

XII Meeting. State of the Art in

HEART FAILURE

CLINICAL PRACTICE AND ORGANIZATIONAL MODELS

Venue: Hotel Meliá María Pita, A Coruña

A Coruña 26-27 September 2025



#ACORUÑAHF2025




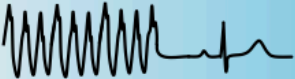



Arrhythmic risk in heart failure. Evaluation and management

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1. VA in Symptomatic HF

2. VA in “Asymptomatic” HF

Ventricular arrhythmias in acute heart failure: a clinical consensus statement of the Association for Acute CardioVascular Care, the European Heart Rhythm Association, and the Heart Failure Association of the European Society of Cardiology

Lower risk		High risk
Decompensated HF normal-high SBP		Systemic hypoperfusion
Single VA episode		Electrical storm
STEMI-non ST ACS		Incessant VA
Reversible cause		Known history advanced HF
Younger age		Older age
Little to no relevant comorbidities		Multiple relevant comorbidities

Structural and functional factors

Electrophysiologic remodelling

- Action potential prolongation
- Altered Ca²⁺ reuptake and SR leak
- Altered K⁺ currents
- Increased automaticity
- Abnormal cell coupling

Structural and functional remodelling

- Fibrosis and scar
- Chamber stretch
- Ventricular dilatation
- Hypertrophy
- Ischaemia

Neurohormonal mechanisms

- Increased adrenergic tone
- Increased RAAS activation
- Electrolyte abnormalities

Drugs interactions

- Anti-arrhythmics, antibiotics, anti-fungal, psychoactive
- Sympathomimetic drugs
- Phosphodiesterase inhibitors¹⁰
- Diuretics
- Digoxin

Arrhythmogenesis mechanism

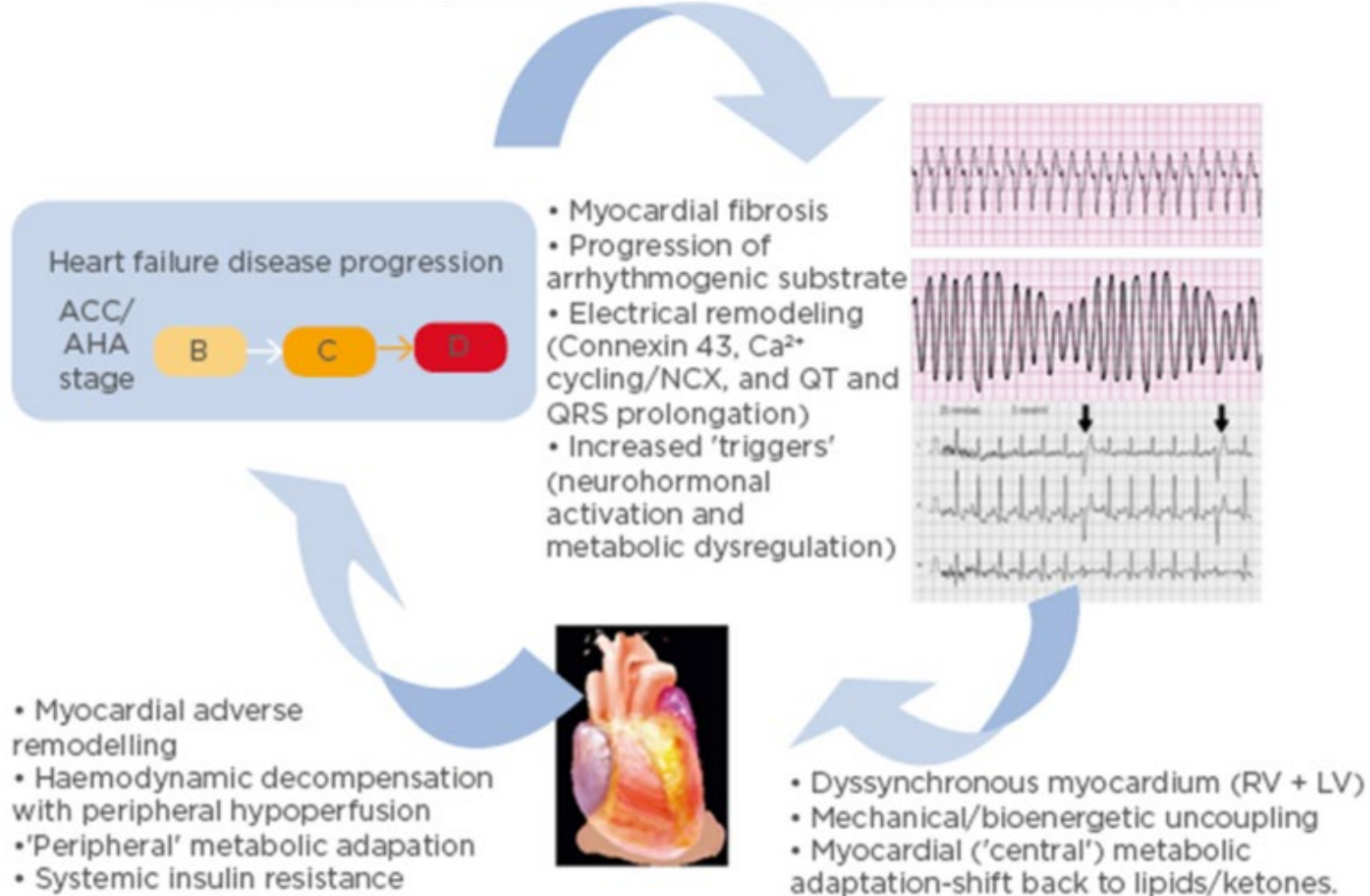
- Early afterdepolarization—focal
- Delayed afterdepolarization—focal
- Increased dispersion of repolarization
- Automatic tachycardias
- Re-entry
- Re-entry

- Re-entry
- Automatism
- Bundle branch re-entry
- Multiple mechanisms
- Multiple mechanisms

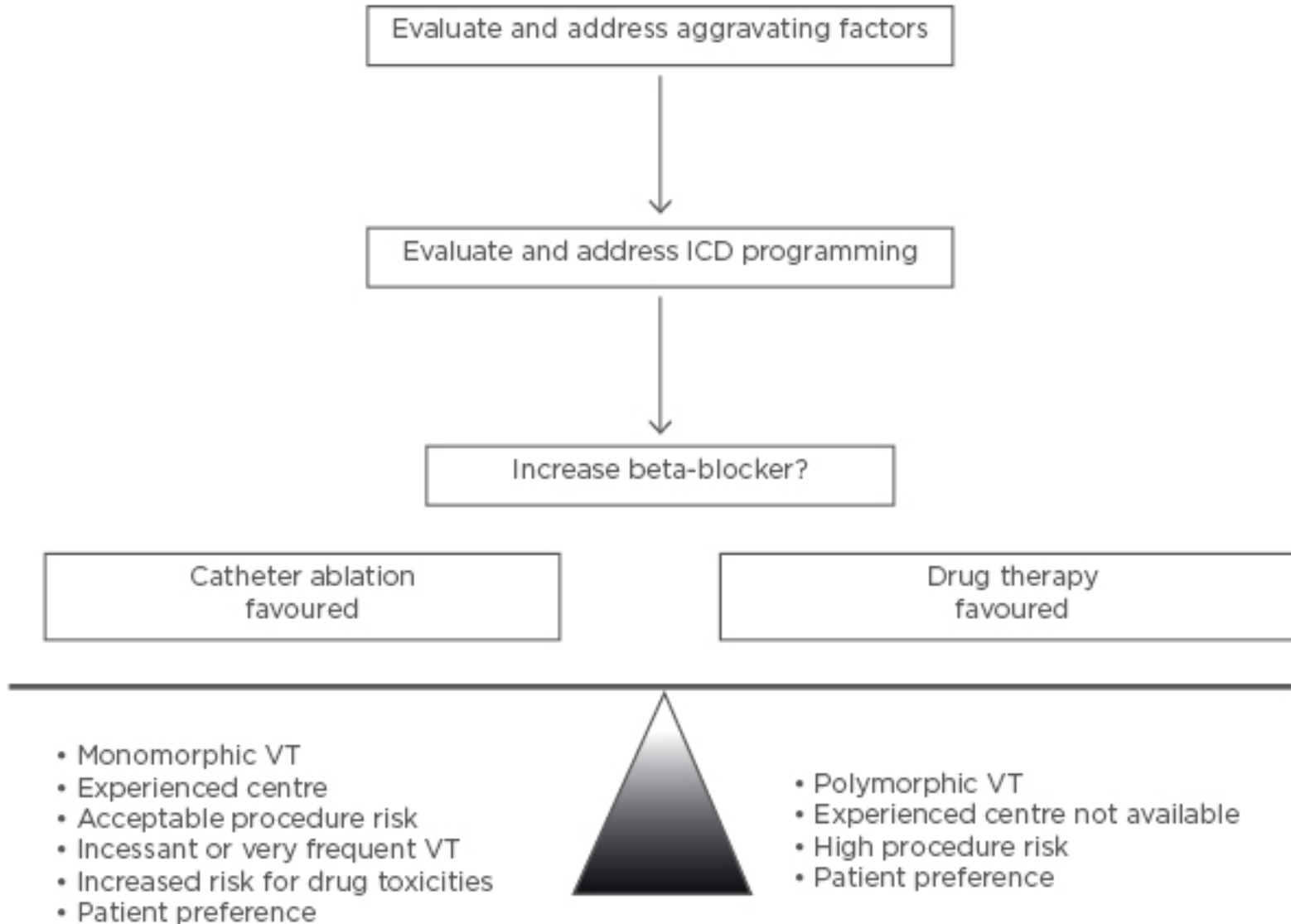
- Multiple mechanisms and enhancing arrhythmia propensity of other mechanisms

