

Original Research

Check for updates

Is It Safe to Preserve Left Atrial Appendage During Maze Procedure?

Kyungsub Song , MD, Woo Sung Jang , MD, PhD, Namhee Park , MD, PhD, Yun Seok Kim , MD, PhD, and Jae Bum Kim , MD

Department of Thoracic and Cardiovascular Surgery, Keimyung University Dongsan Center, Keimyung University College of Medicine, Daegu, Korea

AUTHOR'S SUMMARY

Preserving the left atrial appendage (LAA) during the maze procedure was not increase risk of cerebral infarction, readmission for heart failure, and cardiogenic death during the follow-up period. However, as the result of transthoracic echocardiography 1 year after the heart operation, including the maze procedure, there were significantly more patients with effective LA mechanical contraction in the preserving LAA group than obliterating LAA group. As a result, preserving LAA during a maze procedure more improved LA function than obliterating LAA during a maze procedure without increasing the risk of cerebral infarction.

ABSTRACT

Background and Objectives: The left atrial appendage (LAA) can contribute significantly to LA mechanical contraction. Nevertheless, the preventive effect of LAA occlusion during the maze procedure against cerebral infarction remains controversial. In this study, we compared the surgical, cardiac hemodynamic, and neurologic outcomes between LAA preservation and occlusion performed during the maze procedure.

Methods: Between January 2015 and August 2021, 252 patients underwent the maze procedure using cryoablation at our medical center. After excluding patients according to our exclusion criteria (i.e., mechanical prosthesis implantation, preexisting LAA thrombus), LAA was preserved in 113 patients (non-occlusion group) and occluded in 75 patients (occlusion group). Outcomes were compared using propensity score matching (PSM).

Results: PSM did not reveal significant intergroup differences in baseline characteristics between the non-occlusion (n=53) and occlusion (n=53) groups. During a median follow-up of 44 months, 2 patients in the non-occlusion group (3.8%) experienced ischemic strokes. There was no significant difference in the rate of freedom from stroke (p=0.19) and major adverse cardiac events (p=0.43) between the 2 groups. Through echocardiography at 1-year follow-up, a statistically significant difference in LA mechanical contraction was observed between the non-occlusion group and occlusion group (24 of 33 [72.7%] vs. 18 of 37 [48.6%], respectively; p=0.04).

Conclusions: In this study, preservation of the LAA during the maze procedure resulted in better LA function than LAA occlusion, with similar rates of stroke.

Keywords: Atrial fibrillation; Maze procedure; Stroke

OPEN ACCESS

 Received:
 Feb 19, 2023

 Revised:
 Apr 5, 2023

 Accepted:
 May 3, 2023

 Published online:
 Jun 9, 2023

Correspondence to

Woo Sung Jang, MD, PhD

Department of Thoracic and Cardiovascular Surgery, Keimyung University Dongsan Medical Center, Keimyung University College of Medicine, 1035, Dalgubeol-daero, Dalseo-gu, Daegu 42601, Korea. Email: whiteuri@dsmc.or.kr

Copyright © 2023. The Korean Society of Cardiology

This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (https:// creativecommons.org/licenses/by-nc/4.0) which permits unrestricted noncommercial use, distribution, and reproduction in any medium, provided the original work is properly cited.

ORCID iDs

 Kyungsub Song ID

 https://orcid.org/0000-0002-6556-2261

 Woo Sung Jang ID

 https://orcid.org/0000-0003-1576-9472

 Namhee Park ID

 https://orcid.org/0000-0001-8701-2220

 Yun Seok Kim ID

 https://orcid.org/0000-0002-2817-557X

 Jae Bum Kim ID

 https://orcid.org/0000-0002-2817-557X

Funding

The authors received no financial support for the research, authorship, and/or publication of this article.



Conflict of Interest

The authors have no financial conflicts of interest.

Data Sharing Statement

The data generated in this study is available from the corresponding author upon reasonable request.

Author Contributions

Conceptualization: Kim YS, Kim JB, Jang WS; Data curation: Park N, Kim YS, Kim JB, Jang WS; Investigation: Kim JB, Jang WS; Supervision: Song K, Park N, Jang WS; Visualization: Kim YS, Kim JB, Jang WS; Writing - original draft: Song K, Kim JB; Writing review & editing: Song K, Park N, Kim YS.

