ORIGINAL RESEARCH

CATHETER ABLATION - ATRIAL FIBRILLATION

Impact of Low Body Mass Index on Cardiac Tamponade During Catheter Ablation for Atrial Fibrillation

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ABSTRACT

BACKGROUND Cardiac tamponade is a potentially fatal complication of catheter ablation for atrial fibrillation (AF).

OBJECTIVES This study aimed to evaluate the impact of body mass index (BMI) on cardiac tamponade during AF ablation.

METHODS Patients who underwent catheter ablation for AF between April 1, 2016 and March 31, 2018 were analyzed using a Japanese nationwide claims database. Mixed-effects multivariable Poisson regression analysis was performed to investigate the association between BMI and cardiac tamponade.

RESULTS A total of 59,789 hospitalizations (age 65.6 \pm 10.4 years, 29% women) with catheter ablation for AF were analyzed. Cardiac tamponade occurred in 647 patients (1.1%). Multivariable analysis revealed that being underweight (BMI <18.5 kg/m²) was associated with an increased risk of cardiac tamponade (relative risk [RR]: 1.42; 95% CI: 1.03-1.95) when compared with having a normal weight (BMI \geq 18.5 and <25 kg/m²). Other characteristics that were associated with an increased risk of cardiac tamponade sex, and a history of heart failure, hypertension, diabetes, and dialysis treatment.

CONCLUSIONS In this analysis of a large nationwide database of patients with AF who underwent ablation, being underweight was independently associated with an increased risk of cardiac tamponade during AF ablation. Clinicians should consider the higher risk of cardiac tamponade in the underweight population and take appropriate measures to reduce this risk. (J Am Coll Cardiol EP 2023;9:200-208) © 2023 by the American College of Cardiology Foundation.

atheter ablation for atrial fibrillation (AF) is widely accepted and has become an effective alternative to drug therapy.^{1,2} Cardiac tamponade is among the most common, potentially life-threatening complications of AF ablation and has been employed as a specific electrophysiological performance measure by the Heart Rhythm Society.³ Recent reports from the United States and Europe have indicated the incidence of cardiac tamponade to be 0.6%-1.9%,⁴⁻⁷ and risk factors have also been reported.⁸ Obesity is a known risk factor for AF development and has recently been reported to be a risk

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factor for cardiac tamponade.^{1,8} On the other hand, the risk of cardiac tamponade during AF ablation in patients who are underweight has never been investigated; a reason for this may be that the proportion of patients who are underweight is low in Western countries.⁹⁻¹¹ Recent reports have revealed that being underweight (body mass index [BMI] <18.5 kg/m²) is associated with a higher risk of bleeding in patients with AF who are taking anticoagulants.¹²⁻¹⁴ We hypothesized that being underweight may also be associated with an increased risk of cardiac tamponade during AF ablation because of the higher risk of bleeding.

The present study aimed to evaluate the association between BMI and cardiac tamponade during AF ablation using a large nationwide database.

METHODS

This retrospective cross-sectional study used the JROAD-DPC (Japanese Registry of All Cardiac and Vascular Diseases and the Diagnosis Procedure Combination), which has been previously described in detail.^{15,16} JROAD-DPC is an administrative database that covered more than 1,500 Japanese Circulation Society-certified training hospitals during the study period. The JROAD-DPC includes the following information for each patient: age; sex; height; weight; Barthel index score; primary diagnoses, comorbidities, conditions arising after admission based on the International Classification of Diseases-10th Revision (ICD-10) codes; drugs used; diagnostic and therapeutic procedures; length of stay; and discharge status. BMI was calculated using height and weight and was divided into 4 categories: 1) underweight (BMI <18.5 kg/m²); 2) normal weight (BMI \geq 18.5 <25 kg/m²); 3) overweight (BMI ≥ 25 and and <30 kg/m²); and 4) obese (BMI \geq 30 kg/m²). Regarding diagnosis, in addition to the ICD-10 codes, detailed names of diagnoses are listed; therefore, detailed names of diseases that cannot be identified using the ICD-10 codes alone could be determined. The procedure codes, ICD-10 codes, and other definitions used for baseline characteristics are shown in Supplemental Table S1. The association between hospital-level characteristics and cardiac tamponade was also investigated, and detailed methods and results are described in the Supplemental Appendix.

