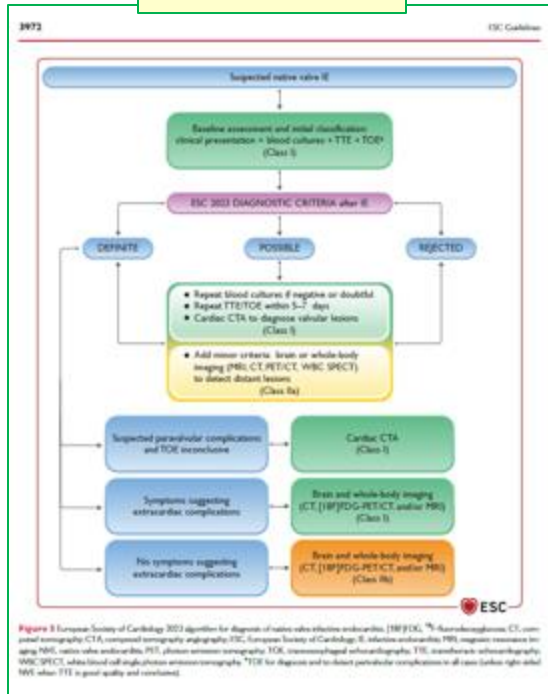
The 2023 new European guidelines on infective endocarditis Imazio 719

Fig. 1

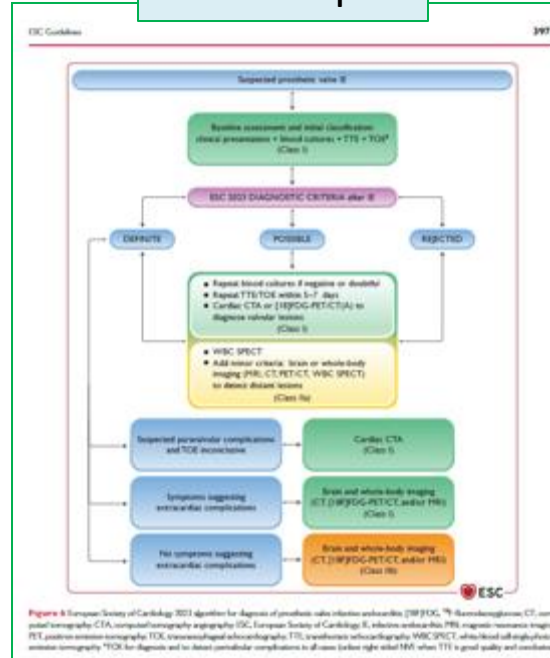


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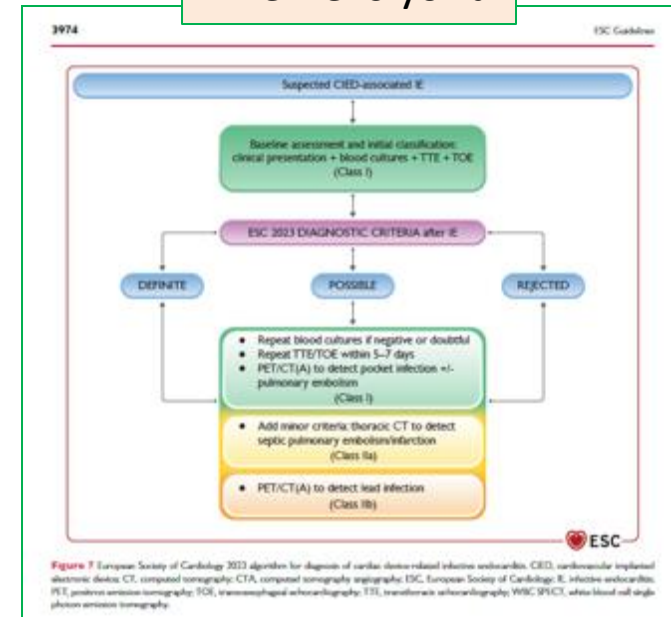
Nativ kapak



Protez kapak

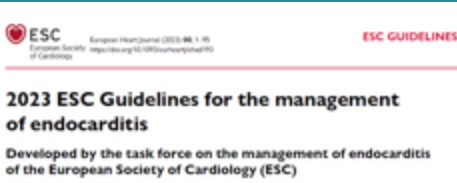


Pil enfeksiyonu



Antibiyotik Tedavisi

Ayaktan Parenteral Tedavi



Section 7. Recommendation Table 11 — Recommendations for outpatient antibiotic treatment of infective endocarditis

Outpatient parenteral antibiotic treatment should be considered in patients with left-sided IE caused by <i>Streptococcus</i> spp., <i>E. faecalis</i> , <i>S. aureus</i> , or CoNS who were receiving appropriate i.v. antibiotic treatment for at least 10 days (or at least 7 days after cardiac surgery), are clinically stable, and who do not show signs of abscess formation or valve abnormalities requiring surgery on TOE.	IIa	A
Outpatient parenteral antibiotic treatment is not recommended in patients with IE caused by highly difficult-to-treat microorganisms, liver cirrhosis (Child-Pugh B or C), severe cerebral nervous system emboli, untreated large extracardiac abscesses, heart valve complications, or other severe conditions requiring surgery, severe post-surgical complications, and in PWID-related IE.	III	C

Uygun koşullarda düşünülebilir (Sınıf IIa, Düzey A):

Sol kalp tarafı enfektif endokarditi *Streptococcus* spp., *E. faecalis*, *S. aureus* veya koagülaz-negatif stafilokoklar (CoNS) tarafından gelişmiş,
en az **10 gün (veya kardiyak cerrahiden sonra ≥ 7 gün)** uygun IV antibiyotik tedavisi almış,
linik olarak stabil, apsesi olmayan ve **cerrahi gerektiren kapak anomalisi bulunmayan** hastalarda

ayaktan parenteral tedavi düşünülebilir

Class I → “Kesin yapılmalı”

Class IIa → “Yapılması güçlü biçimde önerilir”

Class IIb → “Yapılması düşünülebilir”

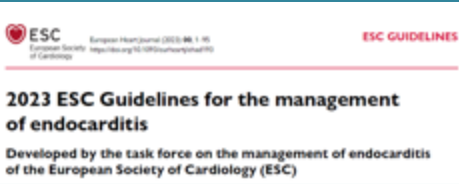
Class III → “Yapılmamalı”

Level A → “Yüksek kanıt (çok merkezli RCT’ler)”

Level B → “Orta kanıt (tek RCT veya gözlemsel veri)”

Level C → “Uzman görüşü / düşük düzey veri”

