

# Heart and Vessels

## Dissociated pulmonary vein activity after cryoballoon ablation and radiofrequency ablation for atrial fibrillation: A propensity score matched analysis

--Manuscript Draft--

<b>Manuscript Number:</b>	HEVE-D-17-00208R3
<b>Full Title:</b>	Dissociated pulmonary vein activity after cryoballoon ablation and radiofrequency ablation for atrial fibrillation: A propensity score matched analysis
<b>Article Type:</b>	Original Article /Clinical Investigation
<b>Keywords:</b>	keyword 1; Dissociated pulmonary vein activity, keyword 2; Cryoballoon ablation, keyword 3; Radiofrequency ablation, keyword 4; Pulmonary vein isolation, keyword 5; atrial fibrillation
<b>Corresponding Author:</b>	Michifumi Tokuda The Jikei University school of medicine Minato, Tokyo JAPAN
<b>Corresponding Author Secondary Information:</b>	
<b>Corresponding Author's Institution:</b>	The Jikei University school of medicine
<b>Corresponding Author's Secondary Institution:</b>	
<b>First Author:</b>	Kenichi Tokutake
<b>First Author Secondary Information:</b>	
<b>Order of Authors:</b>	Kenichi Tokutake Michifumi Tokuda, M.D. Ph.D. Seiichiro Matsuo, M.D. Ph.D. Ryota Isogai, M.D. Kenichi Yokoyama, M.D. Mika Kato, M.D. Ryohsuke Narui, M.D, Shinichi Tanigawa, M.D. Seigo Yamashita, M.D. Ph.D. Satoru Miyanaga, M.D. Michihiro Yoshimura, M.D. Ph.D. Teiichi Yamane, M.D. Ph.D.
<b>Order of Authors Secondary Information:</b>	
<b>Funding Information:</b>	
<b>Abstract:</b>	<p>Cryoballoons (CBs) have proven to be effective for achieving pulmonary vein isolation (PVI) in patients with atrial fibrillation (AF). Dissociated PV activity (DPVA) after successful radiofrequency PVI is sometimes observed inside the PV and has been found to prove the achievement of electrical disconnection from the left atrium. However, little is known about the incidence or characteristics of DPVA after CB-PVI. The aim of this study was to compare the incidence and characteristics of DPVA in patients undergoing CB and radiofrequency (RF) ablation for AF. Two hundred ninety-four propensity score-matched patients from 440 consecutive patients who underwent initial catheter ablation for paroxysmal AF were included in the present study (CB-PVI: 147, RF-PVI: 147). DPVA was more frequently observed after CB-PVI than after RF-PVI (32% vs. 19% of</p>

	<p>the PVs, <math>P &lt; 0.001</math>), especially in the left superior PV (52% vs. 29%, <math>P &lt; 0.001</math>) and left inferior PV (22% vs. 7%, <math>P &lt; 0.001</math>). The AF-free rate after the initial ablation in the patients with and without DPVA was similar in both the CB (<math>P = 0.23</math>) and RF (<math>P = 0.39</math>) groups. During repeat ablation procedures for recurrent AF, PV reconnection was similarly observed in PVs with and without DPVA during initial procedure both in the CB (30% vs. 44%, <math>P = 0.29</math>) and RF (65% vs. 58%, <math>P = 0.41</math>) groups. DPVA was more frequently observed after CB-PVI than after RF-PVI. The presence of DPVA was not related to the ablation outcome or chronic PV reconnection following CB-PVI.</p>
<p><b>Response to Reviewers:</b></p>	<p>Thank you very much for considering our revised manuscript for Heart and Vessels. We thank you for your reviews which have helped us improve the manuscript considerably. Changes in the manuscript are provided in bold below.</p> <p>Response to Comments from Reviewer #2  Reviewer Comment:  The authors addressed my comments sufficiently. However, despite not having been yet compared ("the incidence of DPVA and its impact on the ablation outcome in patients undergoing CB and radiofrequency (RF) ablation for AF", as stated as Motivation), the clinical impact and value of the results is limited.</p> <p>Response:  Thank you. As you mentioned, we did not directly compare the impact of DPVA on the outcome between after CB and RF ablation. We edited as below, "The aim of this study was to compare the incidence and characteristics of DPVA in patients undergoing CB and radiofrequency (RF) ablation for AF."</p> <p>Reviewer Comment:  Furthermore, I do not agree with the conclusions that the larger isolate area in CB PVI can explain the higher incidence of electrical activation.</p> <p>Response:  Thank you. As you mentioned, no previous study did not directly compare the isolated area and incidence of DPVA. However, also no study did not prove that isolated area unrelated to the incidence of DPVA. Even it was not direct comparison, the results of previous studies shows higher incidence of DPVA in wide encircling ablation (12-72%) compared to that in segmental isolation (2-12 %).</p> <p>We deleted these sentences,  "Electrical activity should appear more frequently from the larger isolated muscular sleeve." and "The larger isolated area may affect higher incidence of DPVA after CB-PVI than after individual RF-PVI."  We have edited as below. in Page 11, Line 16.  "The difference of isolated area can relate to the incidence of DPVA. (Figure 7) In this study, the isolated areas after CB-PVI were wider than those after segmental RF-PVI using 3D electroanatomical mapping system. In previous studies, wide encircling ablation resulted in a DPVA rate of 12-72% [5-9], while segmental isolation resulted in 2-12 % [4, 10]."</p>

Thank you very much for considering our revised manuscript for *Heart and Vessels*. We

thank you for your reviews which have helped us improve the manuscript considerably.

Changes in the manuscript are provided in **bold** below.

### *Response to Comments from Reviewer #2*

#### *Reviewer Comment:*

The authors addressed my comments sufficiently.

However, despite not having been yet compared ("the incidence of DPVA and its impact on the ablation outcome in patients undergoing CB and radiofrequency (RF) ablation for AF", as stated as Motivation), the clinical impact and value of the results is limited.

