Definition and management of arrhythmia-induced cardiomyopathy: findings from the European Heart Rhythm Association survey

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Aims
Arrhythmia-induced cardiomyopathy (AiCM) represents a subtype of acute heart failure (HF) in the context of sustained arrhythmia. Clear definitions and management recommendations for AiCM are lacking. The European Heart Rhythm Association Scientific Initiatives Committee (EHRA SIC) conducted a survey to explore the current definitions and management of patients with AiCM among European and non-European electrophysiologists.

Methods and results
A 25-item online questionnaire was developed and distributed among EP specialists on the EHRA SIC website and on social media between 4 September and 5 October 2023. Of the 206 respondents, 16% were female and 61% were between 30 and 49 years old. Most of the respondents were EP specialists (81%) working at university hospitals (47%). While most participants (67%) agreed that AiCM should be defined as a left ventricular ejection fraction (LVEF) impairment after new onset of an arrhythmia, only 35% identified a specific LVEF drop to diagnose AiCM with a wide range of values (5–20% LVEF drop). Most respondents considered all available therapies: catheter ablation (93%), electrical cardioversion (83%), antiarrhythmic drugs (76%), and adjuvant HF treatment (76%). A total of 83% of respondents indicated that adjuvant HF treatment should be started at first HF diagnosis prior to antiarrhythmic treatment, and 84% agreed it should be stopped within six months after LVEF normalization. Responses for the optimal time point for the first LVEF reassessment during follow-up varied markedly (1 day–6 months after antiarrhythmic treatment).

Conclusion
This EHRA Survey reveals varying practices regarding AiCM among physicians, highlighting a lack of consensus and heterogeneous care of these patients.

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Introduction

Arrhythmia-induced cardiomyopathy (AiCM) is an increasingly recognized entity characterized by new cardiac dysfunction in the context of an often persistent cardiac arrhythmia, which is potentially reversible upon its effective control. The potential reversibility of AiCM provides a compelling reason for early diagnosis and intervention, which usually lead to significant improvement in cardiac function and patient outcomes.

The exact pathophysiological mechanisms underlying AiCM are incompletely understood, but it is believed that arrhythmia-induced myo-cardial remodelling plays a central role. This remodelling can lead to adverse structural and functional changes, which may be at least partially reversed with the normalization of heart rate and/or rhythm control. The clinical presentation of AiCM is heterogeneous, ranging from asymptomatic drop in left ventricular ejection fraction (LVEF) to decompensated acute heart failure (HF).

Diagnosis of AiCM is challenging due to the absence of specific diagnostic criteria and the broad differential diagnosis of other HF aetiologies, especially when first manifesting as dilated cardiomyopathy. Traditionally, AiCM has been diagnosed retrospectively after observing an improvement in ventricular function following treatment of the arrhythmia. However, the potential reversibility of the pathology warrants proactive diagnosis particularly with the advent of more sophisticated imaging techniques and biomarkers and effective therapies such as catheter ablation (CA). Recently, a novel predictive model attempted an early identification of LVEF recovery after CA for atrial fibrillation (AF).

Aim

This European Heart Rhythm Association Scientific Committee (EHRA SIC) questionnaire aimed to elucidate the definitions and management of AiCM among European and non-European electrophysiologists.

Methods

Online questionnaire

The questionnaire was developed by the EHRA Scientific Initiatives Committee. The survey electronic link was sent to ~4000 members of EHRA and EHRA Young EP between 4 September 2023 and 5 October 2023, and also promoted via social media. The online-based questionnaire consisted of single- and multiple-choice questions assessing how AiCM is currently defined, the risk factors for AiCM depending on the arrhythmia underlying AiCM [i.e. atrial fibrillation- or premature ventricular contraction (PVC)-induced CM], type and duration of follow-up and monitoring, the clinical and imaging parameters for identification potential AiCM patients, and the treatment and adjuvant treatment of AiCM. The full questionnaire was approved by all investigators and is provided in the Supplementary material online. The response was voluntary, anonymous, and GDPR compliant.

Statistical analysis

Continuous variables are presented as mean ± standard deviation (SD) or median and interquartile range. Categorical variables are expressed as...
numbers and percentages. Test for normality of the distribution was assessed visually. All analyses were performed using R Studio (version 4.2.1, Vienna, Austria).