Ayaktan Parenteral Tedavi



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ayaktan parenteral tedavi düşünülebilir

Önerilmez (Sınıf III, Düzey C):

- Zor tedavi edilen mikroorganizmalar
- Karaciğer sirozu (Child–Pugh B veya C)
- Ağır SSS embolisi
- Tedavi edilmemiş büyük ekstrakardiyak apse
- Kapak komplikasyonu veya cerrahi gerektiren durum
- Ciddi postoperatif komplikasyonlar
- Enjeksiyon yoluyla madde kullananlarda gelişen IE

Class I → “Kesin yapılmalı”

Class IIa → “Yapılması güçlü biçimde önerilir”

Class IIb → “Yapılması düşünülebilir”

Class III → “Yapılmamalı”

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Introduction

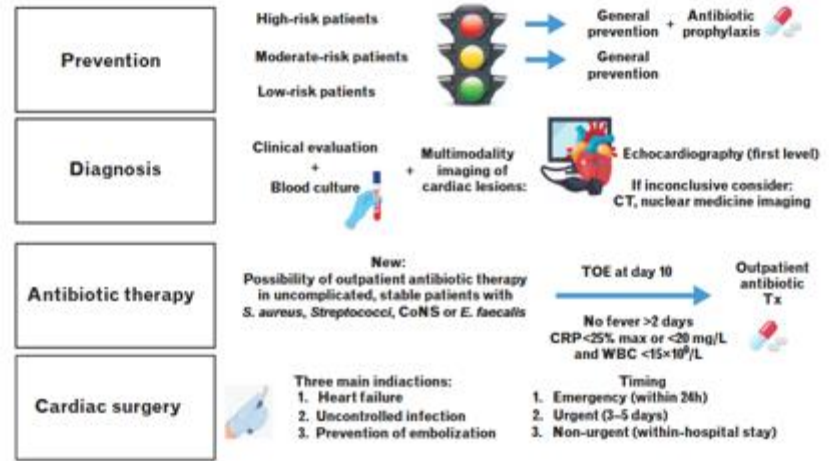
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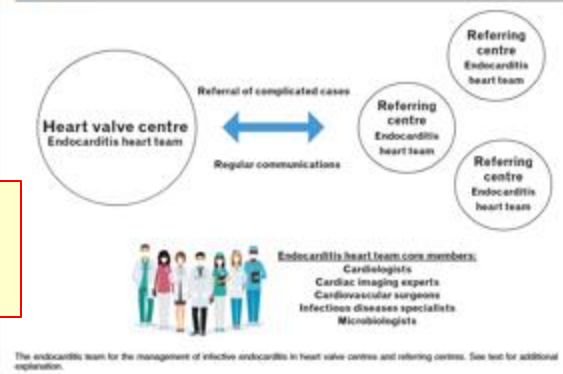
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Fig. 1



724 Journal of Cardiovascular Medicine 2024; Vol 25 No 10

Fig. 2



Oral geçişli antibiyotik tedavisi (IV→oral)
stabil olgularda güvenli ve etkin olabilir

Oral Tedaviye Geçiş Seçenekleri

JAMA Network Open

Consensus Statement | Infectious Diseases

Guidelines for Diagnosis and Management of Infective Endocarditis in Adults A WikiGuidelines Group Consensus Statement

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Abstract

IMPORTANCE: Practice guidelines often provide recommendations in which the strength of the recommendation is dissociated from the quality of the evidence.

OBJECTIVE: To create a clinical guideline for the diagnosis and management of adult bacterial infective endocarditis (IE) that addresses the gap between the evidence and recommendation strength.

EVIDENCE REVIEW: This consensus statement and systematic review applied an approach previously established by the WikiGuidelines Group to construct collaborative clinical guidelines. In April 2022 a call to new and existing members was released electronically (social media and email) for the next WikiGuidelines topic, and subsequently, topics and questions related to the diagnosis and management of adult bacterial IE were crowdsourced and prioritized by vote. For each topic, PubMed literature searches were conducted including all years and languages. Evidence was reported according to the WikiGuidelines charter: clear recommendations were established only when reproducible, prospective, controlled studies provided hypothesis confirming evidence. In the absence of such data, clinical reviews were crafted discussing the risks and benefits of different approaches.

FINDINGS: A total of 51 members from 10 countries reviewed 587 articles and submitted information relevant to 4 sections: establishing the diagnosis of IE (9 questions), multidisciplinary IE teams (3 questions), prophylaxis (2 questions), and treatment (5 questions). Of 17 unique questions, a clear recommendation could only be provided for 1 question. 3 randomized clinical trials have established that oral transitional therapy is at least as effective as intravenous (IV) only therapy for the treatment of IE. Clinical reviews were generated for the remaining questions.

CONCLUSIONS AND RELEVANCE: In this consensus statement that applied the WikiGuidelines method for clinical guideline development, oral transitional therapy was at least as effective as IV only therapy for the treatment of IE. Several randomized clinical trials are underway to inform other areas of practice, and further research is needed.

Key Points

Question: Using the WikiGuidelines approach to the construct of clinical guidelines, how often can clear recommendations be made in the diagnosis and management of adult bacterial infective endocarditis?

Findings: In this consensus statement, a panel of 51 members found that only 1 of 17 questions had sufficiently high quality data to allow for a clear recommendation. Oral transitional therapy is at least as effective as intravenous only therapy for the treatment of infective endocarditis.

Meaning: These findings suggest that a higher quality of evidence needs to be established to guide the diagnosis and management of infective endocarditis.

Supplemental content

Author affiliations and article information are listed at the end of this article.

Table 3. Summary of Oral Transitional Antibiotics Used in Published Clinical Studies^a

Drug	Organism	Dose	References
Amoxicillin	• Sensitive streptococci (with or without combination treatment) • Enterococci (only in combination with rifampin or linezolid)	1 g 4 times daily	Iversen et al, ⁵⁵ 2019; Stamboulis et al, ⁶¹ 1991
Dicloxacillin	Sensitive staphylococci (data from RCT only in combination with rifampin)	1 g 4 times daily	Iversen et al, 2019 ⁵⁵
Levofloxacin ^b	Sensitive staphylococci (only in combination with rifampin)	750 mg once daily	Iversen et al, ⁵⁵ 2019; Heldman et al, ⁶² 1996
Moxifloxacin	Sensitive streptococci, enterococci, or staphylococci (only in combination with amoxicillin, rifampin, or linezolid)	400 mg once daily	Iversen et al, ⁵⁵ 2019
TMP-SMX	Sensitive staphylococci	960 mg or 4800 mg daily in divided doses	Tissot-Dupont et al, ⁶⁶ 2019; Freling et al, ⁴⁷ 2023
Linezolid	Sensitive gram-positive cocci (alone or in combination with rifampin, moxifloxacin, or amoxicillin) ^c	600 mg twice daily	Iversen et al, ⁵⁵ 2019; Tascini et al, ⁶⁸ 2011; Fasagan et al, ⁶⁹ 2006; Colli et al, ⁷⁰ 2007; Muller et al, ⁷¹ 2007; Freling S et al, ⁴⁷ 2023
Rifampin	Only as adjunctive agent (as previously described)	600 mg once or twice daily	Iversen et al, ⁵⁵ 2019; Heldman et al, ⁶² 1996; Accorcello G, ⁷² 1983; Freling et al, ⁴⁷ 2023

Abbreviations: RCT, randomized clinical trial; TMP-SMX, trimethoprim-sulfamethoxazole.

