

Long-Term Outcomes and Predictors of Recurrence in Atrial Arrhythmia Ablations Post-Fontan Procedure: A Retrospective Analysis

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Abstract

Background: Supraventricular tachycardia (SVT) is common in patients who have undergone a Fontan procedure and is poorly tolerated. SVT recurrence rates after catheter ablations are high. Recent data on the outcomes of SVT ablation and the predictors of recurrence in this population are limited.

Methods: Electronic medical records of patients who had undergone a Fontan procedure and SVT ablation between January 1, 1995, and October 1, 2023, at a tertiary care center were reviewed. Demographic, clinical, and outcome variables over 5 years were compared between patients with and without SVT recurrence.

Results: Twenty-five patients (56% male, mean age 31.5 ± 7.2 years) with a mean age of 6.1 ± 3.6 years at Fontan surgery were included. Ablation success rate was 92%. Recurrence occurred in 12 (48%) patients, of whom 7 (28%) required repeat ablations. Thirteen (59%) patients had a cardiac hospitalization, and one patient died in the 5-year follow-up period. Atypical (71%) and typical (33%) right atrial flutter were the most common SVTs. Higher brain natriuretic peptide (BNP) levels pre- (1,702 vs. 242, $P = 0.028$) and post-ablation (862 vs. 112, $P = 0.017$) were associated with recurrence. Atriopulmonary (AP) Fontan type (91% vs. 17%, $P = 0.0006$), number of radiofrequency (RF) applications (48 vs. 14, $P = 0.045$), post-ablation cardiovascular (CV) hospitalizations (82% vs. 36% $P = 0.030$), and post-ablation antiarrhythmic prescriptions (1.8 vs. 1.2, $P = 0.0256$) were more prevalent in patients with recurrence.

Conclusion: Catheter ablation of SVT in patients with Fontan physiology is associated with a high success rate and a high long-term recurrence rate. Recurrence of SVT is associated with markers of severe heart disease and type of Fontan.

Keywords: Supraventricular arrhythmia; Fontan; Ablation; Outcomes; Recurrence

Introduction

Fontan and hemi-Fontan surgery offers a surgical solution for patients with a functional single ventricle, resulting in an 85-95% survival rate at 15 years post-procedure [1, 2]. Despite surgical advances like the total cavopulmonary connection and extra-cardiac (EC) or lateral tunnel (LT) conduits, the burden of atrial or supraventricular arrhythmias after Fontan palliation remains high [3-5]. Atrial arrhythmias are poorly tolerated in Fontan physiology, potentially leading to reduced cardiac output and increased risk of thromboembolic disease [6-8].

Electrophysiology (EP) catheter ablation has been shown to be more effective than antiarrhythmic drugs for managing atrial arrhythmias in patients with Fontan physiology. Multiple studies have confirmed lower arrhythmia burden after the procedure [9-11]. Nevertheless, recurrence rates after ablation are substantial [12-16].

EP studies in patients with Fontan physiology can be challenging due to their complex cardiac anatomies [7, 17]. Little is known about the outcomes of EP ablation in this population in recent years. Previous studies have established the safety of catheter ablation for atrial arrhythmias but have incompletely investigated long-term outcomes [15, 18]. Our study aims to 1) identify clinical characteristics of patients with a history of Fontan palliation surgery who have undergone EP ablations, 2) evaluate the long-term outcomes of ablation in these patients, and 3) determine factors that may influence the recurrence of atrial arrhythmias.

Materials and Methods

This study was deemed exempt from review by the University of Texas (UT) Southwestern Institutional Review Board.

Study population

We conducted a retrospective study of patients at the UT Southwestern Medical Center with history of Fontan or hemi-

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Fontan procedure and EP ablation for SVTs between January 1, 1995, and October 1, 2023. Patients with Current Procedural Terminology (CPT) diagnosis codes for Fontan/hemi-Fontan procedures and EP ablation of SVTs were included in the study. Chart review was subsequently performed to confirm eligibility and abstract baseline data. Patients who had SVT ablations prior to a Fontan procedure were excluded from the cohort.

