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**Novas Fronteiras**  
**em Medicina**  
**Cardiovascular**

22 a 24 de Fevereiro 2019

The Village Praia  
D'El Rey - Óbidos

# IC avançada e IC refractária: de que populações falamos?

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# HF: epidemiology



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**26**  
**million**

Number of heart failure patients worldwide<sup>1</sup>

**1–2%**

Health care expenditure attributed to heart failure  
in Europe and North America<sup>2</sup>

**74%**

Heart failure patients suffering from at least 1 co-morbidity:  
more likely to worsen the patient's overall health status<sup>3</sup>

1. Ambrosy PA et al. J Am Coll Cardiol 2014;63:1123–1133

2. Cowie MR et al. Improving care for patients with acute heart failure. 2014. Oxford PharmaGenesis. ISBN 978-1-903539-12-5.

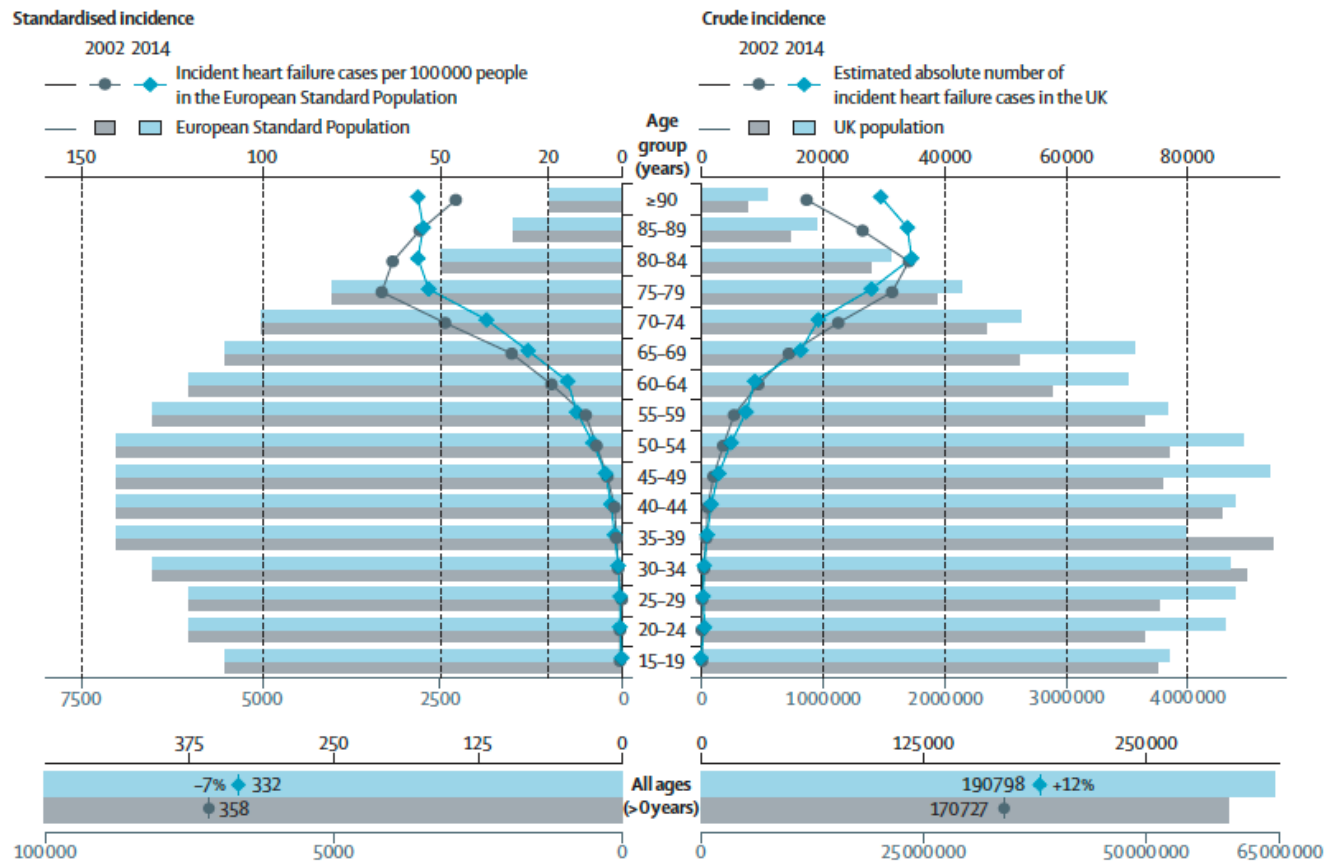
3. van Deursen VM et al. Eur J Heart Fail 2014;16:103–111.