This study was approved by the Institutional Review Board of Jikei University School of Medicine (approval number: 32-323 [10405], approval date: November 9, 2020). The requirement for informed consent was waived because information specific to individuals was not included in the database.

STUDY POPULATION. The flowchart of the study is shown in Figure 1. We initially identified patients aged \geq 20 years who underwent catheter ablation and were diagnosed with AF between April 1, 2016 and March 31, 2018. To exclude ablations with the main purpose of treating other arrhythmias, the following cases were excluded: 1) catheter ablation without septal puncture (n = 5,039); and 2) the presence of atrial flutter, atrial tachycardia, premature atrial/ventricular contractions, and ventricular tachycardia/fibrillation (n = 4,396). Cases of catheter ablation other than radiofrequency or cryoballoon ablation were excluded (n = 4). In addition, patients who died after AF ablation were excluded (n = 18).

OUTCOMES. The primary outcome in this study was the occurrence of cardiac tamponade during AF ablation. Cardiac tamponade was defined as the presence of either: 1) a diagnosis of cardiac tamponade in conditions arising after admission identified by ICD-10 codes (I319 or I971); or 2) procedure codes for pericardiocentesis or pericardial drainage (J048 or J002). Additionally, the diagnosis recorded in text format was carefully reviewed to exclude vague diagnoses other than cardiac tamponade.

STATISTICAL ANALYSIS. Categorical data are presented as frequencies (percentages), whereas continuous data are expressed as mean \pm SD. Emergency hospitalization, BMI, and Barthel index score were missing in 0.002%, 0.7%, and 1.8% of records, respectively; there were no missing data for other patient-level variables. To account for missing data, multiple imputation methods using the chained equation algorithm were applied based on 10 replications, assuming missing random mechanisms.

Mixed-effects multivariable Poisson regression analysis, using institute as the random intercept, was performed to estimate the relative risk (RR) and 95% CI for cardiac tamponade in the overall sample of patients who underwent cryoballoon and radiofrequency ablation. The model assumptions regarding explanatory and response variables were satisfied. Using normal BMI as a reference, the RR and 95% CI for being underweight, overweight, and obese were estimated. All statistical comparisons were 2-sided, with statistical significance set at P < 0.05. All analyses were performed using STATA (version 16.0, StataCorp).