Data collection

We conducted a detailed chart review to collect demographic information, congenital heart disease (CHD) diagnosis, laboratory results, cardiac medication history, electrocardiogram (EKG) findings, transthoracic echocardiogram (TTE) results, right heart catheterization (RHC) data, procedural details, and outcomes measures. Procedural variables, such as length of procedure, number of applications, and location of arrhythmia, were abstracted from procedure notes. Typical right atrial (RA) flutter was defined as macro-reentry around the tricuspid annulus dependent on the cavo-tricuspid isthmus. Atypical RA flutter was defined as any macro-reentrant circuit not dependent on the cavo-tricuspid isthmus and included intra-atrial re-entrant tachycardia. Atrial tachycardia (AT) was defined as automaticity or micro-reentry circuits from sites ectopic to the sinus node. Cardiac medications were defined as antiarrhythmics (class I-V), anti-hypertensives, heart failure medications, diuretics, and anticoagulants including aspirin, warfarin, P2Y12 inhibitors, and direct oral anticoagulants. For pre- and post-ablation variables, only the most recent data within 2 years of the index procedure were recorded.

Outcome measures

Outcome measures of importance included recurrence of arrhythmia after first ablation, repeat ablation, hospitalizations, and death at 5 years. Procedural success was determined by clinician documentation in the ablation procedure note. Recurrence of atrial arrhythmia was defined by documentation of recurrent tachycardia symptoms, EKG confirmation of SVT, or history of repeat atrial arrhythmia ablation. Recurrent arrhythmias were classified as same or different based on EKG analysis upon recurrence or documentation of arrhythmias during repeat ablation. Hospitalizations were identified through the electronic medical record system and were classified as cardiovascular (CV) or non-CV hospitalizations. CV hospitalizations were defined as admission to a medicine or cardiology service with cardiac chief complaint or cardiac primary problem in the initial history and physical documentation. The 5-year follow-up time point was selected to capture a clinically meaningful sample as has been done by Egbe et al, and recurrence rates before and after 2018 were analyzed given a lack of data in the literature on this subject since 2018 [9].

Statistical analysis

Descriptive statistics were used to analyze the variables for the

overall cohort. Student's *t*-tests, Fisher's exact test, and two-sample proportions tests were employed to investigate differences between patients with and without recurrence, with a significance level set at $\alpha < 0.05$.

Results

Twenty-six patients at UT Southwestern had a history of a Fontan procedure and SVT ablation. One patient with ablation before the Fontan procedure was excluded from the study. The 25 participants included had a total of 34 ablations; 12 (48%) had a documented recurrence within 5 years.

Demographics

The mean age ($\pm$ SD) at Fontan surgery was 6.1 ± 3.6 years and the mean age at ablation procedure was 31.5 ± 7.2 years. Fourteen (56%) patients were male. The average body mass index (BMI) of the patients was 26.3 ± 4.7 kg/m². The most common CHDs in the cohort were tricuspid atresia (TA, 40%) and transposition of great arteries (TGA, 35%). AP (48%) and fenestrated LT (36%) were the most common initial Fontan types (Table 1).

Clinical covariates and predictors of recurrence

Forty-four percent of patients who had an ablation after 2018 and 55% of those who had an ablation before 2018 had recurrence ($P = 0.6$). Similarly, 42% of patients who had an ablation after 2020 and 54% of those who had one earlier had recurrence ($P = 0.5$). Of those with an AP approach, 10 (91%) had recurrence while of those with an LT approach, only 1 (11%) had recurrence ($P = 0.0006$ and $P = 0.0094$, respectively). Among the patients with AP Fontan procedure who had a recurrence, two had a revision to EC.

Patients with Fontan palliation undergoing SVT ablation were prescribed on average 4.00 cardiac medications before the ablation and 3.91 medications after their first post-procedural visit. Patients who later developed recurrence had significantly more anti-arrhythmic medications prescribed post-ablation than those without recurrence (1.8 vs. 1.2, $P = 0.03$). The most common post-ablation anti-arrhythmic medications prescribed were metoprolol (52%), digoxin (24%), sotalol (20%), and dofetilide (16%). There were no significant differences in the prescription patterns of these medications between patients with and without recurrence. Six (24%) patients had a history of prior implantable cardioverter defibrillator/pacemaker (ICD/PPM) and 11 (44%) had a history of prior cardioversion (Table 1).