# Incidence is rising



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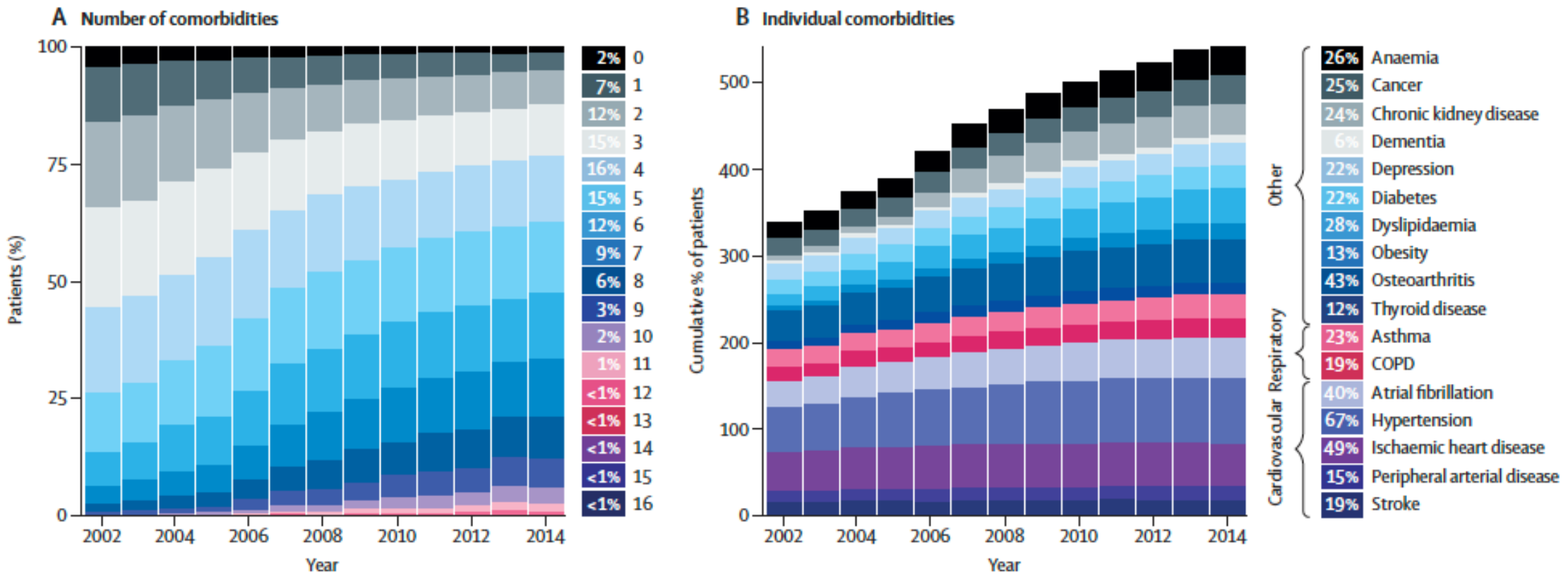