ABBREVIATIONS AND ACRONYMS

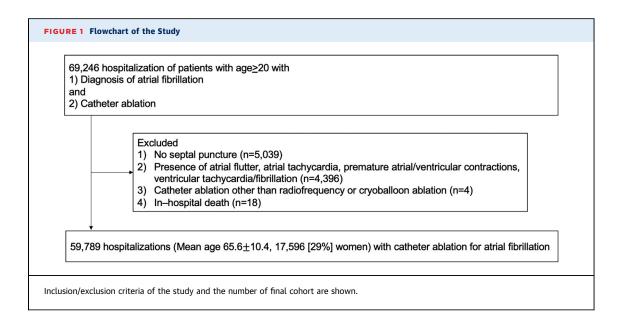
AF = atrial fibrillation

BMI = body mass index

ICD-10 = International Classification of Diseases-10th Revision

ICE = intracardiac echocardiography

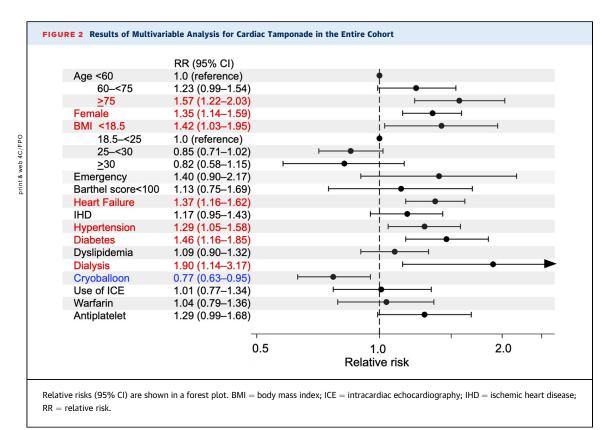
RR = relative risk



		BMI			
	Total (N = 59,789)	<18.5 kg/m ²	\geq 18.5 and <25 kg/m ²	\geq 25 and <30 kg/m ²	≥30 kg/m
Patients		2,302 (3.9)	34,695 (58)	18,681 (31)	4,111 (6.9)
Age, y	65.6 ± 10.4	$\textbf{68.7} \pm \textbf{9.3}$	$\textbf{66.6} \pm \textbf{10.1}$	64.7 ± 10.3	$60.4 \pm 11.$
<60	14,589 (24)	348 (15)	7,260 (21)	5,166 (28)	1,815 (44)
≥60 and <75	33,527 (56)	1,276 (55)	20,034 (58)	10,301 (55)	1,916 (47)
≥75	11,673 (20)	678 (29)	7,401 (21)	3,214 (17)	380 (9.3)
Women	17,596 (29)	1,278 (55)	10,533 (30)	4,430 (24)	1,355 (33)
Emergency admission	1,090 (1.8)	87 (3.8)	666 (1.9)	276 (1.5)	61 (1.5)
Barthel index score <100	2,124 (3.6)	157 (6.8)	1,242 (3.6)	579 (3.1)	146 (3.6)
Comorbidities					
Heart failure	21,600 (36)	940 (41)	12,539 (36)	6,611 (35)	1,510 (37)
Ischemic heart disease	10,020 (17)	373 (16)	5,834 (17)	3,144 (17)	669 (16)
Hypertension	42,311 (71)	1612 (70)	23,668 (68)	13,773 (74)	3,258 (79
Diabetes	5,541 (9.3)	135 (5.8)	2,593 (7.5)	2,001 (11)	812 (20)
Dyslipidemia	15,538 (26)	378 (16)	8,238 (24)	5,487 (29)	1,435 (35
Chronic kidney disease with dialysis	779 (1.3)	102 (4.4)	491 (1.4)	139 (0.7)	47 (1.1)
Procedural factors					
Cryoballoon ablation	14,673 (25)	616 (27)	8,942 (26)	4,280 (23)	835 (20)
Use of ICE	48,381 (81)	1,863 (81)	27,932 (81)	15,151 (81)	3,435 (84
Medications					
Warfarin	4,951 (8.3)	261 (11)	2,884 (8.3)	1,458 (7.8)	348 (8.5)
Antiplatelet therapy	4,947 (8.3)	177 (7.7)	2,823 (8.1)	1,586 (8.5)	361 (8.8)
Length of stay, d					
≤2	5,138 (8.6)	184 (8.0)	3,011 (8.7)	1,610 (8.6)	333 (8.1)
3-5	42,346 (71)	1,492 (65)	24,484 (71)	13,409 (72)	2,961 (72
6-7	7,287 (12)	297 (13)	4200 (12)	2,302 (12)	488 (12)
≥8	5,018 (8.4)	329 (14)	3,004 (8.7)	1,356 (7.3)	329 (8.0)
Complications					
Cardiac tamponade	647 (1.1)	44 (1.9)	393 (1.1)	171 (0.9)	39 (0.9)

Values are n (%) or mean \pm SD.

AF = atrial fibrillation; BMI = body mass index; ICE = intracardiac echocardiography.



RESULTS

A total of 59,789 hospitalizations (age 65.6 ± 10.4 years, 17,596 [29%] women) with catheter ablation for AF were analyzed (**Table 1**, Supplemental Table S2). Cryoballoon ablation was performed in 25% of patients, and radiofrequency ablation was performed in the rest. Intracardiac echocardiography (ICE) was performed in 81% of cases. Cardiac tamponade occurred in 647 patients (1.1%), and pericardiocentesis or pericardial drainage was performed in 619 patients (1.0%).

