

Obesity and Epicardial Fat Associated with Higher Atrial Fibrillation Recurrence After Ablation: Just Coincidence?

Cristiano F. Pisani¹  and Mauricio Scanavacca¹ 

Unidade Clínica de Arritmia do Instituto do Coração (InCor) do Hospital das Clínicas da FM USP (HC-FMUSP),¹ São Paulo, SP – Brazil

Short Editorial related to the article: *The Relationship between Epicardial Fat and Atrial Fibrillation Cannot Be Fully Explained by Left Atrial Fibrosis*

Atrial fibrillation (AF) is the most common sustained arrhythmia in clinical practice, and its prevalence increases with age.¹ It is a condition resulting from multiple pathophysiological aspects, not exclusively electrophysiological, from extrasystoles triggered by the pulmonary veins. It is associated with different cardiovascular and systemic diseases, obesity being a known risk factor.

Overweight and obese individuals are at greater risk for the occurrence of AF. For each unit of increment in BMI, the corrected risk for incidence of AF increases from 3 to 7% concerning individuals with a BMI below 25.^{2,3} The mechanisms that justify this observation are not fully understood and are probably multifactorial. Obese patients often have arterial hypertension⁴ and associated obstructive sleep apnea, factors already considered predisposing to AF.⁵ A recent study demonstrated that implementing a weight control program has an important impact on the clinical control of this arrhythmia.⁶

Thanassoulis et al. highlighted an association between the extent of epicardial fat and greater occurrence of AF (odds ratio = 1.28 per standard deviation of epicardial fat volume). However, this observation does not apply to the amount of intrathoracic and abdominal fat.⁷ Therefore, this association cannot be considered exclusively by the existence of obesity and its consequences. One of the justifications is the atrial inflammation induced by epicardial fat. AF is an arrhythmia associated with increased inflammatory markers, such as C-reactive protein, interleukins and tumor necrosis factor.⁸ These substances can be produced by the atrial adipose tissue, causing inflammation in the adjacent myocardium. Epicardial fat contains higher inflammatory cells levels than subcutaneous fat, including mast cells, macrophages, and lymphocytes.^{9,10}

In an experimental study, 10 sheep fed a hypercaloric diet for 36 weeks were compared with a control group with conventional feeding. The animals were submitted to an electrophysiological study and electroanatomical mapping

of the left atrium. Obese sheep presented greater LA volume and pressure, greater electrical conduction heterogeneity, increased amount of fractionated atrial electrograms, reduced voltage of electrograms in the posterior LA wall and greater voltage heterogeneity, suggesting the presence of an atrial fibrotic substrate such as a possible determinant of this association between obesity and AF.¹¹

In this issue of *Arquivos Brasileiros de Cardiologia*, Matos et al.,¹² studied 68 patients who had undergone coronary CT angiography, cardiac magnetic resonance imaging with 3D enhancement (using ADAS 3D software - Galgo Medical) before a first ablation procedure of atrial fibrillation. They observed a weak but statistically significant correlation between atrial fibrosis on electroanatomical mapping and epicardial adipose tissue measured on tomography in a slice at the left main coronary artery (TAE_{TC}) - Spearman correlation coefficient = 0.40, p=0.001.

Patients were followed up for a median period of 22 months (IQI 12-31), with AF recurrence in 31 patients (46%). Those who had recurrence were more likely to have non-paroxysmal AF, had larger atrial volume and larger volumes of adipose tissue and atrial fibrosis. However, in the multivariate analysis, only TAE_{TC} and non-paroxysmal AF were associated with greater recurrence. This observation is interesting and differs from previous studies,¹³⁻¹⁵ in which recurrence was related to the extent of atrial fibrosis.

These findings corroborate the importance of epicardial fat not only in the generation of fibrosis but also as an inflammatory factor. Furthermore, autonomic influences could contribute to a greater arrhythmia recurrence, as the ganglion plexuses are located in the epicardial fat, which can even be targeted in AF ablation.¹⁶⁻¹⁸

Despite the growing understanding of the pathophysiological mechanisms involved in the occurrence of AF and the constant technical improvement of catheter ablation procedures, many aspects are still not well understood. Among them is its association with obesity and atrial epicardial fat extension.

Keywords

Atrial Fibrillation; Obesity; Catheter Ablation.

Mailing address: Cristiano F. Pisani •

Unidade Clínica de Arritmia do Instituto do Coração (InCor) do Hospital das Clínicas da FM USP (HC-FMUSP) – Av. Dr. Enéas Carvalho de Aguiar, 44.
Postal Code 05403-000, São Paulo, SP – Brazil
E-mail: c.pisani@hc.fm.usp.br

