

Variable	Total study population (<i>n</i> = 257)	Ventricular arrhythmia with clear cause (<i>n</i> = 220)	Ventricular arrhythmia with unclear cause (<i>n</i> = 37)	<i>P</i>-value
Age (years)	64.1 (52.9 – 71.8)	65.2 (56.0 – 72.0)	52.3 (43.4 – 63.9)	<0.001
Female sex	45 (17.5%)	34 (15.5%)	11 (29.7%)	0.035
Presenting ventricular arrhythmia				0.007
VF	173 (67.3%)	141 (64.1%)	32 (86.5%)	
Sustained VT	84 (32.7%)	79 (35.9%)	5 (13.5%)	
OHCA	182 (70.8%)	149 (67.7%)	33 (89.2%)	0.008
Type of ICD				<0.001
S-ICD	12 (4.7%)	4 (1.8%)	8 (21.6%)	
VVI-ICD	121 (47.1%)	102 (46.4%)	19 (51.4%)	
DDD-ICD	88 (34.2%)	78 (35.5%)	10 (27.0%)	
CRT-D	36 (14.0%)	36 (16.4%)	0 (0.0%)	
BMI (kg/m ²)	26.5 (24.4 – 29.4)	26.8 (24.6 – 29.4)	25.3 (23.9 – 30.1)	0.216
NYHA class				0.139
I or II	175 (68.1%)	149 (67.7%)	26 (70.3%)	
III or IV	30 (11.7%)	29 (13.2%)	1 (2.7%)	
Medical history				
Hypertension	124 (48.2%)	108 (49.1%)	16 (43.2%)	0.510
Dyslipidemia	96 (37.4%)	90 (40.9%)	6 (16.2%)	0.004
Myocardial infarction	137 (53.3%)	137 (62.3%)	0 (0.0%)	<0.001
DM	48 (18.7%)	43 (19.5%)	5 (13.5%)	0.384
Family history SCD	24 (9.3%)	22 (10.0%)	2 (5.4%)	0.545
Syncope	24 (9.3%)	20 (9.1%)	4 (10.8%)	0.760
Prior heart surgery	66 (25.7%)	66 (30.0%)	0 (0.0%)	<0.001
Atrial fibrillation	86 (33.5%)	77 (35.0%)	9 (24.3%)	0.203
Non-sustained VT	47 (18.3%)	40 (18.2%)	7 (18.9%)	0.915
CAD				<0.001
No	93 (36.2%)	59 (26.8%)	34 (91.9%)	
1 coronary artery	39 (15.2%)	36 (16.4%)	3 (8.1%)	
2 coronary arteries	54 (21.0%)	54 (24.5%)	0 (0.0%)	
3 coronary arteries	64 (24.9%)	64 (29.1%)	0 (0.0%)	
Echocardiography				
LVEF (%)	45 (34 – 53)	41 (33 – 50)	55 (51 – 58)	<0.001
LAVI (mL/m ²)	35 (29 – 45)	37 (30 – 45)	31 (25 – 38)	0.015
Maximum myocardial wall thickness (mm)	10.9 ± 2.1	10.9 ± 2.1	10.8 ± 2.2	0.883
LVEDD (mm)	55.1 ± 8.5	55.8 ± 8.6	50.9 ± 6.6	0.002
LV mass (g)	213 ± 64	218 ± 64	185 ± 57	0.018
RVF				0.202
Poor	3 (1.2%)	3 (1.4%)	0 (0.0%)	
Mediocre	6 (2.3%)	6 (2.7%)	0 (0.0%)	
Reasonable	36 (14.0%)	34 (15.5%)	2 (5.4%)	
Good	195 (75.9%)	160 (72.7%)	35 (94.6%)	
Mitral insufficiency				0.002
None	153 (59.5%)	121 (55.0%)	32 (86.5%)	
Mild (grade 1)	61 (23.7%)	58 (26.4%)	3 (8.1%)	
Moderate (grade 2)	20 (7.8%)	20 (9.1%)	0 (0.0%)	
Severe (grade 3)	5 (1.9%)	4 (1.8%)	1 (2.7%)	
LGE on CMR				<0.001
Yes	80 (31.1%)	73 (33.2%)	7 (18.9%)	
No	53 (20.6%)	27 (12.3%)	26 (70.3%)	
Electrocardiography				
Rhythm				0.406
Sinus rhythm	221 (86.0%)	187 (85.0%)	34 (91.9%)	