#### *Response:*

Thank you. As you mentioned, we did not directly compare the impact of DPVA on the outcome between after CB and RF ablation. We edited as below,

**“The aim of this study was to compare the incidence and characteristics of DPVA in patients undergoing CB and radiofrequency (RF) ablation for AF.”**

**Reviewer Comment:**

Furthermore, I do not agree with the conclusions that the larger isolate area in CB PVI can explain the higher incidence of electrical activation.

**Response:**

Thank you. As you mentioned, no previous study did not directly compare the isolated area and incidence of DPVA. However, also no study did not prove that isolated area unrelated to the incidence of DPVA. Even it was not direct comparison, the results of previous studies shows higher incidence of DPVA in wide encircling ablation (12-72%) compared to that in segmental isolation (2–12 %).

We deleted these sentences,

“Electrical activity should appear more frequently from the larger isolated muscular sleeve.” and “The larger isolated area may affect higher incidence of DPVA after CB-PVI than after individual RF-PVI.”

We have edited as below. in Page 11, Line 16.

**“The difference of isolated area can relate to the incidence of DPVA. (Figure 7) In this study, the isolated areas after CB-PVI were wider than those after segmental RF-PVI using 3D electroanatomical mapping system. In previous studies, wide encircling ablation resulted in a DPVA rate of 12-72% [5-9], while segmental isolation resulted in 2–12 % [4, 10].”**

[Click here to view linked References](#)

1  
2  
3  
4  
5  
6  
7  
8  
9  
10  
11  
12  
13  
14  
15  
16  
17  
18  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29  
30  
31  
32  
33  
34  
35  
36  
37  
38  
39  
40  
41  
42  
43  
44  
45  
46  
47  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

**Dissociated pulmonary vein activity after cryoballoon ablation and radiofrequency  
ablation for atrial fibrillation: A propensity score matched analysis**

Kenichi Tokutake M.D., Michifumi Tokuda M.D. Ph.D., Seiichiro Matsuo M.D. Ph.D., Ryota  
Isogai M.D., Kenichi Yokoyama M.D., Mika Kato M.D., Ryohsuke Narui M.D., Shinichi  
Tanigawa M.D., Seigo Yamashita M.D. Ph.D., Satoru Miyanaga M.D., and Michihiro Yoshimura  
M.D. Ph.D., Teiichi Yamane M.D. Ph.D.

Department of Cardiology, The Jikei University School of Medicine

Address for correspondence:

Michifumi Tokuda, M.D. Department of Cardiology, The Jikei University School of Medicine

3-25-8, Nishi-shinbashi, Minato-ku, TOKYO 105-8461, JAPAN

Tel: +81.3.3433.1111 Fax: +81.3.3459.6043

E-mail: tokudam@gmail.com

1  
2  
3  
4  
5 Abstract  
6  
7

8 Cryoballoons (CBs) have proven to be effective for achieving pulmonary vein isolation (PVI) in  
9 patients with atrial fibrillation (AF). Dissociated PV activity (DPVA) after successful  
10 radiofrequency PVI is sometimes observed inside the PV and has been found to prove the  
11 achievement of electrical disconnection from the left atrium. However, little is known about the  
12 incidence or characteristics of DPVA after CB-PVI. The aim of this study was to compare the  
13 incidence and characteristics of DPVA in patients undergoing CB and radiofrequency (RF)  
14 ablation for AF.  
15  
16  
17  
18  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29

30 Two hundred ninety-four propensity score-matched patients from 440 consecutive patients who  
31 underwent initial catheter ablation for paroxysmal AF were included in the present study (CB-  
32 PVI: 147, RF-PVI: 147).  
33  
34  
35  
36  
37  
38  
39

40 DPVA was more frequently observed after CB-PVI than after RF-PVI (32% vs. 19% of the PVs,  
41  $P<0.001$ ), especially in the left superior PV (52% vs. 29%,  $P<0.001$ ) and left inferior PV (22%  
42 vs. 7%,  $P<0.001$ ). The AF-free rate after the initial ablation in the patients with and without DPVA  
43 was similar in both the CB ( $P=0.23$ ) and RF ( $P=0.39$ ) groups. During repeat ablation procedures  
44 for recurrent AF, PV reconnection was similarly observed in PVs with and without DPVA during  
45 initial procedure both in the CB (30% vs. 44%,  $P=0.29$ ) and RF (65% vs. 58%,  $P=0.41$ ) groups.  
46  
47  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5  
6  
7  
8  
9  
10  
11  
12  
13  
14  
15  
16  
17  
18  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29  
30  
31  
32  
33  
34  
35  
36  
37  
38  
39  
40  
41  
42  
43  
44  
45  
46  
47  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

DPVA was more frequently observed after CB-PVI than after RF-PVI. The presence of DPVA was not related to the ablation outcome or chronic PV reconnection following CB-PVI.

**Keywords**

Dissociated pulmonary vein activity, Cryoballoon ablation, Radiofrequency ablation, Pulmonary vein isolation, Atrial fibrillation

1  
2  
3  
4  
5 Introduction  
6  
7

8 Pulmonary vein isolation (PVI) is the cornerstone of catheter ablation for atrial  
9 fibrillation (AF), especially in patients with paroxysmal AF [1]. In addition to radiofrequency  
10 (RF) ablation, cryoballoon (CB) ablation has proven to be an effective therapy for PVI in patients  
11 with paroxysmal AF. Two randomized trials have shown the noninferiority of CB ablation in  
12 comparison to RF ablation with respect to the treatment efficacy in patients with drug-refractory  
13 paroxysmal AF [2, 3]. Furthermore, the overall safety of the two types of procedures did not differ  
14 to a statistically significant extent.  
15  
16  
17  
18  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29

30 Dissociated PV activity (DPVA) after successful PVI is observed inside 2-72% of PVs  
31 and has been found to prove the achievement of electrical disconnection from the left atrium [4-  
32 10]. Previous studies demonstrated the occurrence of DPVA after RF-PVI was related to acute  
33 PV reconnection at 30-min after PVI [5], and was shown to improve the long-term ablation  
34 outcome and be an indicator of successful durable isolation of the PVs [7]. The characteristics of  
35 DPVA can differ according to the ablation method such as segmental and circumferential PVI.  
36  
37 Although implication of DPVA after RF-PVI was reported by several reports, little is known about  
38 the characteristics of DPVA in patients undergoing CB-PVI. The aim of the present study was to  
39 compare the incidence and characteristics of DPVA in patients undergoing CB or RF ablation for  
40 AF.  
41  
42  
43  
44  
45  
46  
47  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5 **Materials and methods**  
6

