The 2024 European Society of Cardiology Guidelines for Diagnosis and Management of Atrial Fibrillation: A Viewpoint from a Practicing Clinician's Perspective

Tatiana Potpara^{1,2} Giulio F. Romiti³ Christian Sohns⁴

- ¹ Medical Faculty, University of Belgrade, Belgrade, Serbia
- ² Intensive Care for Arrhythmias, Cardiology Clinic, University Clinical Centre of Serbia, Belgrade, Serbia
- ³ Department of Trans¹ational and Precision Medicine, Sapienza University of Rome, Rome, Italy
- ⁴Clinic for Electrophysiology, Herz-und Diabeteszentrum NRW, Ruhr-Universität Bochum, Med. Fakultät OWL (Universität Bielefeld), Bad Oeynhausen, Germany

Thromb Haemost 2024;124:1087-1094.

Address for correspondence Tatjana Potpara, MD, PhD, University of Belgrade, Belgrade, Serbia Cardiology Clinic, University Clinical Centre of Serbia, Dr Subotica starijeg 13, 1100 Belgrade, Serbia (e-mail: tanjapotpara@qmail.com).

Abstract

Atrial fibrillation (AF) is a complex disease requiring a multidomain and (usually) longterm management, thus posing a significant burden to patients with AF, practitioners, and health care system. Unlike cardiovascular conditions with a narrow referral pathway (e.g., acute coronary syndrome), AF may be first detected by a wide range of specialties (often noncardiology) or a general practitioner. Since timely initiated optimal management is essential for the prevention of AF-related complications, a concise and simple quidance is essential for practitioners managing AF patients, regardless of their specialty. Guideline-adherent management of patients with AF has been shown to translate to improved patient outcomes compared with quidelinenonadherent treatment. To facilitate quideline implementation in routine clinical practice, a good guideline document on AF should introduce only evidence-based new recommendations, while avoiding arbitrary changes, which may be confusing to practitioners. Herein, we discuss the main changes in the 2024 European Society of Cardiology (ESC) AF Guidelines relative to the previous 2020 ESC document. Whether the updates and new recommendations issued by the new quidelines will translate in high adherence in clinical practice (and hence improved prognosis of patients with AF) will need to be addressed in upcoming years.

Keywords

- ► atrial fibrillation
- guidelines
- evidence-based recommendation

Atrial fibrillation (AF) is associated with increased risk of major cardiovascular adverse events, including ischemic stroke/systemic embolism, heart failure, hospitalization, impaired quality of life, and mortality. The arrhythmia is a complex disease requiring a multidomain, integrated, and (usually) long-term management, thus posing a significant

burden to patients with AF, practitioners, and health care system.

Unlike cardiovascular conditions with a narrow referral pathway (e.g., acute coronary syndrome [ACS]), AF may be first detected by a wide range of specialties (often noncardiology ones) or a general practitioner in primary care.

received
September 27, 2024
accepted after revision
October 4, 2024
accepted manuscript online
October 7, 2024
article published online
October 24, 2024

© 2024. Thieme. All rights reserved. Georg Thieme Verlag KG, Rüdigerstraße 14, 70469 Stuttgart, Germany DOI https://doi.org/ 10.1055/a-2434-9244. ISSN 0340-6245. Since timely initiated optimal management is essential for the prevention of AF-related complications, a concise and simple guidance is essential for practitioners managing AF patients, regardless of their specialty.

Guideline-adherent management of patients with AF has been shown to translate to improved patient outcomes compared with guideline-nonadherent treatment.² To facilitate guideline implementation in routine clinical practice, a good guideline document on AF should introduce only evidence-based new recommendations while avoiding arbitrary changes, which may be confusing to practitioners.

Herein we discuss the main changes in the 2024 European Society of Cardiology (ESC) AF Guidelines³ relative to the previous 2020 ESC document¹ and compare the 2024 ESC document with other most recent international AF guidelines.

Integrated Care for Patients with Atrial Fibrillation

Approximately a decade ago, the World Health Organization put forward the concept of integrated care models for chronic diseases in recognition of fragmentation of respective health care services. Thereafter, a structured, patient-centered, multidisciplinary approach to the management of patients with AF (integrating health care professionals, patients, and their family/carers and outlining the main domains of AF care) to improve patient outcomes and adherence to guidelines has been formally proposed in the 2016 ESC AF guidelines (Class IIa, Level of Evidence [LoE] B).

The 2020 ESC AF Guidelines reiterated this recommendation and streamlined the essential domains of care for AF patients across all health care levels and among different specialties into the simple ABC pathway (**Fig. 1**), using gear wheels to emphasize the equal importance of each of the main AF care domains, as follows: "A" Anticoagulation/Avoid stroke, "B" Better symptom management, and "C" Cardiovascular and Comorbidity optimization.¹

The scientific evidence supporting the ABC pathway at that time was already fairly extensive.⁶ There were several observational studies (from retrospective and prospective cohorts) or post hoc analyses of randomized trial cohorts showing a significant association of the ABC pathway implementation with lower health-related costs, lower rates of cardiovascular adverse events, and lower risk of allcause death and composite outcome of stroke/major bleeding/cardiovascular death and first hospitalization in comparison to usual care.8-10 There was also one published prospective cluster randomized mAFA-II trial, which showed a significant 61% risk reduction in the composite outcome of stroke or thromboembolism, all-cause death, and rehospitalization, with ABC pathway management intervention versus usual care. 11 The long-term extension of mAFA-II trial showed a high adherence (over 70%) and persistence (over 90%) with the intervention. 12