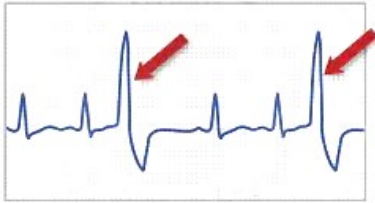
- QT prolongation
- Increased VT/VF risk
- Triggered activity, ischaemia
- Electrolyte disturbances
- Delayed after depolarization

Pathophysiological cycle of ventricular arrhythmias and progressive pump failure



Sustained VT or VF in patients with ICD





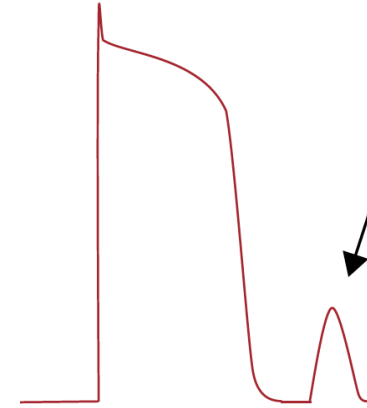
LVEF	PVC burden for 1 week Hotler			
	Infrequent (< 5%) (n = 108)		Frequent (≥ 5% or NSVT) (n = 196)	
	No	%	No	%
Severely reduced LVEF < 30 (Group A)	56	51.9	112	57.1
Reduced LVEF ≥ 30 (Group B)	52	48.1	84	42.9

	Cardiomyopathy			
	Ischemic (n 236)		Non-ischemic (n 68)	
	No.	%	No.	%
Infrequent PVCs (< 5%)	84	35.6	24	35.3
Frequent (≥ 5%)	152	64.4	44	64.7

Sanhoury, M., Mohamed, F., Sadaka, M. *et al.* The impact of asymptomatic ventricular arrhythmias on the outcome of heart failure patients with reduced ejection fraction

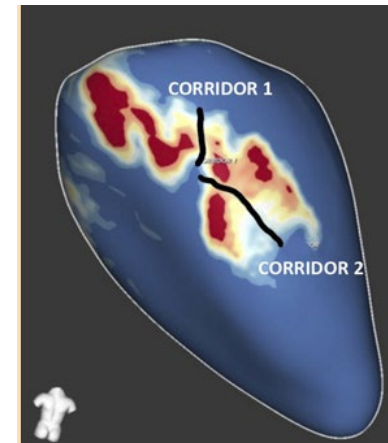
2. VA in “Asymptomatic” HF

Automaticity and triggered activity are achievable in individual cells



HFpEF

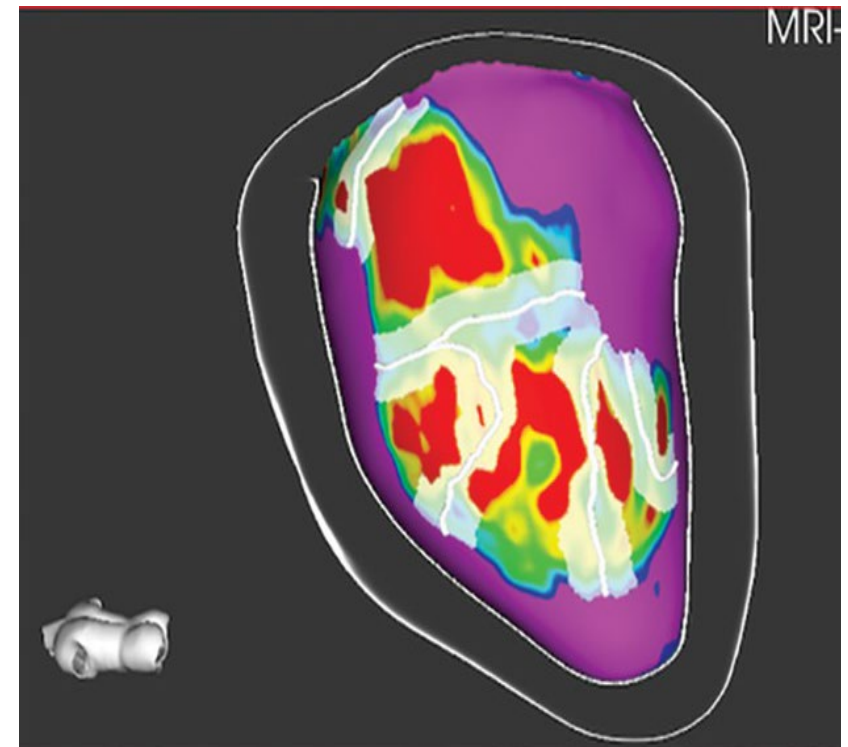
Reentry exists within a mass of myocardial tissue