INTRODUCTION

Atrial fibrillation (AF) accounts for 20% of all stroke cases. The risk of stroke is increased 5-fold in patients with non-rheumatic AF, and 17-fold in those with mitral stenosis and AF.¹) Furthermore, the left atrial appendage (LAA) is a major source of thrombus formation, which is the main cause of embolization and can trigger atrial ectopic focus.²⁻⁴) For this reason, LAA occlusion is routinely performed during the maze procedure. However, the LAA contributes to left atrial (LA) mechanical contraction, which is an important factor for increasing cardiac output.⁵) Moreover, LAA preservation may favorably contribute to appropriate secretion of atrial natriuretic peptides in patients with congestive heart failure.⁶) Whitlock et al.⁷ conducted the Left Atrial Appendage Occlusion Study III (LAAOS III; ClinicalTrials.gov number: NCT01561651) and reported that LAA occlusion during cardiac operation reduced the risk of cerebral infarction. However, the preventive effect of LAA occlusion against cerebral infarction remains controversial. In addition, few clinical studies have investigated the hemodynamic effects of LAA occlusion thus far. Therefore, in this study, we compared the surgical, cardiac hemodynamic, and neurologic outcomes between LAA preservation and occlusion performed during the maze procedure.

METHODS

Ethical statement

This study was approved by the Institutional Review Board of the Keimyung University Dongsan Medical Center (Daegu, Korea) (approval number: 2022-12-016; December 13, 2022). The requirement for informed consent was waived due to the retrospective nature of this study.

Study population

This study included patients who underwent the maze procedure using cryoablation (CryoICE Cryoablation Probes, AtriCure, Mason, OH, USA) at our medical center between January 2015 and August 2021 (n=252). Patients who underwent concomitant valve replacement using a mechanical prosthetic valve, or had pre-operative existing spontaneous echo contrast or intra-cardiac thrombus were excluded. We excluded patients with mechanical prosthetic valve because the mechanical prosthetic valve was a well-established risk factor for cerebral infarction after cardiac operation. Patients were classified into 2 groups according to LAA preservation or occlusion during the maze procedure.

Operative techniques

Most patients in this study underwent a maze procedure via complete sternotomy. The techniques for valve replacement, valve repair, and coronary artery bypass grafting were different depending on the surgical period and preference of the surgeon. In the maze procedure, a modified Cox maze procedure using an argon-based flexible cryo-probe was performed with antegrade or retrograde cardioplegia. Patient selection for the maze procedure was based on the factors known to affect sinus rhythm restoration from previous studies, considering LA size and fibrosis, duration of AF, and combined procedure. The absolute indication for LAA obliteration was a history of cerebral infarction or cardiac thrombus; otherwise, the decision to perform LAA obliteration with a maze procedure was made considering the patients' comorbidity and depending on the surgical period and preference of the surgeon.

Echocardiography and rhythm follow-up

Transthoracic echocardiography (TTE) was performed using commercially available systems with 2.5 MHz transducers (Vivid E95, GE Vingmed Ultrasound AS, Horten, Norway; Acuson SC 2000, Siemens Healthineers, Erlangen, Germany). The results of TTE were reviewed by experienced cardiologists.

During the postoperative hospitalization period, monitoring was performed daily through typical 12-channel surface electrocardiography (ECG). Follow-up TTE was performed prior to discharge and 12 months after surgery. The LA diameter was measured at the apical 4-chamber view or parasternal long-axis view at the end-systole phase. LA volume index and left ventricular volume index were measured using LA or ventricular volume acquired by Simpson's method from a 4-chamber apical view, indexed to the body surface area.

In patients with sinus rhythm at ECG, transmitral peak velocity and the velocity–time integral of the early (E) and late (A) filling waves were measured at the apical 4-chamber view. The absence of mechanical atrial contraction was defined as the absence of an A wave on TTE,⁸⁾ and mechanical atrial contraction was defined as there was an A wave on TTE.

During the follow-up period, ECG was performed in the outpatient clinic at 1, 3, and 6 months postoperatively, and subsequently every 6 months. Sinus rhythm restoration was defined as absence of AF recurrence in at least 2 consecutive ECG examinations. The presence of rhythms other than sinus rhythm, including AF, atrial flutter, ectopic atrial arrhythmia, junctional rhythm, or cardiac rhythm of permanent pacemaker (PMK) without atrioventricular synchrony, denoted the failure of sinus rhythm restoration.

Definitions of endpoint

The primary endpoint was the occurrence of major adverse cardiovascular events (MACEs) and the results of TTE during the follow-up period. MACE included the occurrence of cerebrovascular infarction (stroke) suspected to be of thromboembolic origin at any time during the follow-up period, readmission due to heart failure, and PMK insertion. Any symptoms suggestive of neurologic deficit were evaluated by a neurologist through brain magnetic resonance imaging.

Statistical analysis

Normally distributed variables were presented as means and compared using independent t-tests. Categorical variables were compared using Pearson's χ^2 test or Fisher's exact test and were presented as numbers (percentages). Continuous variables were compared using the 2-sample t-test or Wilcoxon rank-sum test. Continuous data were assessed for normality of distribution using the Shapiro-Wilk test and graphical methods. All statistical tests were 2-sided, with an alpha level of 0.05.