^a Combination regimens were used in the largest RCT; other published regimens have included either monotherapy or combination therapy regimens.³

^b The study used ciprofloxacin rather than levofloxacin, the latter of which was not yet clinically available. However, in ensuing years, ciprofloxacin experienced rapid

emergence of staphylococcal resistance. Levofloxacin or moxifloxacin are preferred to ciprofloxacin due to enhanced in vitro activity against staphylococci, but generally only in combination with a second agent.

^c For most patients in published studies, linezolid was used alone, but in some references,^{3,4,7} linezolid was given in combination with rifampin, moxifloxacin, or amoxicillin.

Oral Tedaviye Geçiş Seçenekleri

- **Streptokoklar: Penisiline duyarlı ($MIC \leq 0.12 \mu\text{g/mL}$)**

Amoksisilin 1 g günde 4 kez (yalnızca nativ kapak enfeksiyonlarında)

Amoksisilin 1 g günde 4 kez + **rifampisin** 600 mg günde 1 kez

Linezolid 600 mg günde 2 kez tek başına veya **rifampisin** 600 mg günde 1 kez ile

Moksifloksasin 400 mg günde 1 kez + **rifampisin** 600 mg günde 1 kez veya **linezolid** 600 mg günde 2 kez

- **Streptokoklar: Penisiline orta duyarlı ($MIC 0.25\text{--}1.00 \mu\text{g/mL}$) veya Enterokoklar**

Amoksisilin 1 g günde 4 kez + **rifampisin** 600 mg günde 1 kez veya **linezolid** 600 mg günde 2 kez

Linezolid 600 mg günde 2 kez tek başına veya **rifampisin** 600 mg günde 1 kez ile

Moksifloksasin 400 mg günde 1 kez + **rifampisin** 600 mg günde 1 kez

- **Streptokoklar: Penisiline dirençli ($MIC \geq 2 \mu\text{g/mL}$) veya amoksisiline dirençli Enterokoklar**

Linezolid 600 mg günde 2 kez tek başına veya **rifampisin** 600 mg günde 1 kez ile

Moksifloksasin 400 mg günde 1 kez + **rifampisin** 600 mg günde 1 kez

- **Staphylococcus spp.**

Levofloksasin 750 mg günde 1 kez + **rifampisin** 600 mg günde 1 kez veya **linezolid** 600 mg günde 2 kez

Linezolid 600 mg günde 2 kez tek başına veya **rifampisin** 600 mg günde 1 kez ile (rifampisin linezolid düzeylerini düşürdüğünden monoterapi mi kombinasyon mu tercih edilmeli, belirsizdir)

TMP-SMX 960 mg veya 4800 mg/gün bölünmüş dozlarda

Dikloksasilin 1 g günde 4 kez + **rifampisin** 600 mg günde 1 kez (yalnızca metisiline duyarlı suşlar için)



Cerrahi Tedavi

Japonya: 2016–2021 Ulusal Kohort Verileri

Journal of the American Heart Association

ORIGINAL RESEARCH

Trends in the Clinical Characteristics and Outcome of Infective Endocarditis: A Nationwide Study From 2016 to 2021

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BACKGROUND: Infective endocarditis (IE) is still a fatal disease, and given its rarity, ongoing updates to patient characteristics and outcomes of IE are essential for providing precision diagnoses and effective treatments. This study sought to examine temporal trends in the clinical characteristics and in-hospital occurrence of adverse outcomes of IE.

METHODS AND RESULTS: Using the Japan nationwide administrative database, we identified patients with IE in Japan from 2016 to 2021. A total of 17 407 patients with IE (37.8% women; median age, 72 [interquartile range, 59–81] years) were identified. The incidence of IE increased from 2.02 per 100 000 population in 2016 to 2.59 per 100 000 population in 2021. The median age of the patients increased from 70 years in 2016 to 73 years in 2021, and patient backgrounds were becoming more complex. The in-hospital mortality rate was 14.5%, which significantly increased from 14.1% in 2016 to 15.4% in 2021. Higher age (odds ratio [OR], 1.34 [95% CI, 1.29–1.40]), heart failure (OR, 1.37 [95% CI, 1.18–1.59]), cerebrovascular complications (OR, 1.78 [95% CI, 1.59–1.99]), renal failure (OR, 2.36 [95% CI, 2.06–2.70]), and sepsis (OR, 2.79 [95% CI, 2.48–3.13]), were independently associated with an increased in-hospital mortality rate. In contrast, cardiac surgery (OR, 0.47 [95% CI, 0.41–0.55]), and a higher number of total cardiac surgeries (OR, 0.77 [95% CI, 0.65–0.92]) independently predicted a lower in-hospital mortality rate.

Cardiac surgery was performed in 4993 (28.7%) patients, of which 2457 (14.1%) were performed within 7 days of admission. The overall rate of cardiac surgeries remained relatively constant over the study period (27.9% in 2016 to 28.8% in 2021; P for trend = 0.47).

CLINICAL PERSPECTIVES

What Is New?

- Contemporary data from a national Japanese administrative database showed that the incidence of infective endocarditis increased over time, patients have been getting older, and there has been an increasing in-hospital mortality rate.
- Cardiac surgery and a high volume of cardiac surgeries independently predicted a lower in-hospital mortality rate.

What Are the Clinical Implications?

- Our study provides valuable insights into the epidemiology, risk factors, and outcomes of infective endocarditis, which may inform health service planning and treatment strategies.