Subsequently, in a systematic review and meta-analysis of 285,000 patients, adherence to the ABC pathway translated to a 58% reduction in all-cause death, a 63% reduction in cardiovascular death, a 45% reduction in ischemic stroke, and a 31% reduction in major bleeding. A retrospective analysis of a large registry-based cohort showed that adherence to all ABC pathway domains resulted in the greatest magnitude of

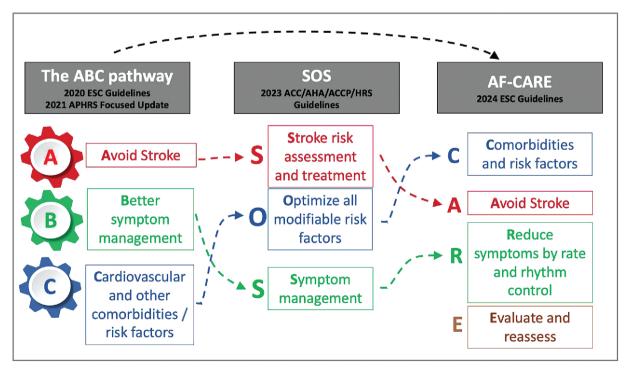


Fig. 1 Integrated management of patients with AF. ^{1,3,20} ACC/AHA/ACCP/HRS, American College of Cardiology/American Heart Association/American College of Chest Physicians/Heart Rhythm Society; AF, atrial fibrillation; ESC, European Society of Cardiology.

risk reduction and the longest event-free survival; also, in patients deemed as "clinically complex" and other analyses have shown the impact of the ABC pathway on patients with multimorbidity and thus at higher baseline risk of adverse outcomes. 15–17

Most recently, the randomized MIRACLE-AF trial was presented as a Late Breaking Trial at the 2024 ESC Congress in London and reported a cluster randomized trial comparison of ABC pathway intervention versus intensified usual care in rural villages in China—this showed a 36% lower rates of the composite outcome (cardiovascular death, stroke, hospitalization due to worsening of heart failure or ACS, and emergency visits due to AF) with the ABC intervention delivered by village doctors (previously called "barefoot doctors") supported by telemedicine (www.escardio.org/Congresses-Events/ESC-Congress/Congress-news/hot-line-9-strokestop-ii-guard-af-and-miracle-af). Secondary outcomes included a significant reduction in stroke and cardiovascular death.

Clearly, the evidence supporting the ABC pathway for integrated AF care to streamline timely optimal management of patients with AF at all health care levels by noncardiologists and cardiologists has been accumulated, fulfilling LoE A. In addition, an ongoing randomized controlled trial is comparing the ABC pathway versus usual care in elderly patients in Europe within the Horizon Europe funded the AFFIRMO programme. ¹⁹

Notwithstanding the significant amount of evidence supporting the ABC pathway, the 2024 ESC AF Guidelines recommended a new, not previously tested acronym AF-CARE, essentially highlighting the same AF care domains as the ABC pathway, although rearranged in a different order (Fig. 1). This change in recommendation, from ABC pathway to AF-CARE, was justified mainly by a concern that the "C" domain (Cardiovascular and Comorbidity management) could be otherwise neglected.

Whether this change, accompanied by a Class I LoE C formal recommendation, will really facilitate the attainment of the "C" domain of AF care, or rather confuse practitioners increasingly familiar with the ABC pathway and compromise guideline implementation in practice, remains to be seen. After all, when managing any cardiovascular disorder, whether AF or other non-AF conditions, it seems common sense that all cardiovascular risk factors and comorbidities should be proactively managed. Hence, the "C" being prioritized is not unique to AF per se.

Of note, the 2023 ACC/AHA/HRS AF Guidelines²⁰ also provided an acronym for integrated AF care, the SOS, streamlining the AF care domains comparably to the ABC pathway (**Fig. 1**).

Overall, the essential principles of care for AF patients worldwide remain the same, and using a particular acronym is probably only a matter of personal preference, as long as the main domains of AF care are optimally addressed.

Prevention of Stroke and Systemic Embolism

The steps essential to effective stroke prevention include: (1) initial stroke risk assessment to identify AF patients at truly

low risk of stroke, (2) initiation of oral anticoagulant (OAC) therapy in all AF patients with one or more stroke risk factors (preferably a non-Vitamin K antagonist oral anticoagulant [NOAC] in NOAC-eligible patients) and bleeding risk assessment, and (3) regular reassessment of stroke and bleeding risk in periodic time intervals, to account for a dynamic changes in the individual patient's risk profile.¹

When tailoring stroke prevention strategy, ethnic differences in stroke and bleeding risk should also be considered.^{21,22}

In the 2024 ESC AF Guidelines, several changes have been made, mostly regarding stroke and bleeding risk assessment.

Stroke Risk Assessment

Notwithstanding that clinical risk factor-based scores generally have a modest ability to predict the clinical event of interest, most international guidelines recommend the clinical stroke risk factor-based CHA₂DS₂-VASc score for initial stroke risk assessment (**Table 1**), as the most validated and widely used stroke risk assessment tool to reliably identify AF patients at sufficiently low risk of stroke so that long-term OAC is not needed (i.e., as long as the score is 0 in male and 1 in female AF patients).^{23,24}

The 2024 ESC AF Guidelines recommend using the CHA₂DS₂-VA score for stroke risk assessment (LoE C), considering that the inclusion of female sex "complicates clinical practice both for health care professionals and patients" and "omits individuals who identify as nonbinary, transgender, or are undergoing sex hormone therapy."³

Indeed, female sex is a stroke risk modifier, rather than a stroke risk factor per se.²⁹ While earlier data showed a greater risk of stroke in female AF patients compared with males (with significant age-dependent interaction between female sex and the presence of additional clinical stroke risk factors)^{30,31} and strokes tended to be more severe in female AF patients compared with males,³² more recent evidence shows that the rates of AF-related strokes are declining in both male and female patients, in the context of decreasing sex-related disparities in OAC use.