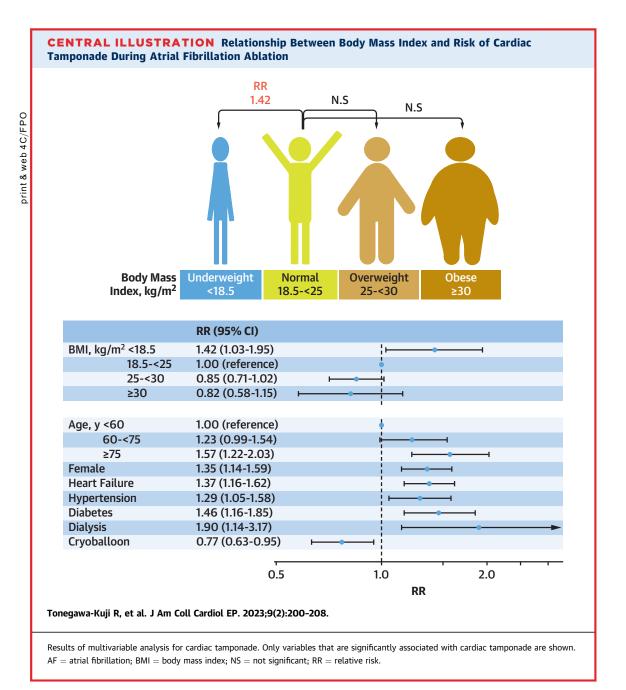
PATIENT-LEVEL CHARACTERISTICS AND CARDIAC TAMPONADE. According to the multivariable analysis, being underweight was associated with an increased risk of cardiac tamponade (RR: 1.42; 95% CI: 1.03-1.95) when compared with having a normal weight (**Figure 2, Central Illustration**). Other factors that were associated with an increased risk of cardiac tamponade were as follows: age \geq 75 years (RR: 1.57; 95% CI: 1.22-2.03); female sex (RR: 1.35; 95% CI: 1.14-1.59); and a history of heart failure (RR: 1.37; 95% CI: 1.16-1.62), hypertension (RR: 1.29; 95% CI: 1.05-1.58), diabetes (RR: 1.46; 95% CI: 1.16-1.85), and dialysis treatment (RR: 1.90; 95% CI: 1.14-3.17). Moreover, the use of cryoballoon ablation (RR: 0.77; 95% CI: 0.63-0.95) was associated with a decreased risk of cardiac tamponade. The use of ICE was not associated with cardiac tamponade.

Multivariable analyses were subsequently performed in subsets of patients who underwent either cryoballoon ablation (n = 14,673) or radiofrequency ablation (n = 45,116) (Table 2). In multivariable analysis, being underweight was associated with an increased risk of cardiac tamponade in the radiofrequency ablation group but not in the cryoballoon ablation group.

The association between hospital-level characteristics and cardiac tamponade was also investigated, and the results are shown in Supplemental Table S3.

DISCUSSION

The main findings obtained from the analysis of the nationwide claims database are as follows: First, the incidence of cardiac tamponade during AF ablation in Japan was 1.1%, which is similar to that in previous reports.^{4,5,7,8} Second, being underweight was associated with an increased risk of cardiac tamponade.



IMPACT OF BMI ON CARDIAC TAMPONADE. In our data, when BMI in the normal range was used as a reference, being underweight was associated with an increased risk of cardiac tamponade. To the best of our knowledge, being underweight has never been reported as a risk factor for complications associated with ablation for AF. Asians tend to be smaller and leaner than non-Asians; thus, patients with low body weight are more common in Asian populations than in non-Asian populations.⁹⁻¹¹ As the number of people

with low BMI is limited in Western countries, the relationship between low body weight and the risk of complications with AF ablation may not have been fully investigated.

The higher risk in patients with a low BMI might be attributed to their higher bleeding risk in the anticoagulant state. A recent observational study that included a large population of patients with AF (\leq 60 kg) taking oral anticoagulants showed that direct oral anticoagulants were associated with lower

