### COMMENTARY

## WILEY

# Atrial fibrillation and 24-h ambulatory blood pressure pattern

## Willem J. Verberk PhD

CARIM School for Cardiovascular Diseases, Maastricht University, Maastricht, the Netherlands

Revised: 25 May 2022

#### Correspondence

Willem J. Verberk, CARIM School for Cardiovascular Diseases, Maastricht University, Maastricht, the Netherlands. Email: willem.verberk@microlife.ch

### 1 | INTRODUCTION

This issue of the journal features a paper on the possibility to predict atrial fibrillation (AF) based on the 24-h blood pressure (BP)  $pattern^1$ 

AF is the most common sustained cardiac arrhythmia and an important risk factor for ischemic stroke.<sup>2</sup> It is estimated that approx 25% of all ischemic strokes are caused by AF and these strokes are often more severe than non-AF related strokes with a higher risk of death and approximately 70% disability at discharge.<sup>3</sup>The population average of AF is estimated at 1-2% of the general population<sup>4,5</sup> but it increases quickly with age to 5% in persons over 65 years and 14% in those over 85 years<sup>6,7</sup>. However, there is evidence that the true prevalence of AF is much higher<sup>8,9</sup>. As an example, Turakhia and associates estimated that 13.1% of all AF cases in US are undiagnosed, an estimation based on a back-calculation methodology among patients identified with incident AF following an ischemic stroke in a 6-year time period.<sup>10</sup> The reasons why AF often remains undetected are straightforward; Approximately one-third of persons with AF have no clear symptoms,<sup>11,12</sup> but even in symptomatic cases, it can be difficult to detect it because, in the case of paroxysmal AF (pxAF), episodes of the arrhythmia may be of short duration<sup>11,13</sup> and not present at the time of EKG performance.

This all has major consequences for patients and society with AFrelated costs having a significant contribution to overall healthcare costs, mainly due to treatment costs for stroke. These costs will exponentially increase unless AF is detected early and effectively treated.<sup>5</sup> On the contrary, AF detection at an early stage followed by the right treatment can reduce the risk of stroke by 64%<sup>14</sup>. This means that any tool that could help to detect AF or predict the occurrence of AF should be carefully considered.

Because nowadays 24-h ambulatory blood pressure monitoring (ABPM) is recommended by most hypertension guidelines for a.o. diag-

nosis of hypertension, a possibility to predict AF based on an ABPM pattern might dramatically improve AF detection. This is supported by the facts that hypertension increases the risk of AF by two-fold and that the majority of AF patients appear to have hypertension.<sup>15</sup> This means that, if hypertension guidelines are followed,<sup>16,17</sup> many patients at high risk of AF will undergo ABPM.

From ABPM it is well known that, particularly night-time blood pressure (BP), is an important cardiovascular risk predictor<sup>18</sup> and reverse dipping, a higher BP at night than during daytime, has been associated with the poorest cardiovascular prognosis.<sup>19</sup>

For the paper in the present issue, the authors performed a study among 412 patients. After a mean follow-up period of almost five years, seven patients developed AF. It appeared that 24-h systolic BP, night-time systolic BP, and presentation of reverse dipping were independently associated with new-onset AF.

The fact that 24-h and night systolic BP were independently associated with new-onset AF should not come as a surprise due to the well-known relationship between hypertension and AF. However, as such a relationship with AF could not be found for office systolic and diastolic BP this suggests that office BP measurement has limited value in diagnosing hypertension. At the same time, this reveals a study limitation as the authors did not use ABPM for the diagnosis of hypertension. The study participants included were classified as hypertensive when having elevated BP in two consecutive visits. Therefore, some white-coat hypertensives might have been included, whereas at the same time masked hypertensives were excluded. In other words, the included patients population might not totally represent true hypertensives. The authors mentioned in their paper that all patients received antihypertensive medication. However, as is well known, office BP measurements may also lead to unnecessary treatment.<sup>20</sup>

The authors found that reverse dipping was the strongest independent predictor for early onset AF. This finding adds another potential

This is an open access article under the terms of the Creative Commons Attribution-NonCommercial-NoDerivs License, which permits use and distribution in any medium, provided the original work is properly cited, the use is non-commercial and no modifications or adaptations are made. © 2022 The Authors. *The Journal of Clinical Hypertension* published by Wiley Periodicals LLC.