Similar observations were made by Nielsen et al in a nationwide cohort study of 158,982 patients with incident AF not on OAC.³³ During the study period 1997 to 2020, the risk of stroke overall has been declining in the last two decades, and the sex difference diminished in most recent years. Whereas the likelihood of prescribing OAC was lower for female patients with AF compared with male AF patients, OAC initiation increased over time, with comparable OAC initiation patterns in male and female AF patients.³³

In a study using UK primary and secondary care data comprising 195,719 patients with AF followed between 1998 and 2016, there was higher thromboembolic events in women compared with men in the population with high CHA₂DS₂-VASc risk scores; however, overall stroke and thromboembolic risk prediction using the CHA₂DS₂-VA and CHA₂DS₂-VASc scores was comparable. Also, the similarity in thromboembolic risk prediction using CHA₂DS₂-VA and CHA₂DS₂-VASc scores was consistent across different ethnicities and socioeconomic status.

Table 1 Thromboembolic and bleeding risk assessment and management in the international atrial fibrillation guidelines

Society	Year	Thromboembolic risk assessment model/score	Recommendation for thrombo-embolic prevention with OAC	Bleeding risk assessment and recommended model/score
NHFA/CSANZ ²⁵ (Australia, New Zealand)	2018	CHA ₂ DS ₂ -VA	$CHA_2DS_2-VA \ge 2$ (Strong)	Identification of reversible bleeding risk factors; no specific score recommended
			CHA ₂ DS ₂ -VA = 1 (Strong)	
APHRS ²⁶ (Asia-Pacific)	2021	CHA ₂ DS ₂ -VASc	CHA ₂ DS ₂ -VASc \geq 2 (males) or \geq 3 (females): recommended	HAS-BLED (to identify modifiable risk factors to be corrected)
			CHA_2DS_2 -VASc =1 (males) or 2 (females): to be considered	
CCS/CHRS ²⁷ (Canada)	2020	CHADS-65 ("CCS algorithm")	Score ≥ 1 (or 65 y) (Strong)	HAS-BLED (to identify high- risk patients and modifiable risk factors)
ESC/EACTS (Europe) ¹	2020	CHA ₂ DS ₂ -VASc	$ \begin{array}{l} CHA_2DS_2\text{-VASc} \geq 2 \; (males) \\ or \geq 3 \; (females) \\ (Class \; I) \end{array} $	HAS-BLED (to identify highrisk patients and address modifiable risk factors)
			CHA ₂ DS ₂ -VASc =1 (males) or =2 (females) (Class IIa)	
ESC/EACTS (Europe) ³	2024	CHA ₂ DS ₂ -VA	CHA_2DS_2 -VA ≥ 2 (Class I)	Assessment and management of modifiable bleeding risk factors; no specific score recommended
			CHA ₂ DS ₂ -VA =1 (Class IIa)	
ACC/AHA/ACCP/HRS (United States) ²⁰	2023	CHA ₂ DS ₂ -VASc (or validated clinical risk scores)	$ \begin{array}{l} CHA_2DS_2\text{-VASc} \geq & 2 \; (males) \\ or \geq & 3 \; (females) \\ (Class \; I) \end{array} $	Identify factors that indicate high risk of bleeding and possible intervention to prevent bleeding; no specific score recommended
			CHA ₂ DS ₂ -VASc = 1 (males) or 2 (females) (Class IIa)	
Chinese Expert Consensus Guidelines ²⁸	2024	CHA ₂ DS ₂ -VASc	CHA ₂ DS ₂ -VASc	HAS-BLED

Abbreviations: ACC, American College of Cardiology; ACCP, American College of Chest Physician AHA, American Heart Association; APHRS, Asia-Pacific Heart Rhythm Society; CCS, Canadian Cardiovascular Society; CHA₂DS₂-VASc, congestive heart failure, hypertension, age ≥75 years (2 points), diabetes mellitus, prior stroke/TIA/thromboembolism (2 points), vascular disease, age 65 to 74 years and female sex; CHS, Canadian Heart Rhythm Society; CSANZ, Cardiac Society of Australia and New Zealand; EACTS, European Association for Cardio-Thoracic Surgery; ESC, European Society of Cardiology; HRS, Heart Rhythm Society; NHFA, National Heart Foundation of Australia; OAC, oral anticoagulant.