Results

From a total of 255 responses, 49 questionnaires were empty (contained no responses to any question) leaving 206 questionnaires that were analysed further. Of the 206 respondents, 32 (16%) were female and most respondents were between 40 and 49 (33%) years old, followed by the age group of 30–39 (28%) and 50–59 (23%). The vast majority was EP specialists (81%), followed by general cardiologists (11%), EP fellows (7%), and Internal Medicine specialists (2%). Most respondents worked at university hospitals (47%), followed by private hospitals (17%), public hospitals (16%), specialized public cardiology centres (16%), and 4% worked in private practices. Most respondents (47%) were practicing in Western Europe, followed by Eastern Europe (23%), Americas (11%), Asia Pacific regions (11%), and Africa and Middle East (8%). A total of 54 (26%) of the responses were gathered from a total of 255 responses, 49 questionnaires were empty (containing no responses to any question) leaving 206 questionnaires that were analysed further. Of the 206 respondents, 32 (16%) were female and most respondents were between 40 and 49 (33%) years old, followed by the age group of 30–39 (28%) and 50–59 (23%). The vast majority was EP specialists (81%), followed by general cardiologists (11%), EP fellows (7%), and Internal Medicine specialists (2%). Most respondents worked at university hospitals (47%), followed by private hospitals (17%), public hospitals (16%), specialized public cardiology centres (16%), and 4% worked in private practices. Most respondents (47%) were practicing in Western Europe, followed by Eastern Europe (23%), Americas (11%), Asia Pacific regions (11%), and Africa and Middle East (8%). A total of 54 (26%) of the responses were gathered using social media, while the rest 152 (72%) were completed through the EHRA platform (Figure 1).

Definition of arrhythmia-induced cardiomyopathy

When asked how the respondents defined AiCM, 73 (35%) defined it as a specific drop/recovery cut-off in LVEF after the occurrence/resolution of the arrhythmia, while 65 (32%) indicated that any % drop in LVEF in relationship with arrhythmia occurrence/treatment should prompt the diagnosis of AiCM. A total of 42 (20%) respondents indicated that a new onset of HF (i.e. either systolic or diastolic) should trigger the diagnosis of AiCM, while 25 (12%) indicated that a LVEF < 50% in the context of a new arrhythmia onset prompts the suspicion of AiCM.

When asked which specific LVEF cut-off should be chosen to define AiCM in the context arrhythmia onset/treatment, most respondents indicated a cut-off of 10% (102, 50%), followed by 20% (35, 17%), 15 (27, 13%), and 5% (17, 8%), while 25 (25, 12%) indicated that any % LVEF drop should trigger the suspicion of AiCM (Figure 1).  

Arrhythmias related to arrhythmia-induced cardiomyopathy

Regarding the arrhythmias that most commonly cause AiCM, AF was chosen by the majority of respondents (190, 92%), followed by high amounts of right-ventricular pacing (133, 65%), atrial flutter (127, 62%), and PVCs (124, 60%). Permanent junctional reciprocating tachycardia was chosen by 76 (37%) of the respondents, atrioventricular nodal re-entry tachycardia in 12%, and ventricular tachycardia by only 3 (1.5%) respondents.

Differential diagnosis in arrhythmia-induced cardiomyopathy

The respondents were asked whether further evaluation for other types of HF with cardiac magnetic resonance (CMR), coronary angiography, or serial brain natriuretic peptide (BNP) measurements was warranted for differential diagnosis of AiCM. Cardiac magnetic resonance was chosen by 24 (16%) respondents, while coronary angiography

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Figure 1 Left—distribution of the respondents in Europe (70% of the total respondents). Right—graph showing the preferred definition of arrhythmia-induced cardiomyopathy (AiCM), single-choice. Indented in the top right corner is the most preferred LVEF drop/improvement in the context of an arrhythmia to be used to trigger diagnosis of AiCM, also single-choice. AiCM, arrhythmia-induced cardiomyopathy; LVEF, left ventricular ejection fraction.
was chosen by 28 (14%) respondents and 100 (49%) respondents use serial BNP measurements.

The respondents were further questioned regarding the known predictors for AiCM (i.e. Antwerp Score, multiple choice): 155 (75%) mentioned the lack of a known HF cause, 124 (62%) left atrial volume, 104 (51%) type of AF, and 37 (18%) QRS duration. Only 21 (10%) of the respondents chose all four answers (Figure 2).