The Kaplan-Meier method was used to estimate stroke-free survival and all-cause mortality; the log-rank test was used for comparisons. Cox proportional hazards regression was used to identify predictors of stroke for the entire patient population tested. Variables with p values <0.20 in the univariable analysis were included in the multivariable analysis.⁹⁾ The results were reported as hazard ratio (HR) and 95% confidence interval (CI).

Propensity score analyses were used to adjust baseline differences between 2 groups. Prespecified covariates (**Table 1**) were included in the propensity score calculation. Patients Table 1. Baseline characteristics of the entire cohort

	Non-occlusion group (n=113)	Occlusion group (n=75)	p value
Sex, male	55 (48.7)	37 (49.3)	0.92
Age (years)	67.3±8.7 65.0±7.8		0.06
Heart disease etiologies			
Rheumatic disease	28 (24.8)	25 (33.3)	0.20
Degenerative	26 (23.0)	8 (10.7)	0.03
Functional	57 (50.4)	45 (60.0)	0.20
Others	6 (5.3)	2 (2.7)	0.48
Concomitant operations with maze procedure			
Mitral valve replacement	24 (21.2)	17 (22.7)	0.82
Mitral valve repair	47 (51.6)	26 (34.7)	0.34
Tricuspid annuloplasty	59 (52.2)	41 (54.7)	0.74
Aortic valve replacement	21 (18.6)	4 (5.3)	0.01
Aortic valve plasty	4 (3.5)	2 (2.7)	>0.99
Coronary artery bypass grafting	17 (15.0)	3 (4.0)	0.02
Aortic replacement	6 (5.3)	1 (1.3)	0.25
Atrial septal defect repair	14 (12.4)	7 (9.3)	0.52
Myxoma removal	3 (2.7)	0	0.28
AF profiles			
Fine (<1 mm) AF wave	59 (52.2)	43 (57.3)	0.49
AF type			0.44
Persistent	69 (61.1)	50 (66.7)	
Paroxysmal	44 (38.9)	25 (33.3)	
Heart failure	24 (21.2)	22 (29.3)	0.21
Coronary artery disease	12 (10.6)	8 (10.7)	0.99
Cerebral infarction	16 (14.2)	18 (24)	0.09
Diabetes mellitus	24 (21.2)	17 (22.7)	0.82
Aortic disease	2 (1.8)	1 (1.3)	>0.99
Hypertension	53 (46.9)	31 (41.3)	0.45
Chronic obstructive pulmonary disease	2 (1.8)	2 (2.7)	0.66
Chronic kidney disease	3 (2.7)	4 (5.3)	0.44
CHA ₂ DS ₂ -VASc score	2.7±1.7	2.7±1.7	0.98
Pre-operative TTE			
EF (%)	55.3±11.5	55.4±12.0	0.99
LAD (cm)	3.1±3.6	5.6±1.0	0.27
LAVI (mL/m ²)	110.9±57.6	13.6±66.3	0.78
LVVI (mL/m ²)	67.2±32.4	57.9±26.1	0.05

Data are presented as number (%) or the mean±standard deviation. Significant p values are shown in italics. Left atrial volume index and left ventricular volume index were measured using left atrial or ventricular volume acquired by Simpson's method from a 4-chamber apical view, indexed to the body surface area. AF = atrial fibrillation; EF = ejection fraction; LAA = left atrial appendage; LAD = left anterior descending artery; LAVI = left atrial volume index; LVVI = left ventricular volume index; TTE = transthoracic echocardiography.

in the non-occlusion and occlusion groups were matched in a 1:1 to 1:4 manner, respectively, using nearest-neighbor matching. After propensity score matching (PSM), the balance between the 2 groups was assessed by calculating the standardized mean difference (SMD) between selected variables, with a standardized mean difference <0.20 suggesting appropriate balance (**Supplementary Figure 1**), and the discrimination of the propensity score model was assessed using C-statistic (C-statistic, 0.798). Thereafter, continuous variables were compared using the paired t-test, and categorical variables were compared using the McNemar test.

Statistical analysis was performed using the SPSS software version 29.0 (IBM Corp., Armonk, NY, USA) and R statistical software version 4.0.2 (R Foundation for Statistical Computing, Vienna, Austria).



RESULTS

Patient characteristics

Between January 2015 and August 2021, a total of 252 patients underwent the maze procedure using cryoablation. Based on the criteria, we excluded 54 (concomitant valve replacement using a mechanical prosthetic valve) and 10 (pre-operative existing spontaneous echo contrast or intra-cardiac thrombus) patients. Eventually, 188 patients were enrolled in the final cohort. The non-occlusion group and occlusion group included 113 and 75 patients, respectively. The median follow-up duration was 44 months (range: 1–94 months).