A most recent retrospective evaluation of temporal trends in the predictive value of the CHA₂DS₂-VASc relative to the CHA₂DS₂-VA score (using nationwide data on AF patients from all levels of care in Finland during the 2007–2018 period) showed that initial differences favoring the CHA₂DS₂-VASc score in early years (when female AF patients were at much higher stroke risk than males) gradually attenuated over time, resulting in no difference in stroke risk prediction or reclassification between the CHA₂DS₂-VASc and CHA₂DS₂-VA scores in the 2017 to 2018 period.³⁴ Thus, recent data from Finland, Denmark, and the United Kingdom found the female to male differences in AF-related strokes were removing the Sc criterion from the CHA₂DS₂-VASc score did not affect its ability to discriminate thromboembolic events in the AF population.^{31,35,36}

Of note, an analysis from the same Finnish dataset and time period showed how female sex was initially associated with lower use of OAC, whereas sex-based disparities attenuated during the study and were finally resolved at the end of the observation.³⁷ Other studies have also shown an increase in the use of OAC among female patients over the last decade.³⁸ This evidence suggests that improved use of OAC in females may have contributed to the decreasing sex-based difference in the incidence of AF-related stroke.

Although the concept of not considering female sex in AF-related stroke risk assessment is not new (the CHA₂DS₂-VA score was first proposed in the 2018 Australian/New Zealand AF Guidelines³⁹ albeit with limited evidence then), the evidence supporting the CHA₂DS₂-VA score remained extremely scarce. Fortunately, the subsequently reported most recent data suggest that adopting the CHA₂DS₂-VA score could potentially simplify stroke risk assessment in AF patients.

Still, some caution is needed, as it is very likely that the use of CHA₂DS₂-VASc score contributed to draw attention to

the risk of stroke in women risk and improved OAC use in female AF patients, in addition to improved overall management of concomitant cardiovascular risk factors and underlying comorbidities. Also, the patterns seen in Finland, Denmark, and United Kingdom may not be evident in other health care systems. Hence, it could still be too early to replace the CHA₂DS₂-VASc score with CHA₂DS₂-VA when assessing the risk of stroke in AF patients.

Bleeding Risk Assessment

All international AF Guidelines recognize the need for bleeding risk assessment (and regular reassessment) in AF patients taking OAC and agree that the estimated bleeding risk itself should not preclude OAC prescription (**-Table 1**). However, the approach to bleeding risk assessment has varied over time in the ESC AF Guideline documents (**-Fig. 2**).

Bleeding risk factors are classified as nonmodifiable (e.g., age >65 years, prior stroke or bleeding), partially modifiable (e.g., renal impairment, anemia), and modifiable (e.g., hypertension, concomitant antiplatelet therapy, alcohol intake). In interaction with modifiable bleeding risk factors, nonmodifiable factors are important drivers of bleeding events, ⁴² hence should not be overlooked.

The importance of reviewing both modifiable and non-modifiable bleeding risk factors to mitigate bleeding risk has been acknowledged in most international AF Guidelines (**Table 1**), and the 2020 ESC AF Guidelines explicitly recommended a structured, clinical risk factor-based bleeding risk assessment (**Fig. 2**), since relying solely on modifiable bleeding risk consideration has been shown to be inferior to formal bleeding risk assessment using a bleeding risk score inclusive of both modifiable and nonmodifiable bleeding risk

factors, $^{43-45}$ such as the HAS-BLED score 23,36,45,46 (\sim **Table 1**, \sim **Fig. 2**).

From the practical perspective, any bleeding (major or minor) is "red flag" for subsequent ischemic events, vet OAC is often discontinued for the bleeding event.⁴⁷ Nevertheless, the guidance on consideration of individual patient bleeding risk in the 2024 ESC AF Guidelines may be confusing, especially for nonexpert clinicians managing AF patients. While the document mentions that patients with nonmodifiable bleeding risk factors should be reviewed more often, or even referred to a multidisciplinary team, the formal recommendation for bleeding risk assessment refers only to the assessment and management of modifiable bleeding risk factors (Class I, LoEB), while the use of bleeding risk scores is not recommended (Class III, LoE B), to avoid underuse of OAC.³ Of note, none of the three references cited in support of the latter examined the effects of bleeding risk scores on OAC underuse, 48,49 and one was the 2014 AHA/ACC/HRS AF Guideline document. 50

Transcatheter Left Atrial Appendage Closure

The evidence supporting nonpharmacological prevention of AF-related stroke using transcatheter left atrial appendage closure (LAAC)⁵¹ has not changed much since consideration of LAAC was recommended in AF patients with a high risk of stroke and contraindications to long-term OAC (Class IIb, LoE B) in the 2012 ESC AF Guideline Update;⁴¹ hence, the recommendation remained unchanged in the 2016 and 2020 ESC AF Guideline documents.^{1,5}

In the 2024 ESC AF Guidelines, the same recommendation is downgraded to LoE C, with the rationale that the available evidence does not refer to patients with contraindications to OAC.³ From the clinicians' practical perspective, the

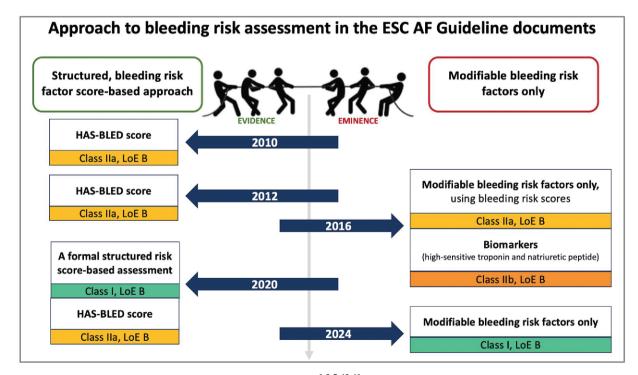


Fig. 2 Bleeding risk assessment in the ESC AF Guidelines 2010 to 2024. ^{1,3,5,40,41} AF, atrial fibrillation; ESC, European Society of Cardiology; HASBLED, Hypertension (uncontrolled, systolic blood pressure >160 mm Hg), Abnormal renal and/or hepatic function, Stroke, Bleeding history or predisposition, Elderly (>65 years), Drugs or excessive alcohol intake; LoE, level of evidence.