HFrEF

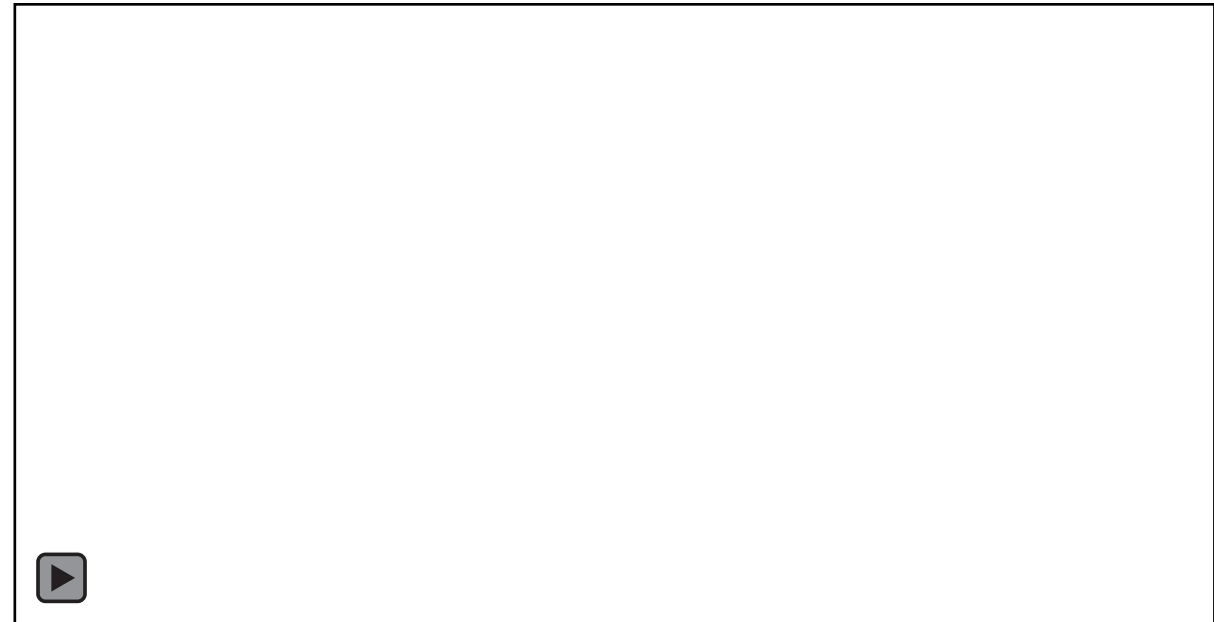
No structural disease

***Scar
cardiomyopathy***



No structural disease

***Scar
cardiomyopathy***

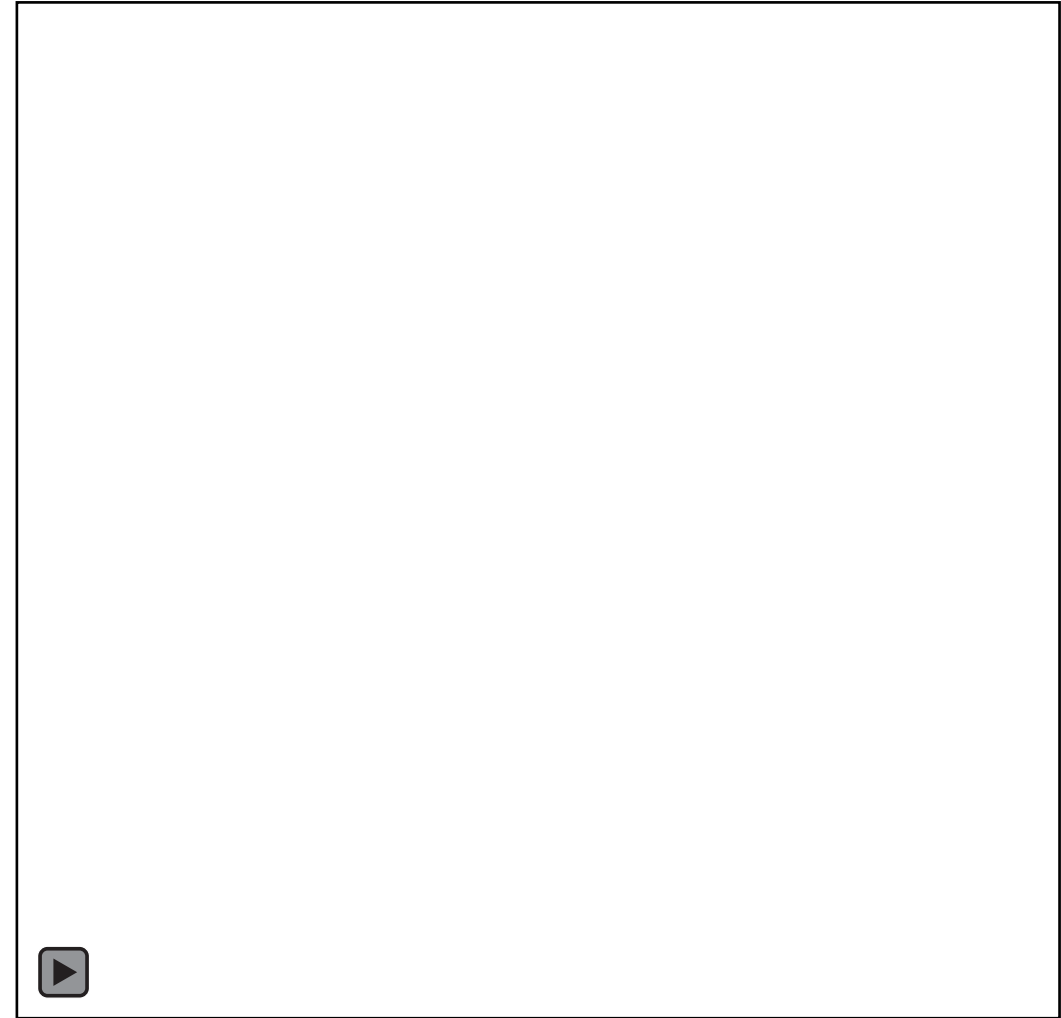
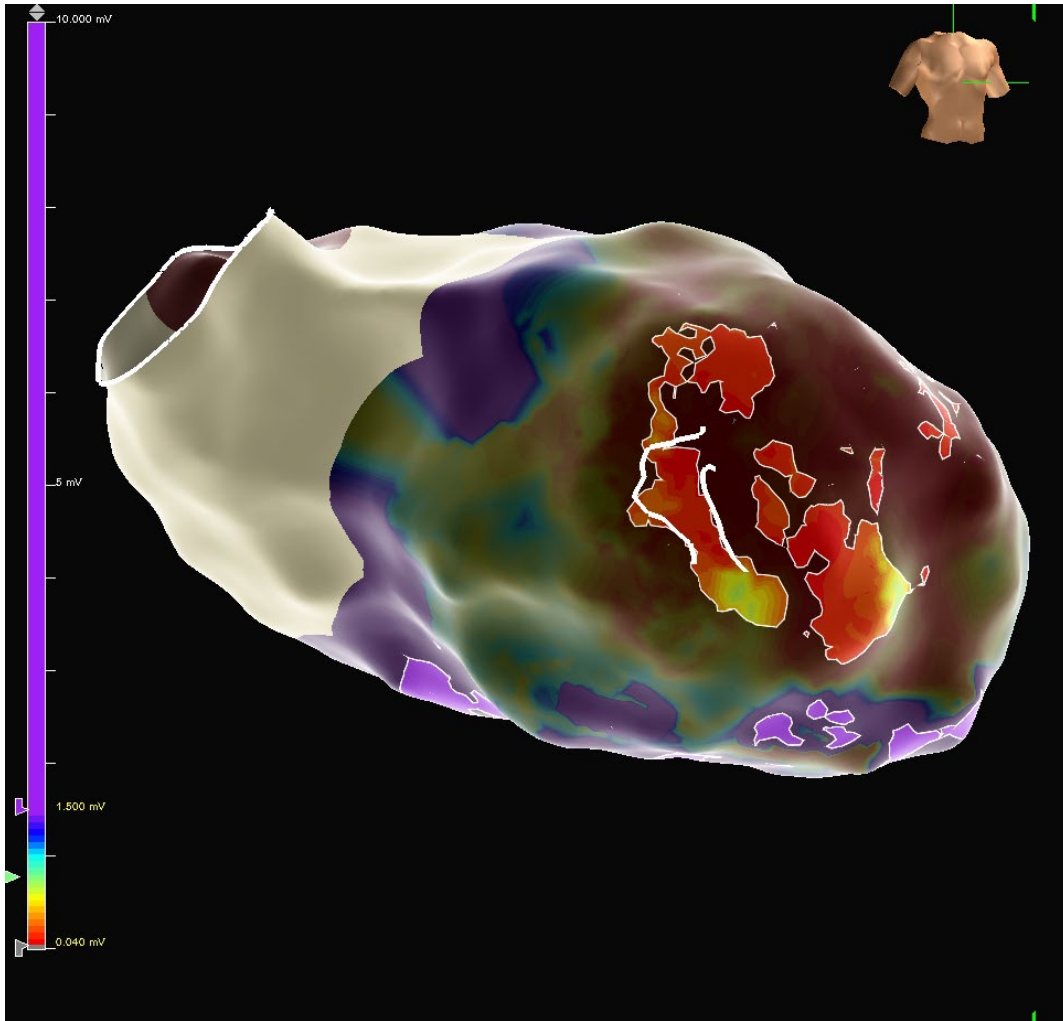