value to reverse dipping and is directly or indirectly supported by other findings; Domenech and associates demonstrated in a small patient study that night-time BP is associated with atrial remodeling,<sup>21</sup> which may eventually lead to AF. Another study showed that sustained nondipping hypertensive patients had a two-fold greater risk of developing AF than dippers.<sup>22</sup> In addition, reverse dipping showed to be independently associated with obstructive sleep apnea syndrome (OSAS),<sup>23</sup> whereas approximately one-third of AF patients appeared to have OSAS.<sup>24,25</sup>

For interpretation of the outcome of the paper, some other items should also be considered; Because the aim of Huang and associates was to investigate AF incidence, patients with a history of AF had to be excluded from the study. However, some patients might have had pxAF which was not yet diagnosed by the time of inclusion and not present at the moment of 12-lead EKG measurement. The same applies to the follow-up measurement. Only if symptoms and/or signs of AF were identified, Holter EKG monitoring was performed. However, since between 25% and 65% of all AF cases are estimated to be pxAF,<sup>26</sup> which are often without symptoms, the chance that the presence of pxAF was missed is high. Seen in that perspective it would have been better if all patients had received 24-h Holter monitoring, although this is costly, less practical and neither guarantees the finding of all pxAF patients.<sup>27</sup>

Finally, another matter of concern is the relatively small sample size and the very small numbers of patients with incident AF. However, as ABPM is implemented more and more in general clinical practice on a global scale, more information may be expected soon regarding the association of ABPM pattern and the risk of AF.

## CONFLICT OF INTEREST

WJV is an employee of Microlife AG, Switzerland.

#### REFERENCES

- Huang PH, Huang CC, Lin SJ, Chen JW. Prediction of Atrial Fibrillation in Patients with Hypertension: a Comprehensive Comparison of Office and Ambulatory Blood Pressure Measurements. J Clin Hypertens. In press.
- Camm AJ, Lip GY, De Caterina R, et al. 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. *Europace*. 2012;14(10):1385–1413. https://doi. org/10.1093/europace/eus305
- Saposnik G, Gladstone D, Raptis R, Zhou L, Hart RG. Investigators of the Registry of the Canadian Stroke Network (RCSN) and the Stroke Outcomes Research Canada (SORCan) Working Group. Atrial fibrillation in ischemic stroke: predicting response to thrombolysis and clinical outcomes. *Stroke*. 2013;44(1):99–104. https://doi.org/10. 1161/STROKEAHA.112.676551
- Stewart S, Hart CL, Hole DJ, McMurray JJ. Population prevalence, incidence, and predictors of atrial fibrillation in the Renfrew/Paisley study. *Heart*. 2001;86(5):516–521.
- Camm AJ, Kirchhof P, Lip GY, et al. 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. https://doi.org/10.1093/eurheartj/ ehq278