approach proposed in the 2023 ACC/AHA/HRS AF Guideline could be more helpful, as the recommendation referring to LAAC is divided to the recommendation on patients with a contraindication to long-term OAC (Class IIa, LoE B-NR) and another one referring to patients with a high risk of both stroke and bleeding (Class IIb, LoE B-R).²⁰

It is very likely that numerous ongoing randomized trials will change the LAAC landscape soon.⁵¹

Rhythm and Rate Control

It is widely accepted that appropriate rate control is an important background therapy in all AF patients. In addition, a large body of evidence supports the consideration of rhythm control in symptomatic patients with AF to improve symptoms and quality of life (Class I, LoE A in the 2020 ESC AF Guidelines and Class IIa, LoE B-R in the 2023 ACC/AHA/ACCP/HRS AF [American College of Cardiology/American Heart Association/American College of Chest Physicians/Heart Rhythm Society Atrial Fibrillation] Guidelines Duidelines Commendation is missing in the 2024 ESC AF Guideline document, being mentioned only in the text.

In line with recently published data,^{52,53} the 2024 ESC AF Guidelines recommended the implementation of a rhythm control strategy within 12 months of diagnosis in selected AF patients at risk of thromboembolism to reduce the risk of cardiovascular death or hospitalization (Class IIa, LoE B).³ However, how to select patients in practice is less clear.

In contrast, the 2023 ACC/AHA/ACCP/HRS AF Guidelines provide a helpful set of goals with rhythm control therapy, including: (1) evaluation of AF contribution to the reduced left ventricular (LV) function in patients with reduced LV function and persistent (high burden) AF (Class I, LoE B-R), (2) symptom improvement in patients with symptomatic AF, (3) reduction in hospitalization, stroke, and mortality in patients recently diagnosed with AF (<1 year), (4) improvement of symptoms and outcomes in patients with AF and heart failure (all Class IIa, LoE B-R), and (5) reduction in AF progression (Class IIa, LoE B-NR).²⁰

Catheter Ablation of Atrial Fibrillation

The recommendation for catheter ablation of AF as the first-line therapy for paroxysmal AF has been upgraded from Class IIa, LoE B¹ to Class I, LoE A³ in the 2024 ESC AF Guidelines, whereas the recommendations regarding AF ablation in patients with heart failure remained unchanged. This is in contrast to the 2023 ACC/AHA/ACCP/HRS AF Guidelines, wherein AF ablation is recommended in appropriate patients with AF and heart failure with reduced ejection fraction to improve symptoms, quality of life, ventricular function, and cardiovascular outcomes (Class I, LoE A).²0 A missed opportunity to upgrade the role of AF ablation in patients with heart failure in the 2024 ESC AF Guidelines could result in the therapy being delayed or withheld from patients who would most benefit from it.⁵4,55

While providing a new recommendation on repeat AF ablation (Class IIa, LoE B), the 2024 ESC AF Guidelines have not addressed AF ablation in asymptomatic AF patients, unlike the 2023 ACC/AHA/ACCP/HRS AF Guidelines where

AF ablation may be considered for reducing progression and complication of AF in younger patients with few comorbidities and moderate-to-high burden of AF (Class IIb, LoE B-NR).²⁰

Other Considerations

Optimal management of patients with so-called "subclinical" AF remains debatable, after the two randomized trials (i.e., ARTESiA and NOAH-AFNET 6) showed reduction in ischemic stroke, at the cost of increased risk of (nonfatal) major bleeding with NOAC versus control (either aspirin in ARTESiA, or placebo in NOAH-AFNET 6) in patients with subclinical AF of short duration. ^{56–58} The 2024 ESC AF Guidelines provided a Class Ilb, LoE B recommendation for considering an NOAC in such patients, excluding those at high risk of bleeding. ³ Questions remain on how to stratify thromboembolic risk and to individualize treatment strategies in these patients.

Concluding Remarks

Overall, the 2024 ESC AF Guidelines claimed 57 new recommendations, of which 17 (29%) were supported with LoE C. Of the latter, some appear rather unlikely to aid management of AF patients in daily practice (e.g., the Class I recommendation, LoE C that "a transthoracic echocardiogram is recommended in patients with an AF diagnosis where this will guide treatment decisions").

There is a strong impression that scientific evidence appreciation was rather unbalanced across some sections, ranging from shifting from an established approach with a significant amount of support evidence (e.g., the ABC pathway) to a new approach (i.e., AF-CARE), which is still to be validated, to meticulous scrutinization of current evidence (e.g., percutaneous LAAC). Whether the updates and new recommendations issued by the new guidelines will translate in high adherence in clinical practice (and hence improved prognosis of patients with AF) will need to be addressed in upcoming years, also taking into account the other changes proposed from previous guidelines (e.g., the ABC pathway vs. the new AF-CARE acronym, CHA₂DS₂-VASc vs. CHA₂DS₂-VA, and removal of the HAS-BLED score).

Clearly, the most striking aspect of the 2024 ESC AF Guideline document is the strong emphasis on the importance of concomitant comorbidity and risk factor management, supported by changing from the ABC pathway to AFCARE acronym. It remains to be seen whether this change will translate into better guideline implementation in practice and improved patients' outcomes, also considering current knowledge on barriers to guidelines implementations in clinical practice. However, it is simply common sense that all cardiovascular risk factors and comorbidities should be proactively managed, and regular review implemented, in patients with heart disease.