**Antiarrhythmic treatment in arrhythmia-induced cardiomyopathy**

When questioned about the therapy options for AiCM (multiple choice), most respondents considered all rhythm control strategies—193 (94%) CA, 171 (83%) electrical cardioversions, as well as 157 (76%) antiarrhythmic drugs. A wearable defibrillator was chosen by 43 (21%, Figure 3).

**Adjuvant treatment in arrhythmia-induced cardiomyopathy**

Most respondents mentioned that they regularly recommend adjuvant HF therapy in AiCM patients (156, 76%). When asked to define adjuvant HF treatment in AiCM (multiple choice), the majority included drug treatment with betablockers—193 (94%) and/or mineralocorticoid receptor antagonists—167 (81%), sodium glucose transport protein 2 (SGLT2) antagonists—161 (78%), angiotensin receptor-neprilysin inhibitor—158 (77%), and angiotensin converting enzyme inhibitors or receptor blockers—142 (69%, Figure 3). Sixty-two (30%) of respondents chose 4 medications, 22 (11%) 3 medications, and 14 (7%) 2 medications.

Most respondents (170, 83%) reported that the adjuvant drug therapy should be started at the time of HF diagnosis, before rhythm control. In contrast, 36 (17%) respondents chose to start the drug therapy only in case of persistent HF after rhythm control.

Most respondents (173, 84%) stopped the adjuvant drug therapy usually within 6 months after HF normalization, while 32 (16%) opted for lifelong treatment. Half of the respondents reported that the follow-up of AiCM patients after LVEF recovery may be ended within the first year—53 (26%) at one year, 30 (15%) at 6 months, and 24 (12%) at 3 months, while the other half of the respondents—98 (48%) usually follow their AiCM patients for over a year. A mean of 10% (±10) of the AiCM patients will receive loop recorders to monitor for early arrhythmia recurrence after restoration of sinus rhythm.

**Premature ventricular contractions-induced cardiomyopathy**

When questioned about predictors of risk factors for PVC-induced CM, 149 (72%) chose the overall PVC burden—with a median PVC burden cut-off of 11% (±7), 137 (67%) chose late gadolinium enhancement (LGE) on cardiac MRI examination, 99 (48%) chose LVEF at presentation, 91 (44%) chose the left ventricular volume at presentation, 90 (44%) the site and origin of the PVCs, and 59 (29%) chose the QRS duration of the PVC. There was uncertainty regarding the assessment of right ventricular (RV) function in addition to LV function in PVC-induced CM patients: 85 (41%) respondents chose to assess the RV function, 74 (36%) respondents assess the RV function in some patients, and 46 (22%) of the respondents do not routinely assess the RV function (Figure 4).

**Discussion**

The purpose of this EHRA SIC survey was to offer a snapshot of current definitions used for AiCM, its risk factors, predictors, therapeutic possibilities, and follow-up strategies. We report that there is a lack of a universally accepted definition for AiCM, with a high degree of uncertainty regarding a specific LVEF drop/recovery cut-off in the context of a new onset/treatment of the culprit arrhythmia.

The questionnaire offered several other important insights: (1) AF is the most commonly identified arrhythmia causing AiCM, followed by high amounts of ventricular pacing, atrial flutter, and PVCs; (2) there is inconsistent use of MRI or coronary angiography for differential diagnosis, with a surprising high reliance on serial BNP measurements; (3) antiarrhythmic treatments, particularly CA, are widely used, alongside adjunctive HF therapy initiated at the time of HF diagnosis. However, one in five respondent chose to start a medical therapy only in case of persistent HF after rhythm control. The time point of stopping adjunctive HF medication is unclear. (4) Follow-up practices after LVEF recovery vary between practitioners, showing an even split between short-term (<1 year) and long-term monitoring (>1 year) for arrhythmia recurrence; (5) PVC burden is recognized as a key risk factor in PVC-induced cardiomyopathy; and 6) there is a great variability in the assessment of right ventricular function in PVC-induced cardiomyopathy.