Atrial fibrillation	28 (10.9%)	26 (11.8%)	2 (5.4%)	0.370
Atrial flutter	2 (0.8%)	2 (0.9%)	0 (0.0%)	
Pacemaker	3 (1.2%)	2 (0.9%)	1 (2.7%)	
Ventricular heart rate (bpm)	67 ± 14	67 ± 14	69 ± 12	
PR interval (ms)	168 (152 – 188)	168 (152 – 190)	164 (148 – 186)	0.338
QRS duration (ms)	108 (98 – 128)	111 (98 – 133)	96 (88 – 106)	<0.001
QRS axis (degrees)	31 (-13 – 65)	30 (-18 – 64)	37 (-1 – 74)	0.072
QT interval (ms)	434 ± 51	441 ± 52	399 ± 33	<0.001
QRS fragmentation				<0.001
Yes	90 (35.0%)	77 (35.0%)	13 (35.1%)	0.062
QRS > 120ms	74 (28.8%)	72 (32.7%)	2 (5.4%)	
QRS morphology				
Normal	140 (54.5%)	111 (50.5%)	29 (78.4%)	
LBBB	25 (9.7%)	24 (10.9%)	1 (2.7%)	
LAFB	38 (14.8%)	34 (15.5%)	4 (10.8%)	
LPFB	4 (1.6%)	3 (1.4%)	1 (2.7%)	
RBBB	14 (5.4%)	13 (5.9%)	1 (2.7%)	
RBBB + LAFB	10 (3.9%)	10 (4.5%)	0 (0.0%)	
RBBB + LPFB	3 (1.2%)	3 (1.4%)	0 (0.0%)	
IVCD	18 (7.0%)	18 (8.2%)	0 (0.0%)	
Pacing	3 (1.2%)	2 (0.9%)	1 (2.7%)	
Early repolarization	15 (5.8%)	12 (5.5%)	3 (8.1%)	
Laboratory values				0.463
Hb (mmol/L)	8.1 ± 1.2	8.1 ± 1.2	8.3 ± 1.1	
Na ⁺ (mmol/L)	140 (138 – 141)	140 (138 – 141)	140 (139 – 141)	
K ⁺ (mmol/L)	4.2 ± 0.4	4.2 ± 0.4	4.2 ± 0.4	
Creatinine (μmol/L)	86 (75 – 100)	88 (77 – 104)	75 (64 – 87)	<0.001
eGFR (mL/min/1.73m ²)	77 ± 23	75 ± 23	91 ± 21	<0.001
Urea (mmol/L)	6.2 (4.9 – 7.7)	6.3 (4.9 – 8.1)	5.3 (4.6 – 6.8)	0.008
NT-proBNP (ng/L)	656 (226 – 1,625)	891 (295 – 1,831)	155 (46 – 330)	<0.001
Medication at baseline				<0.001
ACE-I/ARB	113 (44.0%)	106 (48.2%)	7 (18.9%)	
β-blocker	125 (48.6%)	118 (53.6%)	7 (18.9%)	
Calcium antagonist	43 (16.7%)	39 (17.7%)	4 (10.8%)	
Diuretic	56 (21.8%)	54 (24.5%)	2 (5.4%)	0.007
Statin	111 (43.2%)	107 (48.6%)	4 (10.8%)	<0.001
MRA	22 (8.6%)	22 (10.0%)	0 (0.0%)	0.052
Class 1 antiarrhythmic drugs	1 (0.4%)	1 (0.5%)	0 (0.0%)	1.000
Class 3 antiarrhythmic drugs	15 (5.8%)	15 (6.8%)	0 (0.0%)	0.137
Digoxin	9 (3.5%)	9 (4.1%)	0 (0.0%)	0.364

Data are expressed as *n* (%) in case of categorical data, mean ± standard deviation (SD) in case of normally distributed continuous data and median and interquartile range (IQR) in case of continuous data with a skewed distribution. In some cases, numbers may not add up to 100% due to missing data. The presented *P*-values reflect a comparison between patients with ventricular arrhythmia of clear and unclear cause and are presented in bold if *P* < 0.05. ACE-I, angiotensin-converting enzyme inhibitor; ARB, angiotensin receptor blocker; BMI, body mass index; CAD, coronary artery disease; CMR, cardiac magnetic resonance; CRT-D, cardiac resynchronization therapy ICD; DDD-ICD, dual-chamber ICD; DM, diabetes mellitus; eGFR, estimated glomerular filtration rate; Hb, hemoglobin; ICD, implantable cardioverter-defibrillator; IVCD, intraventricular conduction delay; LAFB, left anterior fascicular block; LAVI, left atrial volume index; LBBB, left bundle branch block; LGE, late gadolinium enhancement; LPFB, left posterior fascicular block; LV, left ventricle; LVEDD, left ventricular end-diastolic diameter; LVEF, left ventricular ejection fraction; MRA, mineralocorticoid receptor antagonist; NT-proBNP, N-terminal pro-brain natriuretic peptide; NYHA, New York Heart Association; OHCA, out-of-hospital cardiac arrest; RBBB, right bundle branch block; RVF, right ventricular function; SCD, sudden cardiac death; S-ICD, subcutaneous ICD; VF, ventricular fibrillation; VT, ventricular tachycardia; VVI-ICD, single-chamber ICD.