Conrad et al. *Lancet* 2018; 391: 572–80

# Patients have more co-morbidities than ever before



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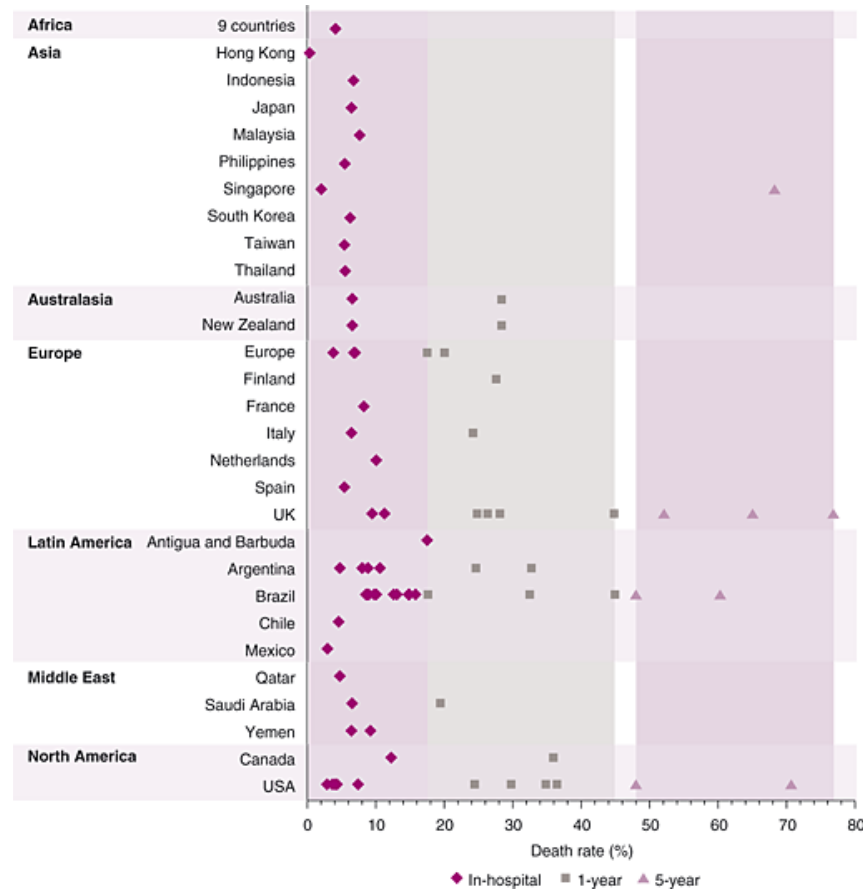