DOI: <https://doi.org/10.36660/abc.20220103>

References

1. Hindricks G, Potpara T, Dagres N, Arbelo E, Bax JJ, Blomström-Lundqvist C, et al. 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(5):373-98. doi: 10.1093/eurheartj/ehaa612.
2. Dublin S, French B, Glazer NL, Wiggins KL, Lumley T, Psaty BM, et al. Risk of New-Onset Atrial Fibrillation in Relation to Body Mass Index. *Arch Intern Med*. 2006;166(21):2322-8. doi: 10.1001/archinte.166.21.2322.
3. Huxley RR, Misialek JR, Agarwal SK, Loehr LR, Soliman EZ, Chen LY, et al. Physical Activity, Obesity, Weight Change, and Risk of Atrial Fibrillation: The Atherosclerosis Risk in Communities study. *Circ Arrhythm Electrophysiol*. 2014;7(4):620-5. doi: 10.1161/CIRCEP.113.001244.
4. Kannel WB, Wolf PA, Benjamin EJ, Levy D. Prevalence, Incidence, Prognosis, and Predisposing Conditions for Atrial Fibrillation: Population-Based Estimates. *Am J Cardiol*. 1998;82(8A):2N-9N. doi: 10.1016/s0002-9149(98)00583-9.
5. Gami AS, Friedman PA, Chung MK, Caples SM, Somers VK. Therapy Insight: Interactions between Atrial Fibrillation and Obstructive Sleep Apnea. *Nat Clin Pract Cardiovasc Med*. 2005;2(3):145-9. doi: 10.1038/ncpcardio0130.
6. Middeldorp ME, Pathak RK, Meredith M, Mehta AB, Elliott AD, Mahajan R, et al. PREvention and regReSsive Effect of weight-loss and Risk Factor Modification on Atrial Fibrillation: The REVERSE-AF Study. *Europace*. 2018;20(12):1929-35. doi: 10.1093/europace/euy117.
7. Thanassoulis G, Massaro JM, O'Donnell CJ, Hoffmann U, Levy D, Ellinor PT, et al. Pericardial Fat is Associated with Prevalent Atrial Fibrillation: The Framingham Heart Study. *Circ Arrhythm Electrophysiol*. 2010;3(4):345-50. doi: 10.1161/CIRCEP.109.912055.
8. Hu YF, Chen YJ, Lin YJ, Chen SA. Inflammation and the Pathogenesis of Atrial Fibrillation. *Nat Rev Cardiol*. 2015;12(4):230-43. doi: 10.1038/nrcardio.2015.2.
9. Mazurek T, Zhang L, Zalewski A, Mannion JD, Diehl JT, Arafat H, et al. Human Epicardial Adipose Tissue is a Source of Inflammatory Mediators. *Circulation*. 2003;108(20):2460-6. doi: 10.1161/01.CIR.0000099542.57313.C5.
10. Shaihov-Teper O, Ram E, Ballan N, Brzezinski RY, Naftali-Shani N, Masoud R, et al. Extracellular Vesicles From Epicardial Fat Facilitate Atrial Fibrillation. *Circulation*. 2021;143(25):2475-93. doi: 10.1161/CIRCULATIONAHA.120.052009.
11. Mahajan R, Lau DH, Brooks AG, Shipp NJ, Manavis J, Wood JP, et al. Electrophysiological, Electroanatomical, and Structural Remodeling of the Atria as Consequences of Sustained Obesity. *J Am Coll Cardiol*. 2015;66(1):1-11. doi: 10.1016/j.jacc.2015.04.058.
12. Matos D, Ferreira AM, Freitas P, Rodrigues G, Carmo J, Costa F, et al. The Relationship between Epicardial Fat and Atrial Fibrillation Cannot Be Fully Explained by Left Atrial Fibrosis. *Arq Bras Cardiol*. 2022; 118(4):737-742. doi: 10.36660/abc.20201083.
13. Marrouche NF, Wilber D, Hindricks G, Jais P, Akoum N, Marchlinski F, et al. Association of Atrial Tissue Fibrosis Identified by Delayed Enhancement MRI and Atrial Fibrillation Catheter Ablation: The DECAAF Study. *JAMA*. 2014;311(5):498-506. doi: 10.1001/jama.2014.3.
14. Correia ETO, Barbeta LMDS, Mesquita ET. Extent of Left Atrial Ablation Lesions and Atrial Fibrillation Recurrence after Catheter Ablation - A Systematic Review and Meta-Analysis. *Arq Bras Cardiol*. 2020;114(4):627-35. doi: 10.36660/abc.20180378.
15. Pisani CF, Scanavacca M. Is Magnetic Resonance Imaging Already an Appropriate Method for Evaluating Patients after Atrial Fibrillation Catheter Ablation? *Arq Bras Cardiol*. 2020;114(4):636-7. doi: 10.36660/abc.20200204.
16. Zhou M, Wang H, Chen J, Zhao L. Epicardial Adipose Tissue and Atrial Fibrillation: Possible Mechanisms, Potential Therapies, and Future Directions. *Pacing Clin Electrophysiol*. 2020;43(1):133-145. doi: 10.1111/pace.13825.
17. Scanavacca M, Pisani CF, Hachul D, Lara S, Hardy C, Darrieux F, et al. Selective Atrial Vagal Denervation Guided by Evoked Vagal Reflex to Treat Patients with Paroxysmal Atrial Fibrillation. *Circulation*. 2006;114(9):876-85. doi: 10.1161/CIRCULATIONAHA.106.633560.
18. Katritsis DG, Pokushalov E, Romanov A, Giazitzoglou E, Siontis GC, Po SS, et al. Autonomic Denervation Added to Pulmonary Vein Isolation for Paroxysmal Atrial Fibrillation: A Randomized Clinical Trial. *J Am Coll Cardiol*. 2013;62(24):2318-25. doi: 10.1016/j.jacc.2013.06.053.



This is an open-access article distributed under the terms of the Creative Commons Attribution License