



**UTIC** | **ANMCO**  
**CLUB** |  
CRITICAL CARE COMMUNITY

# Miocardite acuta: dal sospetto al supporto avanzato in poche ore

*Dott.C.Raineri*

*Divisione di Cardiologia*

*Città della Salute e della Scienza- Ospedale Molinette*

**No Disclosure**

# Clinical Case



**35 years old, male**

**CV risk factors:** none

**Medical history:** multiple sclerosis well controlled on fourth-line therapy with ofatumumab

## Clinical Presentation

Night of 02 Jan 2026: sudden awakening with paroxysmal dyspnea. No prodromal symptoms.

## Management

- Pharmacological CV with amiodarone
- Dobutamine 5  $\mu\text{g}/\text{kg}/\text{min}$  and norepinephrine 0.05  $\mu\text{g}/\text{kg}/\text{min}$  were initiated.
- Coronary arteries without significant disease
- IABP was inserted

## In the Emergency Department

Agitated, confused, cold clammy skin, dyspneic

BP 95/75 mmHg, T 36.8 °C, So2 92%

ECG: Atrial Fibrillation 140 bpm

## Laboratory tests:

WBC 18,000 CRP and PCT negative Creatinine 1.3 mg/dL

AST/ALT 58/59 U/L, Troponin I 1593, Lactate 5 mmol/L

**Echo fast:** DT 42 mm, LVEF 25%, Normal RV function, Mild MR

**“Cardiogenic shock in the setting of suspected fulminant myocarditis»**

**Transfer the patient to our Centre**



# Fulminant myocarditis

**Fulminant myocarditis** is a rare (3% to 9% of adult patients admitted with acute myocarditis), rapidly progressive inflammatory myocardial disease characterized by:

- Abrupt onset of **severe myocardial dysfunction**
- Rapid evolution to **cardiogenic shock**
- Arrhythmias or sudden cardiac death

High short-term mortality without prompt intervention

## **ECG findings are heterogeneous and dynamic:**

- Sinus tachycardia
- Atrial or ventricular arrhythmias
- ST-segment elevation or depression
- Advanced atrio-ventricular block

## **Typical echocardiographic features:**

- **Severe (bi)ventricular systolic dysfunction**
- **Non-dilated ventricles**
- Increased wall thickness (myocardial edema)
- Reduced stroke volume and cardiac output

# Fulminant Myocarditis

*Eur Heart J* 2025; 00, 1–15

Overall study population: 271 cases of fulminant myocarditis



	LM 👤 99	EM 👤 18	NEM 👤 3	GCM 👤 26	Inconclusive EMB 👤 48	No EMB 👤 77
Chest pain, n (%)	29 (29)	5 (28)	1 (33)	11 (42)	9 (19)	24 (31)
Dyspnoea, n (%)	67 (68)	11 (61)	1 (33)	16 (62)	32 (67)	50 (65)
Cardiac arrest, n (%)	13 (13)	2 (11)	0 (0)	4 (15)	12 (25)	20 (26)
Sustained VA, n (%)	10 (10)	2 (11)	0 (0)	5 (19)	2 (4)	16 (21)
LVEF, median [IQR]	17.5 [15.0, 30.0]	23.5 [15.0, 28.8]	15.0 [15.0, 17.5]	20.5 [15.0, 28.8]	20.0 [14.2, 25.0]	20.0 [15.0, 35.0]
RV dysfunction, n (%)	64 (65)	11 (61)	2 (67)	12 (46)	30 (62)	37 (48)
SAPS II, median [IQR]	39 [20, 56]	45 [27, 51]	38 [37, 60]	45 [23, 65]	43 [26, 56]	46 [28, 68]

# Fulminant Myocarditis

Overall study population: 271 cases of fulminant myocarditis

	LM 👤 99	EM 👤 18	NEM 👤 3	GCM 👤 26	Inconclusive EMB 👤 48	No EMB 👤 77
In-hospital mortality, n (%)	24 (24)	4 (22)	0 (0)	13 (50)	12 (25)	32 (42)
One-year mortality, n (%)	25 (25)	5 (28)	1 (33)	14 (54)	13 (27)	33 (43)



Outcomes

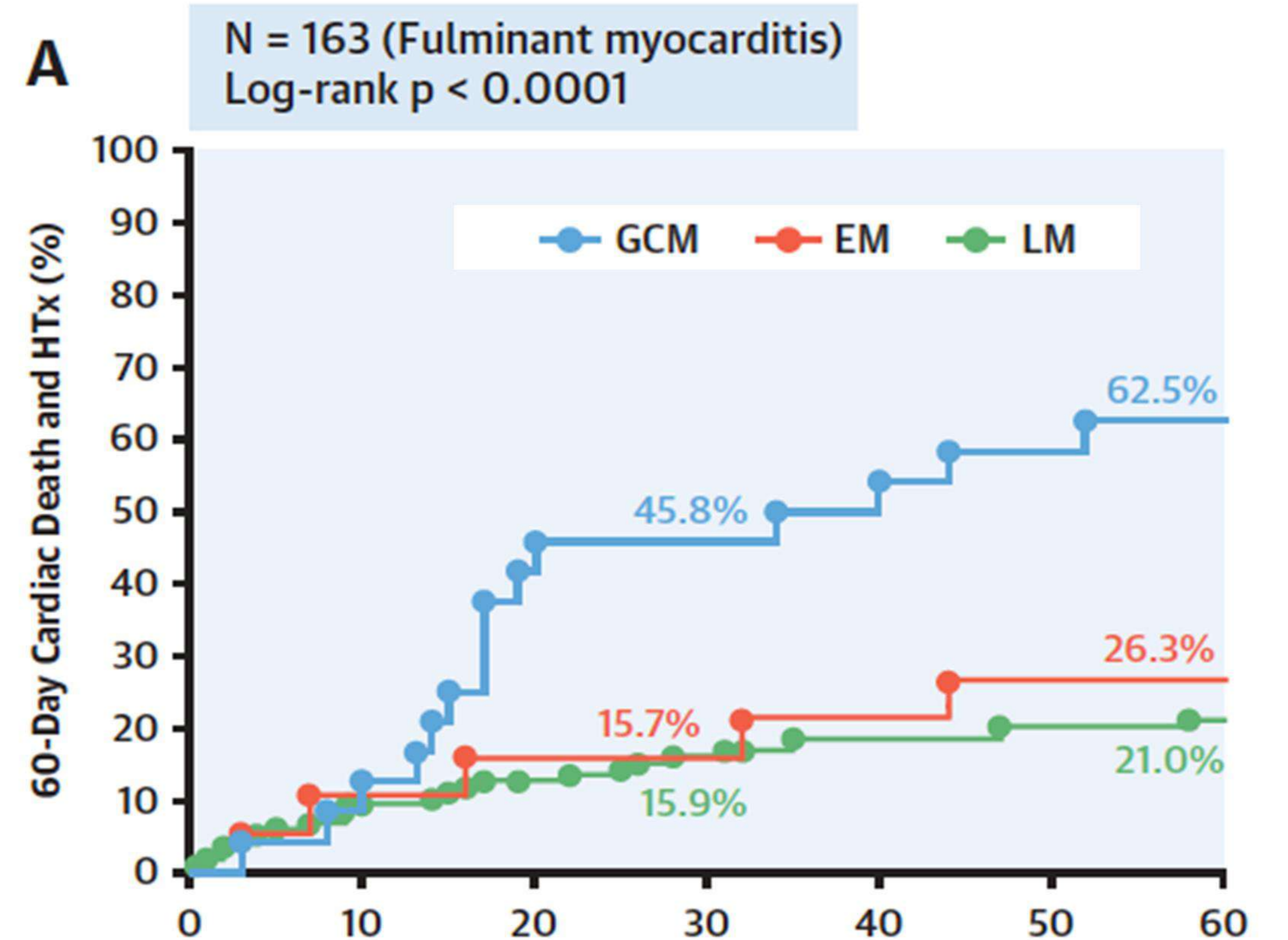
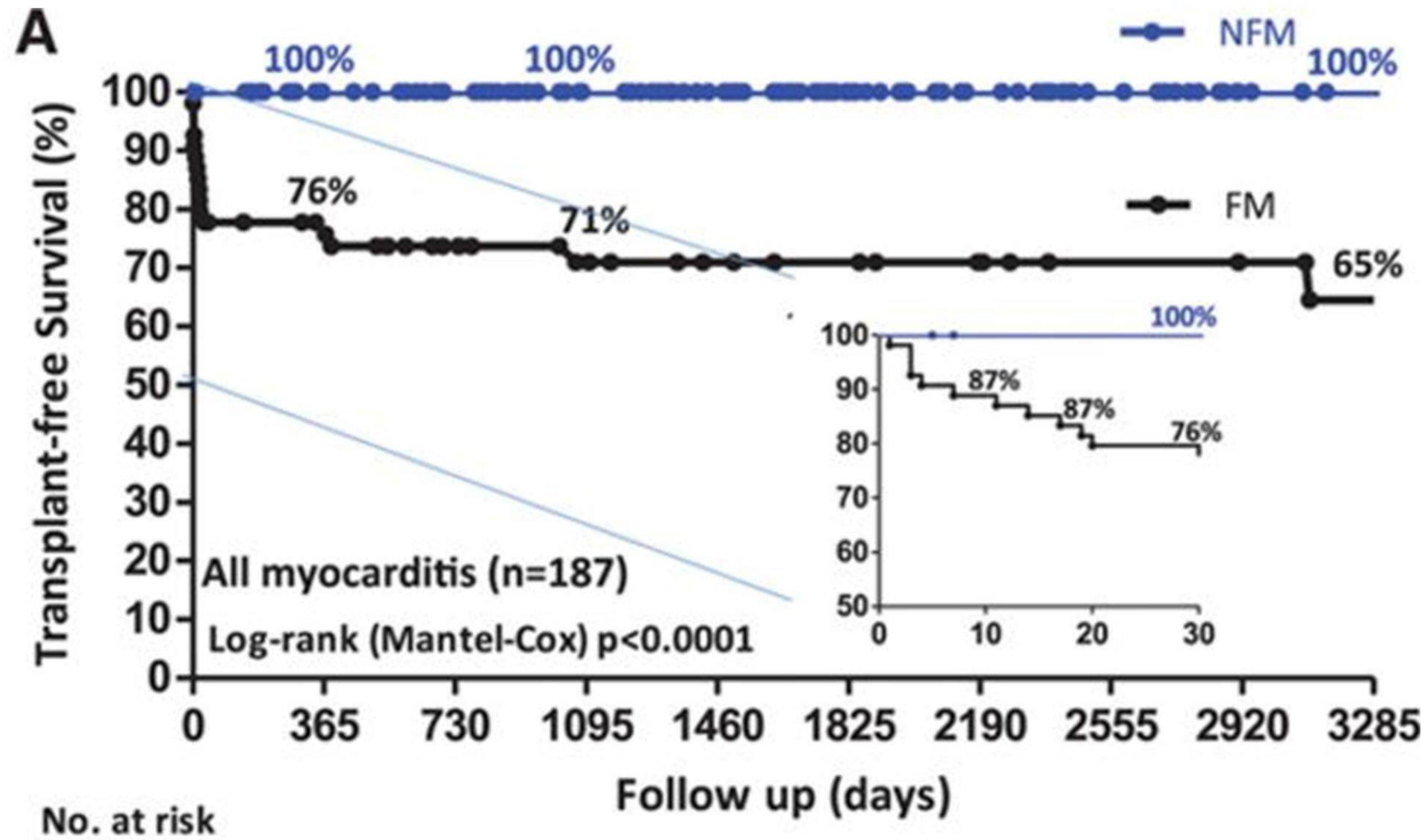
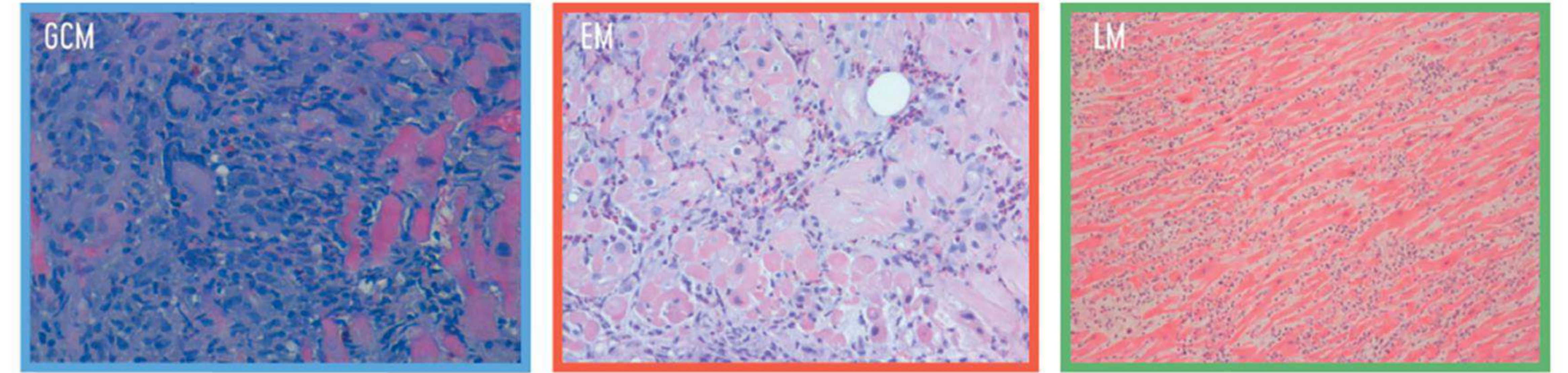
# Challenging diagnosis

- non specific clinical presentation
- potential overlap with other cardiovascular diseases
- general lack of awareness



***Early Suspicion is essential***

# Survival in Fulminant Myocarditis



# ACUTE CLINICAL PRESENTATION

	BP & AHF SYMPTOMS 	LVEF REDUCTION 	VT/VF or AVB 
	Cardiogenic shock (FM)	Severe (<30%)	PRESENT/ ABSENT
	AHF symptoms	Low (30-40%)	PRESENT

**HIGH-RISK**

# INITIAL MANAGEMENT

REFER TO HUB CENTERS 	t-MCS 	EMB 	CMRI 	STEROIDS* 
<input checked="" type="checkbox"/>	<input checked="" type="checkbox"/>	<input checked="" type="checkbox"/>	BEFORE discharge	CONSIDER
<input checked="" type="checkbox"/>	BE prepared	<input checked="" type="checkbox"/>	BEFORE discharge	CONSIDER

	AHF symptoms	Low (30-40%)	ABSENT
	Mild AHF symptoms	Moderate (41-49%)	PRESENT
	Absent	Mild -Normal (>50%)	ABSENT

