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Electrical Storms in Patients with an Implantable Cardioverter Defibrillator

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Purpose: In some patients with an implantable cardioverter defibrillator (ICD), multiple episodes of electrical storm (ES) can occur. We assessed the prevalence, features, and predictors of ES in patients with ICD. Materials and Methods: Eighty-five patients with an ICD were analyzed. ES was defined as the occurrence of two or more ventricular tachyarrhythmias within 24 hours. Results: Twenty-six patients experienced at least one ES episode, and 16 patients experienced two or more ES episodes. The first ES occurred 209 ± 277 days after ICD implantation. In most ES cases, the index arrhythmia was ventricular tachycardia (65%). There were no obvious etiologic factors at the onset of most ES episodes (57%). More patients with a structurally normal heart (p = 0.043) or ventricular fibrillation (VF) as the index arrhythmia (p =0.017) were in the ES-free group. Kaplan-Meier estimates and a log-rank test showed that patients with nonischemic dilated cardiomyopathy (DCMP) (log-rank test, p =0.016) or with left ventricular ejection fraction < 35% (p = 0.032) were more likely to experience ES, and that patients with VF (p = 0.047) were less affected by ES. Cox proportional hazard regression analysis showed that nonischemic DCMP correlated with a greater probability of ES (hazard ratio, 3.71; 95% confidence interval, 1.16-11.85; p = 0.027). Conclusion: ES is a common and recurrent event in patients with an ICD. Nonischemic DCMP is an independent predictor of ES. Patients with VF or with a structurally normal heart are less likely to experience ES.

Key Words: Implantable cardioverter defibrillators, ventricular tachycardia, ventricular fibrillation

INTRODUCTION

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This is an Open Access article distributed under the terms of the Creative Commons Attribution Non-Commercial License (http://creativecommons.org/ licenses/by-nc/3.0) which permits unrestricted noncommercial use, distribution, and reproduction in any medium, provided the original work is properly cited. Sudden cardiac death remains one of the leading causes of death.¹ An implantable cardioverter defibrillator (ICD) prolongs the lifespan of patients when used for primary or secondary prophylaxis of sudden cardiac death.²⁻⁴ Patients with an ICD can develop a so-called electrical storm (ES) during follow-up.⁵⁻⁸ Several studies have assessed the prevalence and possible predictors of ES in patients with an ICD,^{5,8-13} but there are insufficient data about ES in patients with an ICD and espe-

cially about the predictors of ES.^{5,10} We examined the incidence, features, and timing of the occurrence and predictors of ES in patients with an ICD.

MATERIALS AND METHODS

Patient population

The medical records of 85 patients with an ICD implanted from October 1999 to April 2009, who were treated at Samsung Medical Center in Seoul, Korea, were analyzed retrospectively. All patients gave written informed consent for the procedure of ICD implantation. Of these patients, any patient who had had one or more episode of ES was included in the study population. All arrhythmias detected during an ES episode were also reviewed. The study was approved by the regional committee for ethnics in medical research.

Indication of ICD implantation

The indication for ICD implantation was defined as secondary prevention in patients who had experienced aborted sudden cardiac death, sustained ventricular tachyarrhythmia, or in the context of presumed tachyarrhythmic syncopal attacks.¹⁴ The indication for ICD implantation in all other patients was categorized as primary prevention.

Implanted ICDs

The ICDs were manufactured by St. Jude Medical, Inc. (n = 45), Medtronic, Inc. (n = 38), or Guidant Corp. (n = 2). Single chamber devices were implanted in 77 patients, and double chamber devices in 8 patients. A total of 94 procedures were performed in these 85 patients.

Ventricular tachyarrhythmia episodes were appropriately detected by the device. Some devices recorded templates of the ventricular electrogram during sinus rhythm. During a tachyarrhythmia, the device compared the electrograms during the tachycardia to the baseline. Mean detection cutoff for slow ventricular tachycardia (VT) was 380 milliseconds (159 beats/min), and 332 milliseconds (181 beats/ min) for fast VT, and 297 milliseconds (202 beats/min) for ventricular fibrillation (VF). Also, 5 patients were diagnosed to have VT, based on surface electrocardiogram during hospitalization.

We often attempted to tailor therapies based upon the results of electrophysiologic study and/or a patient's arrhythmia history (patient-specific tailored programming). ICDs could be programmed to provide different therapies to tachyarrhythmias in up to three heart rate zones. In each therapy zone, bursts of antitachycardia pacing (APT), cardioversion, or defibrillation could be delivered. Although a variety of algorithms exist, APT was usually programmed to be delivered at a rate that is slightly faster (at a cycle length 10 to 12 percent shorter) than the rate of the detected tachycardia. Devices were often programmed to deliver synchronized cardioversion for tachyarrhythmias in this range (heart rate below 160 or 180 beats/min), and usually programmed to deliver unsynchronized shocks for very rapid ventricular arrhythmias (heart rate > 180 or 200 beats/min).