Patient characteristics are listed in **Table 1**. Patients in the non-occlusion group were older than those in the occlusion group (67.3 ± 8.7 years vs. 65.0 ± 7.8 years, respectively; p=0.06). The percentage of patients with degenerative valve disease was significantly higher in the non-occlusion group (23.0% vs. 10.7%, respectively; p=0.03). Fewer patients in the non-occlusion group underwent aortic valve replacement (5.3% vs. 18.6%, respectively; p=0.01) or coronary artery bypass grafting surgery (4% vs. 15.0%, p=0.02) as concomitant surgery. Moreover, preoperative cerebral infarction history was less frequent in the non-occlusion group (14.2% vs. 24%, respectively; p=0.09). In pre-operative TTE, the left ventricular volume index was significantly larger in the non-occlusion group (67.2 ± 32.4 vs. 57.9 ± 26.1 , respectively; p=0.05).

After PSM, 53 patients from each group were extracted. There were no significant differences in baseline characteristics between the 2 groups (**Table 2**); the mean follow-up period for the non-occlusion and occlusion groups was 50.0±28.8 and 51.1±24.4 months, respectively.

Clinical outcomes

In cohort of propensity-matched patients, hospital stay and postoperative complications were not significantly different between the 2 groups (**Table 3**). Cardiopulmonary bypass (CPB) time was shorter in the non-occlusion group compared with the occlusion group (143.7±41.2 vs. 160.8±47.4 min, respectively; p=0.06). Significantly more patients in non-occlusion group were managed with warfarin at 1 year after operation (34.0% vs. 18.9%, p=0.03). During the follow-up period, 10 patients underwent PMK insertion (3 vs. 7, respectively; p=0.32). At 1, 3, and 5 years after the operation, the percentages of patients in normal sinus rhythm were 74.0% vs. 82.4% (p=0.31), 76.9% vs. 88.5% (p=0.24), and 73.9% vs. 77.8% (p=0.78), respectively (**Figure 1**). Likewise, in results of entire cohort, there was no significant difference in ratio of normal sinus rhythm (**Supplementary Figure 2**).

The results of entire cohort are described in **Supplementary Table 1**. CPB time was significantly longer in occlusion group (162.7±46.9 vs. 146.2±40.8, p=0.01) and more patients of occlusion group were managed with non-vitamin K antagonist oral anticoagulant (35.7% vs. 20.6%, p=0.03) than occlusion group.

Major adverse cardiac events

In propensity-matched patients, there were 2 cases of cerebral infarction in the nonocclusion group, 5 cases of hospital readmission due to heart failure (non-occlusion group: n=3, occlusion group: n=2), and one case of cardiac death in the occlusion group during the follow-up period (**Figure 2**). Of the 2 cases of stroke in the non-occlusion group, one patient underwent mitral valve replacement for rheumatic mitral valve insufficiency (5 years after operation, focal infarction at left corona radiate), while the other patient underwent aortic

	Non-occlusion group (n=53)	Occlusion group (n=53)	p value
Sex, male	27 (50.9)	26 (49.1)	0.846
Age (years)	65.0±8.9 65.5±7.4		0.741
Heart disease etiologies			
Rheumatic disease	16 (30.2)	17 (32.1)	0.834
Degenerative	4 (7.5)	6 (11.3)	0.741
Functional	34 (64.2)	32 (60.4)	0.689
Others	1 (1.9)	1 (1.9)	>0.99
Concomitant operations with maze procedure			
Mitral valve replacement	10 (18.9)	12 (22.6)	0.632
Mitral valve repair	23 (43.4)	18 (34.0)	0.319
Tricuspid annuloplasty	27 (50.9)	27 (50.9)	>0.99
Aortic valve replacement	4 (7.5)	4 (7.5)	>0.99
Aortic valve plasty	2 (3.8)	1 (1.9)	>0.99
Coronary artery bypass grafting	4 (7.5)	2 (3.8)	0.678
Aortic replacement	2 (3.8)	1 (1.9)	>0.99
Atrial septal defect repair	2 (3.8)	4 (7.5)	0.678
Myxoma removal	2 (3.8)	0	0.495
AF profiles			
Fine (<1 mm) AF wave	29 (54.7)	28 (52.8)	0.846
AF type			>0.99
Persistent	35 (66.0)	35 (66.0)	
Paroxysmal	18 (34.0)	18 (34.0)	
Heart failure	12 (22.6)	13 (24.5)	0.819
Coronary artery disease	3 (5.7)	3 (5.7)	>0.99
Cerebral infarction	8 (15.1)	11 (20.8)	0.447
Diabetes mellitus	11 (20.8)	10 (18.9)	0.807
Aortic disease	2 (3.8)	1 (1.9)	>0.99
Hypertension	26 (49.1)	23 (43.4)	0.559
Chronic obstructive pulmonary disease	2 (3.8)	2 (3.8)	>0.99
Chronic kidney disease	0	1 (1.9)	>0.99
CHA ₂ DS ₂ -VASc score	2.5±1.7	2.6±1.7	0.738
Pre-operative TTE			
EF (%)	55.3±11.1	55.6±10.9	0.895
LAD (cm)	5.7±1.1	5.5±1.0	0.410
LAVI (mL/m²)	114.5±63.3	109.9±65.1	0.709
LVVI (mL/m ²)	66.9±28.4	59.5±27.8	0.175