7  
8 *Study Subjects*  
9

10  
11 The study population included a total of 440 patients who had undergone an initial  
12 catheter ablation procedure for paroxysmal AF, between January 2013 and June 2016. The  
13  
14  
15  
16  
17  
18  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29  
30  
31  
32  
33  
34  
35  
36  
37  
38  
39  
40  
41  
42  
43  
44  
45  
46  
47  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

The study population included a total of 440 patients who had undergone an initial catheter ablation procedure for paroxysmal AF, between January 2013 and June 2016. The decision to perform RF ablation versus CB ablation was based on the preference of each individual operator. The patient with common PV and right middle PV was excluded from the study. All of the patients underwent an enhanced CT-scan prior to ablation to establish the anatomy of the LA and PV. Antiarrhythmic drugs were discontinued for at least 5 half-lives prior to ablation. Transesophageal echocardiography was performed prior to the procedure to rule out LA thrombus. In the present study, “paroxysmal AF” was defined as AF that spontaneously terminated within 7 days. The studies and data collection were performed according to protocols that had been approved by the Human Research Committee of the Jikei University School of Medicine.

*The Catheter Ablation of AF*

Each patient gave their written informed consent. Heparin (100 IU/kg body weight) was administered immediately after obtaining venous access; heparinized saline was also infused to maintain an activated clotting time of 250–350 seconds. The procedures were performed under

1  
2  
3  
4  
5 mild sedation with flunitrazepam and propofol. An esophageal temperature probe (Sensitherm, St.  
6  
7  
8 Jude Medical Inc., Minneapolis, MN, USA or Esophastar, Japan Lifeline) was inserted throughout  
9  
10  
11  
12 the ablation procedure to avoid esophageal injury.  
13  
14  
15  
16  
17  
18  
19  
20

### 21 *Radiofrequency PV Isolation*

22

23  
24 An electrophysiological study was performed as previously described under  
25  
26  
27 institutional approval [11]. The LA and PVs were explored through either a patent foramen  
28  
29  
30 ovale or transseptal catheterization with 3 catheters (2 for circumferential PV mapping and 1 for  
31  
32  
33 ablation). The direct visualization of all 4 PVs was performed using selective venography. The  
34  
35  
36  
37 PV antrum was determined using a direct PV visualization 3-dimensional mapping system  
38  
39  
40 (CARTO, Biosense Webster, Diamond Bar, CA, USA, or Ensite NavX, St. Jude Medical). PV  
41  
42  
43 mapping was performed with a 20-polar circular catheter (Lasso<sup>®</sup> 2515 NAV eco variable  
44  
45  
46  
47 catheter, Biosense Webster).  
48

49  
50 All 4 PVs were individually targeted for disconnection from the LA. RF energy was  
51  
52  
53 delivered at the PV antrum for 30–60 seconds using an open irrigated-tip ablation catheter with  
54  
55  
56 a power limit of 25-35 W. The endpoint of the PVI was the establishment of a bidirectional  
57  
58  
59 conduction block between the LA and the PV. The exit block was confirmed by pacing inside  
60  
61

1  
2  
3  
4  
5 the PV with a Lasso catheter. The absence of acute conduction recovery between the PVs and  
6  
7  
8 the LA was confirmed in each PV after waiting at least 30 minutes after the final RF application.  
9

### 10 11 12 13 14 15 *Cryoballoon Ablation* 16

17  
18 A single transseptal puncture was performed using an RF needle (Baylis Medical, Montreal, QC,  
19  
20  
21 Canada) and an 8-Fr long sheath (SL0, AF Division, St. Jude Medical). The transseptal sheath  
22  
23  
24 was exchanged over a guide wire for a 15-Fr steerable sheath (Flexcath Advance, Medtronic). A  
25  
26  
27 20–30 mm circular mapping catheter (Lasso) was used for mapping all of the PVs before and after  
28  
29  
30 the CB to confirm electrical isolation. PVI was performed with a single balloon technique using  
31  
32  
33 a second-generation CB (Arctic Front Advance, Medtronic, Minneapolis, MN, USA). In the CB  
34  
35  
36 group, a 28-mm CB catheter was used for all patients. A spiral mapping catheter (Achieve<sup>®</sup>,  
37  
38  
39 Medtronic) was used to advance the CB and to map the PV potentials. Complete sealing at the  
40  
41  
42 antral aspect of the PV was confirmed by the injection of contrast medium. This was followed by  
43  
44  
45 a freeze cycle of 180 seconds. In order to avoid phrenic nerve injury [12], the diaphragmatic  
46  
47  
48 compound motor action potentials (CMAP) were monitored during phrenic nerve pacing while  
49  
50  
51 all of the CB applications were applied [13]. The procedural endpoint was defined as the  
52  
53  
54 establishment of a bidirectional PV-LA block, which was verified by a 20–30 mm circular  
55  
56  
57 mapping catheter. If electrical isolation was not achieved by a total of 3 CB applications (180  
58  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5 seconds for each application) per vein, additional touch-up ablation was performed with a  
6  
7  
8 conventional RF or cryothermal (Freezer Max, Medtronic) catheter. The voltage map was created  
9  
10 before and after ablation to assess isolated are. The mapping of LA and PV was obtained using  
11  
12 circular mapping catheter with the Ensite Navx (St. Jude Medical Inc.) or CARTO (Biosense  
13  
14 Webster). The scale on the voltage map was set from 0.2 to 1.2 mV. Scar area was defined as the  
15  
16 area with the voltage < 0.2mV. The surface area of the LA and PV were calculated using the  
17  
18 algorithm included in the 3 D mapping system as shown in the previous study [14].  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29  
30  
31  
32  
33