Follow-up

After ICD implantation, the patients were followed up in our outpatient ICD clinic. The devices were interrogated, and the complete set of data (including intracardiac electrograms) was recorded. Blood samples, echocardiography, coronary angiography, reprogramming of ICD, or adjustment of drug therapy were performed as needed according to the events recorded and factors deemed causative.

Definitions

For this study, we defined ES as the occurrence of two or more ventricular tachyarrhythmia within 24 hours, which need immediate electrical therapy (ATP and/or shock) to be terminated, separated by a period of sinus rhythm.^{9,15-17} We divided patients into two groups according to occurrence of ES. Individual demographic and clinical data were entered into a continually updated database. Multiple device activation from supraventricular tachyarrhythmias or other factors not related to ventricular tachyarrhythmic events (defined as "inappropriate therapy"), as judged by the clinical picture and analysis of stored electrograms, were excluded.

Statistical analysis

The data are expressed as mean ± standard deviation or as percentages. Differences in the frequency of characteristics were assessed by the two independent-samples tests for continuous variables. Chi-square statistics (or Fisher's exact test if applicable) were used to analyze discrete variables. We used survival analysis method to investigate the relationship between the significance of clinical outcomes and ES. Cumulative incidence of clinical outcomes was estimated by the Kaplan-Meier method, and differences were assessed with the log-rank test for ES. We used Cox proportional hazard regression analysis with the forward selection method to identify precipitating factors. All tests of statistical significance were two-tailed, and a p value of 0.05 was considered significant. Statistical analyses were performed using SPSS statistical software (version 15.0; SPSS Inc., Chicago, IL, USA).

RESULTS

Clinical and laboratory characteristics of ICD patients

Eighty-five patients with an ICD were followed for a mean of 861 ± 715 days. We compared the baseline characteristics between patients with ES (n = 26, 31%) and without ES (n = 59, 69%) during follow-up. The mean follow-up period

Table 1. Comparison of Patients with and without Electrical Storm

	Electric	Electrical storm		
	Yes $(n = 26)$	No (n = 59)	<i>p</i> value	
Age (yrs)	56 ± 15	51 ± 17	0.283	
Male	22 (85%)	52 (88%)	0.730	
Clinical history				
Hypertension	11 (42%)	19 (32%)	0.369	
Diabetes mellitus	1 (4%)	9 (15%)	0.166	
Atrial fibrillation	7 (27%)	18 (31%)	0.738	
Syncope	16 (62%)	45 (76%)	0.164	
Aborted arrest	7 (27%)	25 (42%)	0.176	
Structural heart disease				
Coronary artery disease	11 (42%)	23 (39%)	0.767	
Nonischemic DCMP	4 (15%)	7 (12%)	0.730	
None (structurally normal heart)	4 (15%)	23 (39%)	0.031	
Concomitant medications				
β-blocker	19 (73%)	37 (63%)	0.353	
ACE inhibitors or ARB	15 (58%)	35 (59%)	0.888	
Amiodarone	3 (12%)	2 (3%)	0.141	
QRS duration \geq 120 (ms)	10 (38%)	15 (25%)	0.224	
LVEF < 35%	10 (38%)	15 (25%)	0.243	
Index arrhythmia				
Ventricular tachycardia	22 (85%)	42 (71%)	0.225	
Ventricular fibrillation	5 (19%)	27 (46%)	0.017	
Indication for ICD implantation			0.133	
Primary prevention	1 (4%)	9 (15%)		
Secondary prevention	25 (96%)	50 (85%)		
Гуре of ICD			0.656	
ICD-single chamber	23 (88%)	54 (92%)		
ICD-dual chamber	3 (12%)	5 (8%)		
ICD manufactured by			0.657	
St. Jude	15 (58%)	30 (51%)		
Medtronic	10 (38%)	28 (47%)		
Guidant	1 (4%)	1 (2%)		
Follow-up duration (days)	990 ± 742	804 ± 702	0.262	
Time to first ES (days)	209 ± 277			
Time to second ES from first ES (days)	251 ± 362			

DCMP, dilated cardiomyopathy; ACE, angiotensin converting enzyme; ARB, angiotensin II receptor blocker; LVEF, left ventricular ejection fraction; ICD, implantable cardioverter defibrillator. ES, electrical storm.