**INTERMEDIATE RISK**

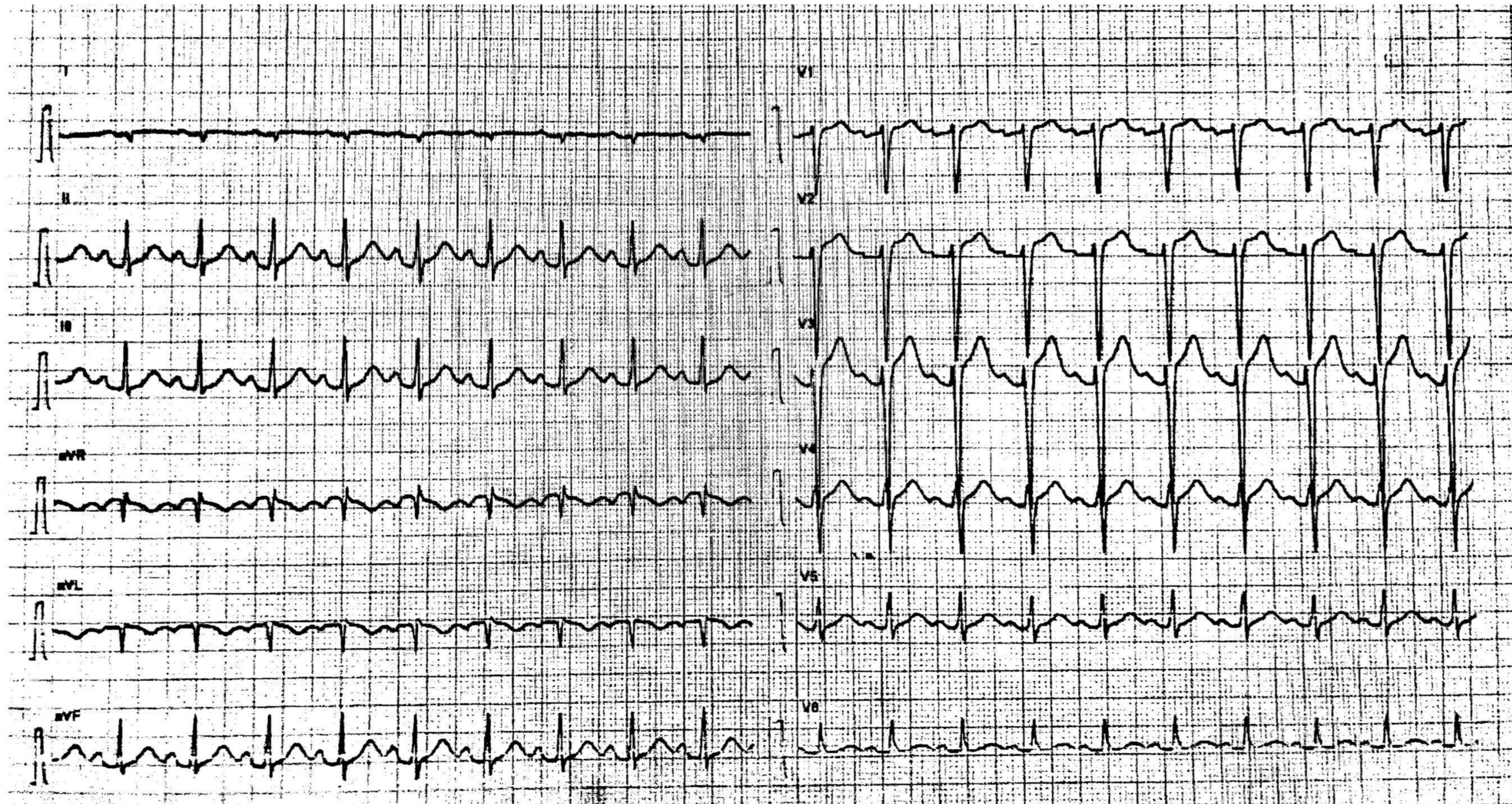
**LOW-RISK**

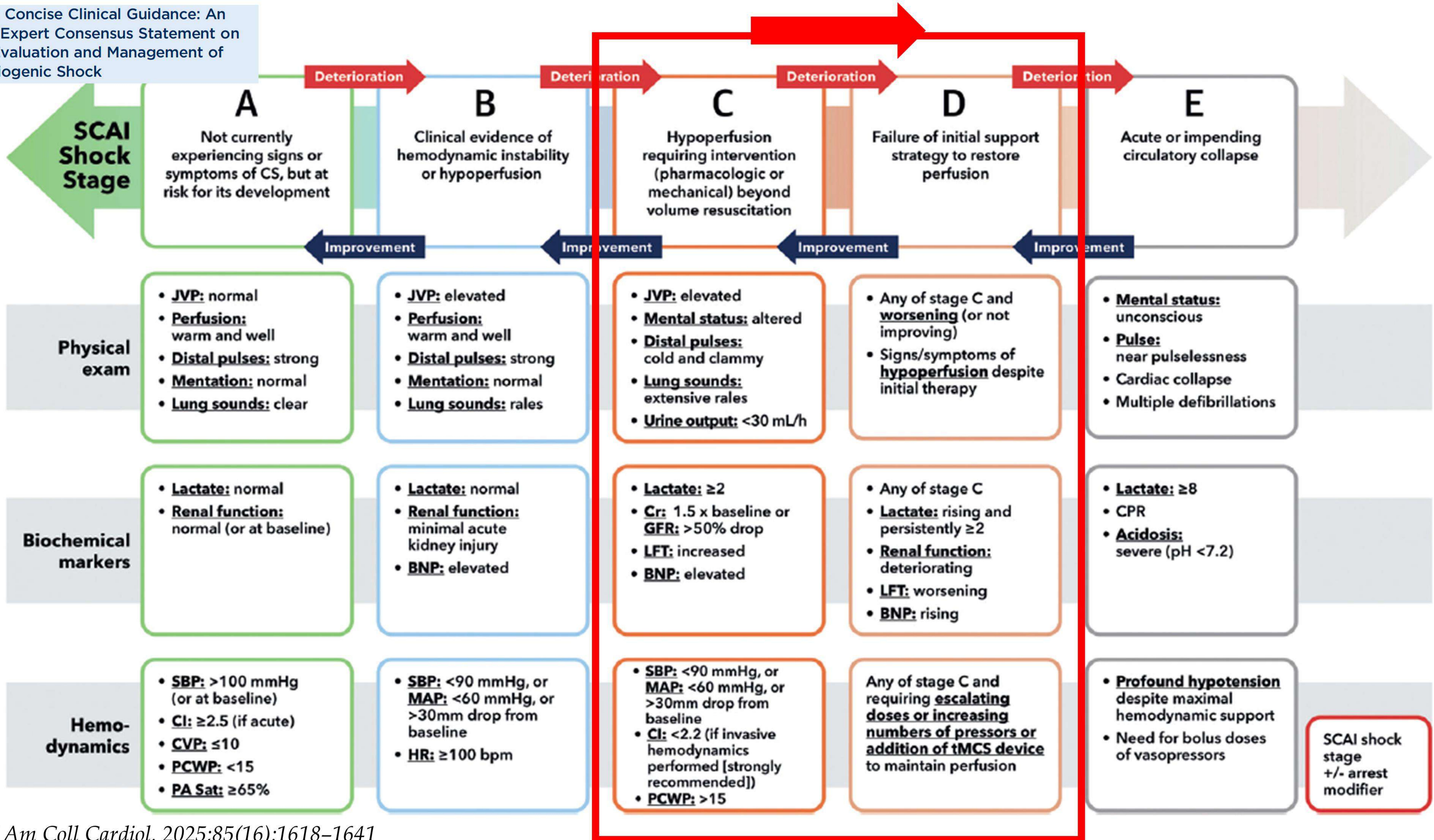
CONSIDER	Rarely needed	CONSIDER	<input checked="" type="checkbox"/>	In specific cases
----------	---------------	----------	-------------------------------------	-------------------

NOT NEEDED	NOT NEEDED	<input type="checkbox"/>	<input checked="" type="checkbox"/>	<input type="checkbox"/>
------------	------------	--------------------------	-------------------------------------	--------------------------

# ....Clinical Case - Initial Management

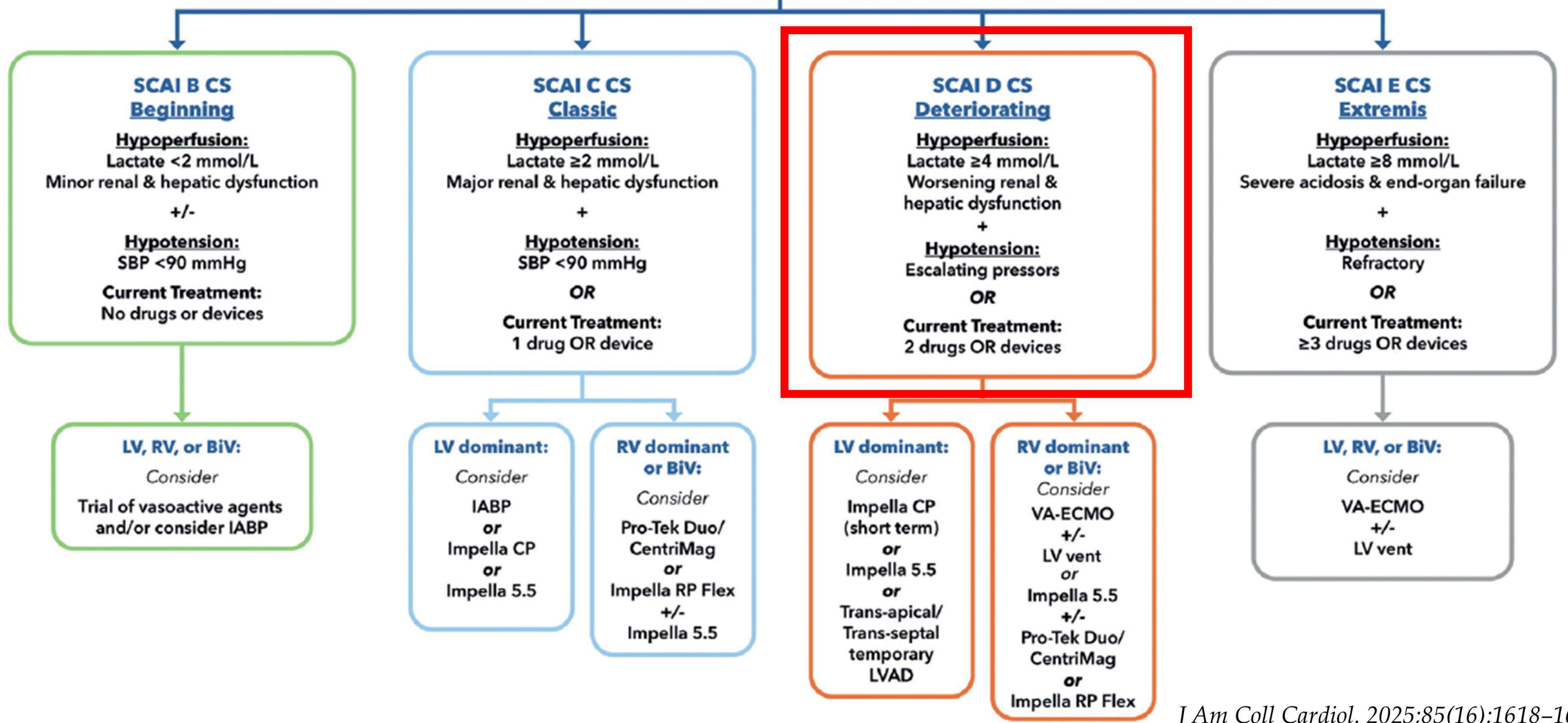
- BP S/D/D<sub>IND</sub> 90/40/105 mmHg
- HR 130 bpm
- Blood gas analysis (FIO<sub>2</sub> 40%): pH 7,33, pO<sub>2</sub> 187, pCO<sub>2</sub> 37, lactate 7,
- LVEF 15%, LVOT VTI 8 cm, TAPSE 16 mm, FAC 32%





**Proposed treatment considerations for HF-CS**

- Shock severity (SCAI stage)
- Shock profile (LV, RV, or BiV)
- Anticipated exit strategy (BTT or BTR)
- Presence of hypoxia
- Presence of arrhythmias
- Anticipated duration of support
- Ability to ambulate
- Contraindications to pMCS



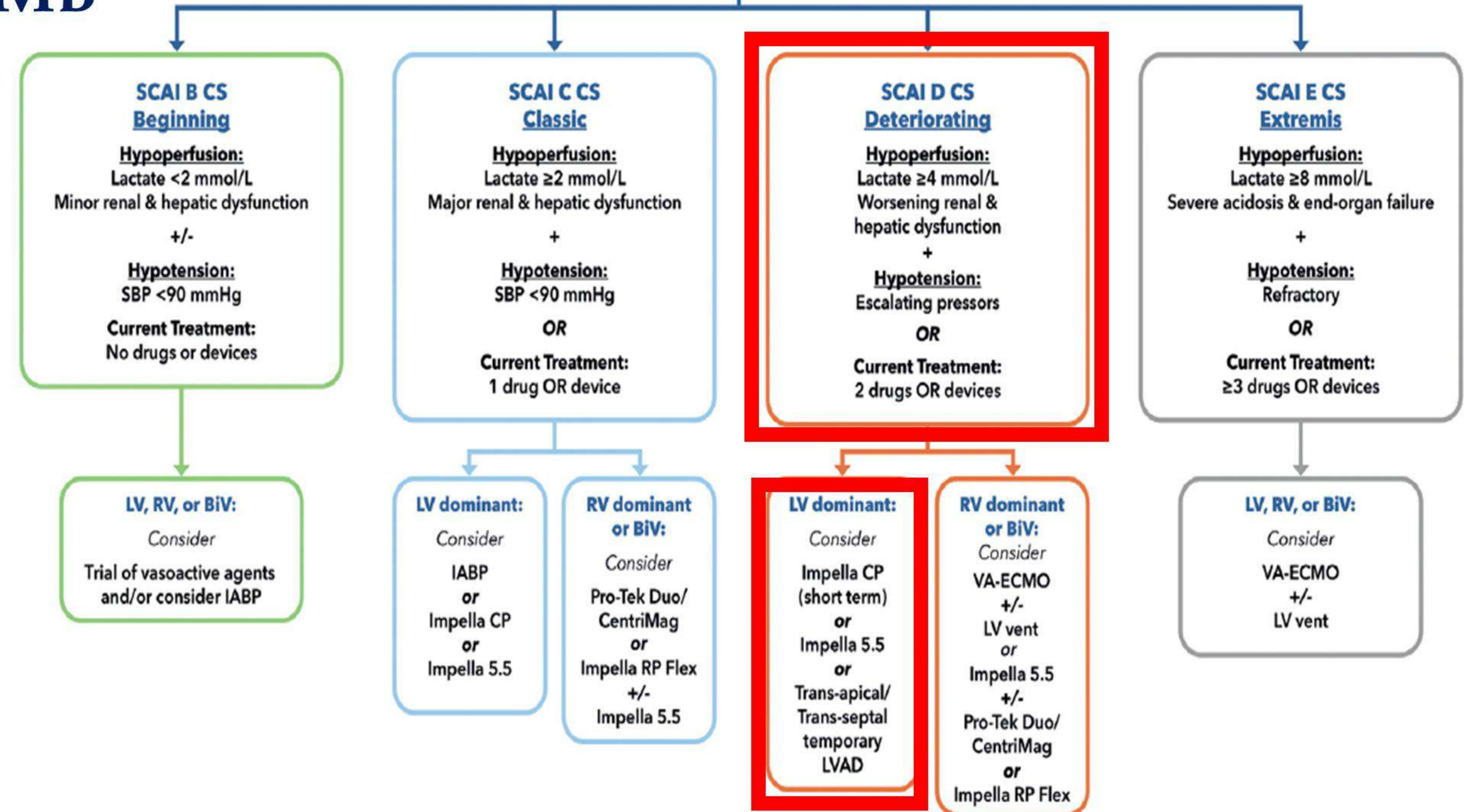
# ...Clinical Case

## Right Heart Right Heart Catheterization and EMB

- PAP 25/13/17 mmHg
- WP 10 mmHg
- RAP 3 mmHg
- CO 3.5 l/min
- CI 2.1 l/min/mq
- PAPI 4
- RAP/WP 0.3

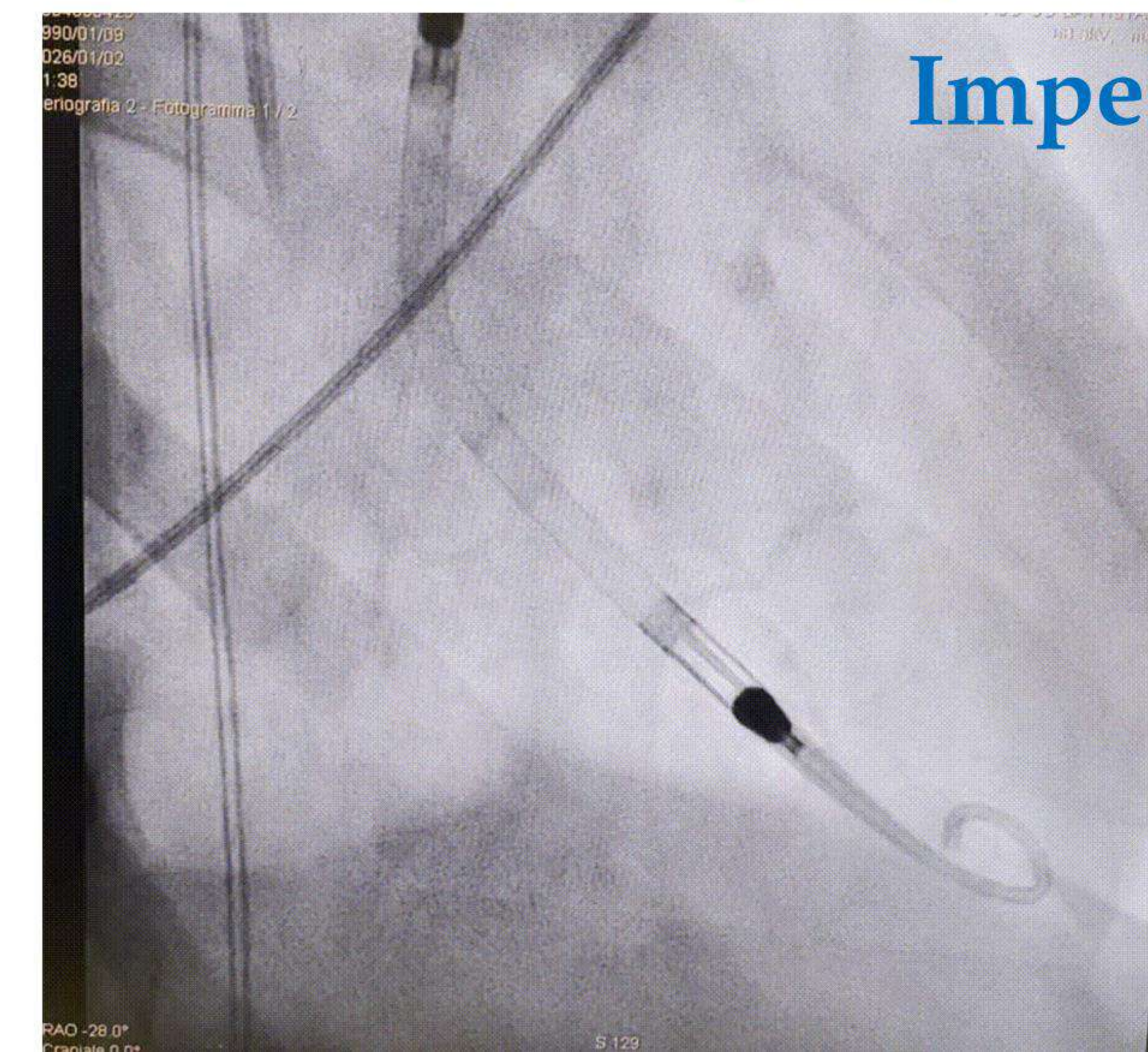
**Proposed treatment considerations for HF-CS**

- Shock severity (SCAI stage)
- Shock profile (LV, RV, or BiV)
- Anticipated exit strategy (BTT or BTR)
- Presence of hypoxia
- Presence of arrhythmias
- Anticipated duration of support
- Ability to ambulate
- Contraindications to pMCS



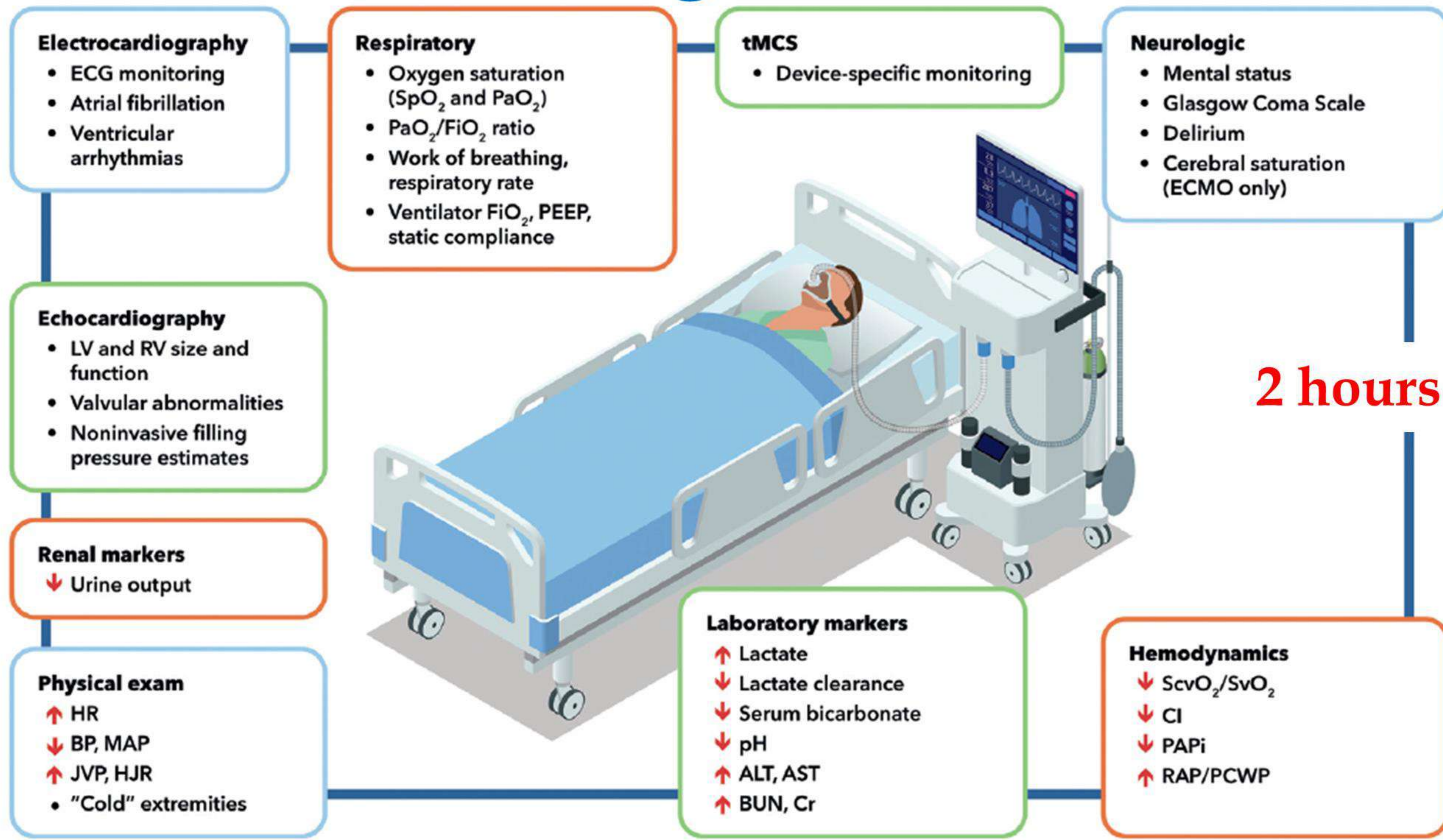
### Medical treatment:

- Epinephrine 0.05 µg/kg/min
- STOP dobutamine and norepinephrine
- Methylprednisolone 1gr ev



Impella CP

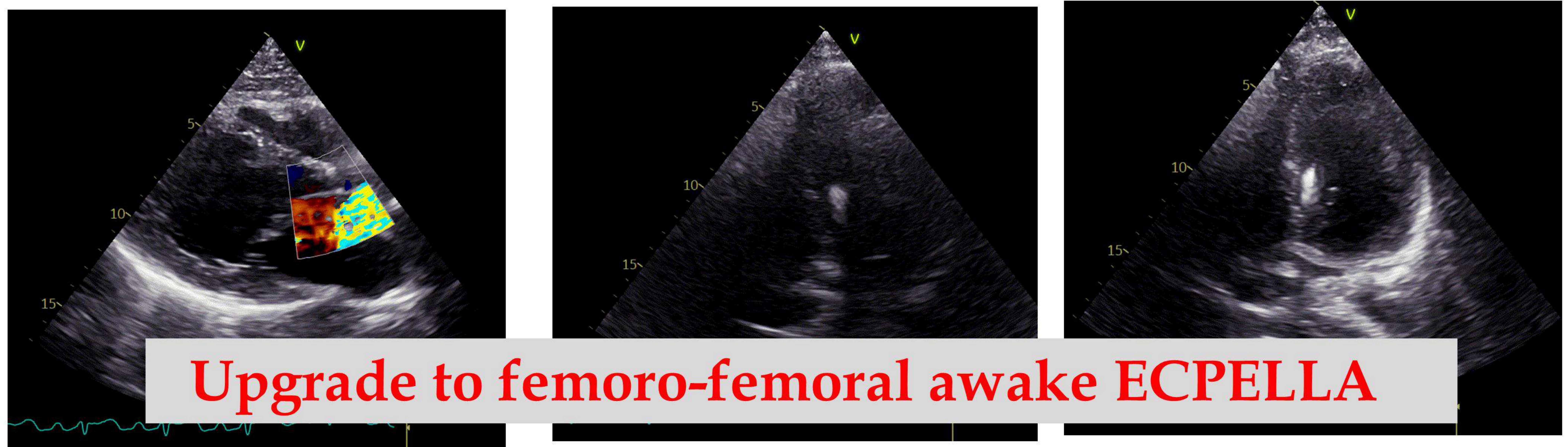
# Monitoring of the CS Patient in the Intensive Care Unit



2 hours after....

- PAP: 20/10/14 mmHg ↓
- PCWP: 6 mmHg ↓
- RAP: 2 mmHg ↓
- CO: 2.6 L/min ↓
- CI: 1.4 L/min/m<sup>2</sup> ↓

- Lactate: 13 mmol/L ↑



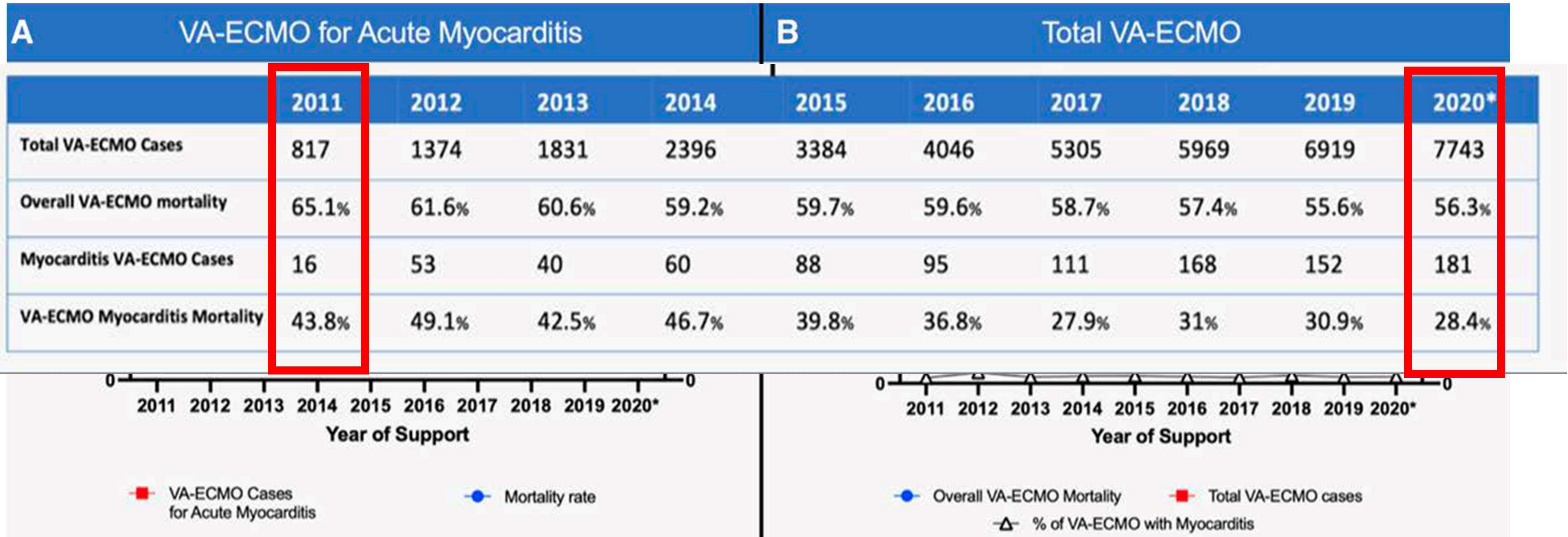
**Upgrade to femoro-femoral awake ECPELLA**

# VA-ECMO for Suspected Acute Myocarditis

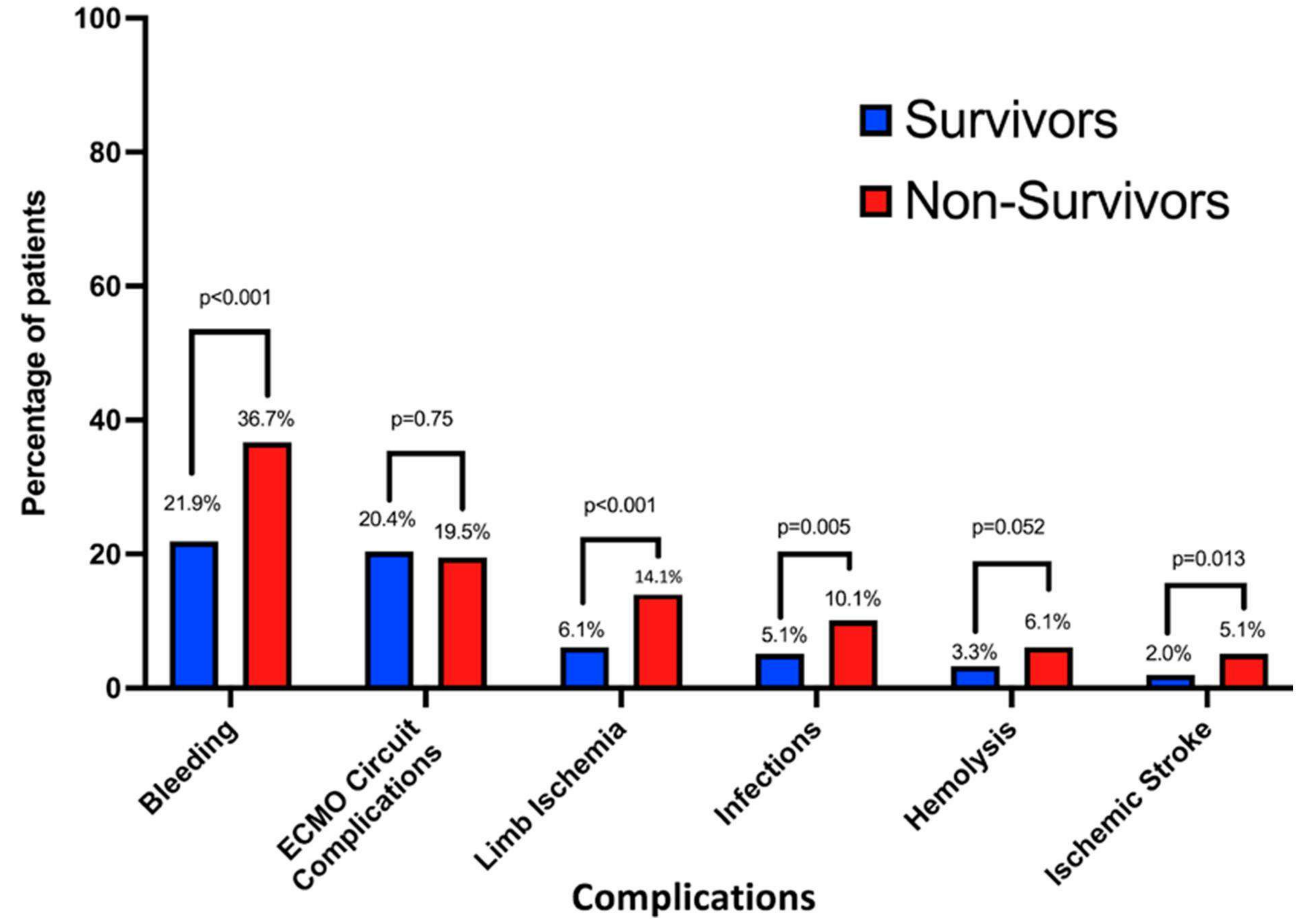
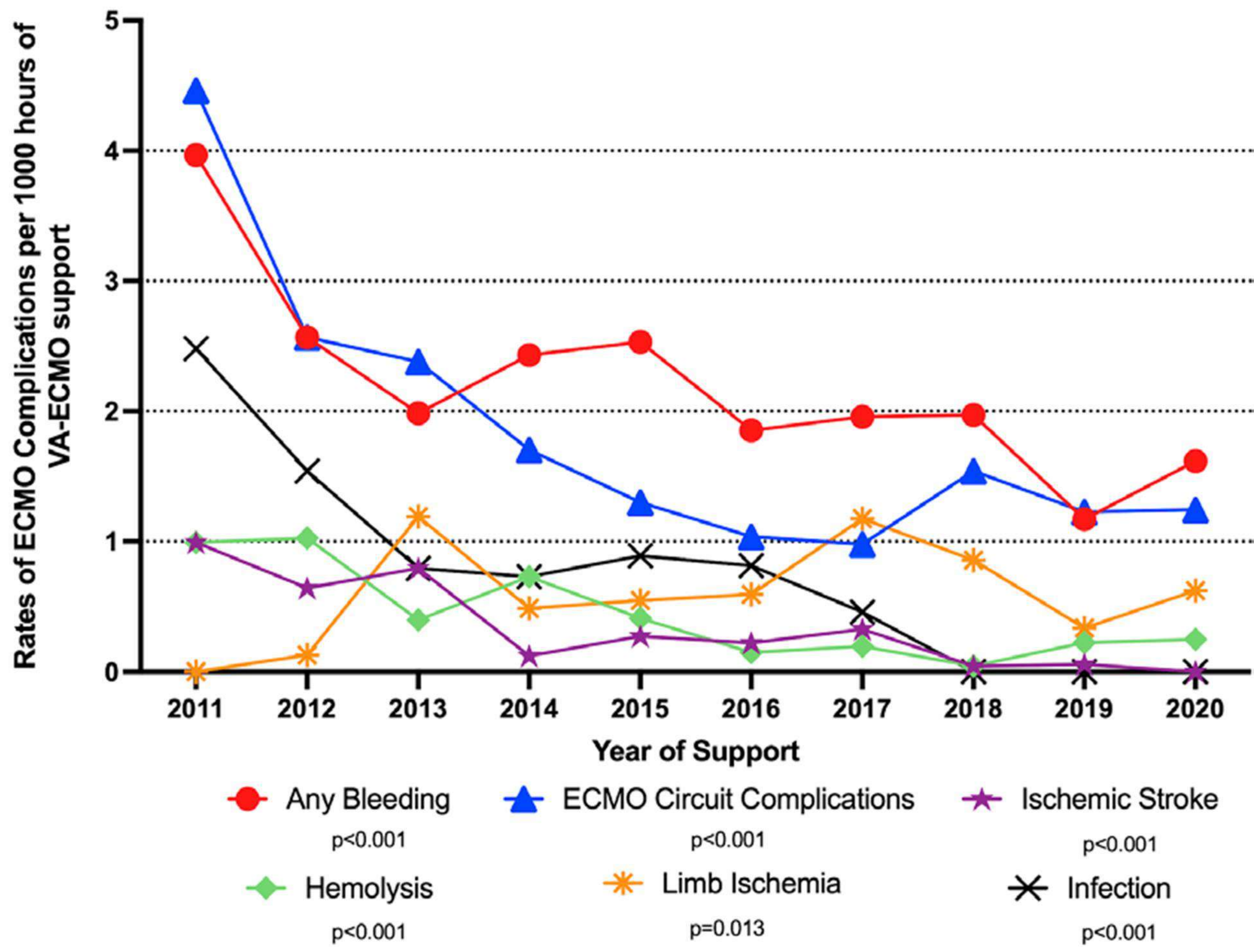
## ELSO registry

- 850 pz
- Mean age 41 years, 52% male
- Mortality 34.9%
- Htx 1.9%
- LVAD 2.4%

*Circ Heart Fail. 2023;16:e010152*



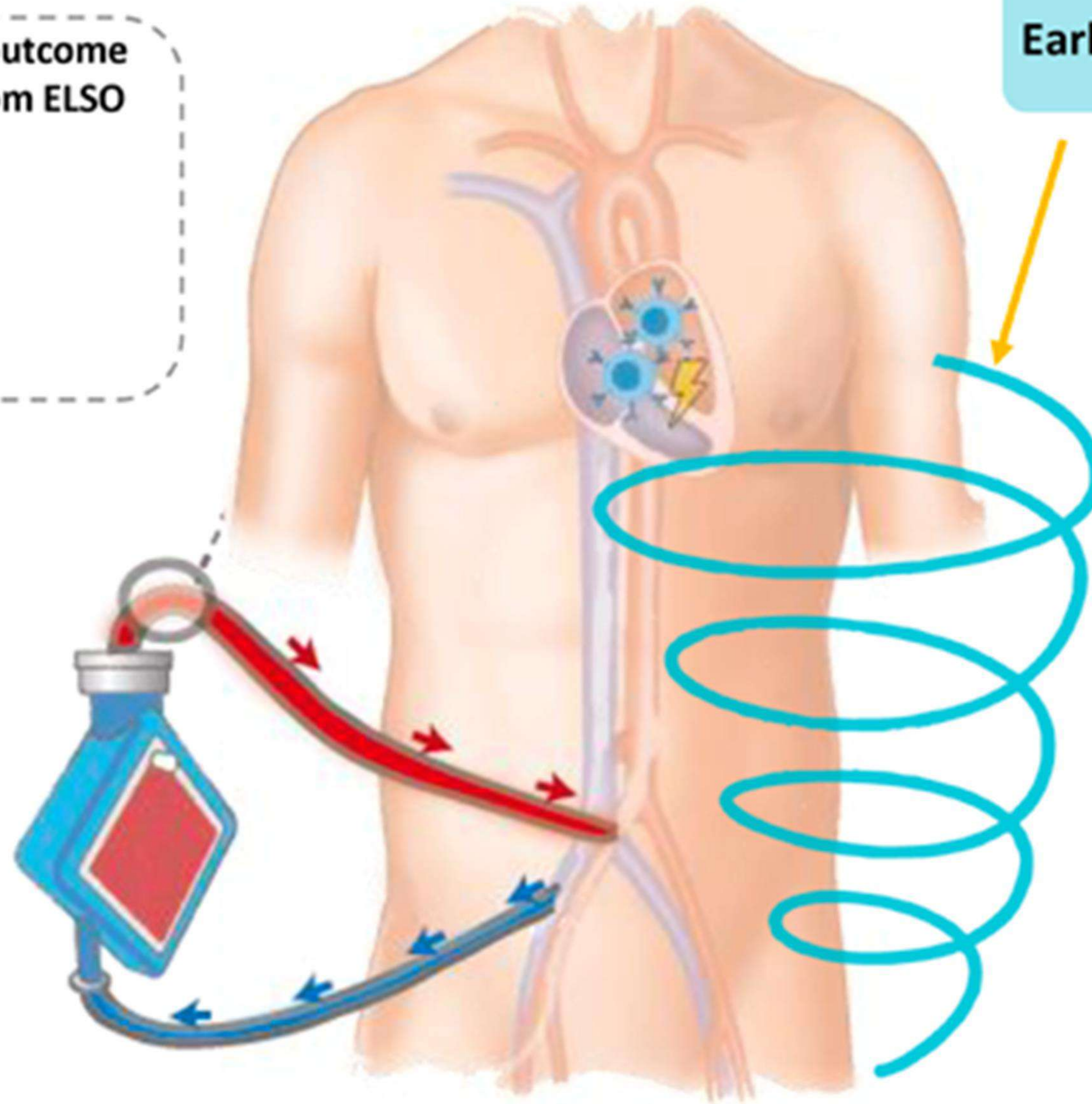
# VA-ECMO for Suspected Acute Myocarditis



# Peripheral VA-ECMO for Fulminant Myocarditis

**Unmodifiable negative outcome predictors for VA-ECMO from ELSO registry in FM:**

- Older age
- Higher weight
- Asian race



**Early recognition and ECMO initiation**

**Negative outcome predictors pre-VA-ECMO from ELSO registry in FM:**

- Low pH / high lactate levels
- ECPR / ongoing resuscitation
- Lower MAP
- Ongoing sepsis