#### 34 *ATP Testing after Pulmonary Vein Isolation*

35  
36

37 All PVs were isolated and the mapping catheter was kept in the PV for at least 5  
38  
39 minutes after isolation. In all patients of both the RF and CB groups—after waiting for at least  
40  
41 30 minutes after the final application of PVI—adenosine triphosphate (20 mg) was rapidly  
42  
43 administered intravenously to induce dormant PV conduction under the continuous infusion of  
44  
45 isoproterenol (20 µg/min) during sinus rhythm (for the right superior [RS] PV and the right  
46  
47 inferior [RI] PV) or coronary sinus pacing (for the left superior [LS] PV and left inferior [LI]  
48  
49 PV). Additional radiofrequency or cryothermal ablation was applied until dormant conduction  
50  
51 was eliminated.  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5  
6  
7  
8  
9 *Definition of Dissociated PV activity*

10  
11 DPVA was defined as at least two PV potentials without propagation to the LA after PVI,  
12  
13 which occurred spontaneously, and which was not induced by the manipulation of catheters. This  
14  
15 phenomenon had to be observed in sinus rhythm at least 30 minutes after the successful isolation  
16  
17 of the veins using conventional circular mapping catheters in each PV for 5 minutes. The rhythms  
18  
19 in the PV were classified according to a previous report [6]: (1) silent, there was no electrical  
20  
21 activity; (2) isolated ectopic beats, there were intermittent potentials without a regular rhythm; (3)  
22  
23 regular ectopic rhythm (Figure 1) or (4) fibrillatory activity (Figure 2).  
24  
25  
26  
27  
28  
29  
30  
31  
32  
33  
34  
35  
36

37 *Patient Follow-up*

38  
39  
40 No antiarrhythmic drugs were prescribed after the procedure. The patients underwent  
41  
42 continuous, in-hospital ECG monitoring for 2–4 days after the procedure. The patients were  
43  
44 carefully observed (2 weeks after discharge, then every month) at the cardiology clinic. The  
45  
46 outcome of AF ablation was evaluated based on the patient’s symptoms, ECG at the periodical  
47  
48 follow-ups, and by periodic 24-hour ambulatory monitoring (at 1, 3, 6, 9, and 12 months after  
49  
50 the procedure). A cardiac event recorder or a portable electrocardiogram was used to define  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5 the causes of symptoms that were suggestive of tachycardia. The recurrence of AF was  
6  
7  
8 defined as AF lasting for more than 60 seconds after a blanking period of 90 days.  
9

10  
11  
12  
13  
14  
15 *Statistical Analysis*  
16

17  
18 Propensity scores were calculated for each of the 414 patients based on a multivariable  
19  
20  
21 logistic regression model. A total of 15 characteristics that were hypothesized to be associated  
22  
23  
24 with the outcome of catheter ablation were assessed for inclusion in the model as independent  
25  
26  
27 variables. All 15 characteristics were retained in the model with stepwise selection and were  
28  
29  
30 subsequently used to generate propensity scores. The selection process used a P value cutoff of  
31  
32  
33 0.05 for a characteristic to be included in the model. In order of stepwise selection, the matching  
34  
35  
36 variables were as follows: sex, age, body mass index, LA diameter, left ventricular ejection  
37  
38  
39 fraction, estimated glomerular filtration rate, B-type natriuretic peptide, history of AF,  
40  
41  
42 hypertension, diabetes mellitus, obstructive sleep apnea, congestive heart failure, coronary  
43  
44  
45 artery disease, the number of AAD and CHADS2 score. Based on their propensity score, the  
46  
47  
48 patients who underwent CB-PVI were matched on a 1:1 basis with patients who underwent RF-  
49  
50  
51 PVI using a nearest neighbor algorithm without replacement using a caliper width equal to 1/5  
52  
53  
54 of the standard deviation of the logit of the propensity score.  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5 Continuous variables were expressed as the mean  $\pm$  standard deviation. An unpaired  
6  
7  
8 Student's *t*-test or the Mann-Whitney U test was used for the analysis of continuous variables.  
9  
10  
11 The categorical variables, which were expressed as numbers or percentages, were analyzed using  
12  
13 the chi-squared test unless the expected values in any cells were  $<5$ , in which case Fisher's exact  
14  
15 test was used. P values of  $<0.05$  were considered to indicate statistical significance. Survival  
16  
17  
18 curves were created using the Kaplan–Meier method and comparisons between groups were  
19  
20  
21 performed using a log-rank test. All of the statistical analyses were performed using the SPSS  
22  
23  
24 software program (version 23.0.0; SPSS, Chicago, IL).  
25  
26  
27  
28  
29  
30  
31  
32  
33

## 34 Results

### 35 36 37 *Patient Characteristics and Procedural Details*

38  
39  
40 CB and RF ablation were performed in 191 and 249 patients, respectively. The  
41  
42  
43 resulting propensity score matched data set included 294 patients—147 underwent CB-PVI and  
44  
45  
46 147 underwent RF-PVI. The mean age of the patients was  $57.3 \pm 10.2$  years and the mean LA  
47  
48  
49 diameter was  $36.6 \pm 5.2$  mm. There were no significant differences between the two groups with  
50  
51  
52 regard to the baseline patient characteristics (Table 1).  
53  
54  
55

56 The total procedure time ( $142 \pm 32$  min vs.  $172 \pm 48$  min,  $P < 0.001$ ) and the fluoroscopic  
57  
58  
59 time ( $62 \pm 18$  min vs.  $99 \pm 33$  min,  $P < 0.001$ ) were significantly shorter in the CB ablation than in  
60  
61  
62

1  
2  
3  
4  
5 the RF ablation (Table 2). While the PV was successfully isolated in all cases in the RF group,  
6  
7  
8 PVI by CB alone failed—thereby necessitating touch-up ablation—in 8% (46/588) of the PVs in  
9  
10  
11 the CB group. Touch-up ablation was most frequently required in the RIPV (LSPV: 6% (9/147  
12  
13  
14 PVs), LIPV: 9% (13/147 PVs), RSPV: 0% and RIPV: 18% (24/147 PVs)). The mean area of  
15  
16  
17 post-ablation scar was larger after CB-PVI than after RF-PVI ( $14.4 \pm 2.2 \text{ cm}^2$  vs.  $11.4 \pm 2.6$   
18  
19  
20  
21  $\text{cm}^2$ ,  $P=0.01$ ).