Mortaliteyi Azaltan Faktörler

- Kardiyak cerrahi uygulanması
- Cerrahi hacmi yüksek merkezlerde tedavi

Profilaksi

Avrupa Endokardit Rehberi- 2023



ESC

European Society
of Cardiology

European Heart Journal (2023) 00, 1–95
<https://doi.org/10.1093/eurheartj/ehad193>

ESC GUIDELINES

2023 ESC Guidelines for the management of endocarditis

Developed by the task force on the management of endocarditis of the European Society of Cardiology (ESC)

Endorsed by the European Association for Cardio-Thoracic Surgery (EACTS) and the European Association of Nuclear Medicine (EANM)

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2. Introduction

Infective endocarditis (IE) is a major public health challenge.¹ In 2019, the estimated incidence of IE was 13.8 cases per 100 000 subjects per year, and IE accounted for 66 300 deaths worldwide.² Due to the associated high morbidity and mortality (1723.59 disability-adjusted life years and 0.87 death cases per 100 000 population, respectively), identification of the best preventive strategies has been the focus of research.^{2,3} Since the publication of the 2015 ESC Guidelines for the management of infective endocarditis,⁴ important new data have been published mandating an update of recommendations. First, the population at risk of IE has increased and new data on IE in different clinical scenarios have arisen.^{5–11} Furthermore, the emerging and increasing antibiotic resistance among oral streptococci is

of concern. The rate of resistance to azythromycin and clarithromycin is higher than that to penicillin.¹² Whether changes in national guidelines on the use of antibiotic prophylaxis have resulted in an increase in the incidence of IE remains unclear.^{13–18} It is likely that the increased use of diagnostic tools to diagnose IE is an important contributor to the increase in the incidence of IE. The use of echocardiography has probably increased in patients with positive blood cultures for *Enterococcus faecalis*, *Staphylococcus aureus*, or streptococci due to the associated increased risk of IE.¹⁹ In addition, computed tomography (CT) and nuclear imaging techniques have increased the number of definite IE cases particularly among patients with prosthetic valves and implantable cardiac devices.^{20–22}

Data on the contemporary characterization of patients with IE have been taken into consideration to update the recommendations on the diagnosis and management of patients with IE.^{5,19,23–41} Furthermore, the recommendations on antibiotic therapy have been updated based on the susceptibility of various microorganisms defined by the European Committee on Antimicrobial Susceptibility Testing (EUCAST) clinical breakpoints.⁴² Recommendations on outpatient parenteral antibiotic therapy (OPAT) or oral antibiotic treatment have been included based on the results of the Partial Oral Treatment of Endocarditis (POET) randomized trial and other trials.^{43–46}

Avrupa Endokardit Rehberi- 2023

Recommendation Table 1 — Recommendations for antibiotic prophylaxis in patients with cardiovascular diseases undergoing oro-dental procedures at increased risk for infective endocarditis

Recommendations	Class ^a	Level ^b
General prevention measures are recommended in individuals at high and intermediate risk for IE.	I	C
Antibiotic prophylaxis is recommended in patients with previous IE. ^{47,84,86}	I	B
Antibiotic prophylaxis is recommended in patients with surgically implanted prosthetic valves and with any material used for surgical cardiac valve repair. ^{47,87-89}	I	C
Antibiotic prophylaxis is recommended in patients with transcatheter implanted aortic and pulmonary valvular prostheses. ⁹¹⁻⁹⁴	I	C
Antibiotic prophylaxis is recommended in patients with untreated cyanotic CHD, and patients treated with surgery or transcatheter procedures with post-operative palliative shunts, conduits, or other prostheses. After surgical repair, in the absence of residual defects or valve prostheses, antibiotic prophylaxis is recommended only for the first 6 months after the procedure. ^{8,47,97,101}	I	C
Antibiotic prophylaxis is recommended in patients with ventricular assist devices. ¹⁰²	I	C
Antibiotic prophylaxis should be considered in patients with transcatheter mitral and tricuspid valve repair. ⁹⁵	IIa	C
Antibiotic prophylaxis may be considered in recipients of heart transplant. ¹⁰⁵⁻¹⁰⁷	IIb	C
Antibiotic prophylaxis is not recommended in other patients at low risk for IE. ^{11,51}	III	C

CHD, congenital heart disease; IE, infective endocarditis.

^aClass of recommendation.

^bLevel of evidence.

Recommendation Table 2 — Recommendations for infective endocarditis prevention in high-risk patients

Recommendations	Class ^a	Level ^b
Antibiotic prophylaxis is recommended in dental extractions, oral surgery procedures, and procedures requiring manipulation of the gingival or periapical region of the teeth. ^{11,49,51,108}	I	B
Systemic antibiotic prophylaxis may be considered for high-risk ^c patients undergoing an invasive diagnostic or therapeutic procedure of the respiratory, gastrointestinal, genitourinary tract, skin, or musculoskeletal systems. ^{6,11}	IIb	C

Sınıf IIb Kanıt Düzeyi C

Zayıf öneri → Kanıt sınırlı, dikkatli uygulanmalı

Avrupa Endokardit Rehberi- 2016



Yüksek riskli kardiyak durumlar

Table 3 Cardiac conditions at highest risk of infective endocarditis for which prophylaxis should be considered when a high-risk procedure is performed

Recommendations	Class ^a	Level ^b
Antibiotic prophylaxis should be considered for patients at highest risk for IE: (1) Patients with any prosthetic valve, including a transcatheter valve, or those in whom any prosthetic material was used for cardiac valve repair. (2) Patients with a previous episode of IE. (3) Patients with CHD: (a) Any type of cyanotic CHD. (b) Any type of CHD repaired with a prosthetic material, whether placed surgically or by percutaneous techniques, up to 6 months after the procedure or lifelong if residual shunt or valvular regurgitation remains. Antibiotic prophylaxis is not recommended in other forms of valvular or CHD.	IIa	C
	III	C

- Protez kapak
- Geçirilmiş İE
- Siyanotik konjenital kalp hastalığı
- Siyanotik kalp hastalığı cerrahisinden sonraki ilk 6 aylık dönem

KLİMİK 2016, 30. YIL KURULUŞU

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Yüksek riskli hastalarda profilaksi önerilen dental işlemler

Table 5 Recommendations for prophylaxis of infective endocarditis in the highest-risk patients according to the type of at-risk procedure

Recommendations	Class ^a	Level ^b
A. Dental procedures		
• Antibiotic prophylaxis should only be considered for dental procedures requiring manipulation of the gingiva or periapical region of the teeth or perforation of the oral mucosa	IIa	C
• Antibiotic prophylaxis is not recommended for local anesthetic injections in non-infected tissues, treatment of superficial caries, removal of sutures, dental X-rays, placement or adjustment of removable prosthodontic or orthodontic appliances or braces or following the shedding of deciduous teeth or trauma to the lips and oral mucosa	III	C

- Jinjival işlemler
- Periapikal işlemler
- Oral mukozada perforasyon

ÖNERİLMİYENLER!