**Negative outcome predictors from other registries on patients with FM:**

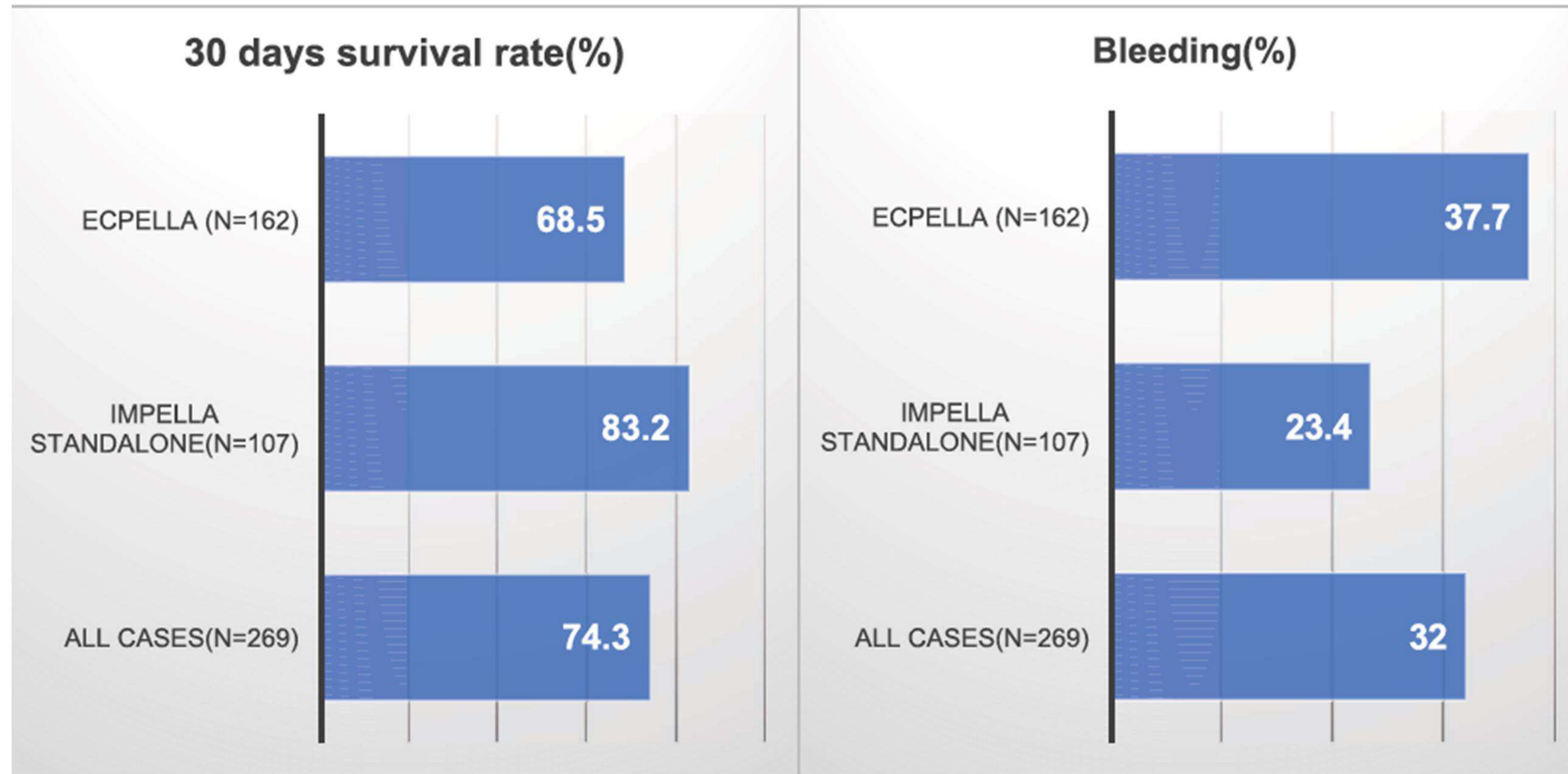
- Giant cell myocarditis
- QRS >120 ms on ECG
- VT/VF on initial ECG

# Impella in Fulminant Myocarditis

269 pts

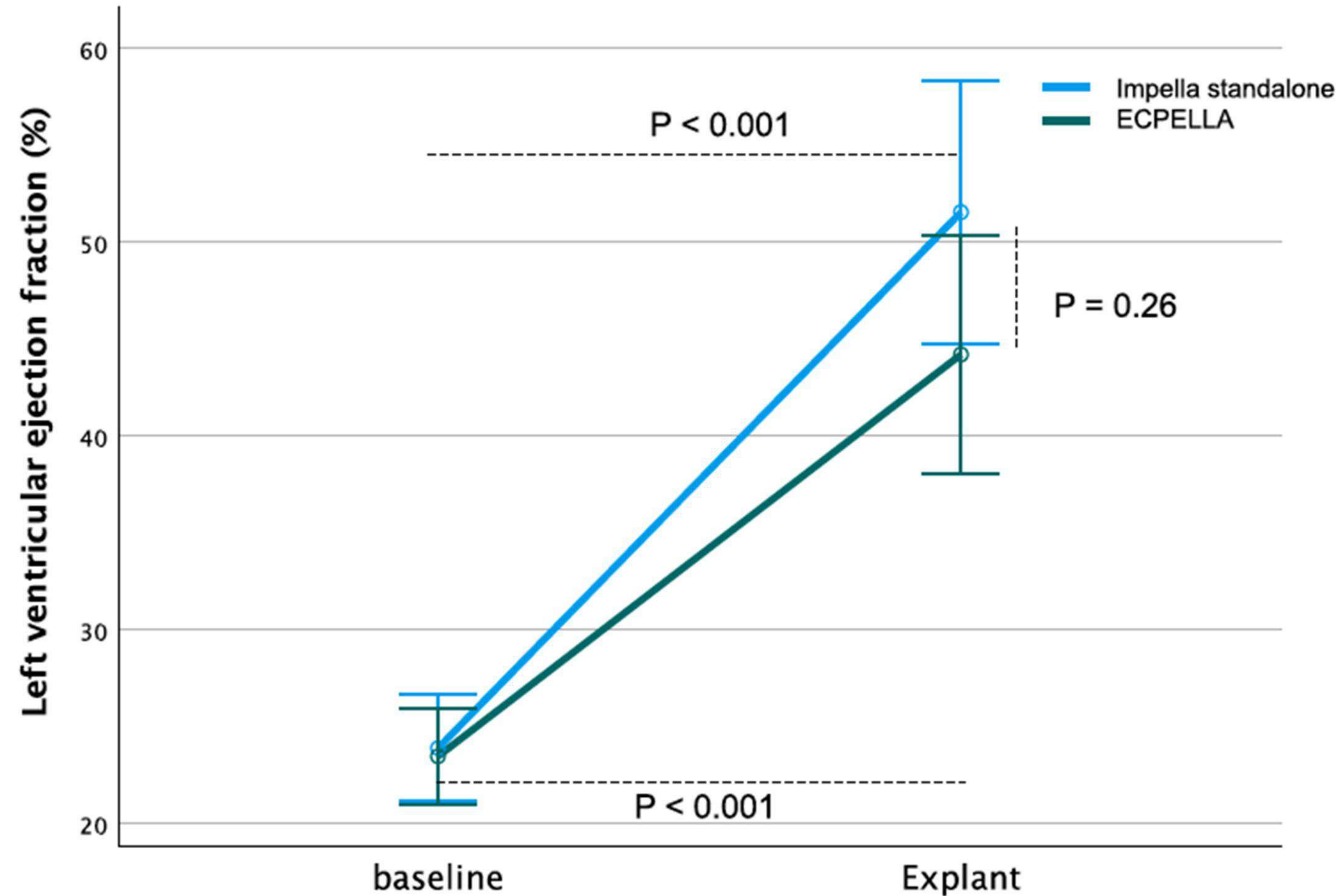
Impella stand-alone 39.8% vs. ECPELLA 60.2%

**Impella device in fulminant myocarditis:  
Japanese Registry for Percutaneous Ventricular  
Assist Device (J-PVAD) registry analysis on  
outcomes and adverse events**



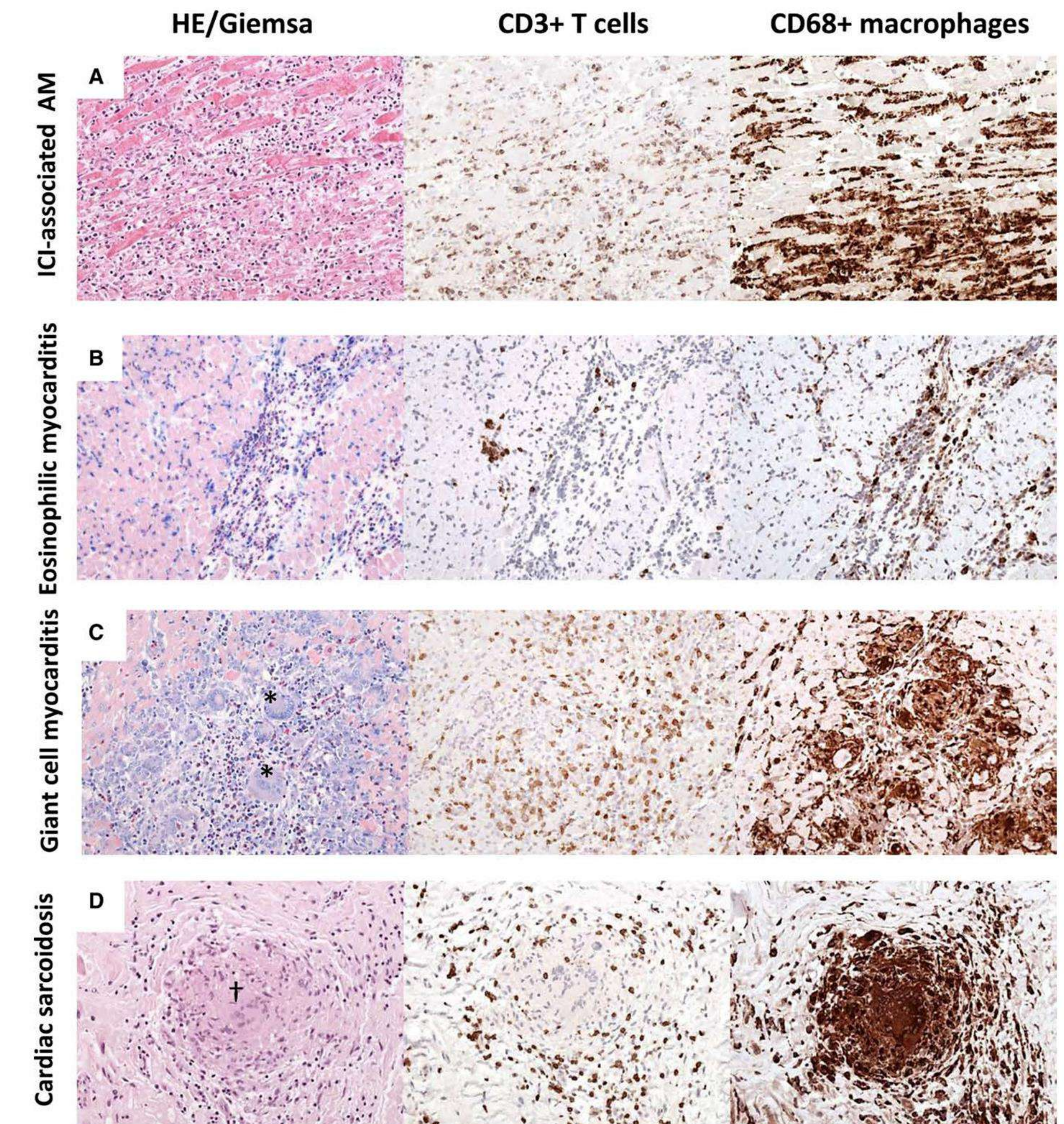
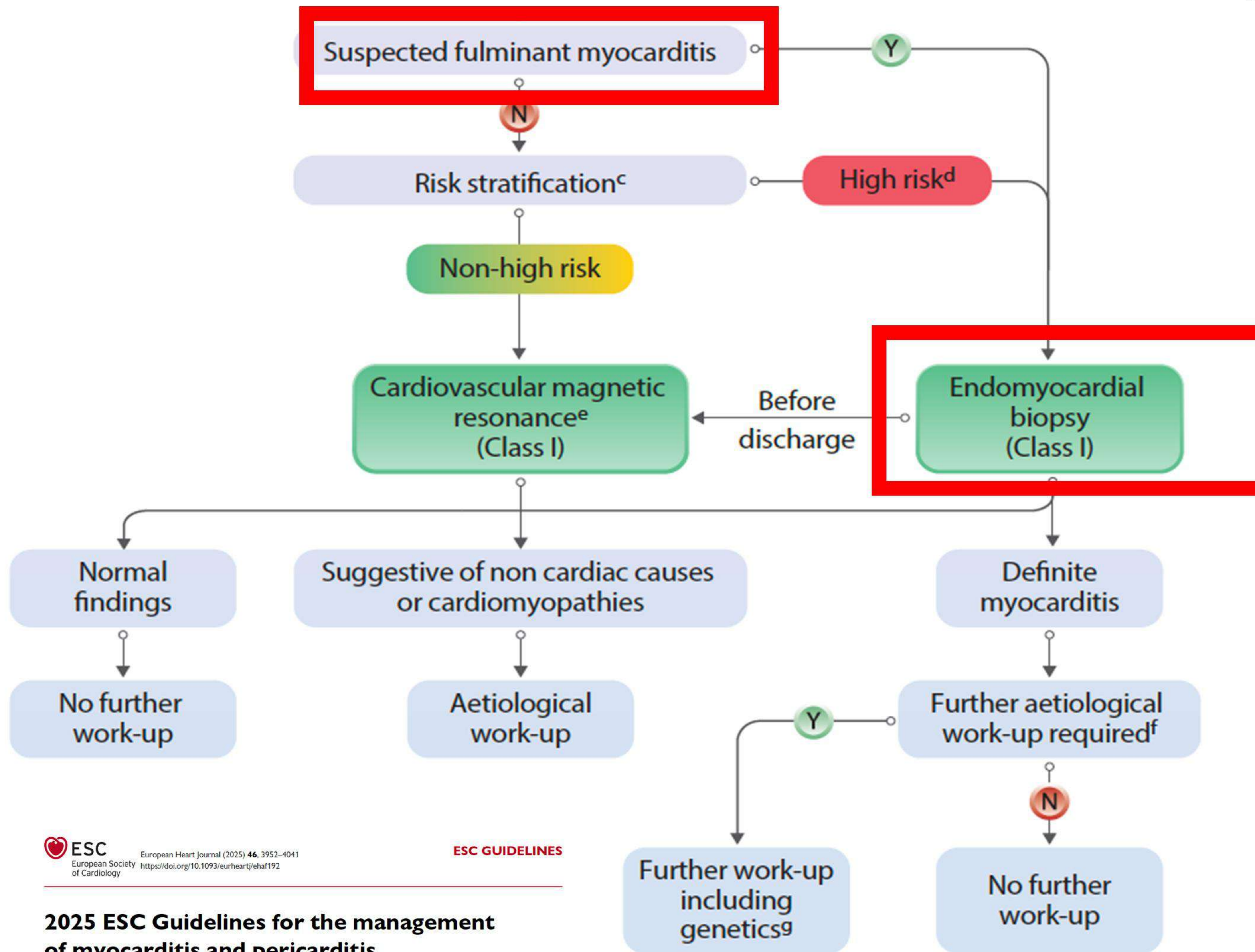
**Impella stand-alone associated with lower mortality and fewer complications**

## Impella device in fulminant myocarditis: Japanese Registry for Percutaneous Ventricular Assist Device (J-PVAD) registry analysis on outcomes and adverse events