#### 22 23 24 25 26 27 *DPVA after PVI*

28  
29  
30 DPVA was observed in 26% of all PVs (1.1 PVs/patient). DPVA was more frequently  
31  
32  
33 observed after CB-PVI than after RF-PVI (32% (1.3 PVs/patient) vs. 19% of the PVs (0.8  
34  
35  
36 PVs/patient),  $P<0.001$ ; Figure 3). After CB-PVI, superior PVs presented DPVA more frequently  
37  
38  
39 than inferior PVs (45% vs. 18%,  $P<0.001$ ), as well as DPVA after RF ablation (30% vs. 10%,  
40  
41  
42  $P<0.001$ ). With regard to the individual PVs, DPVA was more frequently observed after CB-PVI  
43  
44  
45 than after RF-PVI in the LSPV (52% vs. 29%,  $P<0.001$ ) and the LIPV (22% vs. 7%,  $P<0.001$ ),  
46  
47  
48 but was similarly observed in the RSPV (37% vs. 31%,  $P=0.33$ ) and the RIPV (15% vs. 12%,  
49  
50  
51  $P=0.39$ ) (Figure 3). The types of DPVA are shown in Figure 4. Dissociated ectopic beat and  
52  
53  
54  
55  
56  
57  
58 ectopic rhythm were more frequently observed after CB-PVI than after RF-PVI (21% vs. 15%,  
59  
60  
61  $P=0.007$ , 10% vs. 4%,  $P=0.0001$ , respectively). Among the PVs with a regular dissociated rhythm,

1  
2  
3  
4  
5 the mean cycle length of the dissociated rhythm in the RF and CB groups did not differ to a  
6 statistically significant extent ( $2209 \pm 986$  ms vs.  $1869 \pm 825$  ms,  $P=0.16$ ). PV fibrillation was not  
7  
8  
9 observed in the CB group. DPVA was similarly observed in the PVs required touch-up ablation  
10  
11  
12 and these not required ( $13/46$  PVs (28.3%) vs.  $179/542$  PVs (33.0%),  $P=0.51$ ).  
13  
14  
15  
16  
17  
18  
19  
20  
21  
22  
23

#### 24 *Ablation Outcome and durability of PVI*

25  
26  
27 During a mean of  $12 \pm 6$  months follow up, the AF-free rate after the initial procedure in  
28  
29 the patients with and without DPVA did not differ to a statistically significant extent in either the  
30  
31 CB or RF groups ( $P=0.23$  and  $P=0.39$ , respectively; both Figure 5). Among the 294 propensity  
32  
33 score-matched patients, 5 (3%) patients in the CB group and 9 (6%) patients in the RF group  
34  
35 underwent a repeat ablation procedure for recurrent atrial tachyarrhythmia. During the repeat  
36  
37 ablation procedure, PV reconnections were less frequently observed after CB ablation than after  
38  
39 RF ablation (36% vs. 62% of PVs,  $P=0.02$ , Figure 6). PV reconnection was similarly observed in  
40  
41 PVs with and without DPVA in the index ablation procedure, both in the CB (30% vs. 44%,  
42  
43  $P=0.29$ ) and RF groups (65% vs. 58%,  $P=0.41$ , Figure 6). In 16 of 30 PVs without DPVA at the  
44  
45 first procedure, DPVA was observed during repeat procedure. On the other hand, 13 of 25 PVs  
46  
47 with DPVA at first procedure still presented DPVA during repeat procedure.  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5  
6  
7  
8 Discussion  
9

10  
11 *Main Findings*  
12  
13

14  
15 This study resulted in several important findings. First, DPVA was more frequently  
16  
17  
18 observed in the patients who underwent CB-PVI than in those who underwent RF-PVI. Second,  
19  
20  
21 the presence of DPVA was not associated with the ablation outcome in the CB or RF groups.  
22  
23  
24 Third, PV reconnection in repeat procedure was similar between PVs with and without DPVA in  
25  
26  
27 the index procedure. To the best of our knowledge, this is the first study to assess the details of  
28  
29  
30 DPVA after CB-PVI and compare with those after RF ablation.  
31  
32  
33  
34  
35  
36

37 *Incidence of Dissociated PV Firing*  
38  
39

40 Reported incidence of DPVA after RF-PVI showed a broad variation from 2 to 72% of  
41  
42  
43 PVs [4-10]. Differences in the definition of DPVA are probably the main reason of the broad  
44  
45  
46 variations. We observed DPVA in 19% of PVs after RF ablation. DPVA was more frequently  
47  
48  
49 observed in superior PVs than in the inferior PVs. This may be due to the longer and thicker  
50  
51  
52 muscular sleeves of the superior PV [15]. The difference of isolated area can relate to the  
53  
54  
55 incidence of DPVA. (Figure 7) In this study, the isolated areas after CB-PVI were wider than  
56  
57  
58 those after segmental RF-PVI using 3D electroanatomical mapping system. In previous studies,  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5 wide encircling ablation resulted in a DPVA rate of 12-72% [5-9], while segmental isolation  
6  
7  
8 resulted in 2–12 % [4, 10]. The methods of RF-PVI (individual or circumferential) may also  
9  
10 affect the result of this study.  
11  
12

13  
14         The different energy sources of two ablation methods also may affect the incidence of  
15  
16 DPVA. Several previous studies have demonstrated that CB ablation may create lesions that are  
17  
18 more homogeneous and extensive in comparison to those created by RF ablation. The higher  
19  
20 levels of cardiac troponin I and creatinine kinase-MB that are generally present after CB-PVI in  
21  
22 comparison to RF-PVI indicate that a greater degree of inflammation occurs during CB-PVI [16].  
23  
24 Moreover, we previously reported that ATP-induced dormant conduction was less frequently  
25  
26 provoked after CB ablation than after RF ablation [14]. Because the mechanism of ATP-induced  
27  
28 dormant conduction is considered to be the restoration of conduction in damaged—but viable—  
29  
30 myocardial tissues [17], the lower prevalence of dormant conduction indicates that CB-PVI can  
31  
32 make lesions that are more homogeneous and extensive in comparison to those made by RF-PVI.  
33  
34 Greater myocardial injury may affect the activity in the PV myocardial sleeves distal to the  
35  
36 ablation line in the acute period.  
37  
38  
39  
40  
41  
42  
43  
44  
45  
46  
47  
48  
49  
50