Ischemic cardiomyopathy



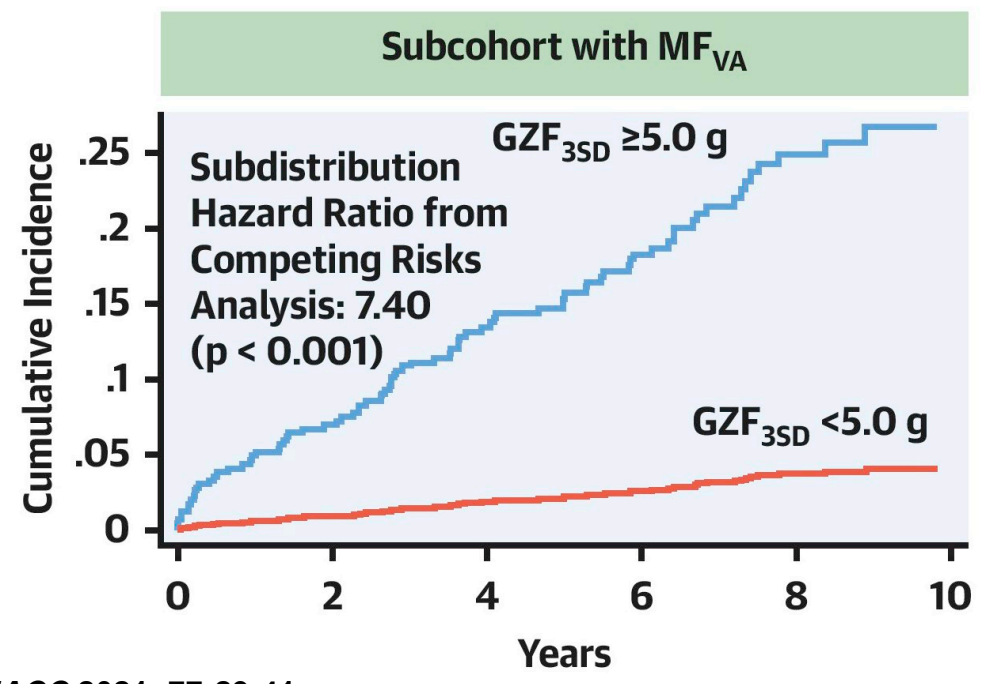
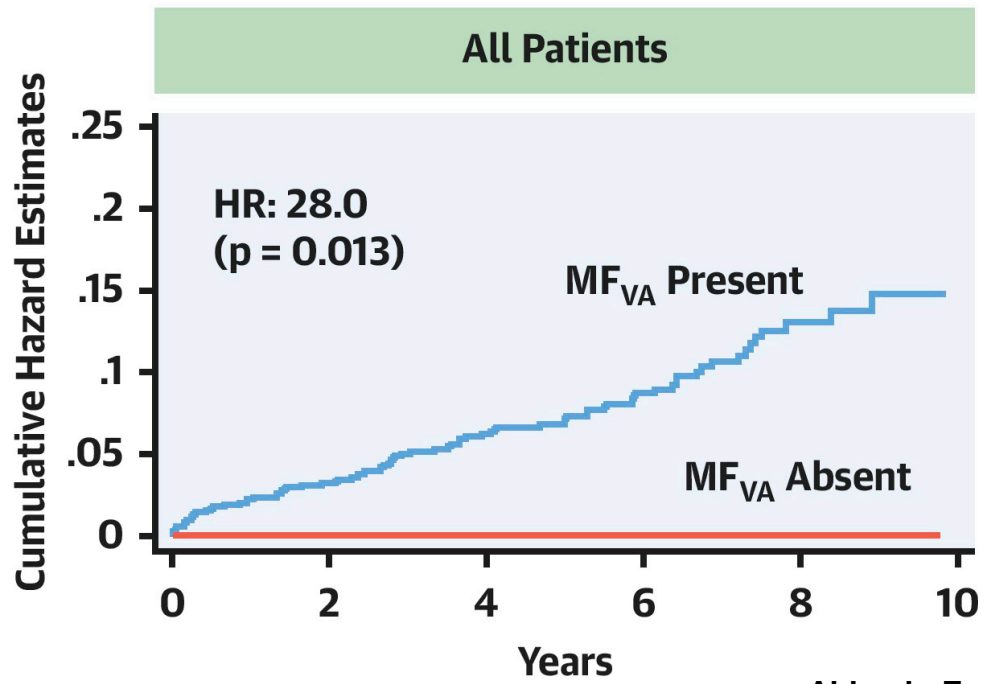
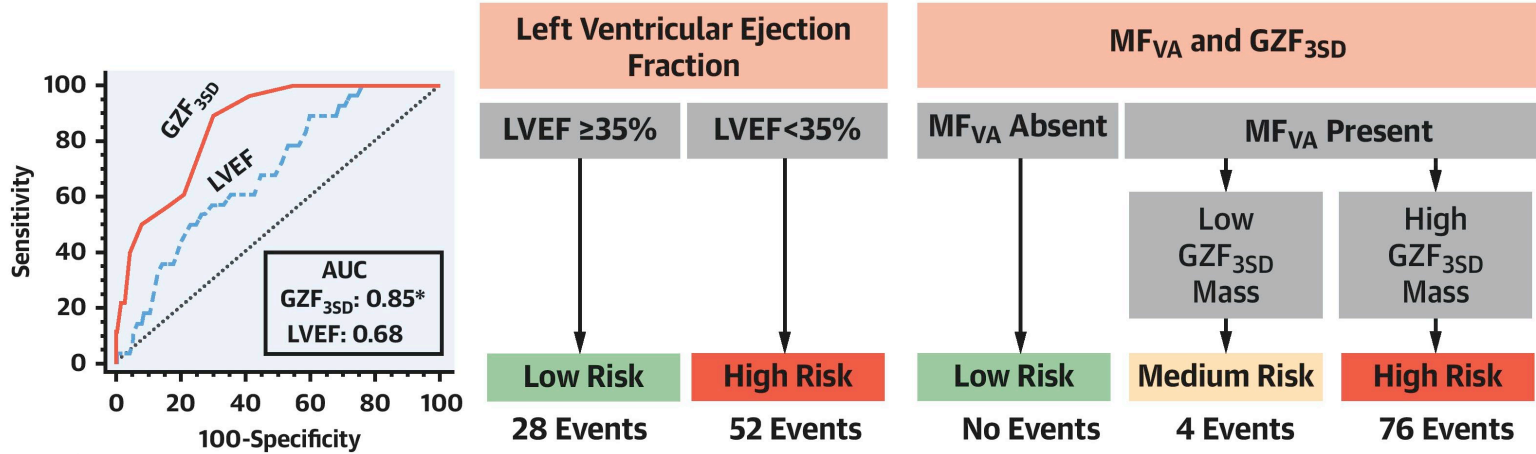
**100%
myocardial
fibrosis**

Ischemic cardiomyopathy



Ischemic cardiomyopathy

2. VA in Asymptomatic HF



Abbasin Zegard et al. JACC 2021; 77:29-41.

Non-ischemic cardiomyopathy



¿?%
**myocardial
fibrosis**

Non-ischemic cardiomyopathy

Sudden death

Table 1 Incidence of sudden death in NICM

Study	Year of publication	Follow-up	SCD	SCD/year in placebo/control group
Von Olshausen <i>et al</i> ²⁸	1984	1 y	7%	7%
Meinertz <i>et al</i> ²⁹	1984	11 m	16%	17.5 %
Costanzo-Nordin <i>et al</i> ³⁰	1985	14.1 m	20%	17%
Stewart <i>et al</i> ³¹	1990	24m	18%	9%
CAT ³²	2002	1 y	0	0 (1st year)
AMIOVIRT ³³	2003	1.8 y	3.80%	2.1%
DEFINITE ³⁴	2004	29m	6%	2.5%
DANISH ³⁵	2017	67.6 m	8.2%	1.46%

AMIOVIRT, Amiodarone vs Implantable Cardioverter-Defibrillator; CAT, Cardiomyopathy Trial; DANISH, Danish Study to Assess the Efficacy of ICDs in Patients With Non-Ischaemic Systolic Heart Failure on Mortality; DEFINITE, Defibrillators in Non-Ischaemic Cardiomyopathy Treatment Evaluation; m, month; NICM, non- ischaemic dilated cardiomyopathy; SCD, Sudden Cardiac Death; y, year.

Cannatà A, et al. Heart 2020;106:656–664.

Non-ischemic cardiomyopathy

EP study

Table 3 Role of electrophysiological studies on the arrhythmic stratification of patients with NICM

Study	Population	N	Definition of positive EPS	Percentage of positive EPS	Result
Meinertz <i>et al</i> , <i>Am J Cardiol</i> , 1985 ²⁹	Non-ischaeamic DCM	42	Electrically-induced VT or VF	7.1%	Positive EPS predictive of SCD*. ✓
Poll <i>et al</i> , <i>Am J Cardiol</i> , 1986 ⁴²	Non-ischaeamic DCM and EF≤45% with major ventricular arrhythmias	47	Electrically-induced sustained VT or VF	53%	Positive EPS not predictive of SCD*. ✗
Brembilla-Perrot <i>et al</i> , <i>Am Heart J</i> , 1991 ⁴³	Non-ischaeamic DCM and EF≤45%	103	Electrically-induced sustained VT or VF	11%	Positive EPS predictive of SCD ✓
Turitto G <i>et al</i> , <i>J Am Coll Cardiol</i> , 1994 ⁴⁴	Non-ischaeamic DCM and EF≤50% without major ventricular arrhythmias	80	Electrically-induced sustained VT or VF	22%	Positive EPS not predictive of MVA ✗
Rankovic V <i>et al</i> , <i>Am J Cardiol</i> , 2002 ⁴⁵	Non-ischaeamic DCM (ICD carriers)	42	Electrically-induced sustained VT or VF	50%	Positive EPS not predictive of MVA ✗
Becker <i>et al</i> , <i>Basic Res Cardiol</i> , 2003 ⁴⁶	Non-ischaeamic DCM and EF ≤50% with documented NSVT	99	Electrically-induced sustained VT or VF	7%	Positive EPS not predictive of MVA ✗
Morgera <i>et al</i> , <i>Ital Heart J</i> , 2004 ⁴⁷	Non-ischaeamic DCM without major ventricular arrhythmias	78	Electrically-induced VT or VF	21%	Positive EPS not predictive of MVA ✗
Daubert <i>et al</i> , <i>Pacing Clin Electrophysiol</i> , 2009 ⁴⁸	Non-ischaeamic DCM (ICD carriers)	204	Electrically-induced sustained VT or VF	14%	Positive EPS predictive of MVA ✓
Gatzoulis <i>et al</i> , <i>Circ Arrhythmia Electrophysiol</i> , 2013 ⁴⁹	Non-ischaeamic DCM	158	Electrically-induced sustained VT or VF	27,8%	Positive EPS predictive of appropriate ICD therapy ✓
Stabile, <i>et al</i> , <i>Hellenic J Cardiol</i> , 2015 ⁵⁰	HFrEF (ICD carriers)	206*	Electrically-induced sustained VT or VF	14%	Positive EP-study not predictive of mortality ✗

Cannatà A, et al. *Heart* 2020;106:656–664.