- Lokal anestezi
- Dolgu
- Film çekimi
- Protodontik veya ortodontik işlemler

KLİMİK 2016, 30. YIL KURULUŞU

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Profilaksi önerilmeyen işlemler

Table 5 Continued

Recommendations	Class ^a	Level ^b
B. Respiratory tract procedures^c		
• Antibiotic prophylaxis is not recommended for respiratory tract procedures, including bronchoscopy or laryngoscopy, or tracheal or endotracheal intubation	III	C
C. Gastrointestinal or urogenital procedures or TÖE^d		
• Antibiotic prophylaxis is not recommended for gastroscopy, colonoscopy, cystoscopy, vaginal or caesarian delivery or TÖE	III	C
D. Skin and soft tissue procedures^e		
• Antibiotic prophylaxis is not recommended for any procedure	III	C

- Bronkoskopi, entübasyon
- Gastroskopi, kolonoskopi, sistoskopi, TÖE, sezeryan, doğum
- Deri ve yumuşak doku ile ilgili işlemler

KLİMİK 2016, 30. YIL KURULUŞU

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Avrupa Endokardit Rehberi- 2023

Recommendation Table 1 — Recommendations for antibiotic prophylaxis in patients with cardiovascular diseases undergoing oro-dental procedures at increased risk for infective endocarditis

Recommendations	Class ^a	Level ^b
General prevention measures are recommended in individuals at high and intermediate risk for IE.	I	C
Antibiotic prophylaxis is recommended in patients with previous IE. ^{47,84,86}	I	B
Antibiotic prophylaxis is recommended in patients with surgically implanted prosthetic valves and with any material used for surgical cardiac valve repair. ^{47,87-89}	I	C
Antibiotic prophylaxis is recommended in patients with transcatheter implanted aortic and pulmonary valvular prostheses. ⁹¹⁻⁹⁴	I	C
Antibiotic prophylaxis is recommended in patients with untreated cyanotic CHD, and patients treated with surgery or transcatheter procedures with post-operative palliative shunts, conduits, or other prostheses. After surgical repair, in the absence of residual defects or valve prostheses, antibiotic prophylaxis is recommended only for the first 6 months after the procedure. ^{8,47,97,101}	I	C
Antibiotic prophylaxis is recommended in patients with ventricular assist devices. ¹⁰²	I	C
Antibiotic prophylaxis should be considered in patients with transcatheter mitral and tricuspid valve repair. ⁹⁵	IIa	C
Antibiotic prophylaxis may be considered in recipients of heart transplant. ¹⁰⁵⁻¹⁰⁷	IIb	C
Antibiotic prophylaxis is not recommended in other patients at low risk for IE. ^{11,51}	III	C

CHD, congenital heart disease; IE, infective endocarditis.

^aClass of recommendation.

^bLevel of evidence.

Recommendation Table 2 — Recommendations for infective endocarditis prevention in high-risk patients

Recommendations	Class ^a	Level ^b
Antibiotic prophylaxis is recommended in dental extractions, oral surgery procedures, and procedures requiring manipulation of the gingival or periapical region of the teeth. ^{11,49,51,108}	I	B
Systemic antibiotic prophylaxis may be considered for high-risk ^c patients undergoing an invasive diagnostic or therapeutic procedure of the respiratory, gastrointestinal, genitourinary tract, skin, or musculoskeletal systems. ^{6,11}	IIb	C

Sınıf IIb Kanıt Düzeyi C

Zayıf öneri → Kanıt sınırlı, dikkatli uygulanmalı

“Değişmeyen tek şey değişimdir”



Herakleitos (iÖ 535-475)

NICE Rehberi- İngiltere

Personal View

Endocarditis prevention: time for a review of NICE guidance

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Summary

In 2023, the European Society of Cardiology (ESC) updated its infective endocarditis (IE) guidelines strongly endorsing antibiotic prophylaxis (AP) before invasive dental procedures (IDPs) for high-risk patients, elevating their recommendation to Class I. The American Heart Association (AHA) is aligned with this view and reaffirmed the need for AP to prevent IE in those at high-risk in its 2021 guidelines. In contrast, the UK's National Institute for Health and Care Excellence (NICE) recommends against routine AP use. Despite considerable new evidence, NICE has not reviewed this recommendation since 2015. In this Personal View, we review the new evidence that has arisen since 2015. Our analysis establishes the association between IDPs and IE and shows that AP is both safe and effective in reducing the IE-risk following IDPs in those at high-risk. Data also show that AP is cost-effective and would result in significant cost savings and health benefits if re-introduced into the UK's National Health Service for high-risk patients. Given these insights, we argue it is time NICE reviewed its guidance so that high-risk patients in the UK receive the same protection against IE that is afforded to patients in the rest of the world.

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Keywords: Infective endocarditis; Antibiotic prophylaxis; Invasive dental procedures; Guidelines; NICE

Introduction

The European Society of Cardiology (ESC) recently reviewed all available evidence in its 2023 guidelines for the prevention, diagnosis and management of infective endocarditis (IE) and concluded that high-risk patients

moderate risk on an individual basis" (Class IIb). Consistent with this guidance, the American Heart Association (AHA) also reviewed the available evidence in 2021 and concluded that high-risk patients should continue to receive AP before IDPs.¹ Sixteen years after

High risk
Previous history of infective endocarditis
Presence of prosthetic cardiac valve (including transcatheter valves)
Prosthetic material used for valve repair (including aortic/bicuspid and transcatheter valve procedures using prosthetic material)
Unrepaired congenital cardiac heart disease
Congenital heart disease intervention using palliative shunts or conduits
Completely repaired congenital heart defect with prosthetic material or device (whether placed by surgical or transcatheter techniques) ²
Patients with ventricular assist devices ³
Antibiotic prophylaxis may be considered in heart transplant recipients ⁴
Moderate risk (also known as intermediate risk)
Rheumatic heart disease
Non-rheumatic valve disease (including mitral valve prolapse)
Congenital valve anomalies (including aortic aneurysm)
Patients with cardiac implantable electronic devices (CIED) e.g. pacemaker or defibrillator ⁵
Hypertrophic cardiomyopathy
Low risk
Patients with none of the above high- or moderate-risk conditions
Notes based on the AHA ¹ and ESC ^{2,3} guideline definitions of those at high-, moderate- or low-risk. IE, infective endocarditis only. ² New moderate- and high-risk conditions added to 2021 ESC guidance. ³
Table 1: Cardiac conditions that identify individuals at high-, moderate- or low-risk of developing IE