51  
52         Adrenergic agents to induce acute PV reconnection and dormant PV conduction must  
53  
54 have an effect on the incidence of DPVA. However, influence of the administered agent was  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5 considered to be small because a similar dose of isoproterenol (20 µg/min) was continuously  
6  
7  
8 infused after PVI in both groups.  
9

### 10 11 12 13 14 15 *Presence of DPVA and the Ablation Outcome* 16

17  
18 In the literature, the association between DPVA and ablation outcomes has been  
19  
20  
21 controversial. Several studies demonstrated that the presence of DPVA was a successful  
22  
23  
24 indicator of durable PVI and was associated with an improved ablation outcome [7, 18, 19].  
25  
26  
27 However, other reports revealed no relationship between this phenomenon and arrhythmia-free  
28  
29  
30 survival [8, 9]. In our study, DPVA was no clinical predictor for AF recurrence after CB or RF  
31  
32  
33 ablation. Most patients with a failed initial ablation procedure have resumption of PV  
34  
35  
36 conduction. No significant relationship between DPVA and the durability of PVI in repeat  
37  
38  
39 ablation procedure might explain the contracting results. Although DPVA after PVI is an  
40  
41  
42 important electrophysiological finding to ensure LA-PV bidirectional conduction block, clinical  
43  
44  
45 implications might be limited.  
46  
47  
48  
49  
50  
51  
52

### 53 **Study limitations** 54 55

56 The present study is associated with several limitations. This study was a non-  
57  
58  
59 randomized, observational, study that was performed in a single institution. The results of  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5 propensity score matching in this study are only generalizable among patients in the range of  
6  
7  
8 propensity scores that were included in the paired analysis, and may not be applicable to patients  
9  
10  
11 who are outside this range. Propensity score methods can reduce bias in causal estimates due to  
12  
13  
14 observed differences between groups, but they are still subject to biases from unobserved  
15  
16  
17 differences. The incidence of DPVA can be affected by time and autonomic nerve system.  
18  
19  
20  
21 Detection of DPVA might be dependent on the positioning of the circumferential mapping  
22  
23  
24 catheter. We usually located circular mapping catheter on PV antrum. PV antrum is defined as  
25  
26  
27 the proximal portion next to the tubular PV as observed on the selective PV angiogram or  
28  
29  
30 intracardiac echocardiogram. Since we pursued to locate the circular mapping catheter in similar  
31  
32  
33 position in both groups, the influence of the catheter position to the results cannot be denied.  
34  
35  
36

37 Further study with longer observation period may improve the reliability of the results.  
38  
39

40 Because the number of patients who underwent repeat ablation procedures was small, the  
41  
42  
43 statistical reliability regarding PV reconnection during repeat ablation procedures was limited.  
44  
45  
46  
47  
48

## 49 Conclusion

50  
51 The DPVA occurred more frequently after CB-PVI than after RF-PVI. The  
52  
53  
54 presence/absence of DPVA was not associated with the ablation outcome or chronic PV  
55  
56  
57 reconnection after the CB or RF ablation procedures.  
58  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5  
6  
7  
8  
9  
10  
11  
12  
13  
14  
15  
16  
17  
18  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29  
30  
31  
32  
33  
34  
35  
36  
37  
38  
39  
40  
41  
42  
43  
44  
45  
46  
47  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

Conflict of interest

The authors declare no conflicts of interest in association with the present study.

Acknowledgements

We are grateful to Dr. Brian Quinn (Japan Medical Communication Inc.) for the comments on the language of the manuscript.

1  
2  
3  
4  
5  
6  
7  
8  
9  
10  
11  
12  
13  
14  
15  
16  
17  
18  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29  
30  
31  
32  
33  
34  
35  
36  
37  
38  
39  
40  
41  
42  
43  
44  
45  
46  
47  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

## References

1. Haïssaguerre M, Jaïs P, Shah DC, Takahashi A, Hocini M, Quiniou G, Garrigue S, Le Mouroux A, Le Métayer P, Clémenty J (1998) Spontaneous initiation of atrial fibrillation by ectopic beats originating in the pulmonary veins. *N Engl J Med* 339:659-666
2. Kuck KH, Brugada J, Fürnkranz A, Metzner A, Ouyang F, Chun KR, Elvan A, Arentz T, Bestehorn K, Pocock SJ, Albenque JP, Tondo C; FIRE AND ICE Investigators (2016) Cryoballoon or Radiofrequency Ablation for Paroxysmal Atrial Fibrillation. *N Engl J Med* 374:2235-2245
3. Luik A, Radzewitz A, Kieser M, Walter M, Bramlage P, Hörmann P, Schmidt K, Horn N, Brinkmeier-Theofanopoulou M, Kunzmann K, Riexinger T, Schymik G, Merkel M, Schmitt C (2015) Cryoballoon Versus Open Irrigated Radiofrequency Ablation in Patients With Paroxysmal Atrial Fibrillation: The Prospective, Randomized, Controlled, Noninferiority FreezeAF Study. *Circulation* 132:1311-1319
4. WeerasooriyaR, Jaïs P, Scavee C, Macle L, Shah DC, Arentz T, Salerno JA, Raybaud F, Choi KJ, Hocini M, Clémenty J, Haïssaguerre M (2003) Dissociated pulmonary vein arrhythmia: Incidence and characteristics. *J Cardiovasc Electrophysiol* 14:1173-1179
5. Jiang CY, Fu JW, Matsuo S, Nault I, He H, Jiang RH, Liu Q, Fan YQ, Sheng X, Zhang ZW,