	Cryoballo	on Ablation	Radiofrequency Ablation		
	Univariate	Multivariable	Univariate	Multivariable	
Age, y					
<60	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	
≥60 and <75	1.47 (0.93-2.34)	1.27 (0.79-2.04)	1.42 (1.12-1.82)	1.22 (0.95-1.56)	
≥75	1.75 (1.03-2.98)	1.37 (0.78-2.40)	2.18 (1.67-2.86)	1.64 (1.23-2.18)	
Female	1.29 (0.91-1.84)	1.15 (0.80-1.66)	1.57 (1.32-1.88)	1.41 (1.17-1.70)	
BMI					
Underweight (<18.5 kg/m²)	1.46 (0.75-2.82)	1.30 (0.67-2.54)	1.68 (1.18-2.40)	1.44 (1.01-2.07)	
Normal (\geq 18.5 and <25 kg/m ²)	1.00 (reference)	1.00 (reference)	1.00 (reference)	1.00 (reference)	
Overweight (\geq 25 and <30 kg/m ²)	0.79 (0.52-1.21)	0.80 (0.53-1.23)	0.83 (0.68-1.02)	0.86 (0.70-1.05	
Obese (≥30 kg/m²)	1.17 (0.56-2.43)	1.18 (0.56-2.47)	0.75 (0.52-1.10)	0.75 (0.51-1.11)	
Emergency hospitalization	2.60 (1.05-6.40)	1.85 (0.70-4.90)	2.00 (1.27-3.16)	1.25 (0.76-2.04)	
Barthel index score <100	1.28 (0.50-3.28)	0.92 (0.34-2.51)	1.77 (1.18-2.64)	1.21 (0.79-1.87)	
Heart failure	1.84 (1.27-2.67)	1.74 (1.20-2.54)	1.38 (1.14-1.66)	1.31 (1.08-1.57)	
Ischemic heart disease	1.20 (0.77-1.86)	0.94 (0.59-1.50)	1.45 (1.17-1.80)	1.24 (0.99-1.57)	
Hypertension	1.51 (0.99-2.30)	1.27 (0.82-1.96)	1.51 (1.21-1.89)	1.30 (1.04-1.64)	
Diabetes	1.30 (0.72-2.38)	1.04 (0.56-1.95)	1.76 (1.38-2.24)	1.57 (1.22-2.03)	
Dyslipidemia	1.45 (0.99-2.13)	1.21 (0.79-1.84)	1.35 (1.11-1.63)	1.05 (0.85-1.29)	
Dialysis	2.96 (1.07-8.17)	2.30 (0.72-7.41)	2.42 (1.44-4.07)	1.86 (1.05-3.31)	
Use of ICE	1.26 (0.64-2.49)	1.24 (0.63-2.43)	1.03 (0.97-1.11)	0.98 (0.73-1.33)	
Warfarin	1.57 (0.84-2.95)	1.08 (0.52-2.23)	1.36 (1.04-1.78)	1.03 (0.77-1.39)	
Antiplatelet	2.07 (1.21-3.52)	1.63 (0.89-2.97)	1.70 (1.31-2.21)	1.22 (0.91-1.64)	

risks of ischemic stroke and major bleeding than warfarin, and these results were largely consistent in patients weighing <50 kg.¹² However, even among patients on direct oral anticoagulants, having a low BMI has been reported to be associated with an increased risk of major bleeding, as compared with having a normal weight or with being overweight or obese.¹³ Because high-dose heparin is additionally used during ablation for AF, patients with a low BMI, who are originally at a higher risk for bleeding, may be more prone to puncture site bleeding and cardiac tamponade than other patients may be.

A lower BMI has been reported to be associated with lower left atrial volume.^{17,18} Although further examination is needed to examine why a low BMI is a risk factor for cardiac tamponade, our hypothesis is that a smaller left atrium in patients with a low BMI may have made it more difficult to manipulate the radiofrequency ablation catheter in the left atrium, causing mechanical trauma to the left atrial wall. In fact, being underweight was not a significant risk factor for cardiac tamponade in patients who underwent cryoballoon ablation. Therefore, cryoballoon ablation should be considered in patients who are underweight. In addition, the use of contact force-sensing catheters was reported to be associated with a lower incidence of tamponade than noncontact forcesensing catheters were.¹⁹ Although the information regarding the use of contact force-sensing catheters was not available in this database and we could not analyze the association of its use and cardiac tamponade, its use may be encouraged in patients considered high-risk, such as those with a low BMI.

Although a recent study from the US Medicare database reported obesity as a risk factor for cardiac tamponade,⁸ our results did not concur. The reason for this difference might be related to the degree of obesity of patients included in the obesity group, as the proportion of people with severe obesity is reported to be lower in Asia than in Western countries.¹¹ In fact, in the present database, the proportion of patients with a BMI >35 kg/m² was only 1.9%. Stratified analysis among patients who are obese may be necessary to further understand the association between obesity and cardiac tamponade.

ABLATION STRATEGIES AND CARDIAC TAMPONADE. Although the notion that cryoballoon ablation might reduce the risk of cardiac tamponade remains controversial, some studies support this idea.^{7,20,21} In the present study, being underweight was associated with an increased risk of cardiac tamponade in the radiofrequency ablation group, but not in the cryoballoon ablation group. However, it might be inappropriate to conclude based only on these results that cryoballoon ablation is a safer option than radiofrequency ablation in patients who are underweight because safety endpoints other than cardiac tamponade and precise ablation strategies could not be considered in the present study. Therefore, further analysis using registry data including other complications and detailed ablation strategies is warranted to determine a better ablation strategy for patients who are underweight.