Non-ischemic cardiomyopathy

Genetic test

Table 2 Role of genetic studies on the arrhythmic risk stratification

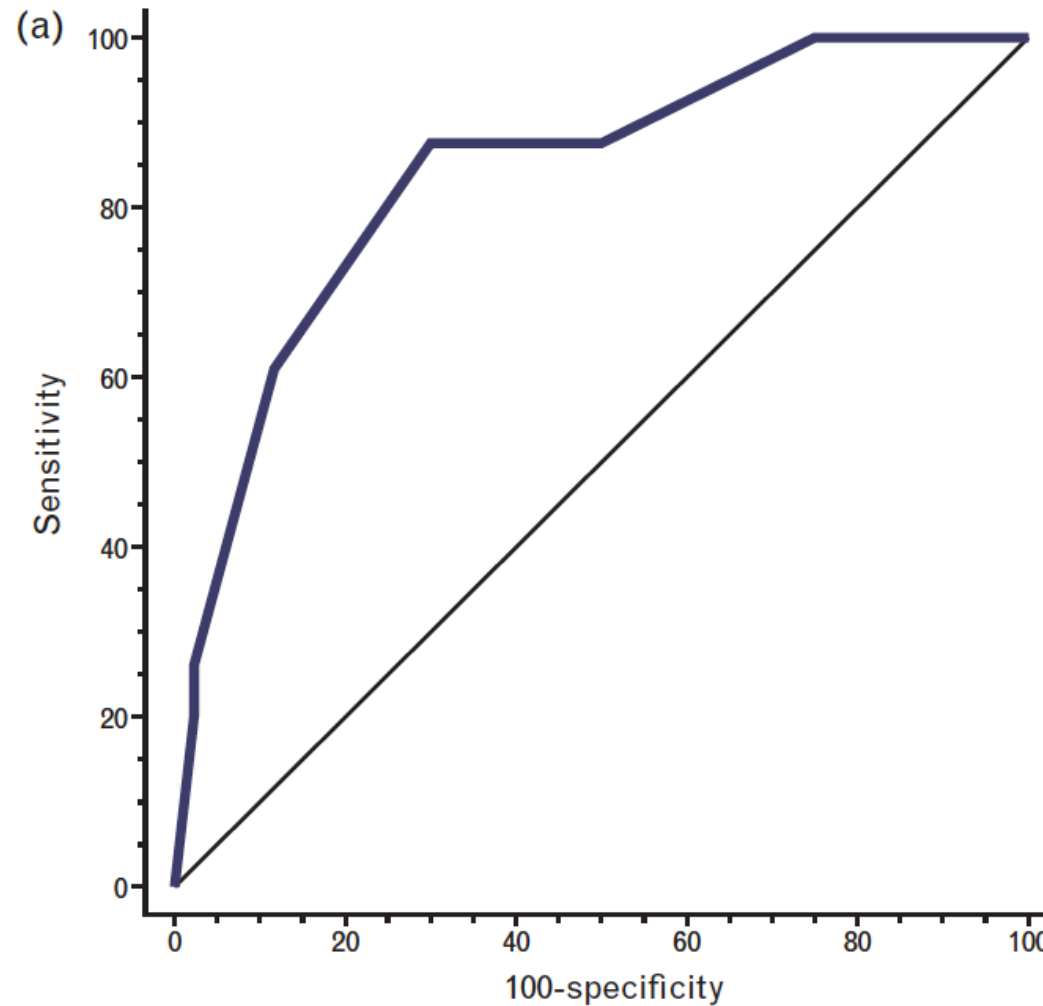
Study	Population	Genes tested	N	Percentage of mutations	Percentage of arrhythmic events
Fatkin <i>et al</i> , <i>N Eng J Med</i> , 1999 ³⁶	DCM and conduction defects (patients and healthy relatives)	LMNA	11 families	NS	44% SCD in patients
Van Rijsingen <i>et al</i> , <i>J Am Coll Cardiol</i> , 2012 ¹⁶	LMNA mutation carriers.	LMNA	269	100%	18% MVA
Van der Zwaag <i>et al</i> , <i>Eur J Heart Fail</i> , 2012 ³⁷	Patients with ARVC and DCM	PLN	354 (257 DCM, 97 ARVC)	11%	46% MVA
Ortiz-Genga <i>et al</i> , <i>J Am Coll Cardiol</i> , 2016 ¹⁷	Inherited cardiovascular disease	FLNC	2177	3,9%	18% MVA 16% SCD
Kumar <i>et al</i> , <i>J Am Coll Cardiol</i> , 2016 ³⁸	LMNA mutation carriers	LMNA	122	100%	19% sustained VT 8% VF
Nishiuchi <i>et al</i> , <i>Circ Cardiovasc Genet</i> , 2017 ³⁹	LMNA mutation carriers	LMNA	77	100%	26% MVA
Hasselberg <i>et al</i> , <i>Eur Heart J</i> , 2018 ⁴⁰	LMNA mutation carriers	LMNA	79	100%	18% MVA
Begay <i>et al</i> , <i>J Am Coll Cardiol</i> , 2018 ⁴¹	DCM (patients and healthy relatives)	FLNC	319	4%	85% MVA or SCD
Verdonschot <i>et al</i> , <i>Eur Heart J</i> , 2018 ²⁰	Patients with DCM	TTNtv	303	13%	24% MVA
Parikh <i>et al</i> , <i>Circ Heart Fail</i> , 2019 ¹⁸	RBM20 mutation carriers	RBM20	74	100%	43% MVA
Gigli <i>et al</i> , <i>J Am Coll Cardiol</i> , 2019 ¹⁹	Patients with DCM	23 genes	487	37%	20% MVA or SCD

Cannatà A, et al. *Heart* 2020;106:656–664.

Non-ischemic cardiomyopathy

score ≥ 3 S 87.5% E 69.8%

- ✓ Hypertension
- ✓ Diabetes
- ✓ chronic renal failure
- ✓ atrial fibrillation
- ✓ COPD
- ✓ NYHA \geq III



AUC = 0.84

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Non-ischemic cardiomyopathy

ARRHYTHMIC PREDICTORS IN DCM	
CLINICAL	<ul style="list-style-type: none"> ■ Etiological characterization ■ Personal and family history (syncope, familial history of SCD)
ECG and Holter ECG	<ul style="list-style-type: none"> ■ Fragmented QRS ■ Long QRS ■ Low QRS amplitude ■ Anterolateral T wave inversion ■ NSVT (≥ 5 beats, ≥ 150 bpm) ■ Frequent PVCs (1000/24h) and couplets ($\geq 50/24h$)
ECHOCARDIOGRAPHY	<ul style="list-style-type: none"> ■ LVEF $\leq 35\%$ ■ Reduced GLS
GENETIC TESTING	<ul style="list-style-type: none"> ■ LMNA ■ FLNC, PLN, RBM20 ■ SCN5A ■ TTN *
ELECTROPHYSIOLOGICAL PARAMETERS	<ul style="list-style-type: none"> ■ Abnormal mTWA test

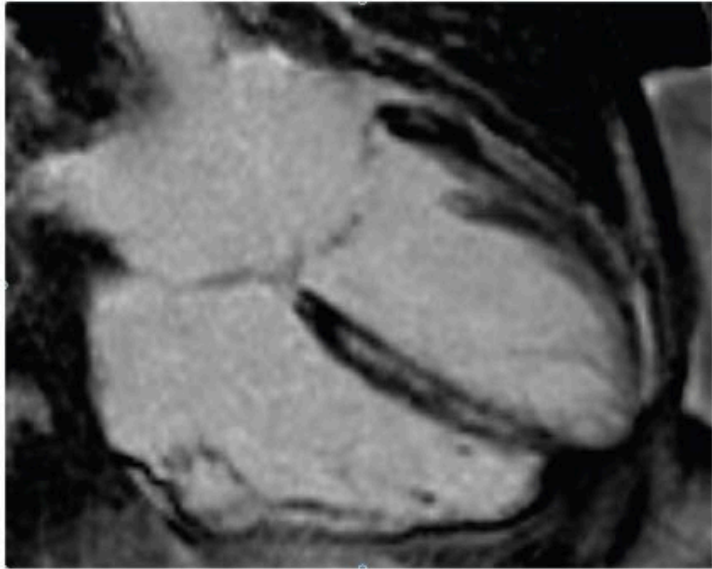
Strong evidence



Weak evidence

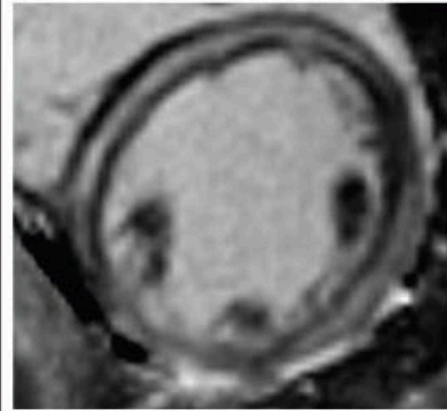
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Non-ischemic cardiomyopathy



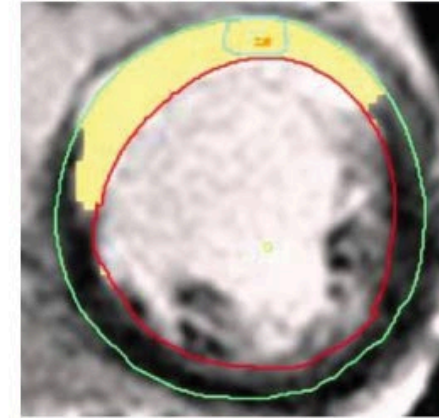
PRESENCE / ABSENCE

Strong association with worse outcomes



LOCALIZATION

Concomitant presence of septal and free wall LGE associated with worse outcome. In the future LGE localization might provide further prognostic impact