was 990 ± 742 days for the patients with ES, and 804 ± 702 days for those without ES. The clinical characteristics in both groups were similar, except for two clinical characteristics: a structurally normal heart and VF as the index arrhythmia for ICD implantation, which were more frequent in the ES-free patients (Table 1). Twenty-six patients experienced at least one ES episode. Overall, 115 ES episodes were recorded (median, 2 ES per patient; range, 1-45), and 16 (62%) patients in the ES group experienced two or more ES episodes. ES occurred first at a mean of 209 ± 277 days after ICD implantation, and the mean duration between the first and second ES occurrence was 251 ± 326 days. Shocks alone were used to treat 28 of the 115 initial ES episodes (n = 28, 24%), and the remaining 87 episodes were treated

with ATP alone (n = 55, 48%) or a combination of shocks and ATP (n = 32, 28%). In most ES cases, the recorded index arrhythmia was ventricular tachycardia (VT, 65%). VF and both VT and VF occurred in 11% and 24% of the ES episodes, respectively. One patient experienced a storm immediately following ICD implantation (on the same day). Details of the clinical characteristics and management of the patients with ES are presented in Table 2. Three patients have had heart transplants; two in the ES group and one in the ES-free group. Nine deaths occurred, five in the ES group and four in the ES-free group.

Etiologic factors of ES

No obvious etiologic factor was evident at the onset of the

Patient no.	Sex	Age (yrs)	Disease	Prevention	Index arrhythmia	LVEF (%)	Time to first ES (days)	Total number of ES episodes
1	М	73	CAD	Secondary	VT	27	1,000	10
2	М	70	CAD	Secondary	VT	39	67	3
3	F	26	SNH	Secondary	VT	72	276	1
4	М	81	CAD	Secondary	VT	38	186	4
5	М	69	CAD	Secondary	VT	30	31	1
6	М	54	NIDCMP	Secondary	VT	29	9	5
7	М	73	ARVD	Secondary	VT	58	950	1
8	М	45	SNH	Secondary	VF	66	43	4
9	М	77	CAD	Secondary	VT	35	262	1
10	М	58	CAD	Secondary	VT	25	0	45
11	М	58	NIDCMP	Secondary	VT	21	60	2
12	М	39	HCMP	Secondary	VF	66	672	2
13	М	46	CAD	Secondary	VT	37	91	1
14	М	49	NIDCMP	Secondary	VT	29	17	3
15	М	35	SNH	Secondary	VF	65	16	1
16	М	64	ARVD	Primary	VT	54	371	2
17	М	59	CAD	Secondary	VT	30	227	1
18	F	72	NIDCMP	Secondary	VT	32	100	2
19	F	37	HCMP	Secondary	VT	76	78	2
20	М	47	HCMP	Secondary	VT	15	14	13
21	М	61	HCMP	Secondary	VT	70	3	1
22	М	42	HCMP	Secondary	VT	53	48	2
23	М	35	SNH	Secondary	VT	59	162	2
24	М	47	HCMP	Secondary	VT	39	138	4
25	F	72	CAD	Secondary	VT	25	147	1
26	М	72	CAD	Secondary	VF	39	475	1

Table 2. Clinical and Laboratory Characteristics of Patients Presenting with Electrical Storm

LVEF, left ventricular ejection fraction; ES, electrical storm; CAD, coronary artery disease; SNH, structurally normal heart; NIDCMP, nonischemic dilated cardiomyopathy; ARVD, arrhythmogenic right ventricular dysplasia; HCMP, hypertrophic cardiomyopathy; VT, ventricular tachycardia; VF, ventricular fibrillation. storm in 65 (57%) ES episodes. Worsening congestive heart failure was evident as an etiologic factor in 19 (17%) cases. Fifteen (13%) storms occurred in the context of other medical illnesses. Six (5%) ES episodes were associated with either non-compliance or adjustment of antiarrhythmics or other medications. Three (3%) ES episodes occurred during and around periods of excess alcohol consumption, and four (3%) storms followed periods of unusual physical or emotional stress. One ES episode was probably caused by hyperkalemia (5.9 mmol/L) and one ES episode was caused by malposition of an ICD lead (Fig. 1).

Predictors of ES

Chi-square tests showed that more patients with a structurally normal heart (p = 0.043) or VF as the index arrhythmia for ICD (p = 0.017) were in the ES-free group. The odds ratios of a structurally normal heart and VF as the index arrhythmia for ICD were 0.285 [95% confidence interval (CI): 0.087-0.933] and 0.273 (95% CI: 0.091-0.824), respectively. These data suggest that having VF as the index arrhythmia or a structurally normal heart seemed to protect against ES. We intended to find predictive factors for ES. The Kaplan-Meier estimate and log-rank test showed that patients with nonischemic DCMP as the underlying heart disease (logrank test, p = 0.016) or with left ventricular ejection fraction (LVEF) < 35% (log-rank test, p = 0.032) were more likely to experience ES, and that patients with VF as the index arrhythmia (log-rank test, p = 0.047) were less affected by ES (Table 3). The Cox proportional hazard regression analysis showed that only nonischemic DCMP as the underlying heart disease (hazard ratio, 3.71; 95% CI: 1.16-11.85; p =0.027) was significantly and independently associated with a greater probability of ES during follow-up.