Brain natriuretic peptide (BNP) levels decreased after ablation (1,054 to 524). Those with recurrence had higher BNP levels both before (1,702 vs. 242, $P = 0.03$) and after the ablation (862 vs. 112, $P = 0.02$). Thirteen (52%) patients had a history of heart failure and 10 of the 21 with echocardiography data had an ejection fraction (EF) $< 50\%$, but neither metric was significantly different between those with and without recurrence (Table 1). Fourteen patients had RHC performed within 2 years of

Table 1. Patient Demographic and Clinical Correlates in Patients With and Without Arrhythmia Recurrence

	Overall (n = 25)	Recurrence (n = 12)	No recurrence (n = 13)	P-value
Age at Fontan (years)	6.1	5.6	6.6	0.525
Age at ablation (years)	31.5	33.3	29.9	0.253
Male, n (%)	14 (56)	9 (75)	5 (42)	0.111
BMI	26.3	27.8	25	0.18
Type of Fontan (initial)				
Atriopulmonary, n (%)	12 (48)	10 (91)	2 (17)	0.0006
Lateral tunnel, n (%)	9 (36)	1 (9)	8 (67)	0.0094
Extra-cardiac, n (%)	2 (8)	0 (0)	2 (17)	0.378
Congenital heart disease ^a				
TA, n (%)	10 (40)	8 (67)	2 (17)	0.0154
D-TGA/L-TGA, n (%)	9 (35)	4 (33)	5 (42)	1
DORV, n (%)	3 (12)	0 (0)	3 (25)	0.22
DILV, n (%)	3 (12)	1 (8.3)	2 (17)	1
HLHS, n (%)	4 (16)	0 (0)	4 (33)	0.0957
PA-IVS, n (%)	4 (16)	3 (25)	1 (8.3)	0.3
Other, n (%)	3 (12)	1 (8.3)	2 (17)	1
Pre-ablation cardiac medications	4	4.5	3.8	0.269
Antiarrhythmic	1.7	1.8	1.6	0.444
Anti-coagulation	0.96	1.09	0.92	0.205
Post-ablation cardiac medications	3.91	4.6	3.58	0.061
Antiarrhythmic	1.41	1.8	1.2	0.0256
Anti-coagulation	0.92	1.1	0.92	0.468
Pre-BNP	1,054	1,703	242	0.028
Post-BNP	524	862	111.9	0.017
Normal VSF (EF > 50%), n (%)	11 (52)	6 (54)	5 (50)	0.687
Mildly reduced VSF (EF 40-50%), n (%)	5 (24)	3 (27)	2 (20)	1
Significantly reduced VSF (EF < 40%), n (%)	5 (24)	2 (18)	3 (30)	1
Comorbidities				
Prior HF, n (%)	13 (52)	7 (58)	6 (46)	0.695
Prior thrombosis, n (%)	4 (16)	0 (0)	4 (31)	0.0957
Prior PPM/ICD, n (%)	6 (24)	4 (33)	2 (12)	0.378
Prior cardioversion, n (%)	11 (44)	7 (58)	5 (38)	0.434
CCI (mean)	1.27	1.09	1.38	0.59

^aCongenital heart diseases were not in isolation and those such as TGA were associated with other lesions. BMI: body mass index; BNP: brain natriuretic peptide; CCI: Charlson Comorbidity Index; DILV: double inlet left ventricle; DORV: double outlet right ventricle; D-TGA/L-TGA: dextro-transposition of the great arteries/levo-transposition of the great arteries; EF: ejection fraction; HF: heart failure; HLHS: hypoplastic left heart syndrome; PA-IVS: pulmonary atresia with intact ventricular septum; PPM/ICD: pacemaker/implantable cardioverter defibrillator; TA: tricuspid atresia; VSF: ventricular systolic function.

the EP ablation procedure. There was no significant difference in the RHC or TTE data between participants with and without recurrence (Supplementary Material 1, cr.elmerpub.com).