EXTENSION

Quantitative assessment might provide further prognostic impact in the future

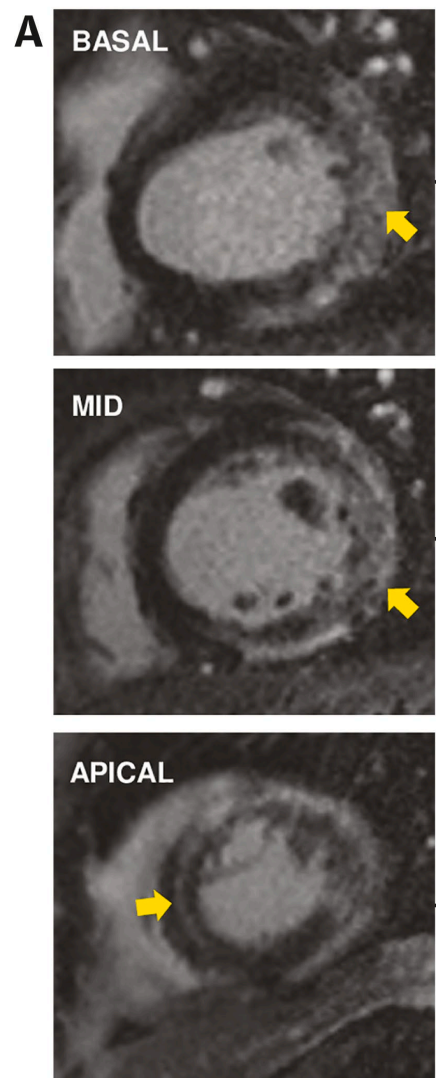
Strong evidences in large studies and metaanalyses

Conflicting evidences from smaller studies

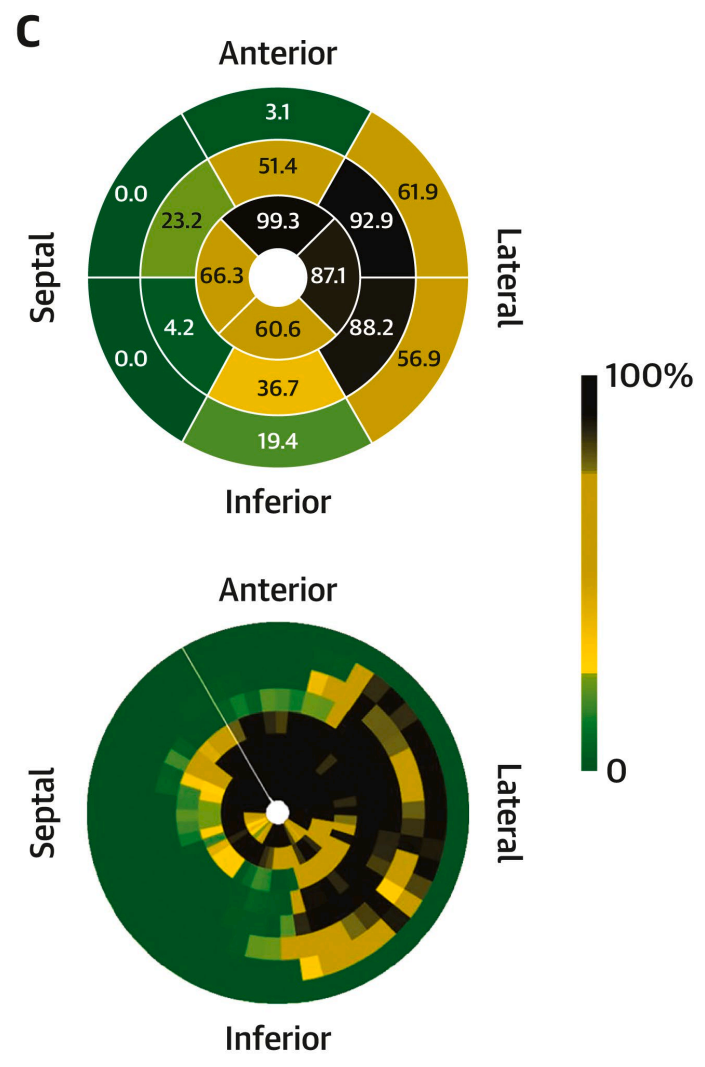
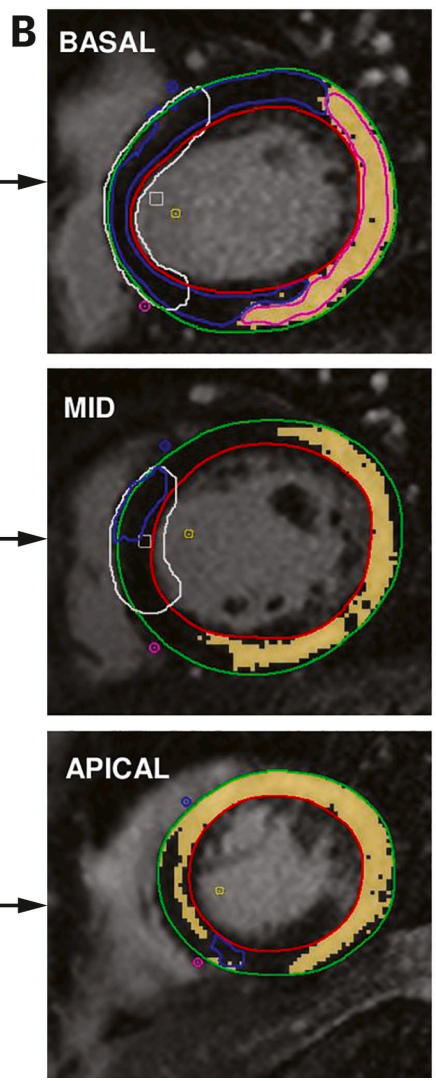
Cannatà A, et al. Heart 2020;106:656–664.