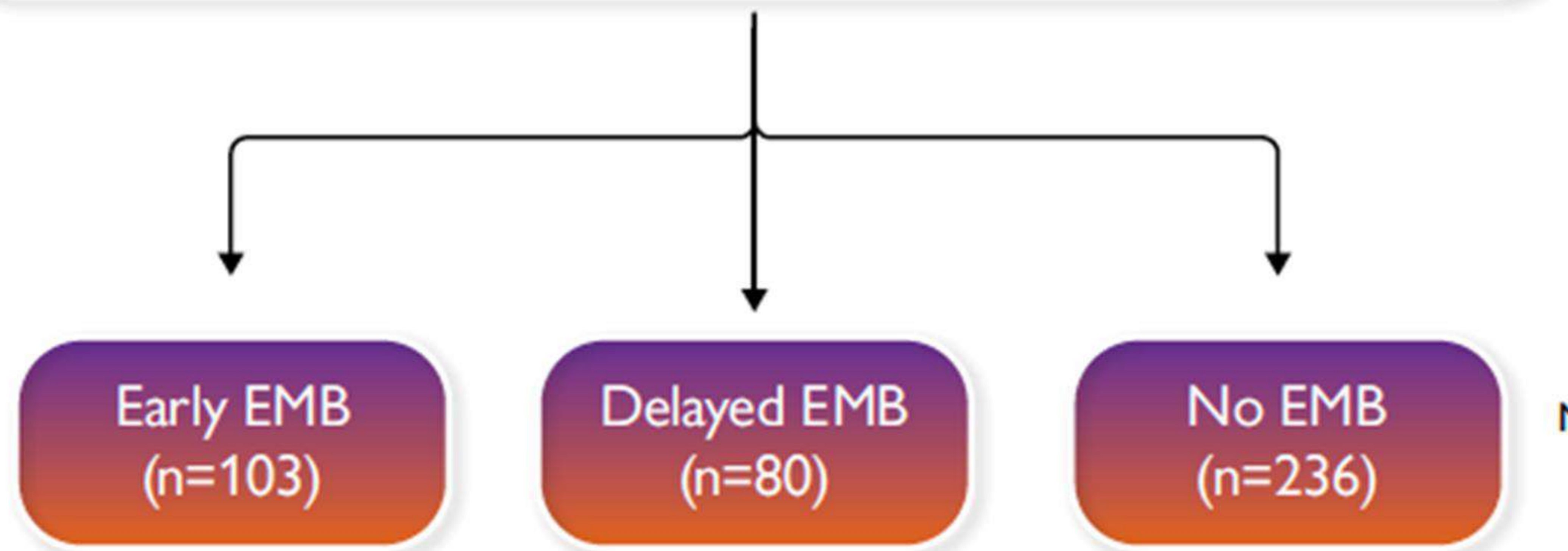
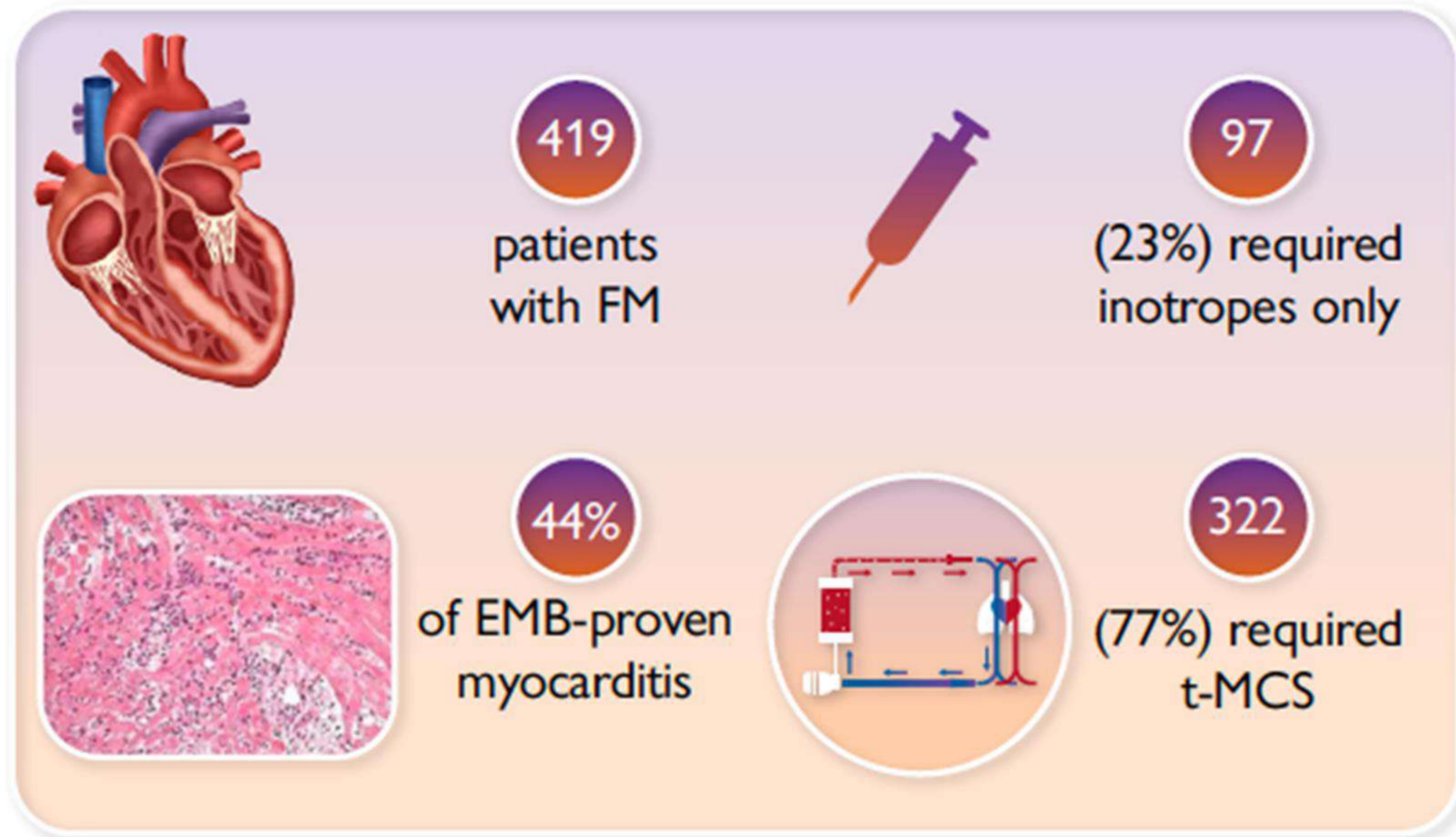


Risk	High risk	Intermediate risk	Low risk
<b>Myocarditis</b>	<ul style="list-style-type: none"> <li>Acute HF/cardiogenic shock</li> <li>Dyspnoea NYHA III-IV refractory to medical therapy</li> <li>Cardiac arrest/syncope<sup>a</sup></li> <li>Ventricular fibrillation/sustained ventricular tachycardia<sup>a</sup></li> <li>High-level AV block<sup>a</sup></li> </ul>	<ul style="list-style-type: none"> <li>New/progressive dyspnoea</li> <li>Non-sustained ventricular arrhythmias</li> <li>Persistent release or relapsing troponin</li> </ul>	Stable symptoms or oligosymptomatic

EMB is crucial in detecting the specific histotype of the AM and assessing the immunologic and virologic status of the myocardium through immunohistochemical and biomolecular PCR analyses.

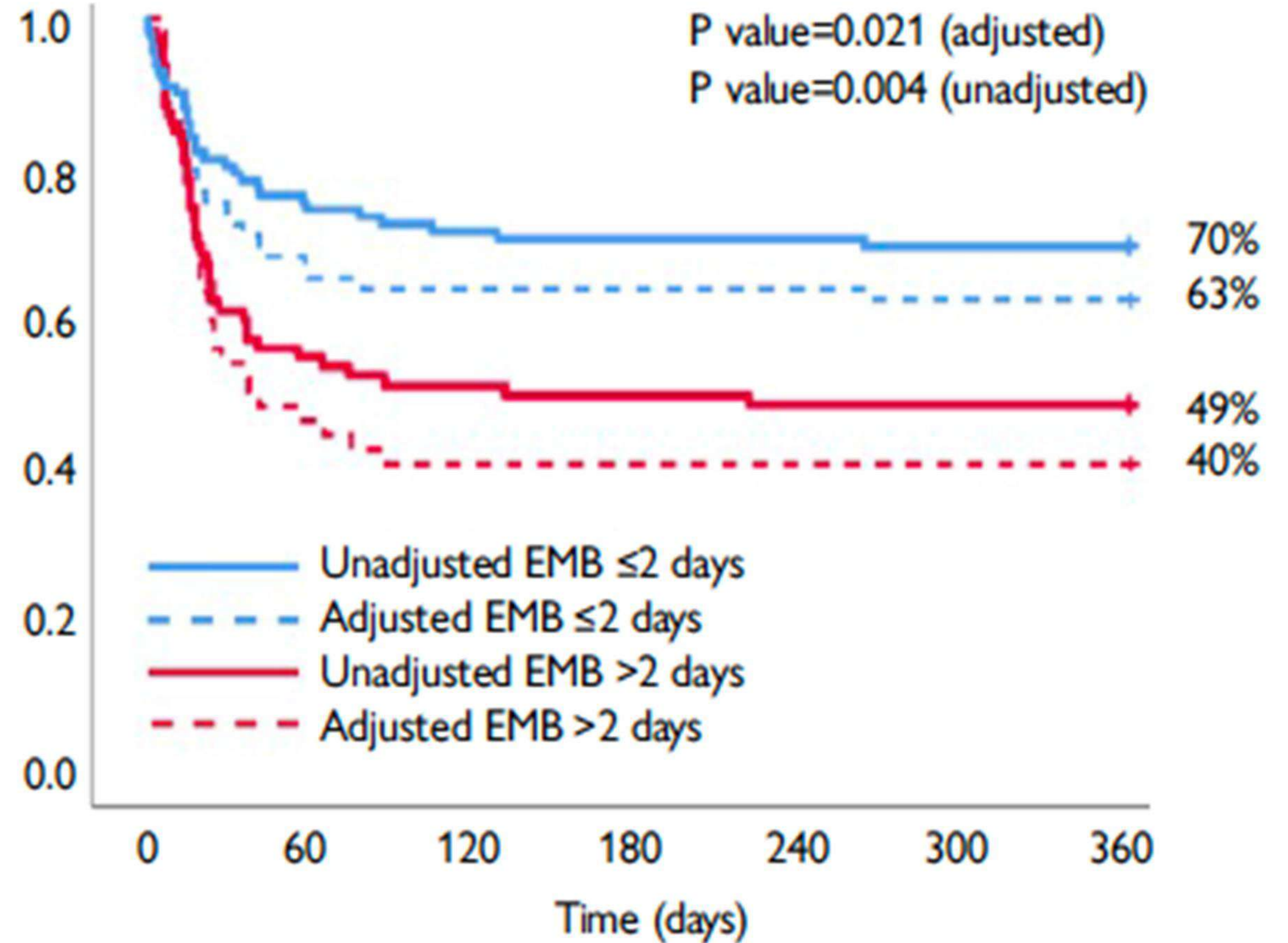


# Endomyocardial Biopsy in Fulminant Myocarditis: Timing and Clinical Impact



Overall survival free of heart transplantation or LVAD at one year after ICU admission: 65%

Survival free of Heart transplant or LVAD



## 2025 ESC Guidelines for the management of myocarditis and pericarditis

### Immunosuppressive therapy

Corticosteroids should be considered in patients with fulminant, non-infectious forms of myocarditis to stabilize the patients.

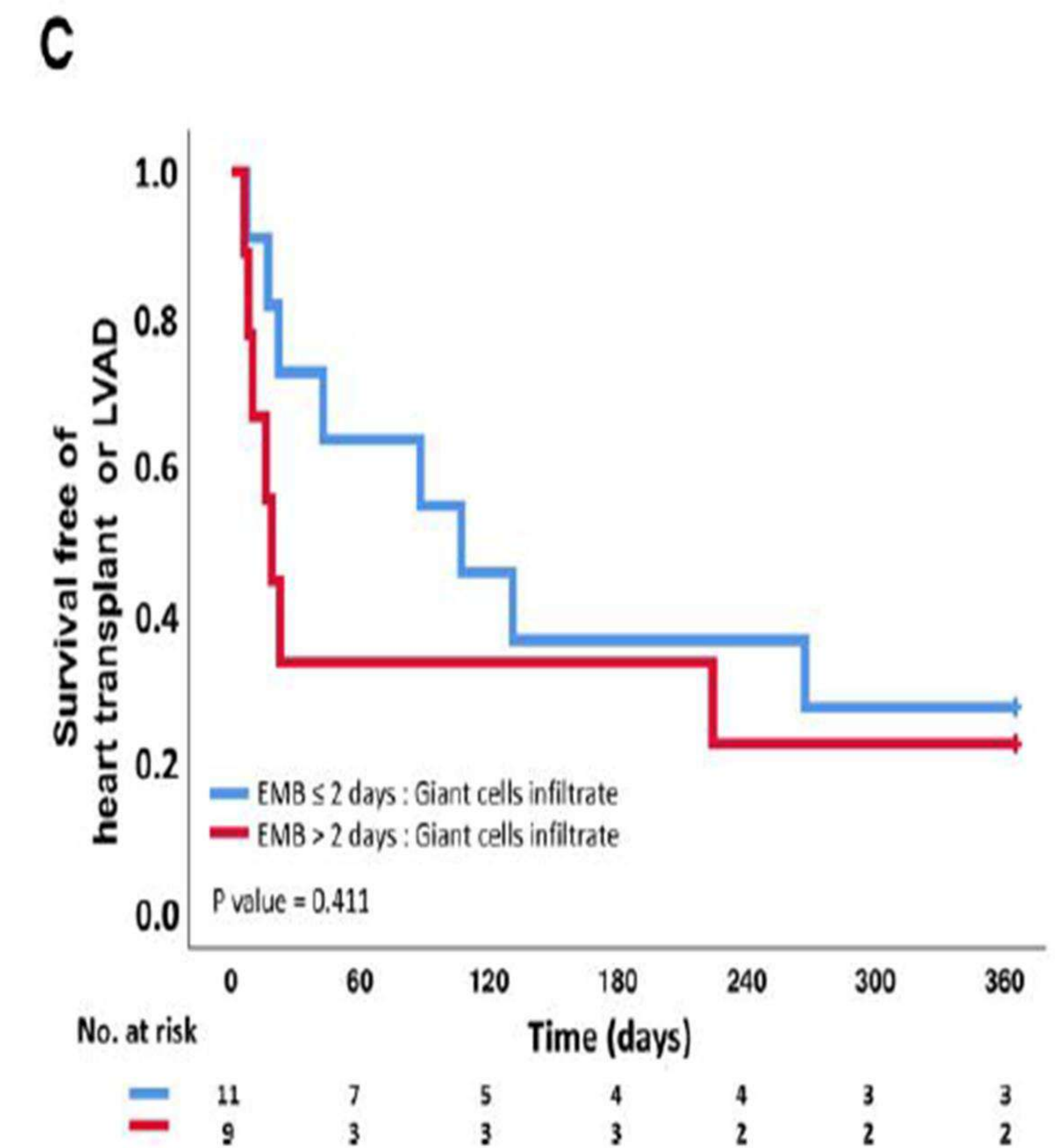
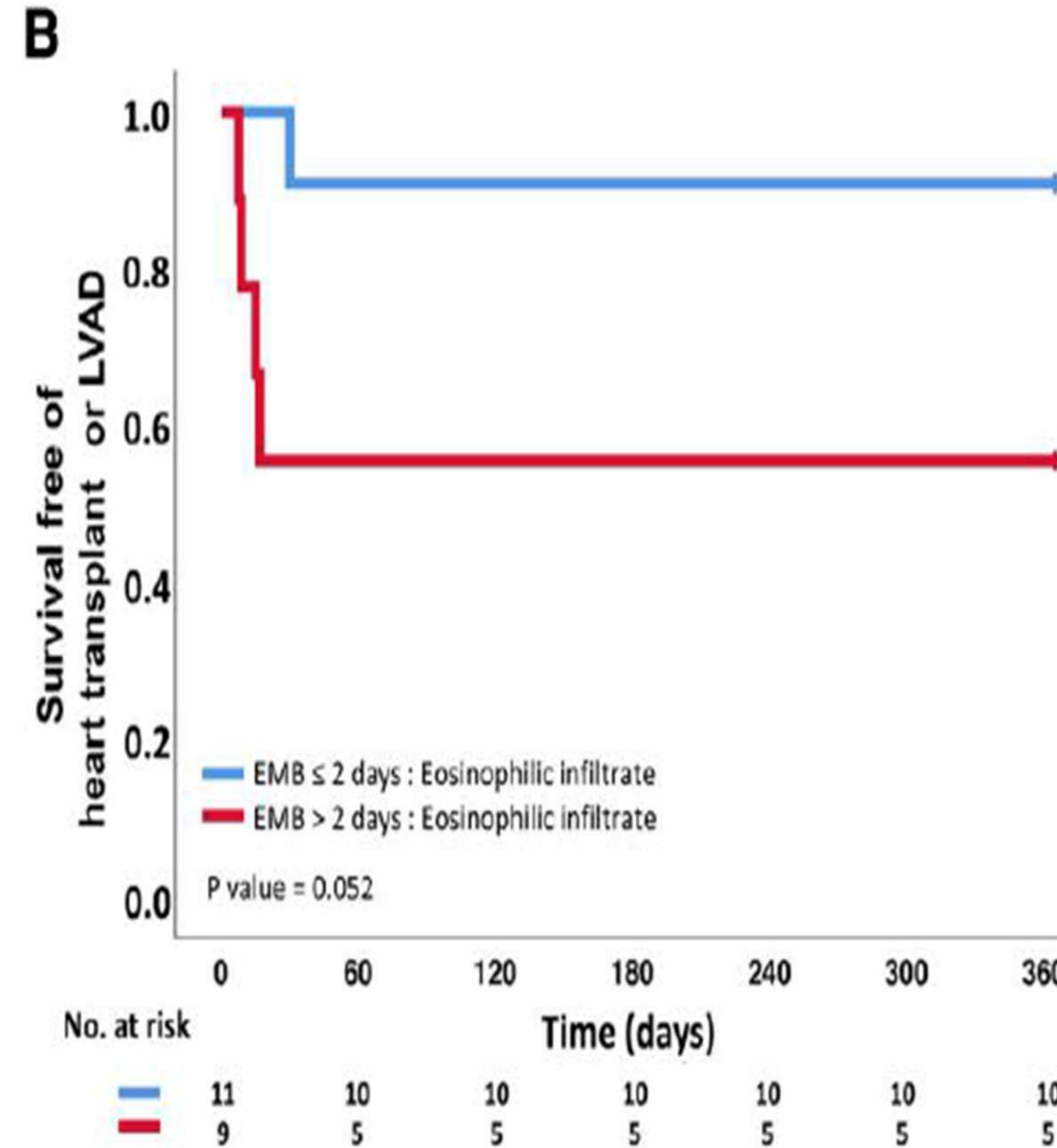
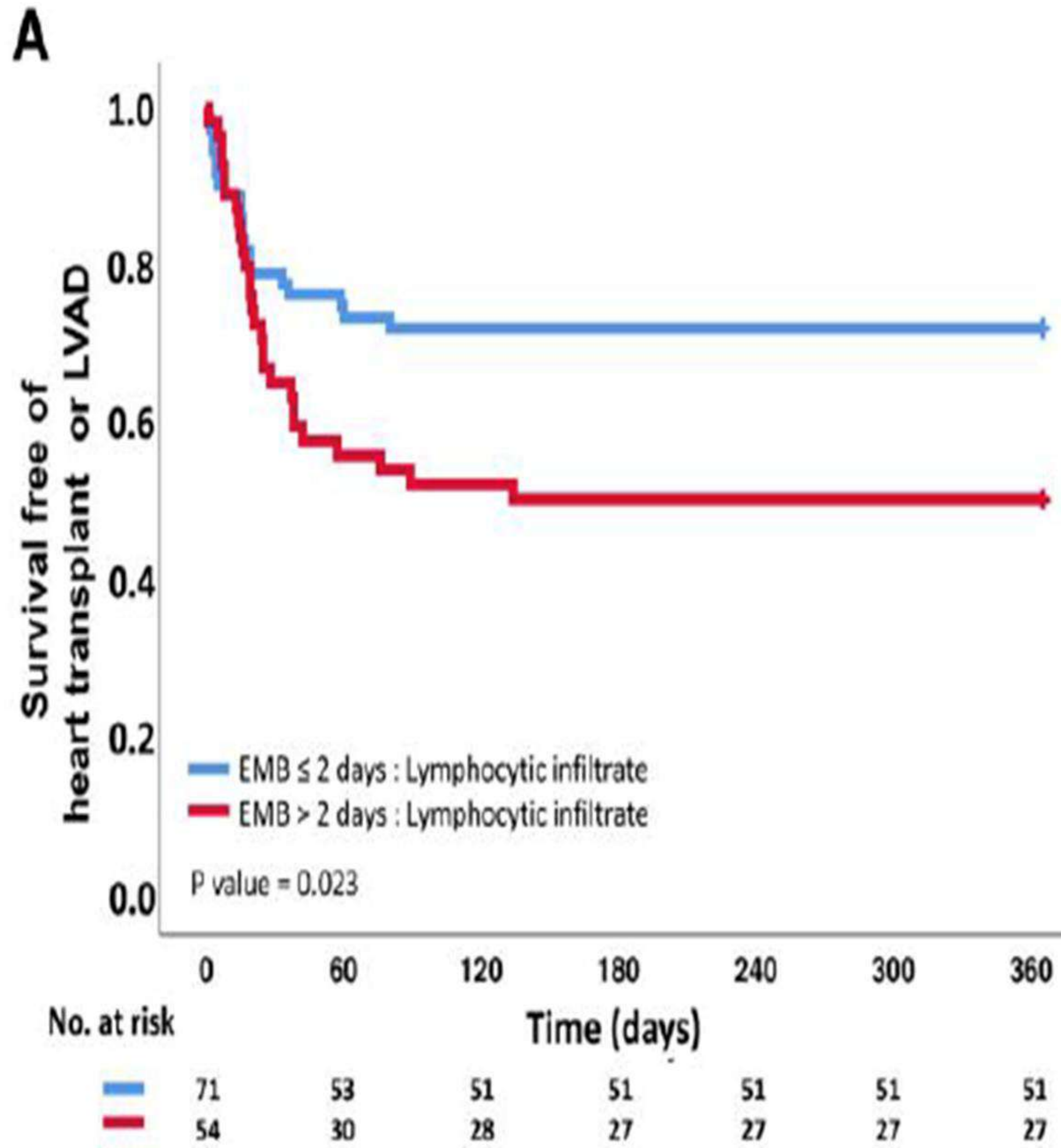
**Ia**