Conclusions and call for action

Most studies described herein have been published since the last NICE guideline review nine years ago. They establish the association between IDPs and IE and demonstrate that AP is safe, cost-effective, and efficacious. They also provide important new evidence that AP can reduce the IE risk for high-risk patients undergoing IDPs and support current ESC and AHA guidance. Whilst a well-designed RCT would be the ideal way to establish AP efficacy, this is unlikely to happen, and the large observational studies described here are as close as we are likely to come. In countries where guidelines recommend AP, randomisation of high-risk patients to placebo or no AP cover is regarded as ethically unacceptable due to the risk of developing IE (particularly with its attendant 30% first year mortality). Attempts to perform an RCT in the UK (where AP is not routinely recommended) have also failed due to the large numbers of high-risk patients that would need to be enrolled to achieve the statistical power required to detect a clinically significant effect, the difficulty of randomising these patients to AP or placebo in the thousands of dental practices across the country where IDPs are performed, and the consequent very high-cost of an RCT. Funders have felt unable to justify prioritising the high cost of funding an RCT of AP prevention of IE over the need to support other less expensive life-saving clinical research, particularly when compelling observational data on AP efficacy already exist.¹⁶

In our view, a review of NICE guidance is now essential so that high-risk patients in the UK can benefit from the same protection afforded these patients in the rest of Europe and the world. Ideally, NICE would adopt similar recommendations to those of the 2023 ESC guidance. If NICE recommended that clinicians in the UK adopt the ESC guidelines, then this would result in a uniform approach across Europe. Ultimately, the ESC and AHA (whose recommendations are very similar) could come together to universalise their IE prevention guidance (perhaps under the auspices of the WHO) so that patients and clinicians world-wide could benefit from a common, evidence-based approach to IE prevention that would minimise unnecessary antibiotic use and the development of bacterial resistance.

- Son yıllarda yapılan çalışmalar, diş tedavi girişimleri sonrası IE riskinin antibiyotik profilaksisi ile anlamlı biçimde azaldığını göstermiştir
- Bu bulgular, ESC ve AHA kılavuzlarını desteklemektedir
- Randomize kontrollü çalışmalar etik ve ekonomik nedenlerle zor olsa da, mevcut veriler yüksek riskli hastalarda profilaksinin yeniden önerilmesi gerektiğini göstermektedir
- Bu yaklaşımın DSÖ gibi kurumlarca da benimsenmesi, antibiyotik kullanımını rasyonelleştirirken direnç gelişimini en aza indirecek bilim temelli bir stratejinin benimsenmesini sağlayacaktır

NICE National Institute for Health and Care Excellence

Guidance, Standards and Indicators, Clinical Knowledge Summaries (CKS), British National Formulary (BNF), British National Formulary for Children (BNF-C), Life sciences, More from NICE

NICE guidance

Evidence-based recommendations for the health and social care sector, developed by independent committees, including professionals and lay members, and consulted on by stakeholders.

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Endokardit Ekibi

2023 Avrupa Endokardit Rehberi'nde neler değişti?

Review article

OPEN

The 2023 new European guidelines on infective endocarditis: main novelties and implications for clinical practice

Massimo Imazio^{a,b}

The 2023 European Society of Cardiology (ESC) guidelines for the management of infective endocarditis update the previous 2015 guidelines with main novelties in five areas: (1) antibiotic prevention for high-risk patients, and prevention measures for intermediate-risk and high-risk patients; (2) diagnosis with emphasis on multimodality imaging to assess cardiac lesions of infective endocarditis; (3) antibiotic therapy allowing an outpatient antibiotic treatment for stabilized, uncomplicated cases; (4) cardiac surgery with an emphasis on early intervention without delay for complicated cases; and (5) shared management decision by the endocarditis team. Most evidence came from observational studies and expert opinions. The guidelines strongly support a patient-centred approach with a shared decision process by a multidisciplinary team that should be implemented either in tertiary referral centres, becoming heart valve centres, and referral

centres. A continuous sharing of data is warranted in the hospitals' network between heart valve centres, which are used for referrals for complicated cases of infective endocarditis, and referral centres, which should be able to manage uncomplicated cases of infective endocarditis.

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Keywords: antibiotic therapy, cardiac surgery, heart team, infective endocarditis, prevention

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Introduction

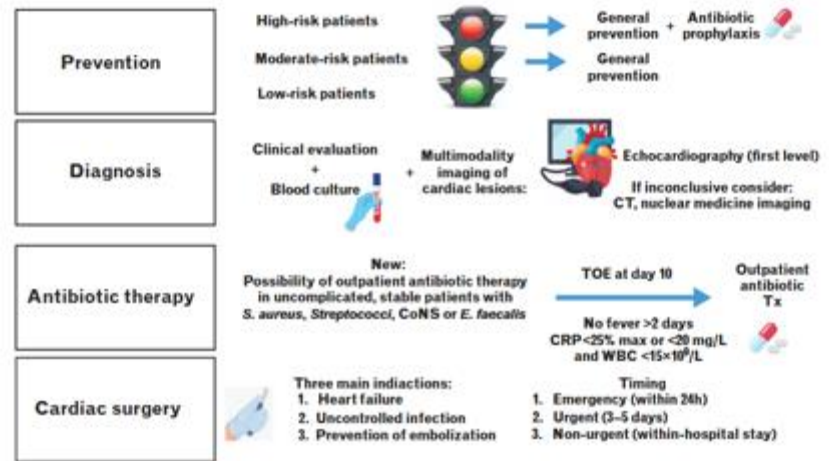
The 2023 European guidelines for the management of infective endocarditis¹ update the old 2015 guidelines² providing new important novelties in five main areas: prevention of infective endocarditis; diagnosis with multimodality imaging; antibiotic therapy and possibility of outpatient antibiotic therapy for stabilized, uncomplicated cases; timing of cardiac surgery; and role of the endocarditis team with a network of heart centres and referral centres (Fig. 1).

on antibiotic prophylaxis based on previous guidelines, have also contributed to this increase in the incidence of infective endocarditis.^{1,5,6}

The development of infective endocarditis requires a predisposing condition, pathogens entering the bloodstream (from oral cavity, skin, gastrointestinal or genitourinary tract), and host's immune response with a reduced capacity for clearance of the infectious agent. The role of pre-