Procedural variables

Procedural data were obtained for 22 of the 25 patients. Three

patients had procedures done at other facilities where detailed procedural notes were unavailable. The most common arrhythmias identified were atypical RA flutter (71%) and typical RA flutter (33%). One patient had left atrial flutter, two had focal ATs, and one had an atrioventricular nodal re-entrant tachycardia. There was no significant difference in procedure length, arrhythmia types, and arrhythmia location between the recurrent and nonrecurrent subgroups. Patients with recurrence had

Table 2. Procedural Correlates in Patients With and Without Arrhythmia Recurrence

	Overall (n = 23)	Recurrence	No recurrence	P-value
Length of procedure (min)	204	217	194	0.601
Number of applications	28.1	48	14	0.045
Mean cycle length (s)	725	761	691	0.642
Multiples arrhythmia, n (%)	18 (82)	10 (83)	8 (73)	
Initial arrhythmia				
Typical RA flutter	7 (33)	4 (40)	3 (27)	0.7
Atypical RA flutter	15 (71)	8 (80)	7 (63)	0.6
Type of ablation catheter				
Biosense Thermocool SmartTouch SFTM, n (%)	16 (73)	8 (73)	8 (73)	1
TactiCath AbbottTM, n (%)	6 (23)	3 (23)	3 (23)	1
Type of mapping catheter				
CARTO 3D® System, n (%)	12 (55)	6 (55)	6 (55)	1
Abbott®, n (%)	3 (14)	2 (18)	1 (9.1)	1
Other, n (%)	3 (14)	1 (9)	2 (18)	1
Location of arrhythmia				
Lateral RA/RA scar	7 (28)	4 (40)	3 (27)	0.659
Cavotricuspid isthmus, n (%)	6 (24)	3 (30)	3 (27)	1
Peri-IVC, n (%)	2 (8.0)	2 (20)	0 (0)	0.214
Posterior RA, n (%)	3 (12)	1 (10)	2 (18)	1
Anterior-septal RA, n (%)	4 (16)	3 (30)	1 (9.1)	0.59
Inferior RA, n (%)	1 (8.0)	1 (10)	0 (0)	1
Between SVC and RA, n (%)	3 (12)	0 (0)	3 (27)	0.214
SVC/SVC scar, n (%)	1 (8.0)	0 (0)	1 (9.1)	1
Around the sinus node, n (%)	1 (8.0)	0 (0)	1 (9.1)	1
Left atrial appendage/mitral annulus, n (%)	1 (8.0)	1 (10)	0 (0)	1

Not all values were available for all included patients; values shown are calculated from available data. IVC: inferior vena cava; RA: right atrium; SVC: superior vena cava.

significantly more applications than those who did not (48 vs. 14, $P = 0.045$, Table 2). Six (23%) patients had documented transeptal puncture during the procedure, of which four had LT, one had EC, and one had AP that was revised to EC Fontan type.

Outcomes

Medium follow-up period was 3.3 ± 3.7 years. Ablation was noted to be successful in 23 of the 25 cases (92%). Two patients had minor complications after the ablation, one with a groin hematoma and the other with mild post-discharge femoral bleeding which was managed conservatively. Of the 12 patients who experienced SVT recurrence, 10 (83%) had a different arrhythmia.

The mean time to recurrence was 1.4 ± 0.93 years. Two patients (8.0%) had a recurrence at 6 months and four (16.0%) had a recurrence by 1 year. Seven (28%) patients had a repeat ablation and the median time to repeat ablation was 2.70 years.

One patient in the cohort died in the 5-year follow-up period.

Thirteen (59%) of the 22 patients with follow-up hospitalization data available had cardiac hospitalizations within 5 years, while seven (32%) had non-cardiac hospitalization. At 5 years, nine patients (82%) in the recurrence and four patients (36%) in the nonrecurrence groups had cardiac hospitalizations (Table 3, $P = 0.030$).

Discussion

Atrial arrhythmias are a significant concern in patients who have undergone Fontan palliation given their high prevalence and potential for adverse outcomes. Our study aimed to investigate long-term outcomes in Fontan patients undergoing EP ablation and determine factors that are associated with recurrence. We found that nearly half of the patients with SVT ablation after Fontan surgery had recurrence and over a quarter had repeat ablations at 5 years. We identified elevated BNP levels, more RF applications, and AP Fontan type to be associated

Table 3. Hospitalization Outcomes in Patients With and Without Arrhythmia Recurrence