1  
2  
3  
4  
5 Fu GS (2011) Dissociated pulmonary vein rhythm may predict the acute pulmonary vein  
6 reconnection post-isolation in patients with paroxysmal atrial fibrillation. *Europace* 13:949-  
7  
8  
9 954  
10

11  
12  
13  
14  
15 6. Kabra R, Heist EK, Barrett CD, Donaldson D, Blendea D, Beinart R, Koruth J, Singh S,  
16  
17  
18 Ruskin J, Mansour M (2010) Incidence and electrophysiologic properties of dissociated  
19  
20  
21 pulmonary vein activity following pulmonary vein isolation during catheter ablation of atrial  
22  
23  
24 fibrillation. *J Cardiovasc Electrophysiol* 21:1338 –1343  
25

26  
27  
28 7. Hussein AA, Ozaki A, Martin DO, Bhargava M, Baranowski B, Dresing T, Callahan T, Kanj  
29  
30  
31 M, Tchou P, Natale A, Lindsay BD, Saliba WI, Wazni OM (2013) Spontaneous dissociated  
32  
33  
34 firing from the pulmonary veins during ablation of paroxysmal atrial fibrillation:  
35  
36  
37 implications and impact on arrhythmia-free survival. *Pacing Clin Electrophysiol* 36:988-993  
38  
39

40  
41 8. Buiatti A, Ammar S, Reents T, Semmler V, Kathan S, Hofmann M, Bourier F, Telishevsk  
42  
43  
44 M, Koch-Büttner K, Kaess B, Lennerz C, Kolb C, Hessling G, Deisenhofer I (2015)  
45  
46  
47 Dissociated pulmonary vein activity after pulmonary vein isolation for paroxysmal atrial  
48  
49  
50 fibrillation: a predictor for recurrence? *J Cardiovasc Electrophysiol* 26:7-13  
51

52  
53 9. Miyazaki S, Kuwahara T, Kobori A, Takahashi Y, Takei A, Sato A, Isobe M, Takahashi A  
54  
55  
56 (2011) Prevalence, electrophysiological properties, and clinical implications of dissociated  
57  
58  
59 pulmonary vein activity following pulmonary vein antrum isolation. *Am J Cardiol* 108:1147-  
60  
61

- 8  
9 10. Marrouche N, Wazni OM, Martin DO, Rossillo A, Saliba W, Erciyas D, Schweikert R,  
10  
11 Khaykin Y, Burkhardt D, Bhargava M, Verma A, Abdul-Karim A, Natale A (2005) Response  
12  
13 to pharmacological challenge of dissociated pulmonary vein rhythm. J Cardiovasc  
14  
15 Electrophysiol 16:122-126  
16  
17  
18  
19  
20  
21 11. Yamane T, Date T, Kanzaki Y, Inada K, Matsuo S, Shibayama K, Miyanaga S, Miyazaki H,  
22  
23 Sugimoto K, Mochizuki S (2007) Segmental pulmonary vein antrum isolation using the  
24  
25 "large-size" lasso catheter in patients with atrial fibrillation. Circ J 71:753-760  
26  
27  
28  
29  
30  
31 12. Sacher F, Monahan KH, Thomas SP, Davidson N, Adragao P, Sanders P, Hocini M, Takahashi  
32  
33 Y, Rotter M, Rostock T, Hsu LF, Clémenty J, Haïssaguerre M, Ross DL, Packer DL, Jaïs  
34  
35 (2006) Phrenic nerve injury after atrial fibrillation catheter ablation: Characterization and  
36  
37 outcome in a multicenter study. J Am Coll Cardiol 47:2498-2503  
38  
39  
40  
41  
42  
43 13. Franceschi F, Dubuc M, Guerra PG, Khairy P (2011) Phrenic nerve monitoring with  
44  
45 diaphragmatic electromyography during cryoballoon ablation for atrial fibrillation: the first  
46  
47 human application. Heart Rhythm 8:1068-1071  
48  
49  
50  
51  
52  
53 14. Tokuda M, Matsuo S, Isogai R, Uno G, Tokutake K, Yokoyama K, Kato M, Narui R,  
54  
55 Tanigawa S, Yamashita S, Inada K, Yoshimura M, Yamane T (2016) Adenosine testing  
56  
57 during cryoballoon ablation and radiofrequency ablation of atrial fibrillation: A propensity  
58  
59  
60  
61  
62  
63  
64  
65

- 1  
2  
3  
4  
5 score-matched analysis. *Heart Rhythm* 13:2128-2134  
6  
7  
8  
9 15. Saito T, Waki K, Becker AE (2000) Left atrial myocardial extension onto pulmonary veins  
10  
11 in humans: Anatomic observations relevant for atrial arrhythmias. *J Cardiovasc*  
12  
13 *Electrophysiol* 11:888-894  
14  
15  
16  
17  
18 16. Casella M, Dello Russo A, Russo E, Al-Mohani G, Santangeli P, Riva S, Fassini G, Moltrasio  
19  
20 M, Innocenti E, Colombo D, Bologna F, Izzo G, Gallinghouse JG, Di Biase L, Natale A,  
21  
22 Tondo C (2014) Biomarkers of myocardial injury with different energy sources for atrial  
23  
24 fibrillation catheter ablation. *Cardiol J* 21:516-523  
25  
26  
27  
28  
29  
30  
31  
32 17. Datino T, Macle L, Qi XY, Maguy A, Comtois P, Chartier D, Guerra PG, Arenal A,  
33  
34 Fernández-Avilés F, Nattel S (2010) Mechanisms by which adenosine restores conduction in  
35  
36 dormant canine pulmonary veins. *Circulation* 121:963-972  
37  
38  
39  
40  
41  
42  
43 18. Ouyang F, Bänsch D, Ernst S, Schaumann A, Hachiya H, Chen M, Chun J, Falk P, Khanedani  
44  
45 A, Antz M, Kuck KH (2004) Complete isolation of left atrium surrounding the pulmonary  
46  
47 veins: New insights from the double-Lasso technique in paroxysmal atrial fibrillation.  
48  
49 *Circulation* 110:2090–2096  
50  
51  
52  
53  
54  
55  
56 19. Chen H, Yang B, Ju W, Zhang F, Hou X, Chen C, Zhai L, Wang J, Cao K, Chen M, Tse HF  
57  
58 (2010) Long-term clinical implication of the occurrence of dissociated pulmonary vein  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5  
6  
7  
8  
9  
10  
11  
12  
13  
14  
15  
16  
17  
18  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29  
30  
31  
32  
33  
34  
35  
36  
37  
38  
39  
40  
41  
42  
43  
44  
45  
46  
47  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

activities after circumferential left atrial ablation in patients with paroxysmal atrial  
fibrillation. *Circ J* 75:73–79

1  
2  
3  
4  
5  
6  
7  
8  
9  
10  
11  
12  
13  
14  
15  
16  
17  
18  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29  
30  
31  
32  
33  
34  
35  
36  
37  
38  
39  
40  
41  
42  
43  
44  
45  
46  
47  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

Figure legends

Figure 1.