**C**

### ESC guidelines

- 500 copies/ $\mu\text{g}$  of DNA in EMB considered as the threshold for virus-related myocardial inflammation
- In cases of acute cardiac and systemic virus infection, immunosuppressive therapy must be avoided

# Endomyocardial Biopsy in Fulminant Myocarditis: Timing and Clinical Impact



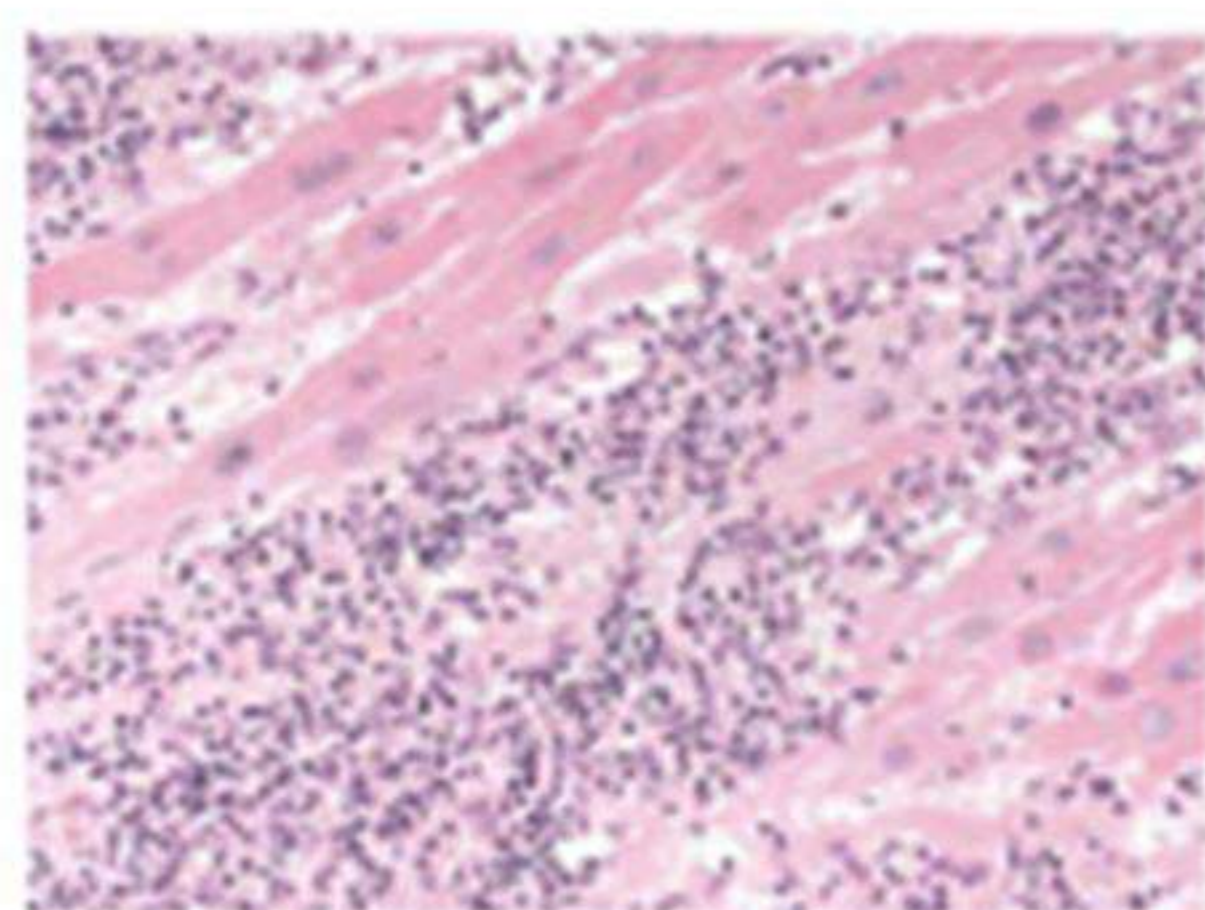
# Immune Suppressive Treatment

SUSPECTED FULMINANT OR COMPLICATED ACUTE MYOCARDITIS

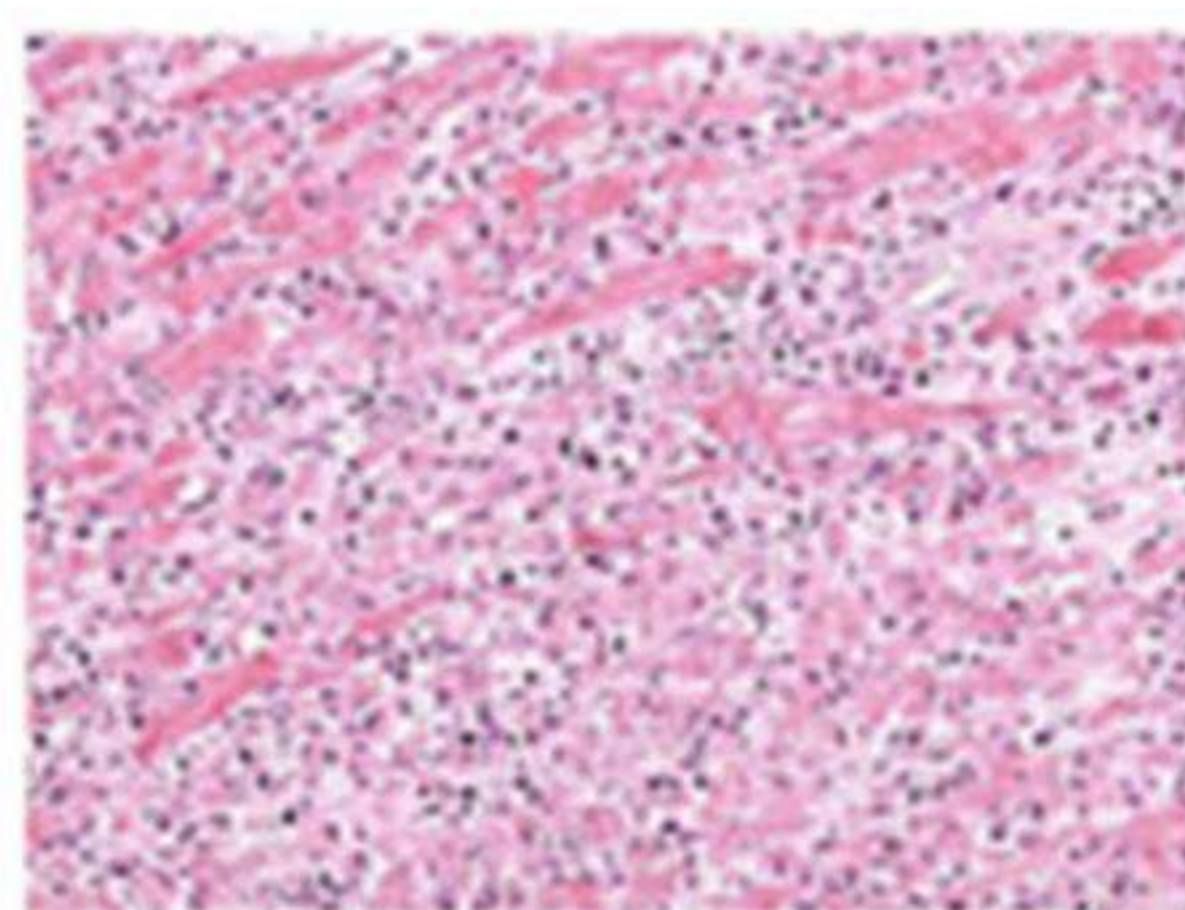


i.v. pulse methylprednisolone 7–14 mg/kg/day for 3 d, then 1 mg/kg/day

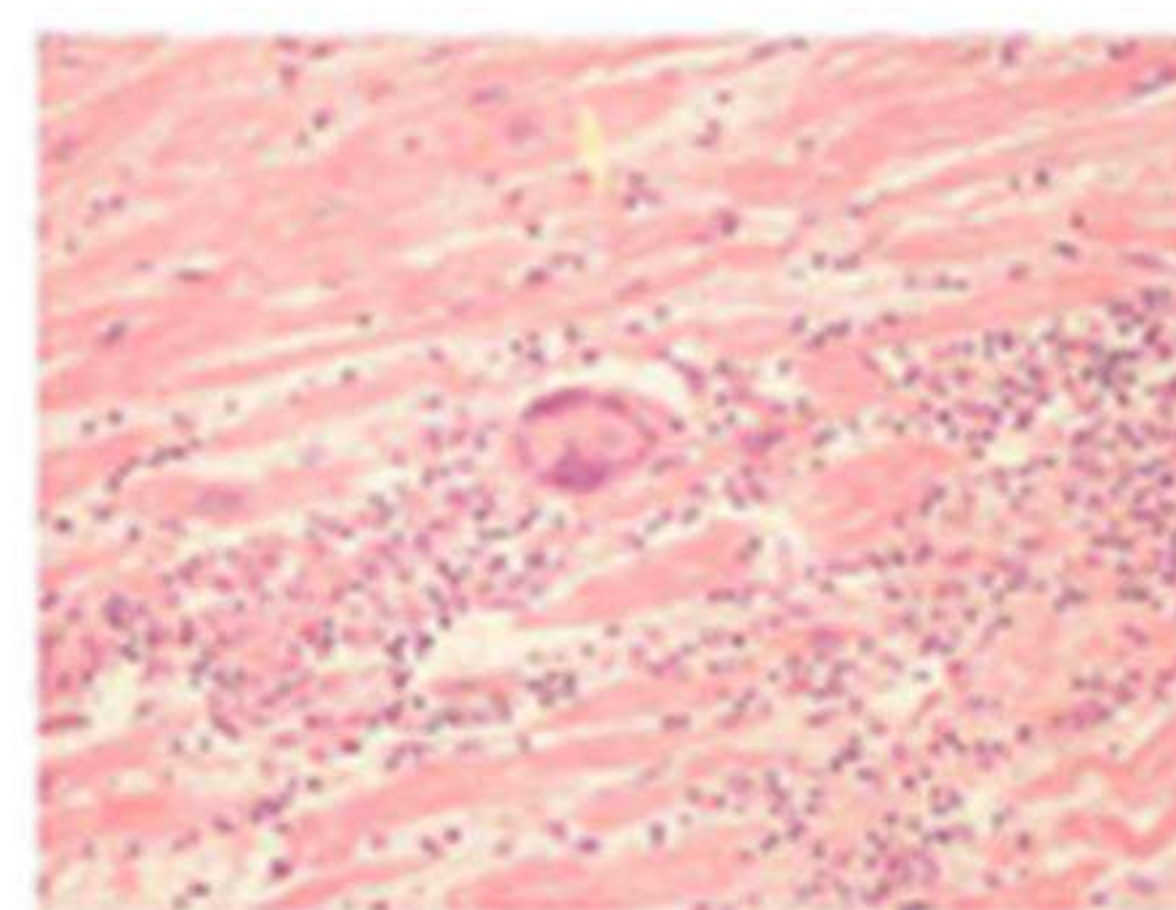
FIRST-LINE



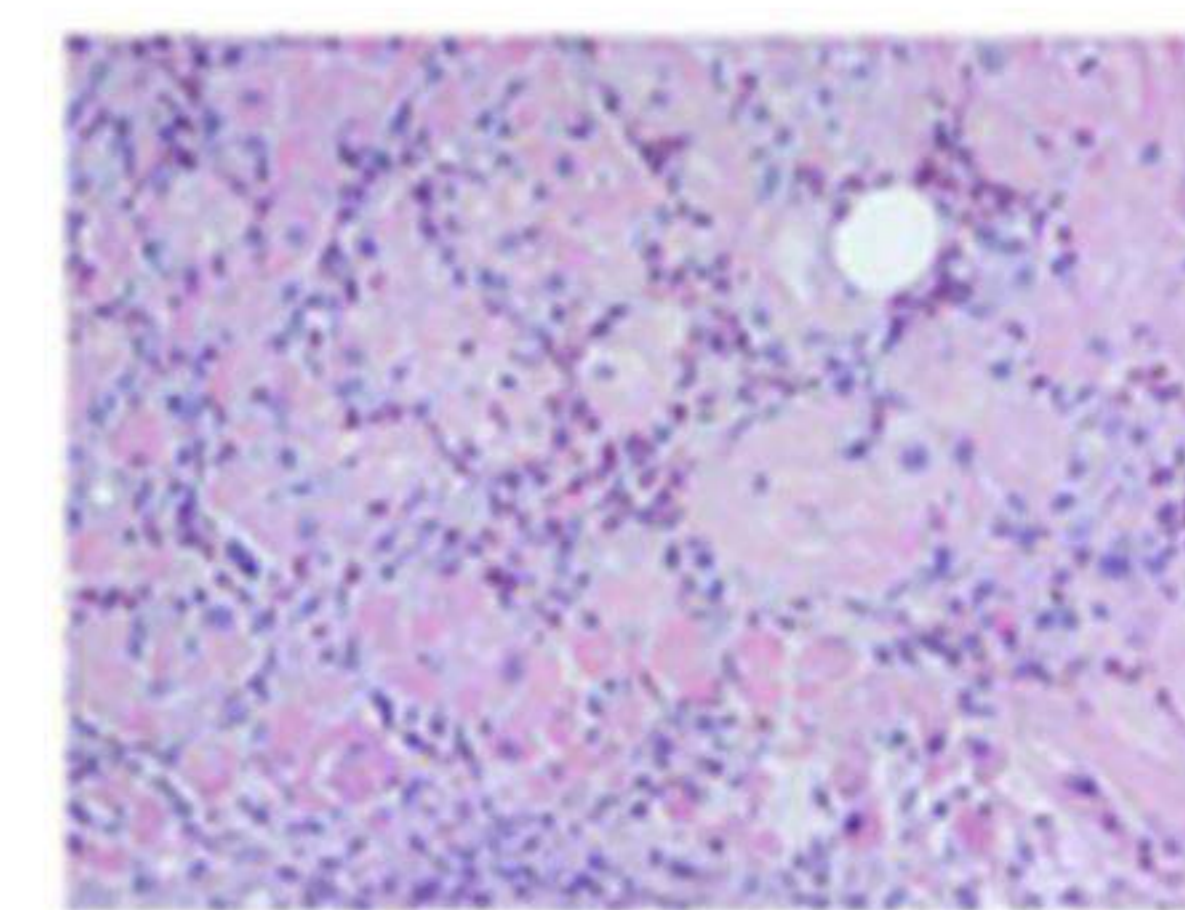
**LYMPHOCYTIC**



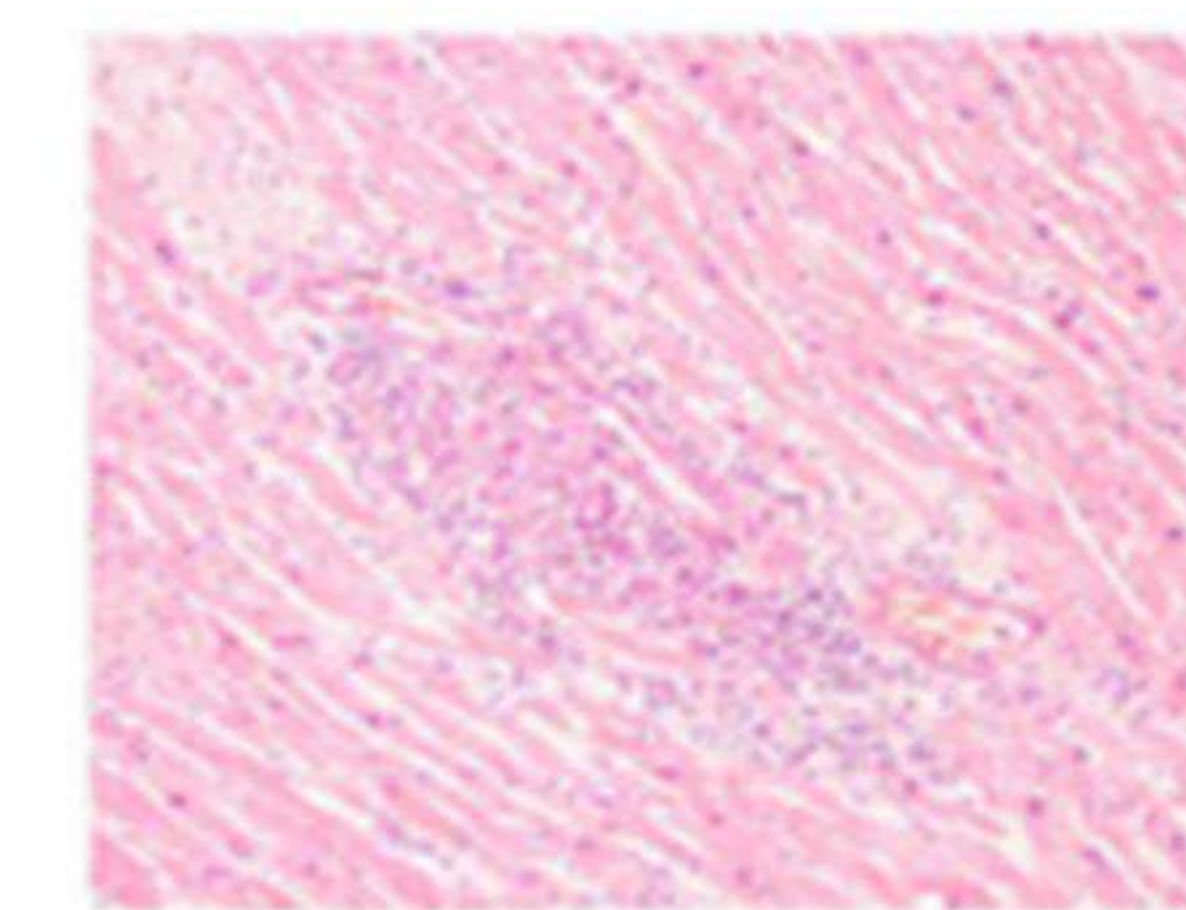
**ICI-ASSOCIATED**



**GIANT CELL**



**EOSINOPHILIC**



**SARCOIDOSIS**

ADDITIONAL

- If associated systemic autoimmune disorders (eg. SLE and APS): add aggressive treatment of associated conditions