Cumulative hospitalizations data	Overall (n = 22)	Recurrence (n = 11)	No recurrence (n = 11)	P-value
CV hospitalization				
Within 6 months, n (%)	6 (27)	3 (27)	3 (3)	1
By 1 year, n (%)	8 (36)	4 (36)	4 (36)	1
By 5 years, n (%)	13 (59)	9 (82)	4 (36)	0.03
Non-CV hospitalization				
Within 6 months, n (%)	1 (4.5)	0 (0)	1	0.308
By 1 year, n (%)	2 (9.1)	0 (0)	2	0.139
By 5 years, n (%)	7 (32)	3 (27)	4 (36)	0.645

CV: cardiovascular.

with increased risk of arrhythmia recurrence.

Limited studies conducted prior to 2018 on this patient population have shown that freedom from atrial arrhythmia recurrence at 60 months was 41% and that 29% of patients underwent repeat ablation [9, 11]. This contemporary study mirrors these findings, revealing a freedom from recurrence at 60 months of 52% and a repeat ablation rate of 28%. A procedural success rate of 92% falls in the range of these past studies as well (83-94%). Consistent with prior studies, the risk of recurrence seems to increase with time. The similarities lend validity to our findings (despite the small sample size) and re-demonstrate that catheter ablation is safe and effective in managing SVT in this patient population.

This study found no significant difference in recurrence between ablations before and after 2018 or 2020, though there was notably a 10-15% reduction of recurrence in recent years. Although the small sample may have been insufficient to capture this difference, these results indicate that there has not been a major change in outcomes in recent years. Rates of hospitalization post-ablation are substantial, with a 5-year CV hospitalization rate of 59%. This was the first study to report that patients with recurrence were more likely to experience CV hospitalization post-ablation than those without. Whether arrhythmia recurrence contributes to CV hospitalization or is simply a consequence of severe underlying disease from deteriorating Fontan hemodynamics and atrial stress is unknown and requires further investigation.

It is known that AP Fontan type is associated with an increased risk of atrial arrhythmia [4]. This study expands on these findings by showing that an AP approach is also strongly associated with a much higher rate of recurrence after ablation than the LT approach. With 90% of those with an AP Fontan having recurrence of arrhythmias, upfront rhythm control with cardioversion or ablation, as well as earlier consideration of transplant workup may be clinically prudent in these patients.

BNP levels were associated with arrhythmia recurrence. The lower BNP levels post-ablation may indicate the effect of ablation on reducing the deleterious effects of arrhythmias on cardiac hemodynamics [19]. Patients with recurrence had significantly higher baseline BNP pre- and post-ablation, suggesting that BNP may be a predictor of outcomes in this population as it has shown to be in other atrial arrhythmias [20].

Patients with recurrence had more procedural applications during the ablation and were prescribed more antiarrhythmic medications post-ablation than those without recurrence. These findings complement prior studies demonstrating that ablating more circuits was associated with unfavorable outcomes, likely related to the presence of more complex arrhythmia substrate in general [9, 18]. The predictive factors noted here (BNP, antiarrhythmic medications, RF applications, etc.) may be helpful in identifying patients with more severe heart disease that may have a greater risk of arrhythmia recurrence.

While atypical flutters are generally associated with higher recurrence rates after ablation (7-53%) compared to typical flutter (11%), this study reveals a high recurrence rate in patients with typical (57%) and atypical flutter (53%) in this cohort [21]. This unexpected similarity in recurrence rates may be explained by the complex hemodynamic and atrial anatomy of patients with a Fontan as opposed to patients in the general population undergoing ablation (Figs. 1 and 2). Most patients in this cohort had 3D high density mapping performed during their EP study, especially in recent years, but a significant difference in recurrence rates with their use was not observed. While the use of these novel technologies has significantly aided ablation procedures by more accurately identifying and successfully ablating arrhythmia foci with less fluoroscopy exposure, their impact on long-term outcomes remains unclear. Future studies should evaluate the long-term benefits of novel mapping and ablation techniques (including pulsed field ablation), exploring how they may impact outcomes in patients with a Fontan and atrial arrhythmias.

