Age-related dynamic embolic risk in patients with nonvalvular atrial fibrillation

Riesgo embólico dinámico asociado con la edad en pacientes con fibrilación auricular no valvular

To the Editor,

According to the current European Society of Cardiology guidelines for patients with atrial fibrillation (AF), oral anticoagulation should be prescribed when the CHA_2DS_2 -VASc¹ score is greater than or equal to 2, provided that the patient does not have mitral stenosis, mechanical prostheses, or hypertrophic cardiomyopathy. This treatment is not recommended for men scoring 0 or women scoring 1.² As patient age is a factor incorporated in this scale (1 point is added for age > 65 years), a dilemma may arise when deciding whether to administer coagulation to patients approaching the age of 65 years who have had an episode of AF.

With the aim of analyzing the impact of starting oral anticoagulation at 65 years of age, we used data from the Cardio CHUVI-AF registry.^{3,4} From January 2014 to January 2018, the registry included 16 202 patients with a diagnosis of AF in the Vigo health care area. The study was approved by the local ethics committee, and because of its retrospective and anonymized nature, the researchers were exempted from obtaining informed consent from participants. Patients with mechanical prostheses, moderate-severe mitral stenosis, or hypertrophic cardiomyopathy

were excluded. All patients with a CHA₂DS₂-VASc score of 0 for men and 1 for women (n = 861; 5.3% of the total) were identified, and those reaching the age of 65 years during the follow-up period were included. Thus, the study population consisted of 389 AF patients whose CHA₂DS₂-VASc score during follow-up (8.1 ± 1.5 years) increased from 0 to 1 point (in the case of men) or from 1 to 2 points (in the case of women), because they reached the age of 65 years. The mean age of these patients was 60.9 ± 2.7 years and 149 (38.3%) were women.

Before reaching the age of 65 years, 187 patients (48.1%) were receiving anticoagulation therapy. After reaching that age, this figure increased to 261 patients (67.1%), indicating a 19.0% increase in the anticoagulation rate at the age of 65 years. The analysis of patients younger than 65 years showed that oral anticoagulation did not lead to a lower risk of ischemic stroke or a higher risk of bleeding. However, at age 65 years, an increase in ischemic stroke was observed in non-anticoagulated patients compared with those receiving this therapy, while bleeding rates were similar (figure 1). Due to the relatively small sample size and limited number of episodes, subgroup analysis by sex was not feasible.

These findings, in which anticoagulation increased by only 19% in AF patients with a CHA₂DS₂-VASc score of 0 or 1 upon reaching the age of 65 years, indicate that up to 32.9% of patients are not provided this therapy (approximately 1 in every 3 patients). This has meaningful prognostic implications: anticoagulation in patients reaching 65 years was associated with a significant reduction in the ischemic stroke rate, with no increase in the bleeding risk. These

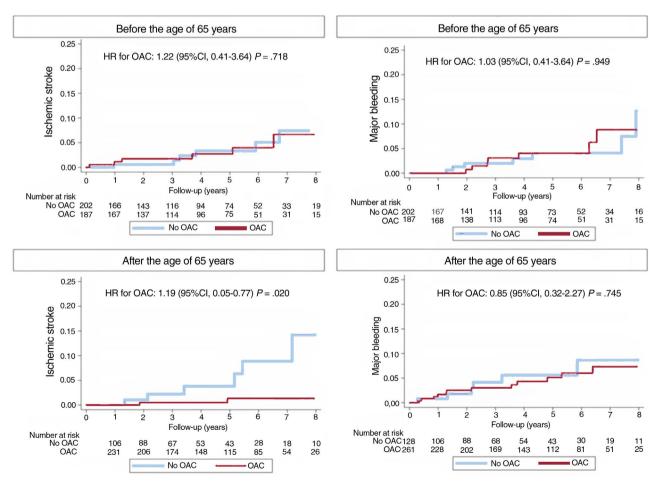


Figure 1. Ischemic stroke and major bleeding events in atrial fibrillation patients before and after the age of 65 years. OAC, oral anticoagulation; HR, hazard ratio; 95%CI, 95% confidence interval.

data suggest that there is a need for systematic reassessment of the indication for anticoagulation in these patients, either by considering initiation of this therapy around age 65 years or, ideally, scheduling follow-up visits at the time patients reach that age.