Non-ischemic cardiomyopathy

Visual Assessment

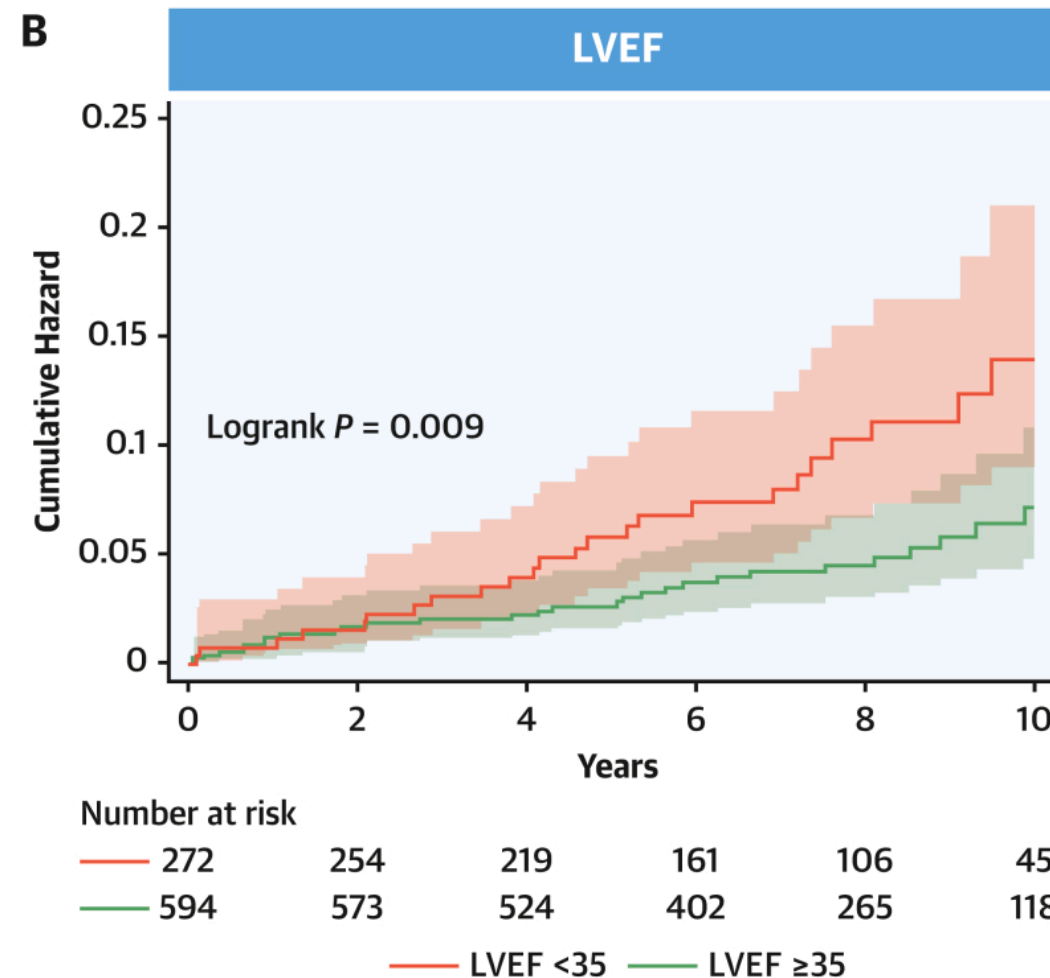
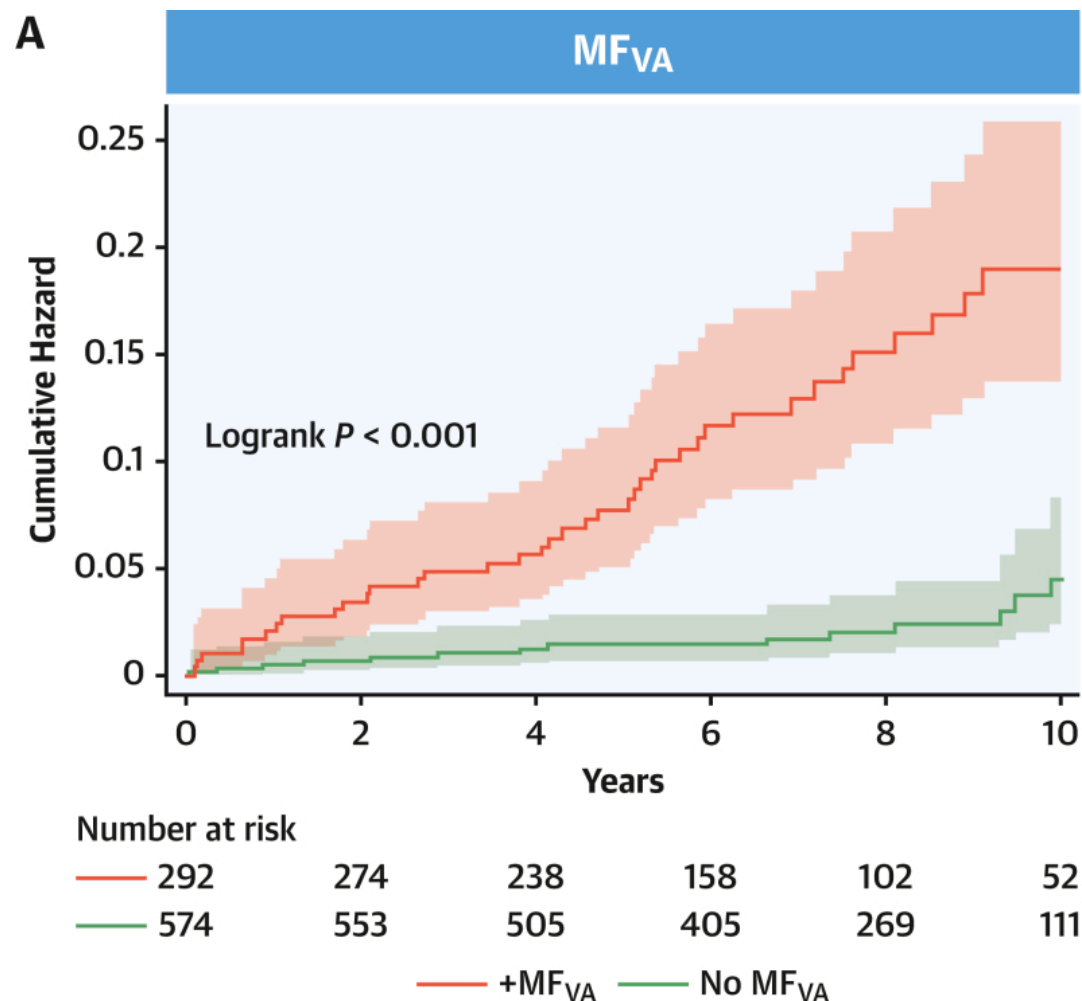


Quantification



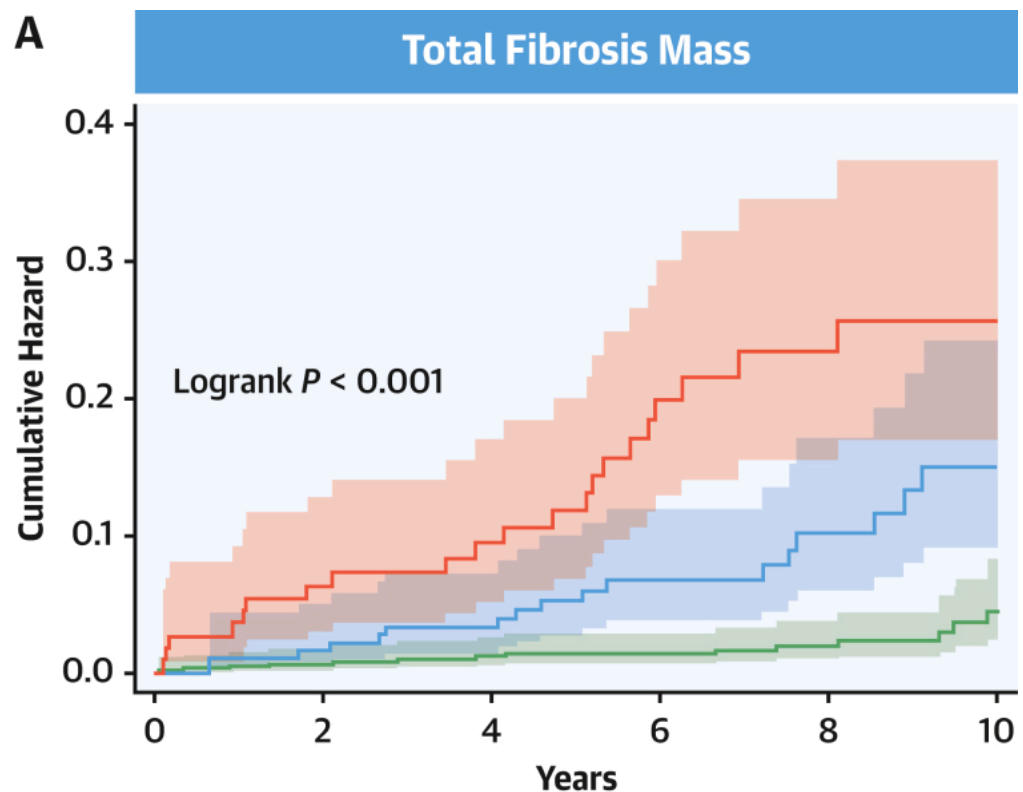
Hammersley et al JACC VOL. 84, NO. 15, 2024CMR and Arrhythmias in NICM OCTOBER 8, 2024

Non-ischemic cardiomyopathy



Hammersley et al JACC VOL. 84, NO. 15, 2024CMR and Arrhythmias in NICM OCTOBER 8, 2024