Limitations

This observational retrospective study is limited by low power given a small sample size at a single tertiary center, incomplete follow-up, and descriptive nature of clinical variables. The characteristics of patients who were seen at this center may not be representative of all patients with a Fontan undergoing SVT ablations. Outcome estimates may be underestimated as it is possible that patients had hospitalizations, procedures, or follow-up at outside facilities that were not captured in our electronic health record. Multicenter prospective studies are needed to better assess outcomes after SVT ablation in patients who have undergone the Fontan procedure.

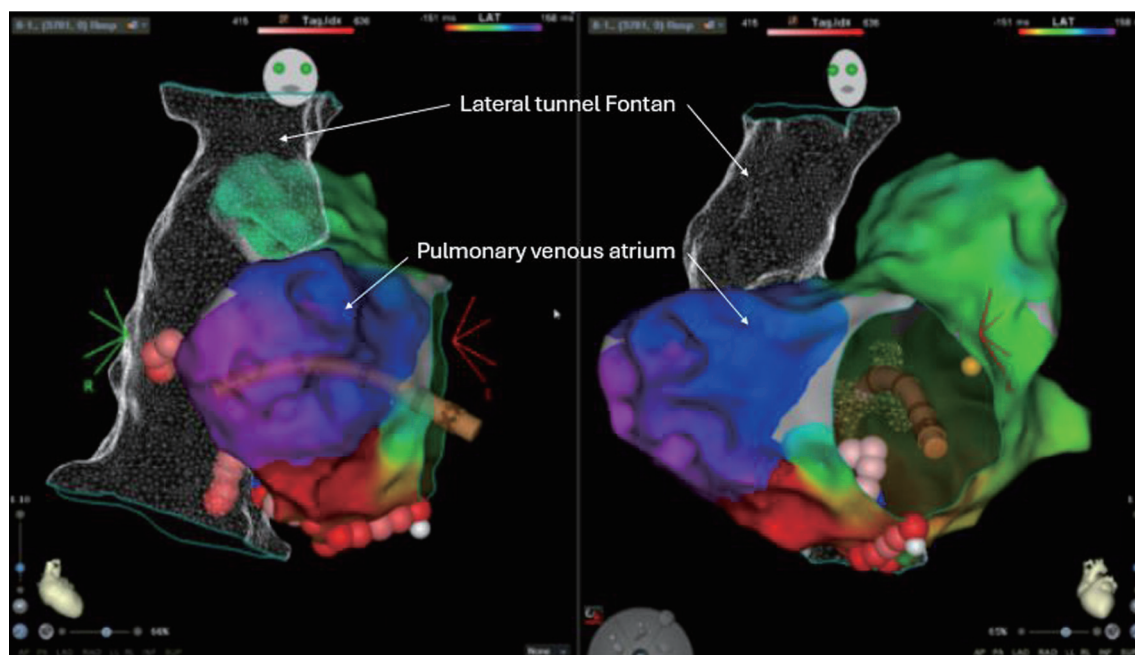


Figure 1. Electroanatomic map for a 26-year-old female with history of hypoplastic left heart syndrome with a lateral tunnel Fontan who underwent electrophysiology study and ablation of atrial flutter. Anterior/posterior (left) and left anterior oblique (right) projections show the lateral tunnel Fontan and the pulmonary venous atrium, which was accessed with trans-baffle puncture and a Vizigo sheath (Carto). Activation map showed counterclockwise cavotricuspid isthmus dependent atrial flutter, which was successfully ablated with ablation (red and pink dots) on both the pulmonary venous atrial and Fontan portion of the cavotricuspid isthmus.