Table 2. Baseline characteristics of propensity-matched patients

Data are presented as number (%) or the mean±standard deviation. Left atrial volume index and left ventricular volume index were measured using left atrial or ventricular volume acquired by Simpson's method from a 4-chamber apical view, indexed to the body surface area.

AF = atrial fibrillation; EF = ejection fraction; LAA = left atrial appendage; LAD = left anterior descending artery; LAVI = left atrial volume index; LVVI = left ventricular volume index; TTE = transthoracic echocardiography.

Table 3. Operative outcomes of propensity-matched patients

	Non-occlusion group (n=53)	Occlusion group (n=53)	p value
Hospital stay after operation (days)	14.9±8.0	13.4±7.0	0.30
Aortic cross clamp time (minutes)	102.4±34.5	110.5±36.0	0.25
Cardiopulmonary bypass time (minutes)	143.7±41.2	160.8±47.4	0.06
Early hospital mortality	1 (1.9)	1 (1.9)	>0.99
Bleeding control	2 (3.8)	2 (3.8)	>0.99
NSR at discharge from hospital	38 (71.7)	37 (69.8)	0.83
Results at 1-year follow-up (44 of non-occlusion and 49 patients of occlusion group)			
NSR	37 (69.8)	42 (79.2)	0.83
Anti-platelet drug usage	21 (39.6)	20 (37.7)	0.54
Anticoagulation drug usage	30 (56.6)	25 (47.2)	0.09
NOAC	12 (22.6)	15 (28.3)	0.72
Warfarin	18 (34.0)	10 (18.9)	0.03

Data are presented as number or the mean±standard deviation. Significant p values are shown in italics. NOAC = non-vitamin K antagonist oral anticoagulant; NSR = normal sinus rhythm.

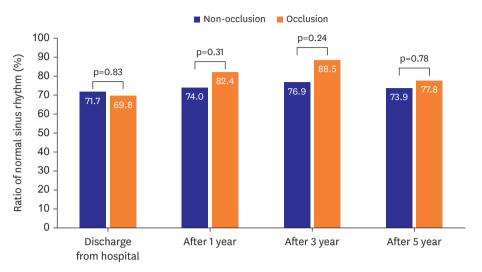


Figure 1. The ratio of patients with normal sinus rhythm after maze procedure of propensity-matched patients.

valve replacement for degenerative aortic valve stenosis (7 months after operation, tiny infarction at both postero-cerebral artery territories). Both patients are currently alive, with mild complications from the cerebral infarction.

The results of the Kaplan-Meier analysis for long-term freedom from MACE, stroke did not show significant differences between the groups in propensity-matched patients and entire cohort (**Figure 2**). There was no significant difference in the rate of freedom from stroke (p=0.19), readmission due to heart failure (p=0.74), or permanent PMK insertion (p=0.24) between the 2 groups in propensity-matched patients. The rate of freedom from cardiac death was also not significantly different in propensity-matched patients (p=0.93) and entire cohort (p=0.61) (**Figure 3**).

In the entire cohort of patients, multivariable analysis (**Table 4**) showed that age of patient (HR, 1.085; 95% CI, 1.005–1.172; p=0.04) and rheumatic heart disease (HR, 4.509; 95% CI, 1.115–11.494; p=0.04) were significant risk factors for the occurrence of cerebral infarction.

Hemodynamic effects of left atrial appendage obliteration

Figure 4 shows the TTE results of patients with normal sinus rhythm in propensity-matched patients. As shown by postoperative follow-up TTE, in those patients, the rate of mechanical LA contraction maintenance was not significantly different between the non-occlusion and occlusion groups (51.4% in both groups; p>0.99). However, this rate at 1-year follow-up was significantly higher in the former group (72.7% vs. 48.6%, respectively; p=0.04). The result of entire cohort (**Supplementary Figure 3**) was similar, and more patients of non-occlusion group showed effective mechanical LA contraction than occlusion group at 1 year after operation (80.9% vs. 60%, p=0.02).