DISCUSSION

Main findings

Our study reveals several findings about the occurrence of ES in ICD patients. Twenty-six (31%) patients who underwent ICD placement experienced at least one ES episode, and a substantial proportion (62%) of patients experienced repeated ES episodes. Most ES episodes involved VT and occurred usually without obvious etiologic factors (57%). Our data suggest that patients with nonischemic DCMP as the underlying heart disease are at greater risk for ES. The

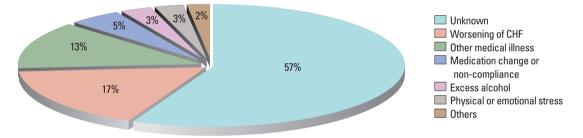


Fig. 1. Presumed etiologic factors of electrical storm. Cause assignment was made by the physician at the time the electrical storm occurred. CHF, congestive heart failure.

Covariates	Median survival (days)	Chi-square	Significance level
Nonischemic DCMP		5.79	0.016
No	1,638		
Yes	1,160		
LVEF		4.60	0.032
\geq 35%	1,638		
< 35%	1,292		
VF as index arrhythmia		3.94	0.047
No	1,441		
Yes	1,974		

 Table 3. Survival Analysis of the Univariate Predictors of Electrical Storm (Kaplan-Meier Estimate and Log-Rank Test)

DCMP, dilated cardiomyopathy; LVEF, left ventricular ejection fraction; VF, ventricular fibrillation.

finding that patients with VF as the index ventricular tachyarrhythmia or with a structurally normal heart were at lower risk of ES is interesting.

Comparison the previous studies on ES

Several studies have attempted to identify the risk factors for the development of ES. Verma, et al.9 demonstrated that coronary artery disease as the underlying heart disease was an independent predictor of ES in 2,028 patients with ICDs. However, the Cox proportion hazard regression analysis of our data indicated that nonischemic DCMP as the underlying heart disease was an independent predictor of ES. The incidence of ES in our study is similar to 10-28% range reported during follow-up durations of 13-33 months.^{5,10,18,19} Our study and previous reports together suggest that most storm episodes (86-97%) are caused by monomorphic VT. VF alone accounts for 1-21% of ES, mixed VT/VF for 3-14%, and polymorphic VT 2-8%.5,18,20-22 Our findings are consistent with these values. In our study, most of the index arrhythmias causing ES were VT (65%). Identifiable triggers such as aggravation of congestive heart failure (31%) or electrolyte disorders (20%) have been reported.²³ Most (57%) of the ES episodes in our study had no obvious etiologic factors, although we found other identifiable precipitating factors, such as worsening congestive heart failure (17%), acute medical illness (13%), and physical or emotional stress (3%).

Clinical profiles of the ES-free group

Chi-square analysis with odds ratios, the Kaplan-Meier estimate, and the log-rank test identified two protective factors against ES: VF as the presenting arrhythmia and a structurally normal heart. Consistent with our trial, Exner, et al.¹⁰ reported that the majority of initial ES episodes were related to episodes of VT (77 of 90 initial episodes; 86%), and the residual 14% of episodes were attributed to VF or a mixture of VT and VF. Raitt, et al.24 also showed that patients who were included in study after an episode of VT, were more likely to experience an episode of VT (73.5% vs. 30.1%, p < 0.001), and were less likely to have an episode of VF (18.3% vs. 28.0%, p = 0.013) than those who enrolled after an episode of VF in the AVID trial, which suggested that there are significant differences in the electrophysiologic features of these 2 groups. On the other hand, Verma, et al.9 described that 208 (10%) patients presented with ES in 2,028 patients who were evaluated in the ICD clinic, and VF was the reason of ES in 99 of 208 patients, for an overall incidence of 48%.

Limitations of this study

Our study is limited by the relatively small sample size comprising a selected patient population and the retrospective, observational single-center design. Our results should be confirmed in a large, prospective study.

Conclusions

ES is a relatively frequent complication that may occur at any time after ICD implantation and can become a recurrent event in ICD recipients. Our data provide evidence to indicate that nonischemic DCMP as the underlying heart disease is associated with an increased risk of ES, and that patients with VF as the index arrhythmia or a structurally normal heart are less likely to experience ES.

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