**Hold ICI** therapy  
Confirm ICI-myocarditis via definitive imaging and/or endomyocardial biopsy

- If hemodynamically unstable pts: **ATG**, from 1 mg/kg, usually single-dose to 300 mg in 3 days or (alternative) i.v. **alemtuzumab** (anti-CD52 antibody) single dose of 30 mg plus oral **CyA**, BID, target trough levels 150–250 ng/mL  
- If hemodynamically stable pts: only oral **CyA**, BID, target trough levels 150–250 ng/mL

- If EGPA: consider i.v. **cyclophosphamide** (especially in ANCA-positive pts), 600 mg<sup>m<sup>2</sup></sup> at days 1, 15, and 30  
- If clonal (myeloproliferative) HES: **imatinib** 100–400 mg OD  
- If helminthic infection: **albendazole** 400 mg BID for 2–4 wk  
- If hypersensitivity reaction: **withdraw suspected drug**

SECOND-LINE

**IVIg** (2 g/kg), single continuous infusion in 24–48 h or divided in 4 d or **plasmapheresis**, 3–5 sessions in 5–10 d

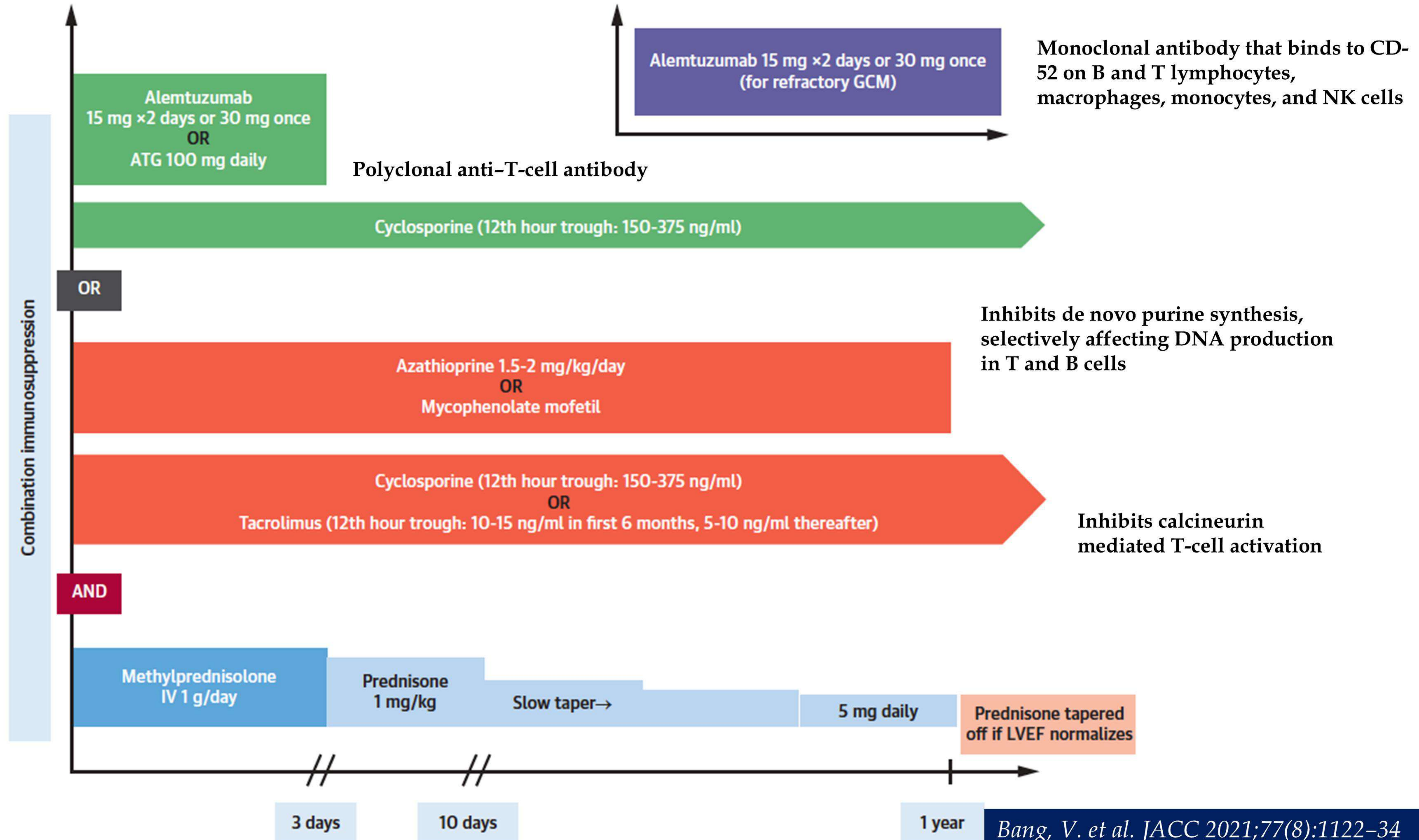
i.v. **abatacept** (a CTLA-4 agonist) or **ATG**, 1 mg/kg, usually single dose or i.v. **alemtuzumab** (anti-CD52 antibody), 30 mg, single dose

i.v. **rituximab** 375 mg<sup>m<sup>2</sup></sup> (BSA) mg (once a wk for 4 wk and then every 4 mo as maintenance therapy) for 1 yr

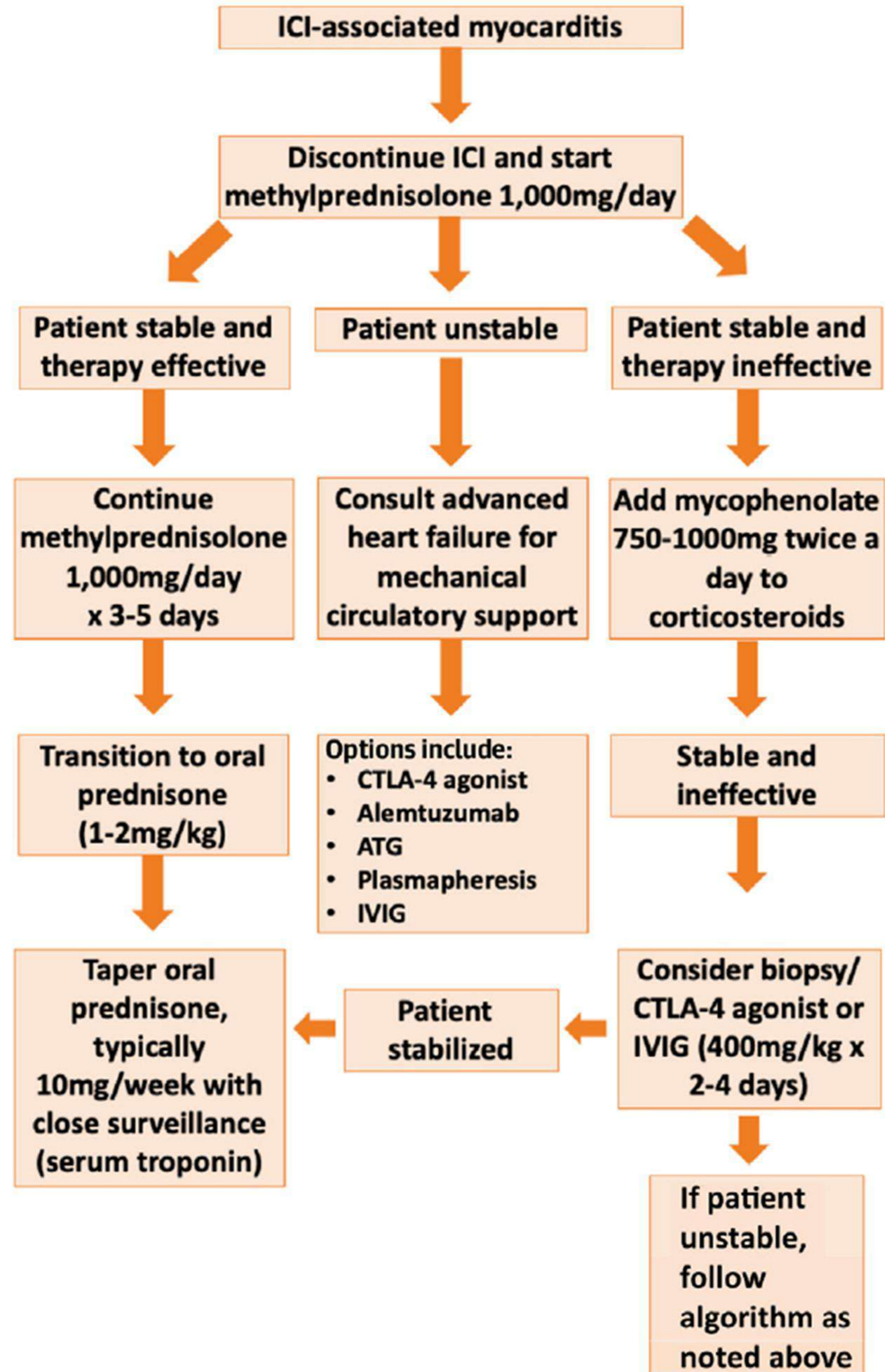
- If DRESS, EGPA or idiopathic HES: anti-IL5 agents (e.g., **benralizumab** 30 mg s.c./4–8wk or **mepolizumab** 100–300 mg/4wk)

s.c. **methotrexate** 15–20 mg/wk or i.v. **infliximab** 5 mg/kg (up to 500 mg) at time 0 and after 2 and 4 wk and then every 6–8 wk or s.c. **adalimumab** 40 mg/2wk

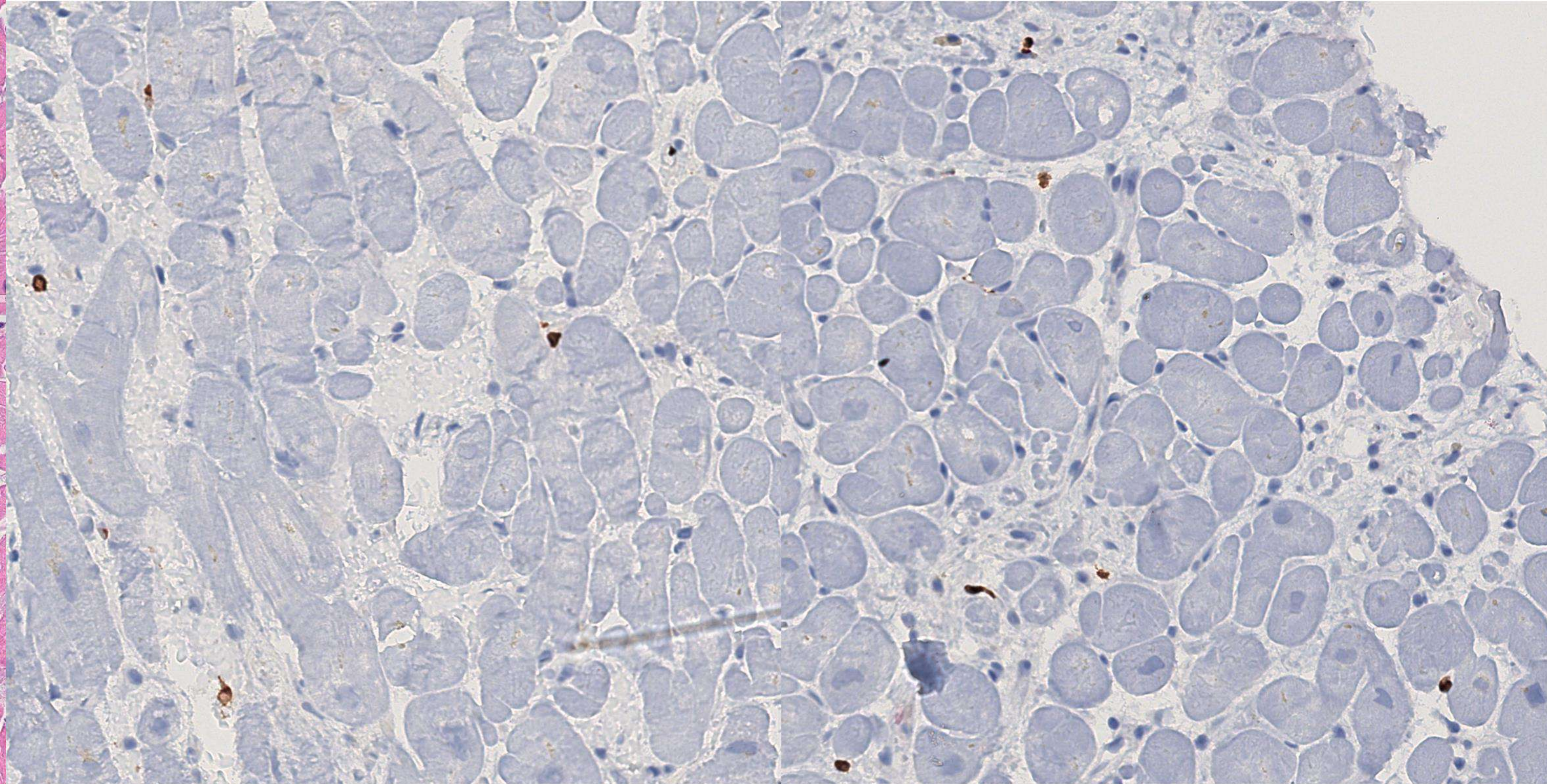
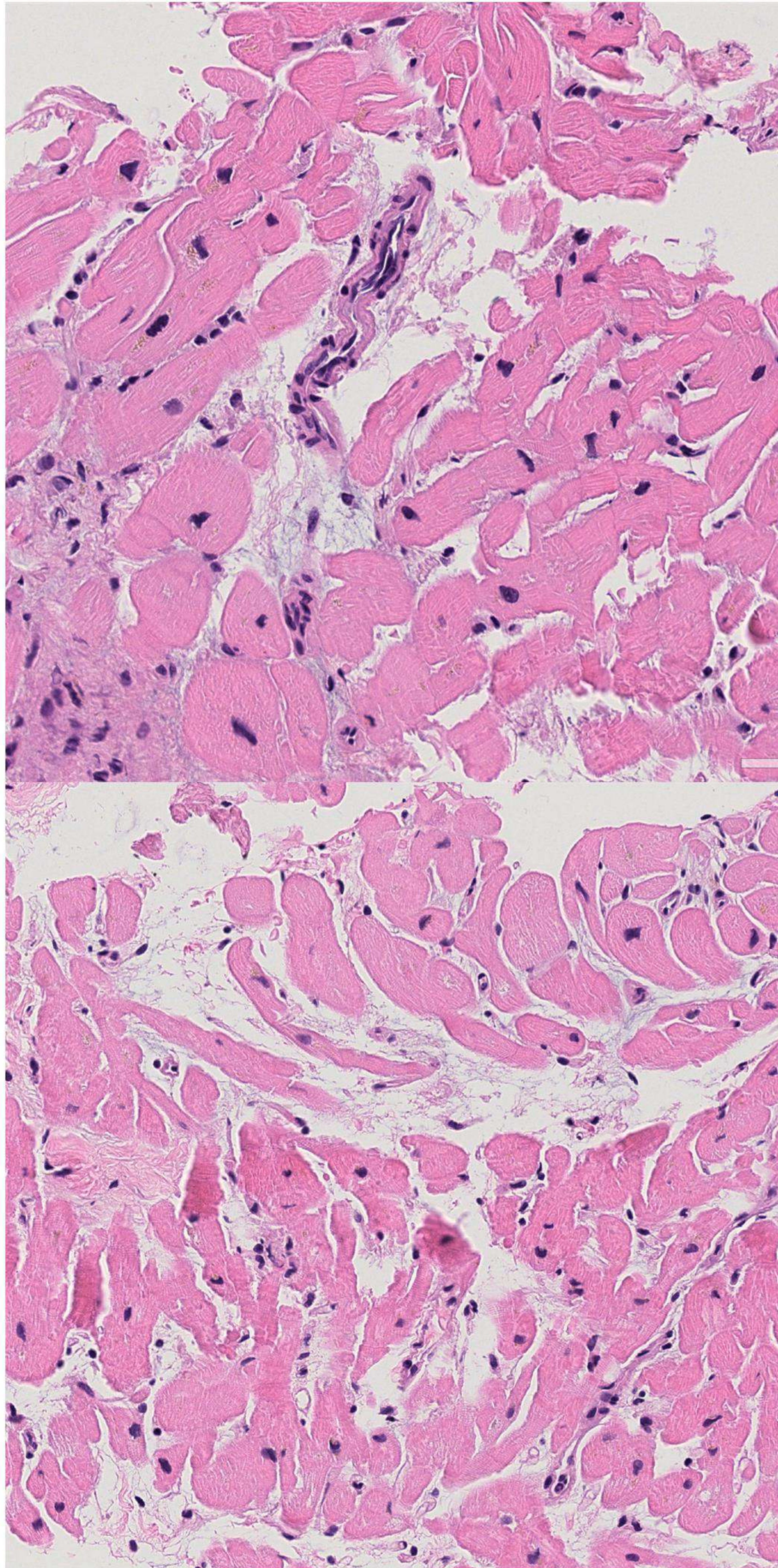
# Combination and Duration of Immunosuppressive Therapy in GCM