Conflict of Interest

T.P. declared no conflict of interest; G.F.R. reports consultancy for Boehringer Ingelheim and an educational grant

from Anthos, outside the submitted work. No fees were directly received personally.

Referencest

- 1 Hindricks G, Potpara T, Dagres N, et al; ESC Scientific Document Group. 2020 ESC Guidelines for the diagnosis and management of atrial fibrillation developed in collaboration with the European Association for Cardio-Thoracic Surgery (EACTS): The Task Force for the diagnosis and management of atrial fibrillation of the European Society of Cardiology (ESC) Developed with the special contribution of the European Heart Rhythm Association (EHRA) of the ESC. Eur Heart J 2021;42(05):373-498
- 2 Lip GY, Laroche C, Popescu MI, et al. Improved outcomes with European Society of Cardiology guideline-adherent antithrombotic treatment in high-risk patients with atrial fibrillation: a report from the EORP-AF General Pilot Registry. Europace 2015; 17(12):1777-1786
- 3 Van Gelder IC, Rienstra M, Bunting KV, et al; ESC Scientific Document Group. 2024 ESC Guidelines for the management of atrial fibrillation developed in collaboration with the European Association for Cardio-Thoracic Surgery (EACTS). Eur Heart J 2024;45(36):3314-3414
- 4 Bhat A, Khanna S, Chen HHL, et al. Integrated care in atrial fibrillation: a road map to the future. Circ Cardiovasc Qual Outcomes 2021;14(03):e007411
- 5 Kirchhof P, Benussi S, Kotecha D, et al; ESC Scientific Document Group, 2016 ESC guidelines for the management of atrial fibrillation developed in collaboration with EACTS. Eur Heart J 2016;37
- 6 Stevens D, Harrison SL, Kolamunnage-Dona R, Lip GYH, Lane DA. The atrial fibrillation better care pathway for managing atrial fibrillation: a review. Europace 2021;23(10):1511-1527
- 7 Luo X, Xu W, Ming WK, et al. Cost-effectiveness of mobile healthbased integrated care for atrial fibrillation: model development and data analysis. J Med Internet Res 2022;24(04):e29408
- 8 Proietti M, Romiti GF, Olshansky B, Lane DA, Lip GYH. Improved outcomes by integrated care of anticoagulated patients with atrial fibrillation using the simple ABC (atrial fibrillation better care) pathway. Am J Med 2018;131(11):1359-1366.e6
- 9 Proietti M, Lip GYH, Laroche C, et al; ESC-EORP Atrial Fibrillation General Long-Term Registry Investigators Group, Relation of outcomes to ABC (Atrial Fibrillation Better Care) pathway adherent care in European patients with atrial fibrillation: an analysis from the ESC-EHRA EORP Atrial Fibrillation General Long-Term (AFGen LT) Registry. Europace 2021;23(02):174-183
- 10 Romiti GF, Proietti M, Vitolo M, et al; ESC-EHRA EORP-AF General Long-Term Registry Investigators. Clinical complexity and impact of the ABC (atrial fibrillation better care) pathway in patients with atrial fibrillation: a report from the ESC-EHRA EURObservational Research Programme in AF General Long-Term Registry. BMC Med 2022;20(01):326
- 11 Guo Y, Lane DA, Wang L, et al; mAF-App II Trial Investigators. Mobile health technology to improve care for patients with atrial fibrillation. J Am Coll Cardiol 2020;75(13):1523-1534
- 12 Guo Y, Guo J, Shi X, et al; mAF-App II Trial investigators. Mobile health technology-supported atrial fibrillation screening and integrated care: a report from the mAFA-II trial long-term extension cohort. Eur J Intern Med 2020;82:105-111
- 13 Romiti GF, Pastori D, Rivera-Caravaca JM, et al. Adherence to the 'atrial fibrillation better care' pathway in patients with atrial fibrillation: impact on clinical outcomes-a systematic review and meta-analysis of 285,000 patients. Thromb Haemost 2022;122
- 14 Romiti GF, Proietti M, Bonini N, et al; GLORIA-AF Investigators. Adherence to the atrial fibrillation better care (ABC) pathway and the risk of major outcomes in patients with atrial fibrillation: a