FUNDING

None.

ETHICAL CONSIDERATIONS

We accept the responsibility outlined by the International Committee of Medical Journal Editors. The study has been approved by the local ethics committee, which, given the retrospective and anonymized nature of the research, has waived the need for consent. Potential biases related to sex and gender have been taken into account.

STATEMENT ON THE USE OF ARTIFICIAL INTELLIGENCE

Artificial intelligence was not used.

AUTHORS' CONTRIBUTIONS

S. Raposeiras-Roubín, E. Abu-Assi, and A. Íñiguez-Romo designed the study; S. Raposeiras-Rubín and D. González-Fernández wrote the manuscript; D. González-Fernández, A. González-García, and C. Iglesias-Otero collected the data. All authors reviewed and approved the final draft.

CONFLICTS OF INTEREST

S. Raposeiras-Roubín has received presentation fees from the following companies: Amgen, Abbott, Sanofi, Novartis, AstraZeneca, Daichii, Pfyzer-BMS, Bayer, and Boehringer. The remaining authors declare no conflicts of interest.

C-reactive protein in patients with acute heart failure and preserved ejection fraction

Concentraciones de proteína C reactiva en pacientes con insuficiencia cardiaca aguda y fracción de eyección conservada

To the Editor,

In recent years, increasing evidence has emerged to support the distinct biological behavior of patients with heart failure (HF) in the upper ranges of left ventricular ejection fraction (LVEF). Indeed, among patients with HF and preserved ejection fraction (HFpEF), those with higher LVEF have been termed as having "supranormal" ejection fraction (HFsnEF).^{1–3}

There are well-known differences among patients with reduced, mid-range, and preserved ejection fraction.^{1,3} However, the factors associated with those patients with higher systolic function remain poorly understood. Along this line, heightened inflammatory activity has emerged as a crucial pathophysiological mechanism and potential therapeutic target in HFpEF.⁴ For instance, an ongoing trial is evaluating the efficacy of ziltivekimab

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Available online 15 December 2023

REFERENCES

- 1. Lip GY, Nieuwlaat R, Pisters R, Lane DA, Crijns HJ. Refining clinical risk stratification for predicting stroke and thromboembolism in atrial fibrillation using a novel risk factor-based approach: the euro heart survey on atrial fibrillation. *Chest.* 2010;137:263–272.
- 2. Hindricks G, Potpara T, Dagres N, et al. 2020 ESC Guidelines for the diagnosis and management of atrial fibrillation developed in collaboation 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;42:373–498.
- Ledo Piñeiro A, Raposeiras-Roubín S, Abu-Assi E, et al. Suboptimal anticoagulation with vitamin K antagonists: the need to change the national therapeutic positioning report. *Rev Esp Cardiol.* 2023;76:197–198.
- 4. Raposeiras-Roubín S, Abu-Assi E, Fernández Sanz T, et al. Bleeding and embolic risk in patients with atrial fibrillation and cancer. *Rev Esp Cardiol.* 2023;76:344–352.

https://doi.org/10.1016/j.recesp.2023.10.010

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vs placebo in patients with ambulatory HF, LVEF > 40%, and high-sensitivity C-reactive protein (hs-CRP) > 2 mg/L (NCT05636176). No prior studies have evaluated the inflammatory status profile along the continuum of LVEF, especially when LVEF \geq 50%. In this study, we aimed to examine whether circulating hs-CRP at presentation differs along the continuum of LVEF in patients with acute HF (AHF) and LVEF \geq 50%.

We conducted a retrospective study of an ongoing multicenter registry of patients admitted with AHF from January 2010 to January 2021 that enrolled 5246 patients. Patients with evidence of LVEF < 50% during hospitalization (n = 2433), evidence of infection at admission (n = 113), missing values of hs-CRP (n = 312), early deaths without assessment of LVEF (n = 38) were excluded from this analysis. No patients received inotropes at presentation. The final study sample included 2350 patients. Clinical and biochemical characteristics, including hs-CRP, were assessed at presentation. Echocardiographic assessments, including LVEF, were performed during hospitalization (72 \pm 24 hours after admission). LVEF was assessed by 2-dimensional echocardiography using the Simpson method. The association between hs-CRP and LVEF was evaluated along the continuum of LVEF or dichotomized (< 65% vs