- Majeed A, Moser K, Carroll K. Trends in the prevalence and management of atrial fibrillation in general practice in England and Wales, 1994–1998: analysis of data from the general practice research database. *Heart*. 2001;86(3):284–288.
- Naccarelli GV, Varker H, Lin J, Schulman KL. Increasing prevalence of atrial fibrillation and flutter in the United States. *Am J Cardiol.* 2009;104(11):1534–1539. https://doi.org/10.1016/j.amjcard. 2009.07.022
- Gladstone DJ, Spring M, Dorian P, et al. Atrial fibrillation in patients with cryptogenic stroke. N Engl J Med. 2014;370(26):2467–2477. https://doi.org/10.1056/NEJMoa1311376
- Sanna T, Diener HC, Passman RS, et al. Cryptogenic stroke and underlying atrial fibrillation. N Engl J Med. 2014;370(26):2478–2486. https:// doi.org/10.1056/NEJMoa1313600
- Turakhia MP, Shafrin J, Bognar K, et al. Estimated prevalence of undiagnosed atrial fibrillation in the United States. *PLOS ONE*. 2018; 13(4):e0195088. https://doi.org/10.1371/journal.pone.0195088
- 11. Savelieva I, Camm AJ. Clinical relevance of silent atrial fibrillation: prevalence, prognosis, quality of life, and management. *J Interv Card Electrophysiol*. 2000;4(2):369–382.
- Furberg CD, Psaty BM, Manolio TA, Gardin JM, Smith VE, Rautaharju PM. Prevalence of atrial fibrillation in elderly subjects (the Cardiovascular Health Study). *Am J Cardiol*. 1994;74(3):236–241.
- Lau YC, Lane DA, Lip GY. Atrial fibrillation in cryptogenic stroke: look harder, look longer, but just keep looking. *Stroke*. 2014;45(11):3184– 3185. https://doi.org/10.1161/strokeaha.114.006862
- Hart RG, Pearce LA, Aguilar MI. Meta-analysis: antithrombotic therapy to prevent stroke in patients who have nonvalvular atrial fibrillation. *Ann Intern Med.* 2007;146(12):857–867.
- 15. Manolis AJ, Rosei EA, Coca A, et al. Hypertension and atrial fibrillation: diagnostic approach, prevention and treatment. Position paper of the Working Group "Hypertension Arrhythmias and Thrombosis" of the European Society of Hypertension. J Hypertens. 2012;30(2):239–252. https://doi.org/10.1097/HJH.0b013e32834f03bf
- Stergiou GS, Palatini P, Parati G, et al. 2021 European Society of Hypertension practice guidelines for office and out-of-office blood pressure measurement. J Hypertens. 2021;39(7):1293–1302. https:// doi.org/10.1097/HJH.00000000002843
- Whelton PK, Carey RM, Aronow WS, et al. 2017 ACC/AHA/AAPA/ ABC/ACPM/AGS/APhA/ASH/ASPC/NMA/PCNA Guideline for the Prevention, Detection, Evaluation, and Management of High Blood Pressure in Adults: executive Summary: a Report of the American College of Cardiology/American Heart Association Task Force on Clinical Practice Guidelines. J Am Coll Cardiol. 2018;71(19):2199–2269. https://doi.org/10.1016/j.jacc.2017.11.005
- Fagard RH, Celis H, Thijs L, et al. Daytime and nighttime blood pressure as predictors of death and cause-specific cardiovascular events in hypertension. *Hypertension*. 2008;51(1):55–61. https://doi.org/10. 1161/HYPERTENSIONAHA.107.100727
- Cuspidi C, Sala C, Tadic M, et al. Clinical and prognostic significance of a reverse dipping pattern on ambulatory monitoring: an updated review. J Clin Hypertens. 2017;19(7):713–721. https://doi.org/ 10.1111/jch.13023
- Verberk WJ, Kroon AA, Lenders JWM, et al. Self-measurement of blood pressure at home reduces the need for antihypertensive drugs: a randomized, controlled trial. *Hypertens Dallas Tex* 1979. 2007; 50(6):1019–1025. https://doi.org/10.1161/HYPERTENSIONAHA. 107.094193
- Doménech M, Berruezo A, Molina I, Mont L, Coca A. Nighttime ambulatory blood pressure is associated with atrial remodelling and neurohormonal activation in patients with idiopathic atrial fibrillation. *Rev Espanola Cardiol Engl Ed.* 2013;66(6):458–463. https://doi.org/10. 1016/j.rec.2012.11.011
- 22. Pierdomenico SD, Lapenna D, Cuccurullo F. Risk of atrial fibrillation in dipper and nondipper sustained hypertensive patients.

Blood Press Monit. 2008;13(4):193–197. https://doi.org/10.1097/MBP. 0b013e3282feea70

850

- Genta-Pereira DC, Furlan SF, Omote DQ, et al. Nondipping Blood Pressure Patterns Predict Obstructive Sleep Apnea in Patients Undergoing Ambulatory Blood Pressure Monitoring. *Hypertension*. 2018;72(4):979–985. https://doi.org/10.1161/HYPERTENSIONAHA. 118.11525
- 24. Marulanda-Londoño E, Chaturvedi S. The Interplay between Obstructive Sleep Apnea and Atrial Fibrillation. *Front Neurol.* 2017;8:668. https://doi.org/10.3389/fneur.2017.00668
- Calkins H, Hindricks G, Cappato R, et al. 2017 HRS/EHRA/ECAS/ APHRS/SOLAECE expert consensus statement on catheter and surgical ablation of atrial fibrillation. *Heart Rhythm*. 2017;14(10):e275e444. https://doi.org/10.1016/j.hrthm.2017.05.012

- Lip GY, Hee FL. Paroxysmal atrial fibrillation. QJM. 2001;94(12):665– 678.
- Gahungu N, Trueick R, Coopes M, Gabbay E. Paroxysmal atrial fibrillation. BMJ. 2021;375:e058568. https://doi.org/10.1136/bmj-2021-058568

How to cite this article: Verberk WJ. Atrial fibrillation and 24-h ambulatory blood pressure pattern. *J Clin Hypertens*. 2022;24:848–850. https://doi.org/10.1111/jch.14522