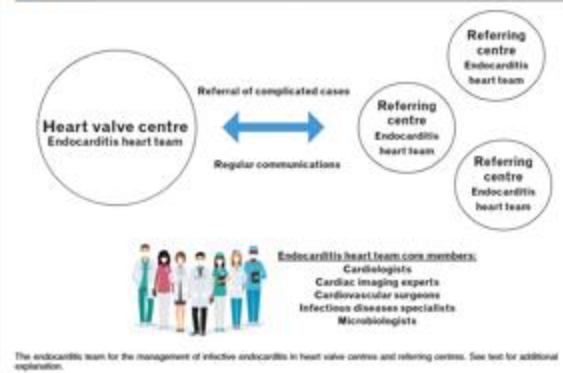
The 2023 new European guidelines on infective endocarditis Imazio 719

Fig. 1



726 Journal of Cardiovascular Medicine 2024; Vol 25 No 10

Fig. 2



Avrupa Endokardit Rehberi- 2023

4. The Endocarditis Team

The importance of an Endocarditis Team in the diagnosis, management, and clinical outcomes of patients with IE has been demonstrated in several observational studies.^{36-41,122-126} Establishing multidisciplinary endocarditis teams according to the European Society of Cardiology (ESC) and the American College of Cardiology/American Heart Association Guidelines^{4,127,128} has resulted in earlier and more accurate diagnosis of the primary disease and its complications,^{5,22,31,40,129} uniform antibiotic treatment,^{36,40,123} and optimized timing for surgical intervention.^{36,37,40,123} A variety of scenarios of patients presenting with IE justifies a multidisciplinary approach.^{5,25,27,28,130-135} Furthermore, the clinical presentation may vary significantly depending on the characteristics of the host and virulence of the microorganism. Accordingly, the concept of the Endocarditis Team needs to embrace a multidisciplinary approach that must adapt according to the patient's clinical needs and the local epidemiology to ensure prompt diagnosis and treatment.

The members of the Endocarditis Team should include the specialists with direct involvement in the diagnostic and therapeutic processes (Table 7), and may vary depending on the type of centre. In the

Table 7 Members of the Endocarditis Team

	Heart Valve Centre
Core members	• Cardiologists. • Cardiac imaging experts. • Cardiovascular surgeons. • Infectious disease specialist (or internal medicine specialist with expertise in infectious diseases). • Microbiologist. • Specialist in outpatient parenteral antibiotic treatment.
Adjunct specialties	• Radiologist and nuclear medicine specialist. • Pharmacologist. • Neurologist and neurosurgeon. • Nephrologist. • Anaesthesiologists. • Critical care. • Multidisciplinary addiction medicine teams. • Geriatricians. • Social worker. • Nurses. • Pathologist.

Recommendations for the Endocarditis Team

Diagnosis and management of patients with complicated IE are recommended to be performed at an early stage in a Heart Valve Centre, with immediate surgical facilities and an 'Endocarditis Team' to improve the outcomes.

I

B

For patients with uncomplicated IE managed in a Referring Centre, early and regular communication between the local and the Heart Valve Centre endocarditis teams is recommended to improve the outcomes of the patients.

I

B

İnfektif Endokardit: Bir ekip işi



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https://doi.org/10.1093/eurheartj/ehaf219

STATE OF THE ART REVIEW
Valvular heart disease

Infective endocarditis: it takes a team

Lawrence Lau^{1,*}, Larry Baddour², Nùria Fernández Hidalgo^{3,4,5}, Thomas D. Brothers^{6,7}, William K. F. Kong⁸, Michael A. Berger⁹, Xavier Duval^{10,11}, Christophe Tribouilloy¹², Jean-Francois Obadia¹³, Mehrdad Golian¹, Vicente F. Corrales-Medina¹⁴, Francois Auclair¹⁴, Mikael Mazighi¹⁵, Kwan Leung Chan¹, Bernard Prendergast¹⁶, Gilbert Habib¹⁷, Fraser D. Rubens¹⁸, and David Messika-Zeitoun^{1,*}

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Graphical Abstract

The endocarditis team

Core team

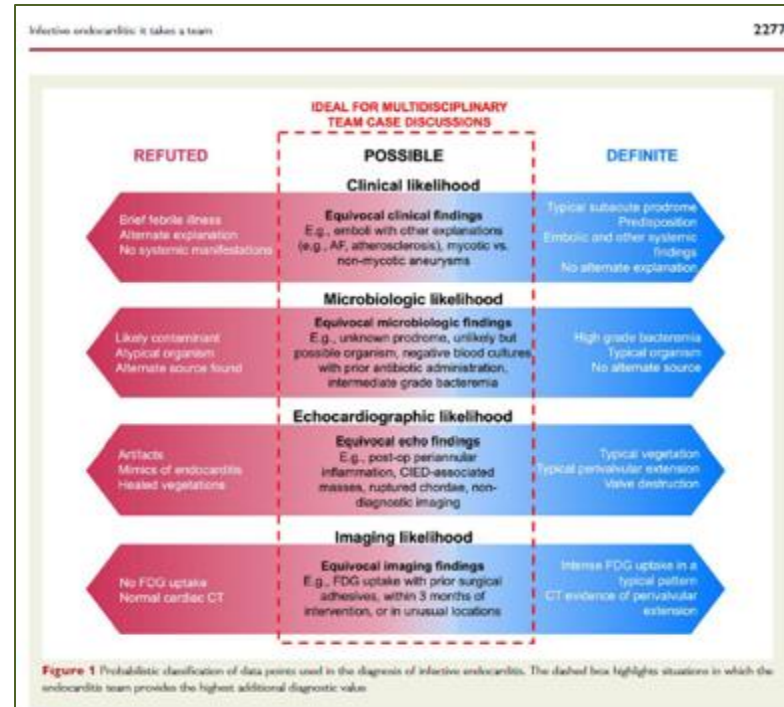
- Cardiologist (infectious expert)
- Infectious disease specialist
- Cardiac surgeon
- ECG/echocardiography specialist

Additional endocarditis team members as required

- Bacteriologist
- Radiologist
- Neurologist
- Congenital heart disease specialist
- Nuclear medicine specialist
- Pharmacist
- Intensivist
- Additional members as required by regional expertise, endocarditis
- Additional medicine specialist
- Social worker
- Hand hygienist

Overarching goals of the endocarditis team

- Patients and clinicians (diagnosis uncertainty)
- Define clinical suspicion
- Integrate multidisciplinary findings within the clinical context
- Address uncertainties in special populations with endocarditis
- Implementing care
- Adult congenital heart disease
- Neurological complications
- Prosthetic heart valves
- Device recommendations based on endocarditis team consensus
- Antibiotic type, duration, and peroral anti-thrombotic therapy
- Extraction, targeting, and replacement of cardiac implantable electronic devices
- Endocarditis or surgical treatment of neurologic complications
- Time and timing of cardiac surgery
- Prosthetic and/or pacemaker/ICD therapy



Conclusions

Infective endocarditis is a life-threatening infection characterized by both cardiac and extra-cardiac complications. The diagnosis, investigation and management of IE is increasingly complex and requires expertise in numerous disparate fields. Evidence suggests that ET may improve IE outcomes. Whereas many models of ET exist, flexibility to adapt to institutional and regional practices is paramount. The formation of an ET provides opportunities for mutual learning at a time when cardiovascular care is increasingly specialized and segregated and a framework for research and patient education in an area where these are often lacking.