- post-hoc analysis from the prospective GLORIA-AF Registry. EClinicalMedicine 2022;55:101757
- 15 Krittayaphong R, Winijkul A, Methavigul K, Chichareon P, Lip GYH. Clinical outcomes of patients with atrial fibrillation in relation to multimorbidity status changes over time and the impact of ABC pathway compliance: a nationwide cohort study. J Thromb Thrombolysis 2024
- Yao Y, Guo Y, Lip GYHmAF-App II Trial investigators. The effects of implementing a mobile health-technology supported pathway on atrial fibrillation-related adverse events among patients with multimorbidity: the mAFA-II randomized clinical trial. JAMA Netw Open 2021;4(12):e2140071
- Romiti GF, Guo Y, Corica B, Proietti M, Zhang H, Lip GYHmAF-App II trial investigators. Mobile health-technology-integrated care for atrial fibrillation: a win ratio analysis from the mAFA-II randomized clinical trial. Thromb Haemost 2023;123(11): 1042-1048
- Li M, Chu M, Shen Y, et al. A novel model of integrated care of older patients with atrial fibrillation in rural China. JACC Asia 2024
- Johnsen SP, Proietti M, Maggioni AP, Lip GYH. A multinational European network to implement integrated care in elderly multimorbid atrial fibrillation patients: the AFFIRMO consortium. Eur Heart | 2022;43(31):2916-2918
- 20 Joglar JA, Chung MK, Armbruster AL, et al; Peer Review Committee Members. 2023 ACC/AHA/ACCP/HRS guideline for the diagnosis and management of atrial fibrillation: a report of the American College of Cardiology/American Heart Association Joint Committee on clinical practice guidelines. Circulation 2024;149(01): e1-e156
- 21 Kang DS, Yang PS, Kim D, et al. Racial differences in ischemic and hemorrhagic stroke: an ecological epidemiological study. Thromb Haemost 2024;124(09):883-892
- 22 Kang DS, Yang PS, Kim D, et al. Racial differences in bleeding risk: an ecological epidemiological study comparing Korea and United Kingdom subjects. Thromb Haemost 2024;124(09):842-851
- Borre ED, Goode A, Raitz G, et al. Predicting thromboembolic and bleeding event risk in patients with non-valvular atrial fibrillation: a systematic review. Thromb Haemost 2018;118(12): 2171-2187
- 24 Potpara TS, Polovina MM, Licina MM, Marinkovic JM, Prostran MS, Lip GY. Reliable identification of "truly low" thromboembolic risk in patients initially diagnosed with "lone" atrial fibrillation: the Belgrade atrial fibrillation study. Circ Arrhythm Electrophysiol 2012;5(02):319-326
- Brieger D, Amerena J, Attia J, et al; NHFA CSANZ Atrial Fibrillation Guideline Working Group. National Heart Foundation of Australia and the Cardiac Society of Australia and New Zealand: Australian clinical guidelines for the diagnosis and management of atrial fibrillation 2018. Heart Lung Circ 2018;27(10):1209-1266
- 26 Chao TF, Joung B, Takahashi Y, et al. 2021 focused update consensus guidelines of the Asia Pacific Heart Rhythm Society on stroke prevention in atrial fibrillation: executive summary. Thromb Haemost 2022;122(01):20-47
- Andrade JG, Aguilar M, Atzema C, et al; Members of the Secondary Panel. The 2020 Canadian Cardiovascular Society/Canadian Heart Rhythm Society comprehensive guidelines for the management of atrial fibrillation. Can J Cardiol 2020;36(12):1847-1948
- Wang Y, Guo Y, Qin M, et al; Expert Reviewers. 2024 Chinese Expert Consensus Guidelines on the diagnosis and treatment of atrial fibrillation in the elderly, endorsed by Geriatric Society of Chinese Medical Association (Cardiovascular Group) and Chinese Society of Geriatric Health Medicine (Cardiovascular Branch): executive summary. Thromb Haemost 2024;124(10):897-911
- Nielsen PB, Skjøth F, Overvad TF, Larsen TB, Lip GYH. Female sex is a risk modifier rather than a risk factor for stroke in atrial fibrillation: should we use a CHA2DS2-VA score rather than CHA₂DS₂-VASc? Circulation 2018;137(08):832-840

- 30 Wagstaff AJ, Overvad TF, Lip GY, Lane DA. Is female sex a risk factor for stroke and thromboembolism in patients with atrial fibrillation? A systematic review and meta-analysis. QJM 2014;107(12):955–967
- 31 Corica B, Lobban T, True Hills M, Proietti M, Romiti GF. Sex as a risk factor for atrial fibrillation-related stroke. Thromb Haemost 2024;124(04):281–285
- 32 Lang C, Seyfang L, Ferrari J, et al; Austrian Stroke Registry Collaborators. Do women with atrial fibrillation experience more severe strokes? Results from the Austrian Stroke Unit Registry. Stroke 2017;48(03):778–780
- 33 Nielsen PB, Brøndum RF, Nøhr AK, Overvad TF, Lip GYH. Risk of stroke in male and female patients with atrial fibrillation in a nationwide cohort. Nat Commun 2024;15(01):6728
- 34 Teppo K, Lip GYH, Airaksinen KEJ, et al. Comparing CHA₂DS₂-VA and CHA₂DS₂-VASc scores for stroke risk stratification in patients with atrial fibrillation: a temporal trends analysis from the retrospective Finnish AntiCoagulation in Atrial Fibrillation (FinA-CAF) cohort. Lancet Reg Health Eur 2024;43:100967
- 35 Teppo K, Airaksinen KEJ, Jaakkola J, et al. Ischaemic stroke in women with atrial fibrillation: temporal trends and clinical implications. Eur Heart J 2024;45(20):1819–1827
- 36 Yoshimura HPR, Finan C, Schmidt AF, Lip GYH. Refining the CHA2DS2VASc risk stratification scheme: shall we drop the sex category criterion? Europace 2024; available at: https://doi.org/10.1093/europace/euae280
- 37 Teppo K, Airaksinen KEJ, Jaakkola J, et al. Temporal trends of gender disparities in oral anticoagulant use in patients with atrial fibrillation. Eur J Clin Invest 2024;54(01):e14107
- 38 Gadsbøll K, Staerk L, Fosbøl EL, et al. Increased use of oral anticoagulants in patients with atrial fibrillation: temporal trends from 2005 to 2015 in Denmark. Eur Heart J 2017;38(12):899–906
- 39 Brieger D, Amerena J, Attia JR, et al. National Heart Foundation of Australia and Cardiac Society of Australia and New Zealand: Australian clinical guidelines for the diagnosis and management of atrial fibrillation 2018. Med J Aust 2018;209(08):356–362
- 40 Camm AJ, Kirchhof P, Lip GY, et al; European Heart Rhythm Association European Association for Cardio-Thoracic Surgery. Guidelines for the management of atrial fibrillation: the Task Force for the Management of Atrial Fibrillation of the European Society of Cardiology (ESC). Eur Heart J 2010;31(19):2369–2429
- 41 Camm AJ, Lip GY, De Caterina R, et al; ESC Committee for Practice Guidelines (CPG) 2012 focused update of the ESC Guidelines for the management of atrial fibrillation: an update of the 2010 ESC Guidelines for the management of atrial fibrillation. Developed with the special contribution of the European Heart Rhythm Association. Eur Heart J 2012;33(21):2719–2747
- 42 Chao TF, Lip GYH, Lin YJ, et al. Incident risk factors and major bleeding in patients with atrial fibrillation treated with oral anticoagulants: a comparison of baseline, follow-up and delta HAS-BLED scores with an approach focused on modifiable bleeding risk factors. Thromb Haemost 2018;118(04):768–777
- 43 Esteve-Pastor MA, Rivera-Caravaca JM, Shantsila A, Roldán V, Lip GYH, Marín F. Assessing bleeding risk in atrial fibrillation patients: comparing a bleeding risk score based only on modifiable bleeding risk factors against the HAS-BLED Score. The AMADEUS trial. Thromb Haemost 2017;117(12):2261–2266
- 44 Guo Y, Zhu H, Chen Y, Lip GYH. Comparing bleeding risk assessment focused on modifiable risk factors only versus validated bleeding risk scores in atrial fibrillation. Am J Med 2018;131(02):185–192