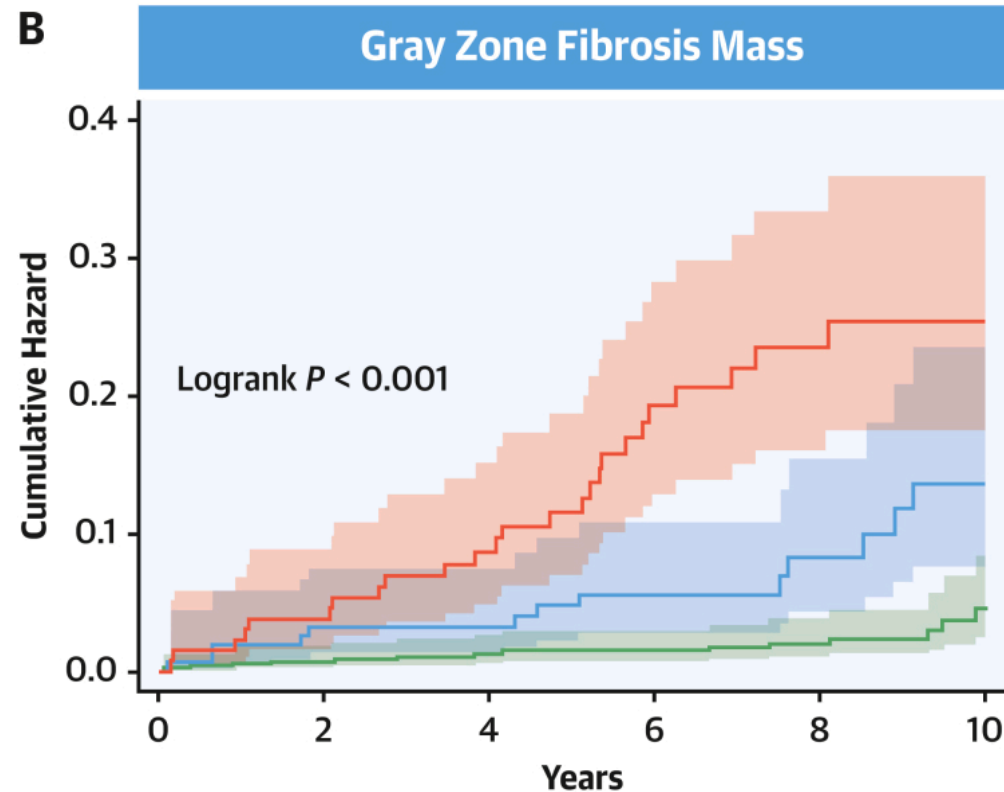
Non-ischemic cardiomyopathy



Number at risk

— 112	99	81	54	35	15
— 180	175	157	104	67	37
— 574	553	505	405	269	111

— High TF Mass — Low TF Mass — No MF_{VA}



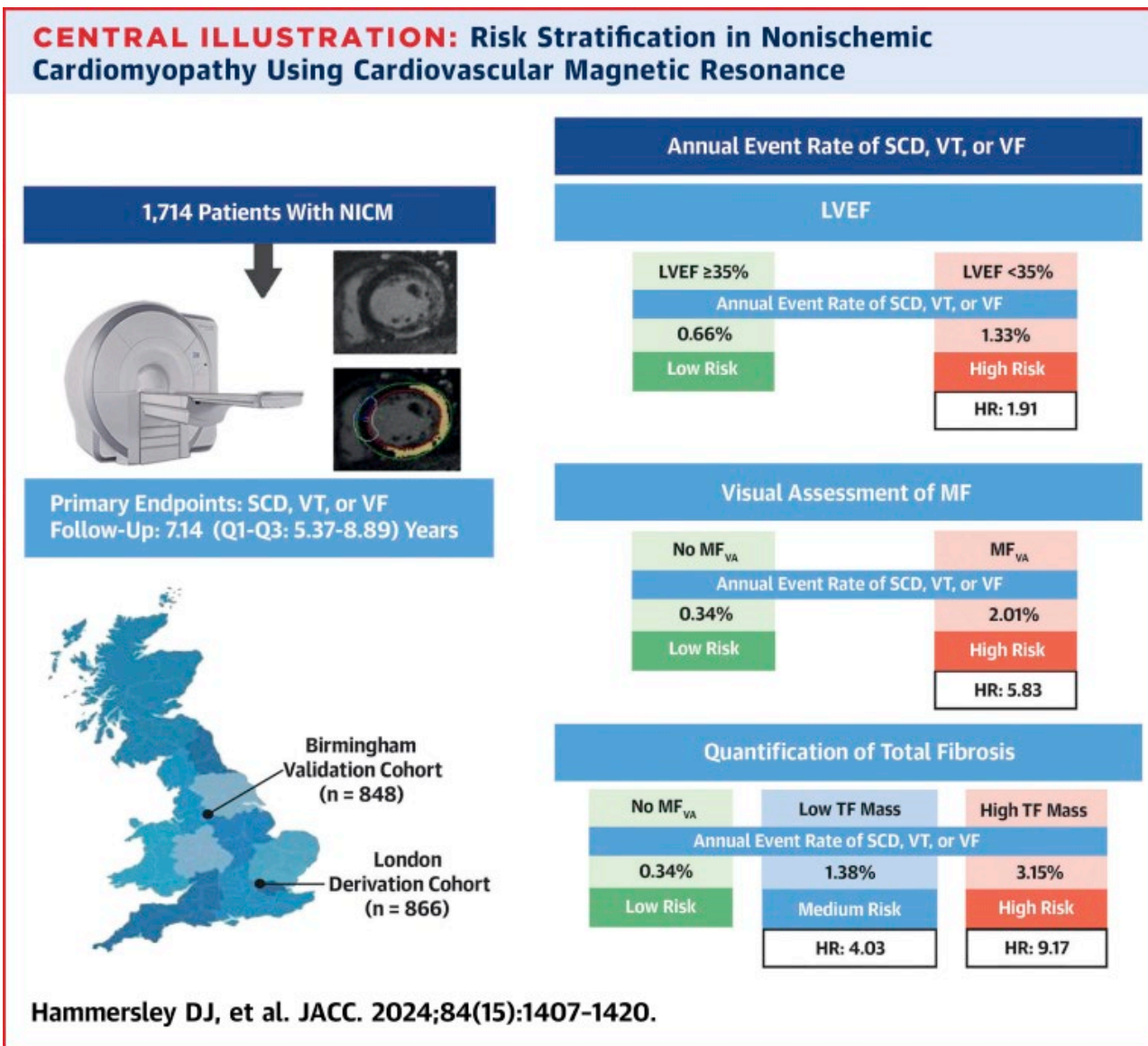
Number at risk

— 135	125	101	67	40	18
— 157	149	137	91	62	34
— 574	553	505	405	269	111

— High GZF Mass — Low GZF Mass — No MF_{VA}

Hammersley et al JACC VOL. 84, NO. 15, 2024CMR and Arrhythmias in NICM OCTOBER 8, 2024

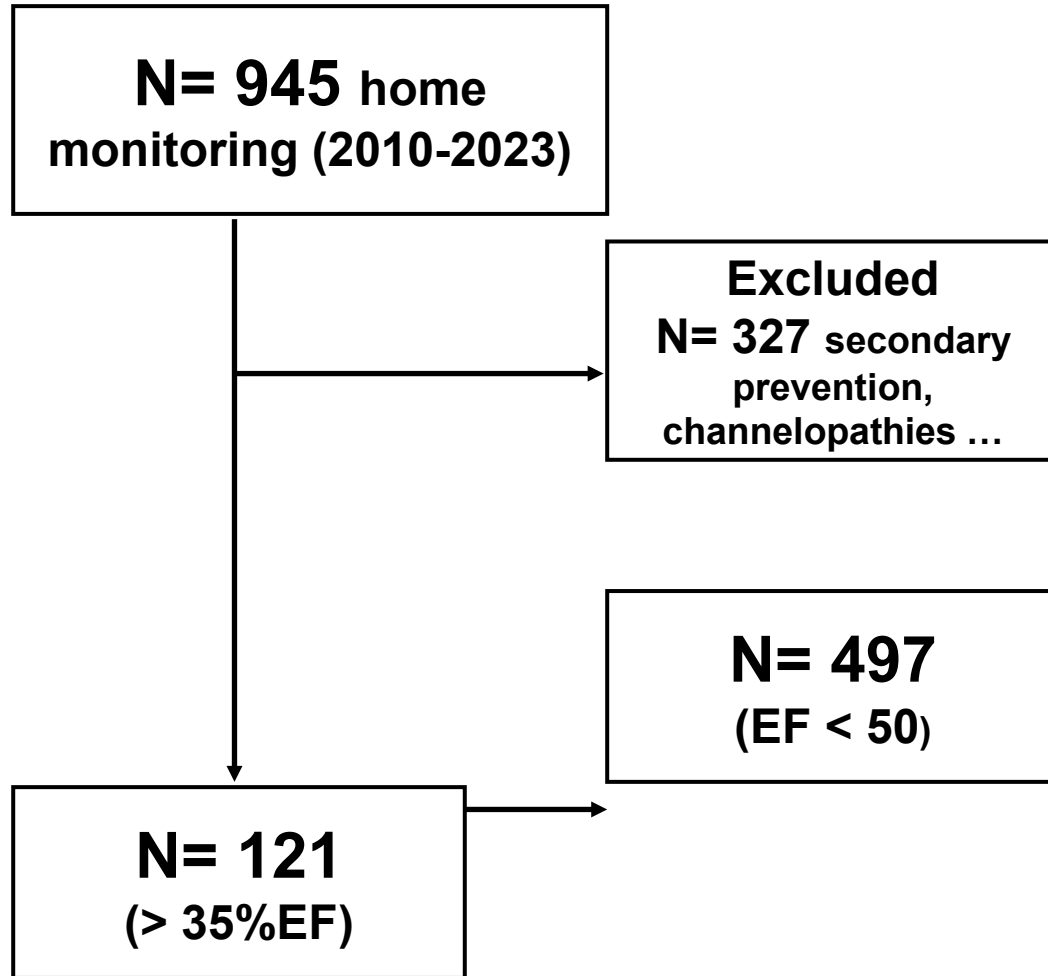
Non-ischemic cardiomyopathy



Hammersley et al JACC VOL. 84 , NO . 15 , 2024CMR and Arrhythmias in NICM OCTOBER 8, 2024

VT in recovered EF

Centro referencia: Unidad de Arritmias Hospital Universitario de A Coruña, Galicia. España.
Investigador principal: Jorge Rodríguez Garrido, MD.



6 (4.95%) VA
Follow up: 4 y



100% myocardial fibrosis
4 (66%) ischemic

Summary



- Different ways ventricular arrhythmias develop in patients with acute heart failure.
- Myocardial fibrosis can predict ventricular events and is especially useful for ruling them out.
- However, it is not included in ESC or AHA/ACC guidelines because:
 - Lack of robust prospective validation
 - No standardized measurement methods
 - No trials proving clinical benefit from fibrosis-guided decisions