# Proposed Treatment Algorithm for ICI-Associated Myocarditis



.....Clinical case - EMB

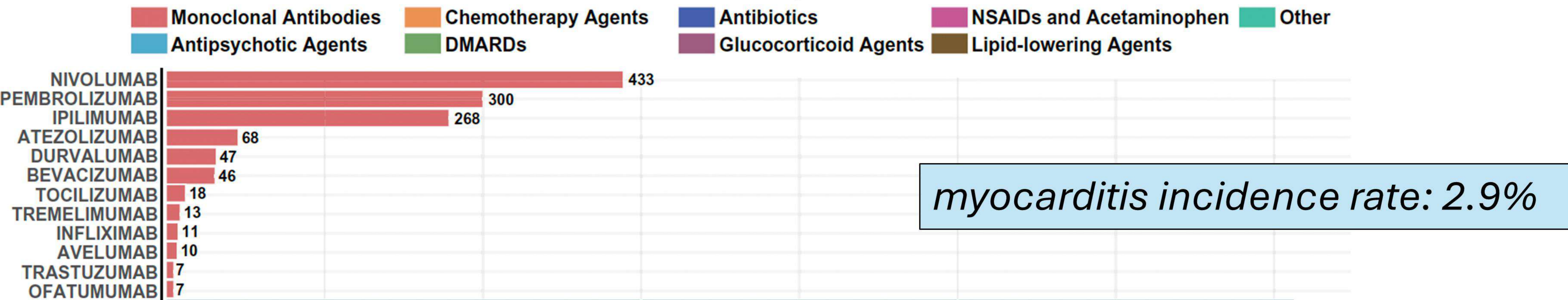


**Virus negative**

**ORIGINAL RESEARCH**

# Evolving Trends and Drug Class Dynamics in Drug-Induced Myocarditis: A 2-Decade Comparative Analysis of Pharmacovigilance Data

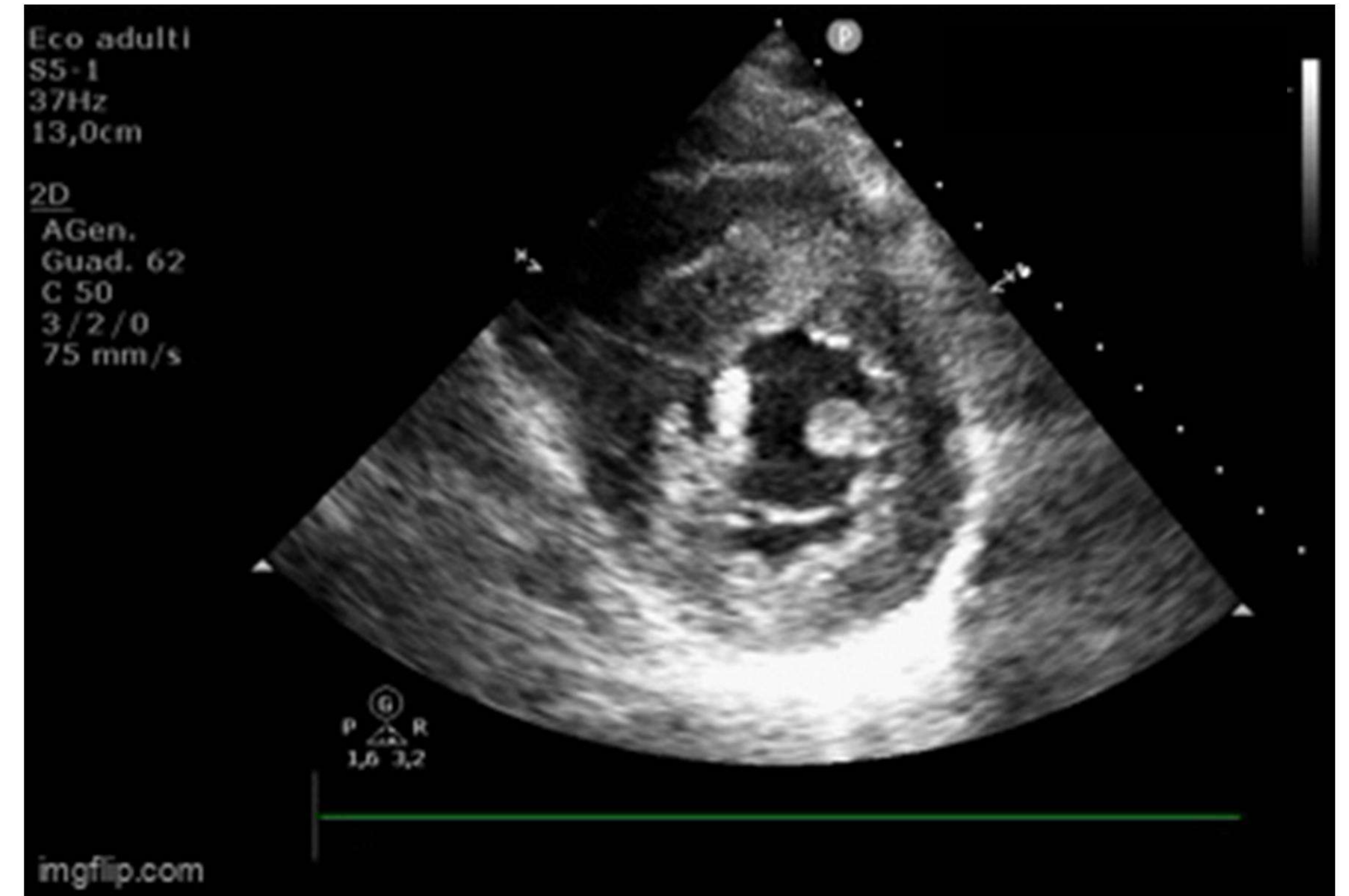
**Total Number of Myocarditis Cases**



**Table 2. Top 50 Drugs Associated With Reports of Myocarditis to the JADER From 2004 to 2023**

Drug name and rank	Total adverse event reports	Myocarditis reports (‰)	Maximum annual reports (y)	Median, y [IQR]*	Female sex (%), n=3638 <sup>†</sup>
45. Ofatumumab	237	7 (29.54)	7 (2014)	2014 [2014–2014]	0 (0.00)

## Clinical case - ECMO WEANING (DAY 8)



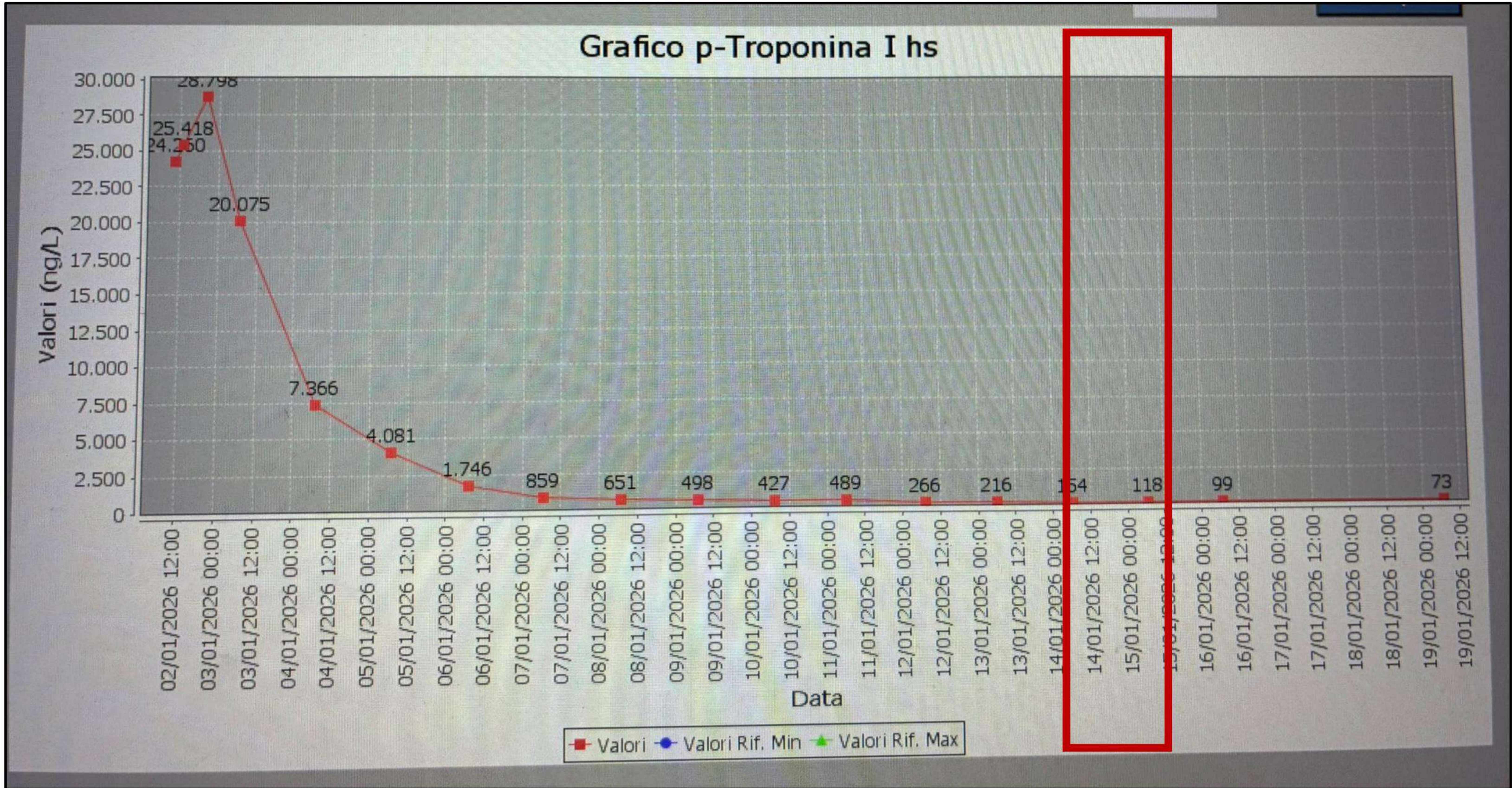
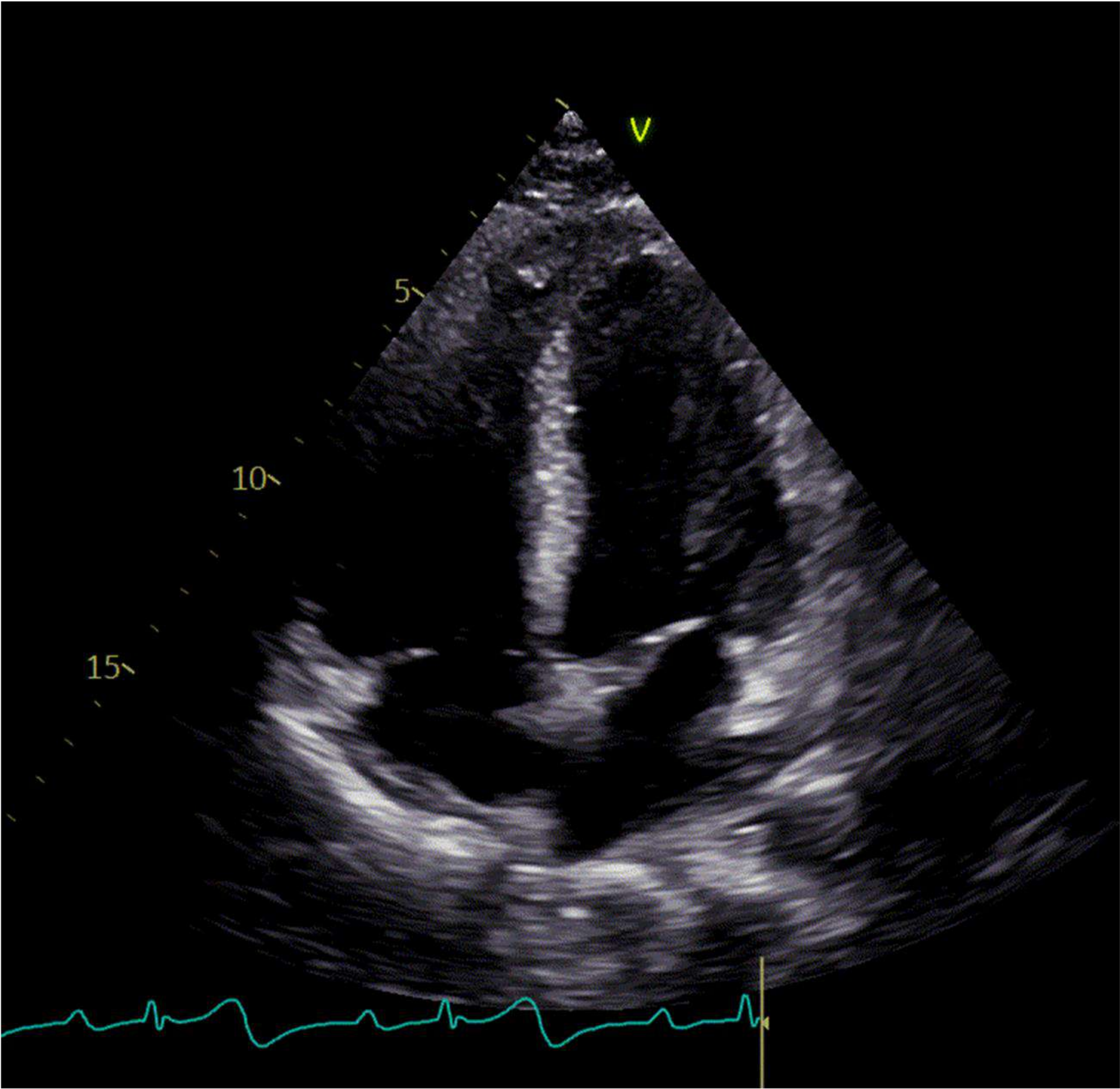
- Epinephrine 0.05  $\mu\text{g}/\text{kg}/\text{min}$
- Impella P4

Right femoral access hematoma

# Clinical case - Impella weaning (day 11)

## Day 14

- Troponine 118, creatinine 0.7
- Ramipril 2.5 mg, Bisoprololo 5 mg, Colchicina 1 mg



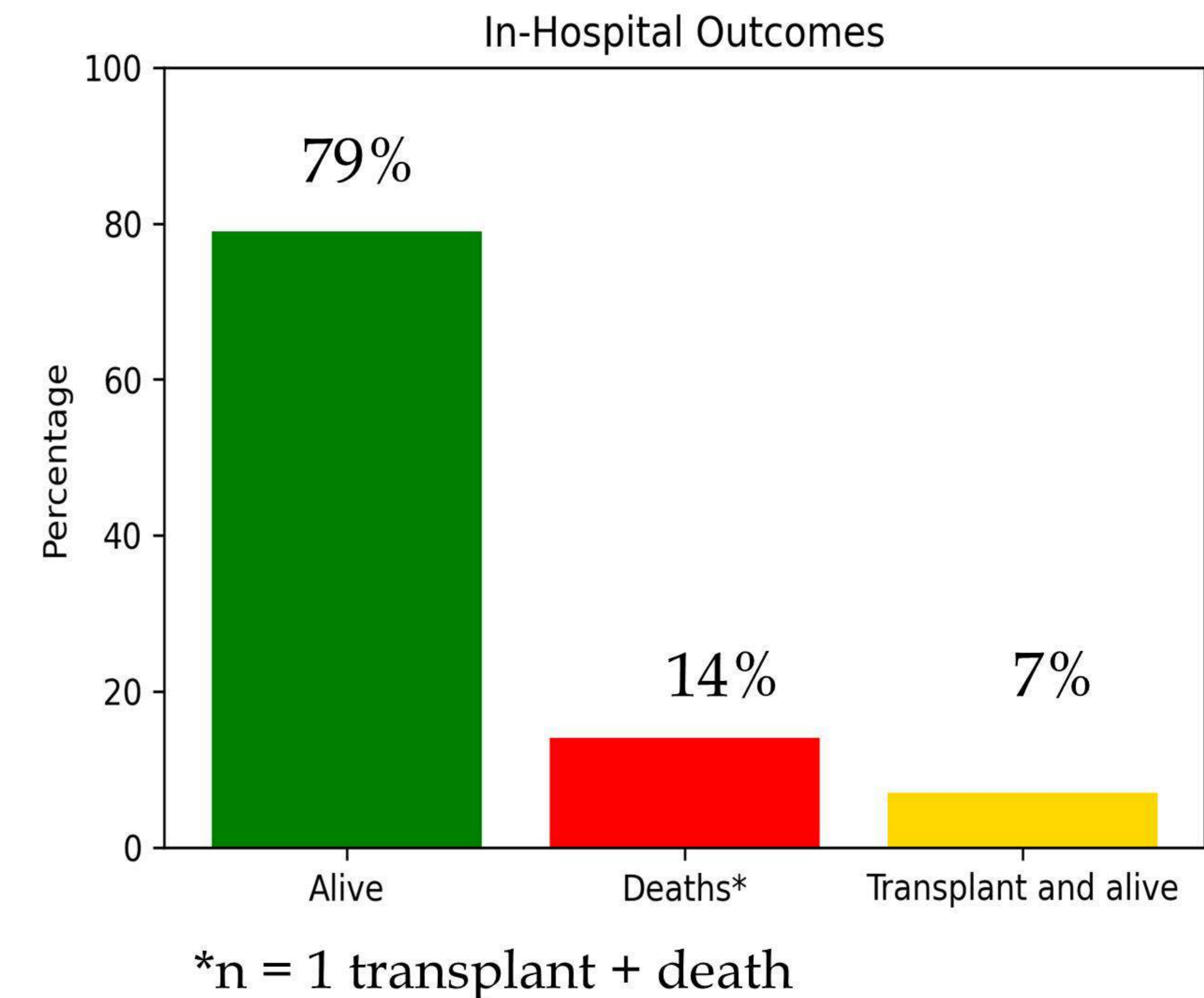
- T2: 68 msec, (v.n. <55 msec)
- ECV 40.5% (v.n. 20-30%)
- Mild midwall LGE infero-lateral

# Fulminant Myocarditis Hub-and-Spoke Network (Molinette) - Real-World Data 2024-2025

	N=14
Baseline Characteristics	
Male sex	57% (8)
Age	50 anni (IQR 40-56)
LVEF at presentation	20% (IQR 15-30%)
Lactate	4 mmol/L (IQR 3-12)
Clinical management	
Endomyocardial biopsy	93% (13)
Inotropes	100% (14)
MCS	57% (8)
IABP	36% (7)
Impella	39% (4)
ECMO	57% (8)
Invasive mechanical ventilation	43% (6)
Immunosuppressive therapy	86% (12)
Corticosteroids	86% (12)
Other agents:	36% (5)

Centre	N=14
Molinette	5
Aosta	1
Moncalieri	2
Ciriè	1
Gradenigo	1
Ivrea	1
Biella	1
Verduno	1
Mondovi	1

Diagnosi	N=13
Lymphocytic, virus-negative myocarditis	5
Giant cell myocarditis	1
Eosinophilic myocarditis	2
ICI-related	2
Anti-CD20 related	2
LES-related	1



**Survival to hospital discharge: 86%**  
**LVEF at ICU discharge: 45% (IQR 37-49)**

# Take-Home Messages

- Fulminant myocarditis is a time-critical syndrome with rapidly evolving clinical course
- Outcomes are driven by early clinical suspicion and prompt referral within an integrated Hub-and-Spoke network
- Hemodynamic stabilization and timely deployment of mechanical circulatory support are essential pre-requisites for survival
- Early endomyocardial biopsy is pivotal for precision medicine, enabling etiology-driven therapeutic strategies
- Shock Team-based decision making converts complexity into coordinated, life-saving care