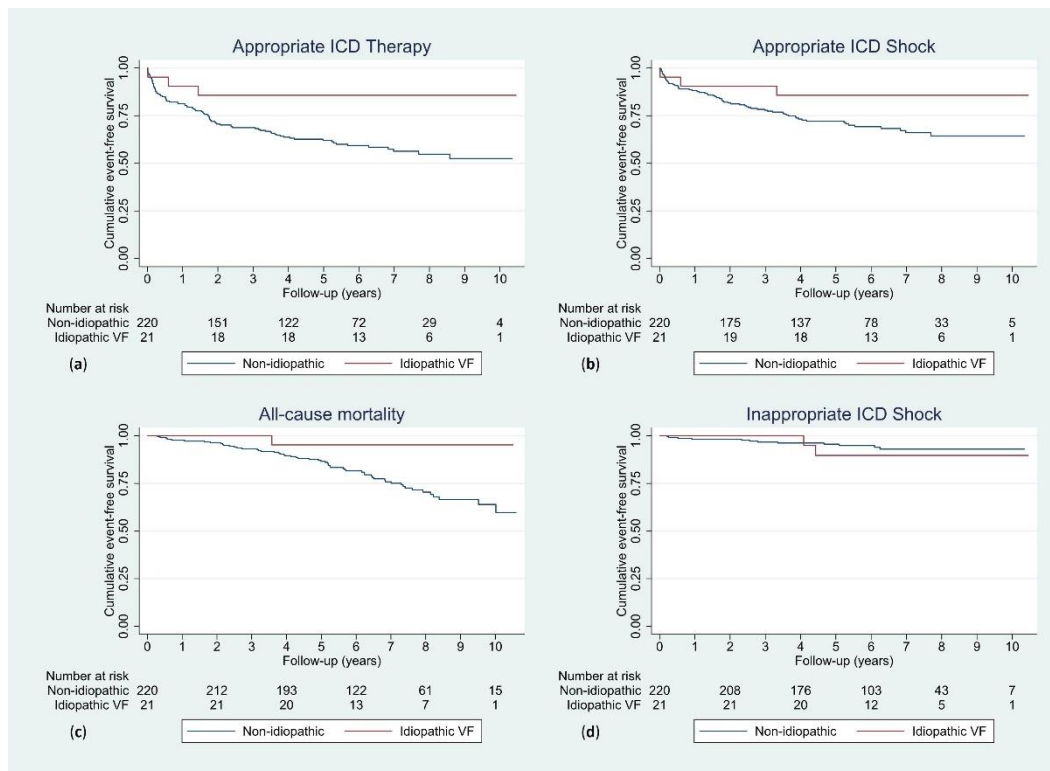


Figure S1. Kaplan-Meier curves displaying cumulative event-free survival for the idiopathic VF and non-idiopathic groups with appropriate ICD therapy (log-rank test $P = 0.022$) (a), appropriate ICD shock (log-rank test $P = 0.109$) (b), all-cause mortality (log-rank test $P = 0.045$) (c) and inappropriate ICD shock (log-rank test $P = 0.529$) (d) as outcomes. ICD, implantable cardioverter-defibrillator; VF, ventricular fibrillation.

Table S2. Cox regression with idiopathic VF as determinant of the outcomes, both univariably and after adjusting for potential confounders.				
	Univariable		Adjusted	
Outcome	HR (95% CI)	P-value	HR (95% CI)	P-value
Appropriate ICD therapy	0.28 (0.09 – 0.90)	0.032	0.48 (0.14 – 1.65)*	0.245
Appropriate ICD shock	0.40 (0.13 – 1.28)	0.122	0.75 (0.21 – 2.72)*	0.659
All-cause mortality	0.17 (0.02 – 1.22)	0.078	0.70 (0.09 – 5.65)§	0.741
Inappropriate ICD shock	1.61 (0.36 – 7.20)	0.532	2.28 (0.44 – 11.80)°	0.328

CI, confidence interval; HR, hazard ratio; ICD, implantable cardioverter-defibrillator; VF, ventricular fibrillation.

* Adjusted for age, sex, BMI, index arrhythmia, history of atrial fibrillation, prior syncope, history of non-sustained VT, history of myocardial infarction, QRS fragmentation, eGFR and left ventricular ejection fraction.

§ Adjusted for age, sex, index arrhythmia, diabetes, history of myocardial infarction, history of heart surgery and eGFR.

° Adjusted for age, sex and rhythm on ECG.

Table S3. Cox regression with idiopathic VF as determinant of the outcomes after adjusting for potential confounders, using multiple imputation.		
	Adjusted	
Outcome	HR (95% CI)	P-value
Appropriate ICD therapy	0.42 (0.12 – 1.42)*	0.162
Appropriate ICD shock	0.62 (0.18 – 2.20)*	0.460
All-cause mortality	0.69 (0.09 – 5.47)§	0.723
Inappropriate ICD shock	2.27 (0.44 – 11.80)°	0.328

CI, confidence interval; HR, hazard ratio; ICD, implantable cardioverter-defibrillator; VF, ventricular fibrillation.

* Adjusted for age, sex, BMI, index arrhythmia, history of atrial fibrillation, prior syncope, history of non-sustained VT, history of myocardial infarction, QRS fragmentation, eGFR and left ventricular ejection fraction.

§ Adjusted for age, sex, index arrhythmia, diabetes, history of myocardial infarction, history of heart surgery and eGFR.

° Adjusted for age, sex and rhythm on ECG.