ICE was used in 80% of cases in our study, which is high compared to that reported in the United States and described in the guidelines.^{1,8,22,23} Additionally, the use of ICE for AF ablation to reduce cardiac tamponade is controversial.^{8,23,24} Although the use of ICE was not associated with complications in this study, this result does not immediately negate the usefulness of ICE to improve safety; cases performed without ICE might have been performed by highly experienced operators, and the usefulness of ICE depends largely on the proficiency of the operators in understanding anatomical structures from ICE images. Further studies are warranted to determine the contexts in which ICE might be useful for improving both the safety and efficacy of AF ablation.

OTHER PATIENT-LEVEL CHARACTERISTICS AND CARDIAC TAMPONADE. Consistent with several previous studies, our study identified higher age, female sex, hypertension, and diabetes as risk factors for cardiac tamponade.^{1,7,8,22} Although little is known about the safety of AF ablation in patients receiving dialysis treatment because of the limited number of such patients in previous reports,25,26 the present study identified dialysis treatment as a risk factor for cardiac tamponade during AF ablation. This might be attributed to the patients' higher risk of bleeding caused by anticoagulant use during dialysis. Another possibility is that dialysis-related cardiac amyloidosis among patients with long-term dialysis treatment might cause some histological change in the left atrial wall, making them more susceptible to cardiac tamponade.

Moreover, a history of heart failure was identified as a risk factor for cardiac tamponade. Further studies are warranted to examine why these comorbidities were important for the occurrence of cardiac tamponade during AF ablation.

STUDY LIMITATIONS. First, the study design was observational and retrospective. Because the DPC data are based on medical claims, data that are not

directly related to the cost, such as ICD-10 diagnosis codes of comorbidities other than the main diagnoses, are not completely validated. However, previous studies have proven the validity of JROAD-DPC diagnoses in comparison with other nationwide databases or in-hospital registries.^{27,28} Second, characteristics of operators, the number of operators in the hospital, procedural details (eg, only pulmonary vein isolation or any additional procedures performed, whether it was the first or repeat ablation, usage of a contact force-sensing catheter, and precise dose of heparin used during ablation), and precise clinical information, such as type of AF (eg, paroxysmal or persistent AF) and laboratory or echocardiology data, were not available. Third, a history of cardiac surgery has been reported as an important risk factor for cardiac tamponade during AF ablation, but this was not available in our database.^{8,29} Fourth, we were only able to identify complications during hospitalization. However, the postoperative hospitalization period is longer than that in Western countries, and almost all ablation procedures are performed in hospitalized settings in Japan. Therefore, we believe that we were able to evaluate complications that may have been overlooked in inpatient reports from countries with shorter hospitalization periods or ablations performed in outpatient settings. Fifth, the precise date of cardiac tamponade occurring during hospitalization was not available in the database. Sixth, although the JROAD-DPC covers >60% of admissions in cardiology or cardiovascular sections of >1,500 Japanese Circulation Societycertified cardiology training hospitals in Japan, admissions to nonteaching hospitals with relatively fewer beds were not included. Lastly, most of the patients included in this study were presumed to be Asian, because most of the Japanese population is of Asian descent. Therefore, further investigation is warranted to determine whether a low BMI is also a risk factor in other races.

CONCLUSIONS

In this analysis of a large nationwide database of patients with AF who underwent ablation and were underweight (BMI <18.5 kg/m²) was independently associated with an increased risk of cardiac tamponade during AF ablation. Clinicians should consider the higher risk of cardiac tamponade in the underweight population and take appropriate measures to reduce this risk.

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PERSPECTIVES

COMPETENCY IN MEDICAL KNOWLEDGE: Being underweight (BMI <18.5 kg/m²) was associated with an increased risk of cardiac tamponade. Moreover, being underweight was associated with an increased risk of cardiac tamponade in the radiofrequency ablation group but not in the cryoballoon ablation group.

TRANSLATIONAL OUTLOOK: Clinicians should consider tamponade risk in the underweight population and take appropriate measures to reduce this risk, such as choosing cryoballoon ablation instead of radiofrequency ablation, if applicable.

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APPENDIX For expanded Methods, Results, and Discussion sections, and supplemental tables, please see the online version of this paper.