Conrad et al. *Lancet* 2018; 391: 572–80

# Deadly no matter where you live



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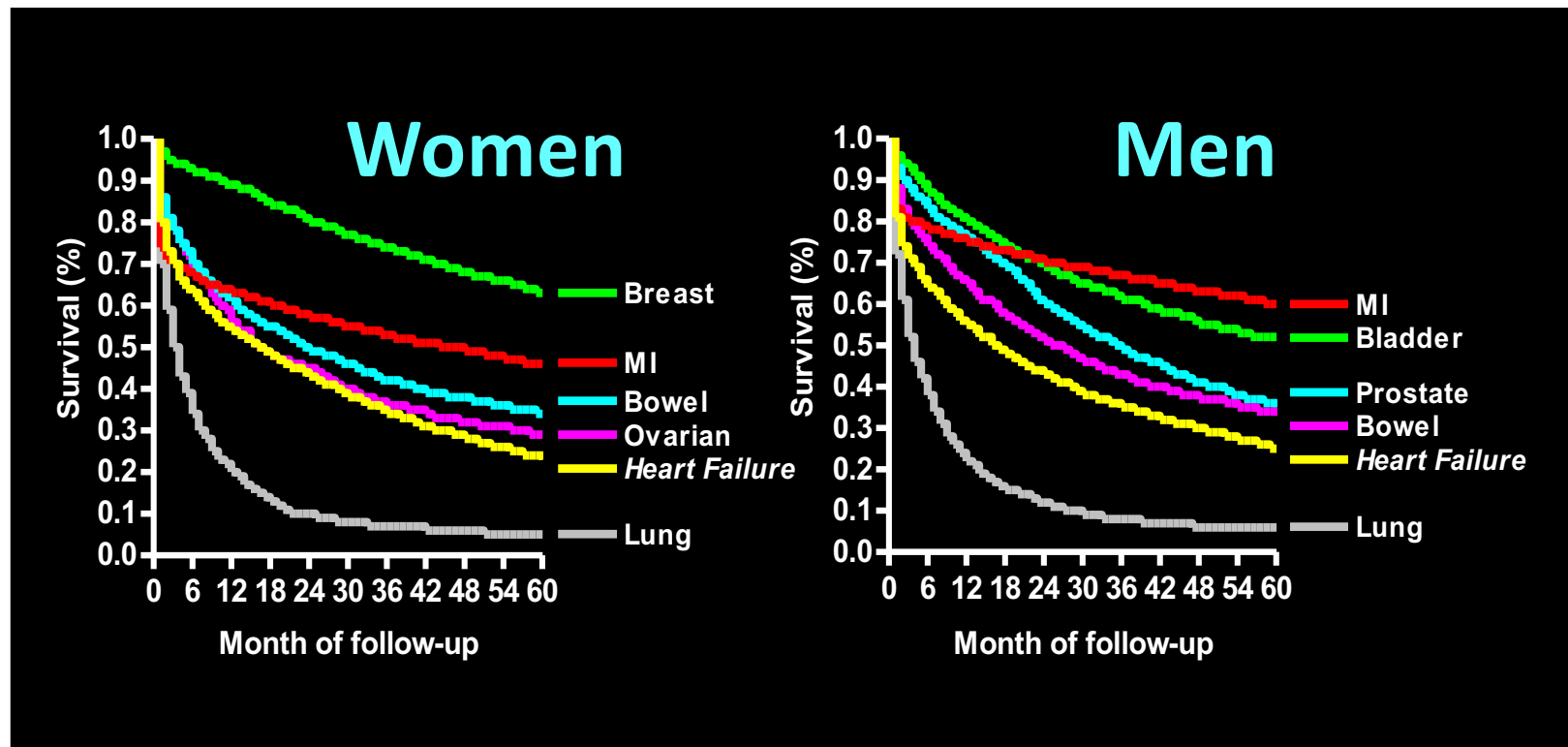
- 17–45% of patients die within 1 year of admission
- majority die within 5 years of admission

Ponikowski et al. *ESC Heart Failure* 2014; 1: 4–25

# More malignant than cancer ?



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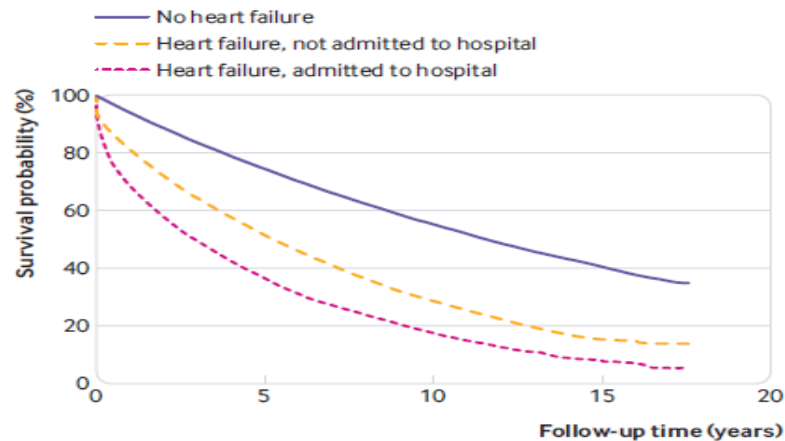


Stewart S et al. *Eur J Heart Fail.* 2001;3(3):315-22.