The survey clearly identifies AF as the leading arrhythmia associated with AiCM, followed by other supraventricular and ventricular arrhythmias. This aligns with existing literature emphasizing the high prevalence of AF in AiCM cases.18,19

In terms of differential diagnosis, the low endorsement for advanced imaging techniques such as CMR and coronary angiography points to a potential underutilization of these modalities. This is rather surprising, since the CAMERA-MRI25 showed that restoration of sinus rhythm via CA results in significant improvement in ventricular function, particularly in patients with absent ventricular fibrosis on CMR. The
Figure 3 Left—distribution of the responses regarding the therapy of arrhythmia-induced cardiomyopathy. Multiple choices were allowed. Right—distribution of the responses regarding adjuvant heart failure therapy in AICM. Multiple choices were allowed. AICM, arrhythmia-induced cardiomyopathy; ACE, angiotensin conversion enzyme antagonist; ARB, angiotensin receptor blocker; ARNI, angiotensin receptor/neprilysin inhibitor; MRA, mineralocorticoid receptor antagonist.

Figure 4 Distribution of the answers regarding the known predictors of premature ventricular contraction-induced cardiomyopathy. LVEF, left ventricular ejection fraction; PVC, premature ventricular contractions.
reliance on serial BNP measurements reflects a non-invasive, cost-effective approach, but may miss structural heart disease that could be elucidated by imaging. There are only limited data available regarding a biomarker-driven approach. Nia et al. assessed 40 patients with supraventricular tachycardia and HF with serial NT-proBNP measurements at baseline, Day 1, and weekly over a period of 4 weeks.

Arrhythmia-induced cardiomyopathy patients revealed higher decrement velocity of NT-proBNP compared to patients with other causes of HF.

The Antwerp score has been recently proposed to identify AICM patients from a population with depressed LVEF undergoing CA for AF showing variable predictive power in different populations. According to the current questionnaire, as only a very small number of the respondents chose all four proposed parameters as predictors for LVEF recovery after CA for AF, we report that the awareness for this predictive model is currently very low. This tool, however, requires prospective validation.

The consensus on antiarrhythmic treatment strategies, particularly the widespread use of CA, suggests a strong preference for rhythm control in AICM, aligning with recent data and current guidelines recommending CA as a first-line treatment when AICM is highly probable independent of symptom status (class I, level of evidence B, ESC Guidelines).

The results also highlight the importance of adjunctive HF therapy, with a notable consensus on initiating treatment at first HF diagnosis irrespective of the relationship with the arrhythmia. However, no clear time point as to whether/when this adjuvant therapy should be ended after LVEF recovery was identified. This ambiguity may have various reasons. One of them might be the increasing evidence, that over time, cardiac arrhythmias can lead to progressive ventricular remodeling via tachycardia and irregular ventricular rhythm. While LVEF can normalize, left ventricular diameter stay dilated and diffuse fibrosis as detected by CMR might persist as a late outcome after AICM and thus might argue for lifelong HF therapy.

The follow-up practices after antiarrhythmic treatment of AICM showed once again a split. There was no consensus with regard to the best time point for LVEF reassessment after antiarrhythmic treatment and with half of the respondents ending the follow-up of AICM patients within a year post-LVEF recovery and the other half extending beyond a year. This divergence underscores the lack of standardized follow-up duration and calls for research to optimize monitoring periods.

For PVC-induced cardiomyopathy, the identification of PVC burden as a primary risk factor echoes current understanding, yet the survey indicates variability in the assessment of right ventricular function, suggesting an area where guidelines could provide clarity.

There is a lack of guidance from the ESC guidelines regarding AICM, and there is no clear algorithm for diagnosis or management. In this context, some responses such as the time to LVEF reassessment might indicate a level IIb indication, while other answers clearly underlie the importance of further research, such as studies evaluating the role of ongoing therapy after rhythm restoration.

As limitations of this survey, we report that most respondents came from Western Europe and that we did not include pacemaker-induced cardiomyopathy into our questionnaire.

Conclusion

This EHRA Survey reveals varying practices in defining and managing AICM among European and non-European physicians. There is an unmet need for a consensus in the definition and management of AICM patients to improve patient care.

Supplementary material

Supplementary material is available at Europace online.

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Data availability

All relevant data are within the manuscript and its Supporting Information files.

References


17. Serban T, du Fay de Lavallaz J, Barker DC, Sticherling C, Kühne M, Badertscher P. Validation of a novel score to predict which patients with atrial fibrillation and depressed left ventricular ejection fraction will respond to catheter ablation. Rev Esp Cardiol (Eng Ed) 2023;76:745–8.