- 45 Chao TF, Lip GYH, Lin YJ, et al. Major bleeding and intracranial hemorrhage risk prediction in patients with atrial fibrillation: attention to modifiable bleeding risk factors or use of a bleeding risk stratification score? A nationwide cohort study. Int J Cardiol 2018:254:157–161
- 46 Gorog DA, Gue YX, Chao TF, et al. Assessment and mitigation of bleeding risk in atrial fibrillation and venous thromboembolism: executive summary of a European and Asia-Pacific expert consensus paper. Thromb Haemost 2022;122(10):1625–1652
- 47 Winijkul A, Kaewkumdee P, Yindeengam A, Lip GYH, Krittayaphong R. Clinical outcomes of patients with atrial fibrillation who survived from bleeding event: the results from COOL-AF Thailand registry. Thromb Haemost 2024;124(11):991–1002
- 48 Hilkens NA, Algra A, Greving JP. Predicting major bleeding in ischemic stroke patients with atrial fibrillation. Stroke 2017;48 (11):3142–3144
- 49 Olesen JB, Lip GY, Lindhardsen J, et al. Risks of thromboembolism and bleeding with thromboprophylaxis in patients with atrial fibrillation: a net clinical benefit analysis using a 'real world' nationwide cohort study. Thromb Haemost 2011;106(04): 739–749
- 50 January CT, Wann LS, Alpert JS, et al; ACC/AHA Task Force Members. 2014 AHA/ACC/HRS guideline for the management of patients with atrial fibrillation: executive summary: a report of the American College of Cardiology/American Heart Association Task Force on practice guidelines and the Heart Rhythm Society. Circulation 2014;130(23):2071–2104
- 51 Potpara T, Grygier M, Häusler KG, et al. Practical guide on left atrial appendage closure for the non-implanting physician: an international consensus paper. Europace 2024;26(04): euae035
- 52 Kirchhof P, Camm AJ, Goette A, et al; EAST-AFNET 4 Trial Investigators. Early rhythm-control therapy in patients with atrial fibrillation. N Engl J Med 2020;383(14):1305–1316
- 53 Chao TF, Chan YH, Chiang CE, et al. Early rhythm control and the risks of ischemic stroke, heart failure, mortality, and adverse events when performed early (<3 months): a nationwide cohort study of newly diagnosed patients with atrial fibrillation. Thromb Haemost 2022;122(11):1899–1910
- 54 Sohns C, Fox H, Marrouche NF, et al; CASTLE HTx Investigators. Catheter ablation in end-stage heart failure with atrial fibrillation. N Engl J Med 2023;389(15):1380–1389
- 55 Marrouche NF, Brachmann J, Andresen D, et al; CASTLE-AF Investigators. Catheter ablation for atrial fibrillation with heart failure. N Engl J Med 2018;378(05):417–427
- Kirchhof P, Toennis T, Goette A, et al; NOAH-AFNET 6 Investigators NOAH-AFNET6 sites and investigators. Anticoagulation with edoxaban in patients with atrial high-rate episodes. N Engl J Med 2023;389(13):1167–1179
- 57 Healey JS, Lopes RD, Granger CB, et al; ARTESIA Investigators. Apixaban for stroke prevention in subclinical atrial fibrillation. N Engl J Med 2024;390(02):107–117
- 58 McIntyre WF, Benz AP, Becher N, et al. Direct oral anticoagulants for stroke prevention in patients with device-detected atrial fibrillation: a study-level meta-analysis of the NOAH-AFNET 6 and ARTESiA trials. Circulation 2024;149(13):981–988
- 59 Cabana MD, Rand CS, Powe NR, et al. Why don't physicians follow clinical practice guidelines? A framework for improvement. JAMA 1999;282(15):1458–1465