Endokardit Ekibi Deneyimi

02837

Endocarditis team: Three years experience

02. Bacterial infection & disease

2b. Severe sepsis, bacteraemia & endocarditis (incl. epidemiology, diagnosis, host biomarkers, treatment and outcome prediction)

Likely attendance

Onsite

Nuran Sarı¹, Emir Karaçağlar², Elif Ateş¹, Bahadır Gültekin³, Seda Kibaroglu⁴, Ayşen Terzi⁵, Feride Kural Rahatlı⁶, Ayşe Aktaş⁷, Özlem Kurt Azap¹, Atilla Sezgin³

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Endokardit Ekibi

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<https://doi.org/10.1007/s44197-025-00413-x>

RESEARCH ARTICLE



Global Trends and Regional Differences in the Burden of Infective Endocarditis, 1990–2021: An Analysis of the Global Burden of Disease Study 2021

Huanhuan Miao¹ · Zhanyang Zhou¹ · Zheng Yin¹ · Xue Li¹ · Yuhui Zhang¹ · Yuqing Zhang¹ · Jian Zhang^{1,2}

4.1 Public Health Implications

The study on the burden of IE had significant public health implications. Given the global increase in the ASIR and varying burdens across regions and demographics, health-care resources must be reallocated. High-ASIR regions, especially those with high SDI, require more advanced diagnostic and treatment facilities, as well as more specialized cardiac care units. The involvement of a multidisciplinary Endocarditis Team is essential for the accurate diagnosis and comprehensive management of IE [18, 29]. While for the low SDI region, resources should be focused on improving basic healthcare infrastructure and access to treatment. Since males experienced a higher burden of IE compared to females, and the peak burden was shifting towards older individuals, resources should be targeted towards these demographics. In particular, screening programs for high-risk groups, such as elderly men with pre-existing heart conditions, should be enhanced. Additionally, the formulation of preventive measures is imperative. Launching public health initiatives to educate the population about the risk factors of IE, such as inadequate oral hygiene, intravenous drug usage, and underlying heart conditions, is crucial [30]. Moreover, the administration of antibiotic prophylaxis before invasive dental procedures is recommended as a preventive measure against IE in individuals at high risk [31, 32]. Future research should focus on clarifying risk factors across diverse regions and evaluating the efficacy of interventions to enhance the management and prevention of IE.

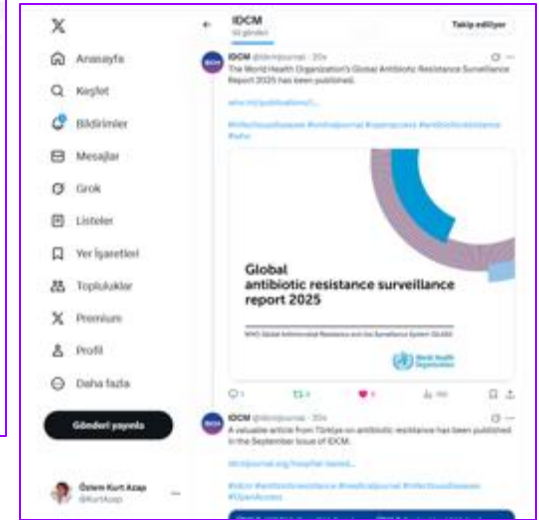
IDCM



The image shows the IDCM (Infectious Diseases and Clinical Microbiology) journal cover and its Indexes page. The cover features the journal title, volume information (SEP 2020, VOLUME 5, ISSUE 3), and a featured article titled "Risk Factors for Mortality in Patient Treated with Colistin Monotherapy for Carbapenem-Resistant *Acinetobacter baumannii* Infections" by Wilson Kulyk and others. The Indexes page lists various databases where the journal is indexed, including PubMed, Web of Science, Scopus, and others.



The image shows the IDCM Aims and Scope page. It defines the journal as an international, scientific, open access periodical published in accordance with independent, unbiased, and double blinded peer review principles. The journal is indexed in various databases and is published in English. The page also includes a section for "Aims and Scope" and a "Submit Online" button.



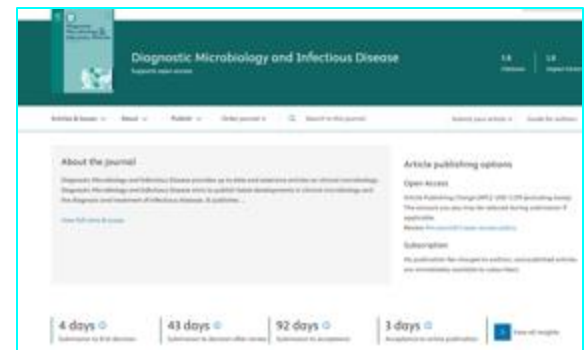
The image shows a Twitter post from IDCM (@idcmjournal). The post announces the publication of the "Global antibiotic resistance surveillance report 2025" by WHO Global Antibiotic Resistance and Use Surveillance System (GLASS). The post includes a link to the report and a mention of the journal's website.



The image shows the cover of the journal "Clinical Microbiology and Infection". The cover features the journal title, volume information (2020, Volume 21, Issue 1), and a featured article titled "Risk Factors for Mortality in Patient Treated with Colistin Monotherapy for Carbapenem-Resistant *Acinetobacter baumannii* Infections" by Wilson Kulyk and others. The cover also includes the ESCMID logo.



The image shows the cover of the journal "International Journal of Infectious Diseases". The cover features the journal title, volume information (2020, Volume 100, Issue 1), and a featured article titled "Risk Factors for Mortality in Patient Treated with Colistin Monotherapy for Carbapenem-Resistant *Acinetobacter baumannii* Infections" by Wilson Kulyk and others. The cover also includes the journal's website and a "Submit Online" button.



The image shows the cover of the journal "Diagnostic Microbiology and Infectious Disease". The cover features the journal title, volume information (2020, Volume 100, Issue 1), and a featured article titled "Risk Factors for Mortality in Patient Treated with Colistin Monotherapy for Carbapenem-Resistant *Acinetobacter baumannii* Infections" by Wilson Kulyk and others. The cover also includes the journal's website and a "Submit Online" button.