DISCUSSION

AF is common in elderly patients¹⁰⁾ and is responsible for approximately a quarter of ischemic strokes¹¹; of those, many are cardioembolic¹²⁾ and originate from the LAA.²⁾ In this study, we analyzed the effects of LAA non-occlusion on hemodynamics and the prevention of cerebral

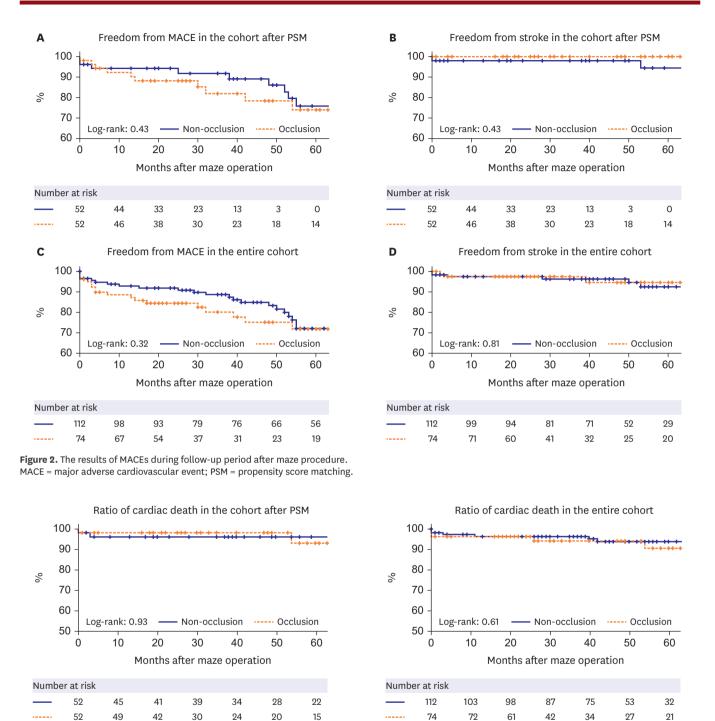


Figure 3. The result of freedom from cardiac death ratio after the maze procedure in both groups. PSM = propensity score matching.

infarction. The key finding of this study was that LAA preservation did not significantly increase the rate of cerebral infarction. In addition, LAA preservation significantly improved LA function.

In a recently reported randomized trial (LAAOS III),⁷⁾ the risk of ischemic stroke or systemic embolism was lower in patients with AF who underwent concomitant LAA occlusion

Variable	Univariate		Multivariate	
variable	HR (95% CI)	p value	HR (95% CI)	p value
Sex, male	0.467 (0.117-1.870)	0.28		
Age	0.947 (0.883-1.015)	0.12	1.085 (1.005-1.172)	0.04
Rheumatic heart disease	4.584 (1.141-18.415)	0.03	0.285 (0.708-11.494)	0.14
Degenerative valve disease	0.638 (0.080-5.112)	0.67		
Functional MR	0.425 (0.106-1.701)	0.23		
Fine (<1 mm) AF wave	1.474 (0.395-5.494)	0.56		
Persistent AF	1.068 (0.282-4.043)	0.92		
Comorbidities				
Heart failure	2.597 (0.697-9.680)	0.16	1.951 (0.503-7.564)	0.33
Cerebral infarction	2.458 (0.613-9.854)	0.20	3.728 (0.824-16.863)	0.09
Diabetes mellitus	2.041 (0.510-8.176)	0.31		
Hypertension	1.023 (0.274-3.821)	0.97		
CHA ₂ DS ₂ -VASc score	1.134 (0.796-1.615)	0.49		
Postoperative TTE				
EF	0.979 (0.925-1.037)	0.48		
LAD	0.990 (0.827-1.186)	0.92		
LAVI	1.001 (0.991-1.012)	0.83		
LVVI	0.990 (0.962-1.019)	0.51		

Table 4. Multivariable analysis for stroke or the entire cohort	(Cox proportional hazards regression model)	1

Significant p values are shown in italics.

AF = atrial fibrillation; CI = confidence interval; EF = ejection fraction; HR = hazard ratio; LAD = left anterior descending artery; LAVI = left atrial volume index; LVVI = left ventricular volume index; MR = mitral regurgitation; TTE = transthoracic echocardiography.