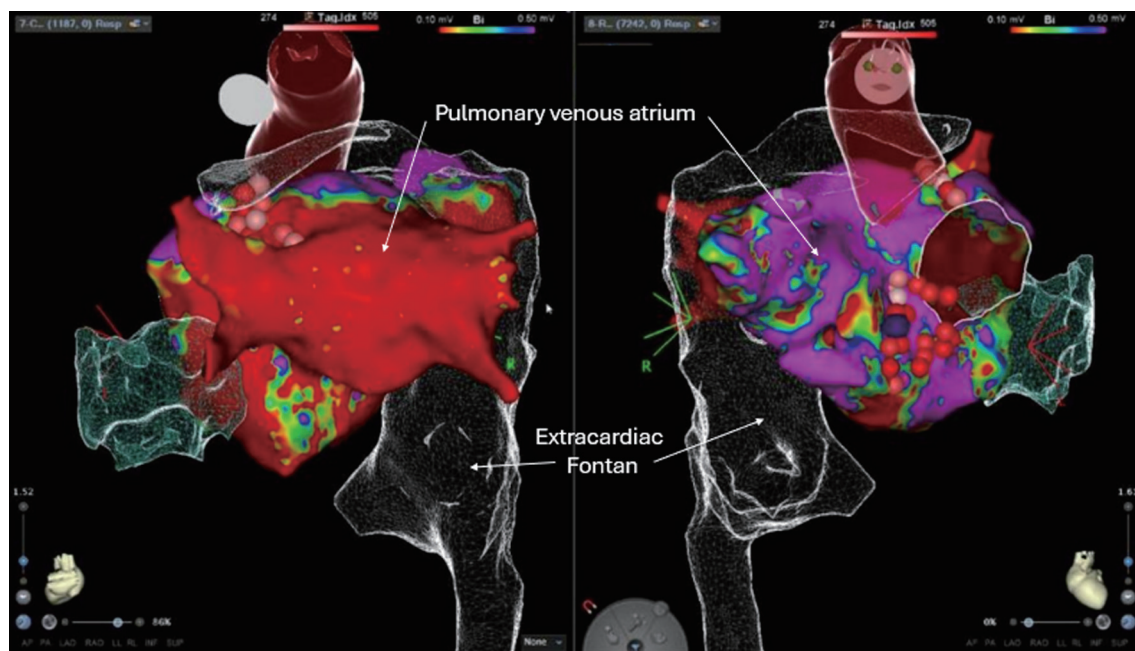


Figure 2. A 30-year-old male with tricuspid and pulmonary atresia status post bidirectional Glenn with anastomosis between the right-sided SVC and right pulmonary artery, classic right atrial appendage-pulmonary artery Fontan procedure, and history of bilateral cryo maze with conversion to extracardiac Fontan, and atrial reduction with septectomy presented for ablation of atypical atrial flutter. Electroanatomical 3D bipolar voltage and activation map created during atrial pacing from the epicardial pacemaker system is shown here. Activation map showed typical flutters with double loop mitral/septal scar re-entry and LA roof/LA appendage reentry that were ablated (red and pink dots). Trans-Fontan baffle puncture and balloon angioplasty of the Fontan baffle was performed for this procedure. LA: left atrial; SVC: superior vena cava.

Conclusions

Catheter ablation remains safe and effective for managing SVTs that develop after the Fontan procedure. Recurrence rates and repeat ablation rates at 5 years after ablation in this population remain high. Elevated pre- and post-ablation BNP levels and history of AP Fontan approach may help clinicians identify patients at higher risk of atrial arrhythmia recurrence after ablation. Patients with recurrence are at a greater risk of CV hospitalization at 5 years after their initial SVT ablation.

Supplementary Material

Suppl 1. Echocardiographic and hemodynamic correlates of atrial arrhythmia recurrence post-ablation in patients with Fontan procedure.

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Financial Disclosure

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Conflict of Interest

The authors have no conflict of interest.

Informed Consent

Informed consent was not required for this IRB approved retrospective study, as it involved the use of de-identified, pre-existing data in accordance with ethical guidelines.

Author Contributions

Khush Kharidia: study design, data collection, data analysis, interpretation of data, manuscript writing and approval. Weiye Tan: study design, interpretation of data, manuscript writing and approval. Nimesh Patel: study design, data analysis, interpretation of data, manuscript writing and approval.

Data Availability

The data supporting the findings of this study are available from the corresponding author upon reasonable request.

Abbreviations

AP: atriopulmonary; AT: atrial tachycardia; BNP: brain natriuretic peptide; CHD: congenital heart disease; CPT: Current Procedural Terminology; CV: cardiovascular; EC: extra-cardiac; EKG: electrocardiogram; EP: electrophysiology; ICD: implantable cardioverter-defibrillator; IQR: interquartile range; LT: lateral tunnel; PPM: pacemaker; RA: right atrial; RHC: right heart catheterization; SVT: supraventricular tachycardia; TGA: transposition of great arteries; TA: tricuspid atresia; TTE: transthoracic echocardiogram

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