A representative case of regular ectopic rhythm in an isolated pulmonary vein (PV).

Regular dissociated potentials were observed in the PV.

CS, coronary sinus; prox, proximal; dis, distal.

Figure 2.

A representative case of fibrillatory activity in an isolated PV.

Fibrillatory firing was observed in the PV. However, it did not conduct to the atrium.

The abbreviations are the same as in the previous figure.

Figure 3.

The incidence of dissociated PV activity (DPVA) after CB-PVI or RF-PVI.

1  
2  
3  
4  
5 PVI, pulmonary vein isolation; LSPV, left superior pulmonary vein; LI, left inferior; RS, right  
6  
7  
8 superior; RI, right inferior. The other abbreviations are the same as in the previous figures.  
9

10  
11  
12 \* Statistically significant ( $P < 0.05$ ).  
13  
14  
15  
16  
17  
18  
19  
20

21 Figure 4.

22  
23  
24  
25 The characteristics of DPVA after PVI.  
26

27  
28  
29 The abbreviations are the same as in the previous figures.  
30  
31  
32  
33  
34  
35  
36  
37

38 Figure 5.

39  
40  
41  
42 The AF-free rate after the initial ablation procedure according to the presence of DPVA after the  
43  
44 index ablation procedure.  
45  
46  
47  
48

49  
50 Kaplan-Meier curves showing the AF-free rate after the initial CB (left panel) and RF (right  
51  
52 panel) ablation procedures according to the presence of DPVA.  
53  
54  
55  
56

57 The other abbreviations are the same as in the previous figures.  
58  
59  
60  
61  
62  
63  
64  
65

1  
2  
3  
4  
5  
6  
7  
8  
9  
10  
11  
12  
13  
14  
15  
16  
17  
18  
19  
20  
21  
22  
23  
24  
25  
26  
27  
28  
29  
30  
31  
32  
33  
34  
35  
36  
37  
38  
39  
40  
41  
42  
43  
44  
45  
46  
47  
48  
49  
50  
51  
52  
53  
54  
55  
56  
57  
58  
59  
60  
61  
62  
63  
64  
65

Figure 6.

Incidence of PV reconnection in repeat ablation procedure.

The abbreviations are the same as in the previous figures.

Figure 7.

Left panel: A representative position of circular catheter in PV.

Middle panel: Ablation sites and the virtual isolation line (dotted red line) of RF PVI.

Right panel: Freezing sites and the virtual isolation line (dotted red line) of CB PVI.

Abbreviations are the same as in the previous figures.

Table 1.

The characteristics of the patients who underwent cryoballoon and radiofrequency ablation

	All N=294	Cryoballoon ablation (n=147)	Radiofrequency ablation (n=147)	<i>P</i> value
Sex (male)	253 (86%)	127 (86%)	126 (86%)	0.87
Age (years)	57.3 ± 10.2	57.0 ± 11.0	57.6 ± 9.6	0.62
BMI	23.7 ± 3.1	23.6 ± 3.1	23.7 ± 3.1	0.57
LA diameter (mm)	36.6 ± 5.2	36.4 ± 4.7	36.7 ± 5.6	0.68
LV ejection fraction (%)	64.8 ± 5.0	64.8 ± 4.8	64.8 ± 5.2	0.99
eGFR (ml/min/1.73m <sup>2</sup> )	75.5 ± 13.1	75.6 ± 12.8	75.5 ± 13.5	0.92
Hypertension	98 (33%)	49(33%)	49 (33%)	1.00
Diabetes Mellitus	26 (9%)	13 (9%)	13 (9%)	1.00
Obstructive sleep apnea	32 (11%)	18 (12%)	14 (10%)	0.67
Congestive heart failure	10 (3.4%)	5 (3.4%)	5 (3.4%)	1.00
Coronary artery disease	14 (4.8%)	6 (4.1%)	8 (5.4%)	0.58
BNP (pg/ml)	41.7 ± 59.9	40.3 ± 59.0	43.0 ± 60.9	0.70
No. of AAD	1.0 ± 0.8	1.0 ± 0.8	1.0 ± 0.8	0.89
CHADS <sub>2</sub> Score	0.6 ± 0.8	0.6 ± 0.8	0.6 ± 0.8	1.00
AF history (years)	3.0 ± 3.7	3.2 ± 4.1	2.8 ± 3.2	0.06

The data are presented as the mean ± SD or n (%).

LA, left atrium; LV, left ventricle; eGFR, estimated glomerular filtration rate; BNP, B-type natriuretic peptide; AAD, anti-arrhythmic drug.

Table 2

The procedural details

	Cryoballoon ablation	Radiofrequency ablation	<i>P</i> value
Total procedure time (min)	143 ± 32	172 ± 48	<0.001
Total fluoroscopic time (min)	62 ± 18	99 ± 33	<0.001
Ablation time (s)			
LSPV	314 ± 163	454 ± 246	<0.001
LIPV	255 ± 122	271 ± 161	0.33
RSPV	254 ± 96	567 ± 280	<0.001
RIPV	259 ± 122	410 ± 239	<0.001

The data are presented as the mean ± SD.

LSPV, left superior pulmonary vein; LI, left inferior; RS, right superior; RI, right inferior. The

other abbreviations are the same as in the previous tables.