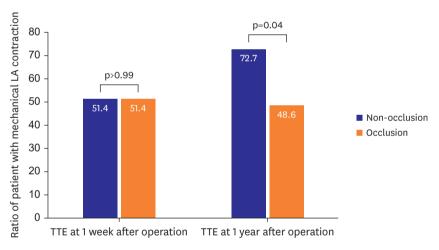


Figure 4. Ratio of patients who have effective left atrial mechanical contraction in patient with normal sinus rhythm (in propensity-matched patients). LA = left atrial; TTE = transthoracic echocardiography.

during the cardiac operation than in those who did not. In the LAAOS III clinical trial, the rate of ischemic stroke or systemic embolism occurrence was 7.0% and 4.8% in the LAA non-occlusion group and occlusion group, respectively. These rates were higher than those recorded in the present investigation (3.8% and 0%, respectively) and another study reported by Lee et al.¹³ (2.5% and 1.7%, respectively). In particular, there was a significant difference in the patients included in the LAA non-occlusion group between the LAAOS III study and the present study. In the LAAOS III trial, only 32.7% of patients (1,562 of 4,770 patients) underwent concomitant surgical ablation of AF. Therefore, although the exact number was not described in this report, most patients may have been at an AF state during the follow-up period. Thus, the LAAOS III trial was focused on patients in an AF state rather than on patients who underwent the maze procedure.

In the multivariate analysis of this study, rheumatic heart disease and advanced age were identified as significant risk factors for stroke, not LAA preservation. In previous studies, rheumatic valve disease and a multiple valvular disease have been linked to worse outcomes, including stroke.⁹⁾⁽¹⁴⁾⁽⁵⁾ However, other reports,⁹⁾⁽¹³⁾ including this study, did not show a significant difference in the rates of stroke between the LAA non-occlusion and occlusion groups. Nevertheless, the relatively small number of patients affected by stroke who are included in studies should be increased to obtain adequate statistical power.

Regarding the effects of LAA occlusion on cardiac hemodynamics, the non-occlusion group showed significantly more frequent LA mechanical contraction on TTE 1 year after the maze procedure compared with the occlusion group. These results showed that preservation of the LAA assisted in the LA booster contribution to ventricular filling, particularly 1 year after the operation. Buber et al.⁹⁾ and Lee et al.¹³⁾ also reported superior results of LA contractile in LAA preservation group than LAA obliteration group. This is because the LAA accounts for 17–30% of the total LA volume, ¹⁴⁻¹⁶⁾ and LAA contractile function contributes greatly to the LA transport function. ⁵⁾⁹⁾¹⁷⁾ The absence of LA mechanical contraction after the maze procedure should alert physicians to the possible risk of future strokes. Buber et al.⁹⁾ reported that the absence of LA contraction and an LA volume index \geq 33 mL/m² resulted in a significant increase in the risk of thromboembolic stroke after the maze procedure for patients with sinus rhythm. This finding suggests that LA mechanical standstill is an independent and powerful risk factor for thromboembolic stroke. This should be taken into account when considering anticoagulation treatment for patients following the maze procedure.

This study has several limitations. First, the data in this study were retrospectively analyzed in a small number of cohort. Although several results showed large differences in ratio including the result of more stroke in LAA preservation group despite a more anticoagulant therapy than non-occlusion group, there were not significant differences due to the limited number of cohort. Also, this was not a randomized controlled trial; thus, this study has limited statistical power. Secondly, follow-up screening for AF recurrence was based on serial 12-lead ECG rather than 24-hour Holter monitoring. Hence, it is possible that our results were overestimated because asymptomatic paroxysmal AF may have been present during the follow-up period without appearing on subsequent ECG examinations carried out in outpatient clinics. Third, evaluation for stroke using brain magnetic resonance imaging was performed only in patients with neurological symptoms rather than in all patients. Therefore, it is possible that asymptomatic cases of stroke may have been missed, resulting in an underestimated occurrence rate of stroke in our cohort. Forth, the shapes of appendages are also an important factor for cerebral infarction and LA contractile.¹⁸⁻²⁰⁾ However, it is difficult to analyze the shape of LAA on cardiac CT or TTE. The most accurate method to investigate the shape of LAA is external appearance observation during cardiac surgery. Unfortunately, this was a retrospective study using operative records, and most of our hospital's records did not describe the external appearance of the LAA shape. Finally, there was heterogeneity in the indication of concomitant cardiac surgery. For this reason, the impact of concomitant cardiac surgery on outcomes of maze procedure was not totally excluded.

In conclusion, preservation of LAA during the maze procedure effectively maintained LA contractile function. The result of stroke outcome was not significantly different between LAA occlusion and LAA preservation during the maze procedure. However, further long-term studies involving larger patient populations are required to verify the present results.



SUPPLEMENTARY MATERIALS

Supplementary Table 1

Operative outcomes of the entire cohort

Click here to view

Supplementary Figure 1

The results of matching. (A) Propensity score distribution of both groups (before and after propensity score matching). (B) A standardized mean difference (SMD) was used to examine the degree of PSM. A threshold of less than 0.1 was considered acceptable.

Click here to view

Supplementary Figure 2

The ratio of patients with normal sinus rhythm after maze procedure of the entire cohort.

Click here to view

Supplementary Figure 3

Ratio of patients who have effective left atrial mechanical contraction in patient with normal sinus rhythm (in the entire cohort).

Click here to view

REFERENCES

- Wolf PA, Dawber TR, Thomas HE Jr, Kannel WB. Epidemiologic assessment of chronic atrial fibrillation and risk of stroke: the Framingham study. *Neurology* 1978;28:973-7.
 PUBMED | CROSSREF
- Blackshear JL, Odell JA. Appendage obliteration to reduce stroke in cardiac surgical patients with atrial fibrillation. *Ann Thorac Surg* 1996;61:755-9.
 PUBMED | CROSSREF
- 3. García-Fernández MA, Pérez-David E, Quiles J, et al. Role of left atrial appendage obliteration in stroke reduction in patients with mitral valve prosthesis: a transesophageal echocardiographic study. *J Am Coll Cardiol* 2003;42:1253-8.

PUBMED | CROSSREF

- Stöllberger C, Schneider B, Finsterer J. Elimination of the left atrial appendage to prevent stroke or embolism? Anatomic, physiologic, and pathophysiologic considerations. *Chest* 2003;124:2356-62.
 PUBMED | CROSSREF
- Yamanaka K, Sekine Y, Nonaka M, et al. Left atrial appendage contributes to left atrial booster function after the maze procedure: quantitative assessment with multidetector computed tomography. *Eur J Cardiothorac Surg* 2010;38:361-5.
 PUBMED | CROSSREF
- Yoshihara F, Nishikimi T, Kosakai Y, et al. Atrial natriuretic peptide secretion and body fluid balance after bilateral atrial appendectomy by the maze procedure. *J Thorac Cardiovasc Surg* 1998;116:213-9.
 PUBMED | CROSSREF
- Whitlock RP, Belley-Cote EP, Paparella D, et al. Left atrial appendage occlusion during cardiac surgery to prevent stroke. N Engl J Med 2021;384:2081-91.
 PUBMED | CROSSREF
- Buber J, Luria D, Sternik L, et al. Left atrial contractile function following a successful modified Maze procedure at surgery and the risk for subsequent thromboembolic stroke. *J Am Coll Cardiol* 2011;58:1614-21.
 PUBMED | CROSSREF



- Greenland S, Mickey RM. Re: "The impact of confounder selection criteria on effect estimation". *Am J Epidemiol* 1989;130:1066.
 PUBMED | CROSSREF
- Go AS, Hylek EM, Phillips KA, et al. Prevalence of diagnosed atrial fibrillation in adults: national implications for rhythm management and stroke prevention: the AnTicoagulation and Risk Factors in Atrial Fibrillation (ATRIA) Study. *JAMA* 2001;285:2370-5.
- Gage BF, van Walraven C, Pearce L, et al. Selecting patients with atrial fibrillation for anticoagulation: stroke risk stratification in patients taking aspirin. *Circulation* 2004;110:2287-92.
 PUBMED | CROSSREF
- 12. Stroke prevention in atrial fibrillation study. Final results. *Circulation* 1991;84:527-39. PUBMED | CROSSREF
- Lee CH, Kim JB, Jung SH, Choo SJ, Chung CH, Lee JW. Left atrial appendage resection versus preservation during the surgical ablation of atrial fibrillation. *Ann Thorac Surg* 2014;97:124-32.
 PUBMED | CROSSREF
- Hondo T, Okamoto M, Yamane T, et al. The role of the left atrial appendage. A volume loading study in open-chest dogs. *Jpn Heart J* 1995;36:225-34.
 PUBMED | CROSSREF
- Yamamoto N. Experimental study of combined left atrium resection for lung cancer. Nippon Kyobu Geka Gakkai Zasshi 1986;34:958-65.
 PUBMED
- Isobe F, Kumano H, Ishikawa T, et al. A new procedure for chronic atrial fibrillation: bilateral appendagepreserving maze procedure. *Ann Thorac Surg* 2001;72:1473-8.
 PUBMED | CROSSREF
- Schneider B, Stollberger C, Sievers HH. Surgical closure of the left atrial appendage a beneficial procedure? *Cardiology* 2005;104:127-32.
 PUBMED | CROSSREF
- Bieging ET, Morris A, Chang L, Dagher L, Marrouche NF, Cates J. Statistical shape analysis of the left atrial appendage predicts stroke in atrial fibrillation. *Int J Cardiovasc Imaging* 2021;37:2521-7.
 PUBMED | CROSSREF
- Ogata J, Yutani C, Otsubo R, et al. Heart and vessel pathology underlying brain infarction in 142 stroke patients. *Ann Neurol* 2008;63:770-81.
 PUBMED | CROSSREF
- Virani SS, Alonso A, Benjamin EJ, et al. Heart disease and stroke statistics—2020 update: a report from the American Heart Association. *Circulation* 2020;141:e139-596.
 PUBMED | CROSSREF