# We are not doing much better

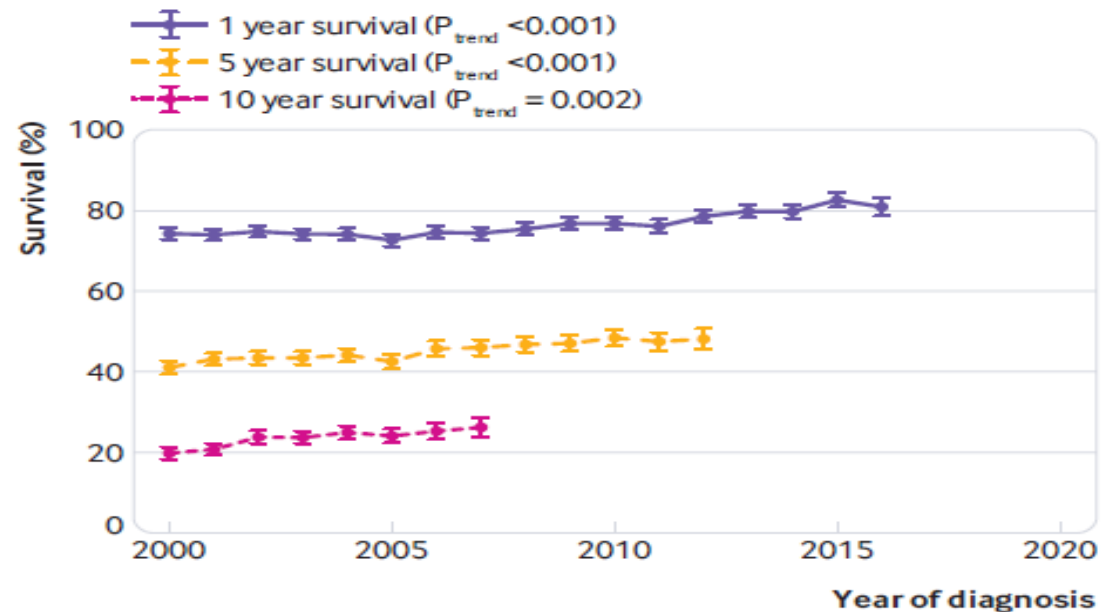


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## No at risk (number censored)

|                                         |           |           |           |  |
|-----------------------------------------|-----------|-----------|-----------|--|
| No heart failure                        |           |           |           |  |
| 278 679                                 | 105 331   | 31 494    | 3 130     |  |
| (0)                                     | (120 343) | (175 326) | (199 292) |  |
| Heart failure, not admitted to hospital |           |           |           |  |
| 31 834                                  | 9 641     | 2 457     | 185       |  |
| (0)                                     | (9 298)   | (13 301)  | (14 920)  |  |
| Heart failure, admitted to hospital     |           |           |           |  |
| 24 125                                  | 4 170     | 729       | 40        |  |
| (0)                                     | (7 505)   | (9 457)   | (9 923)   |  |



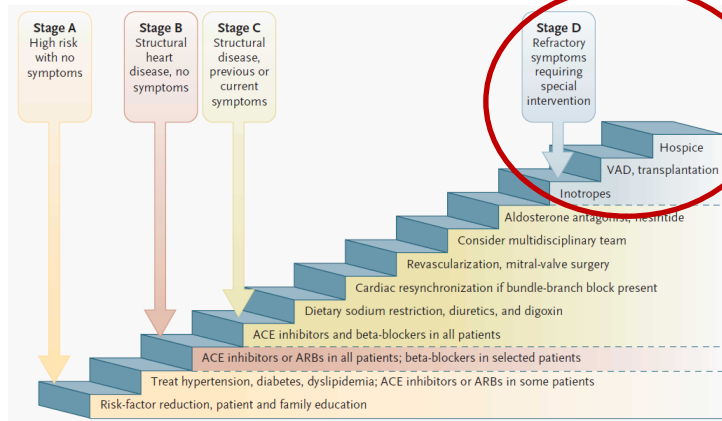
Taylor et al. BMJ 2019; 364: l223

# What is advanced Heart Failure?



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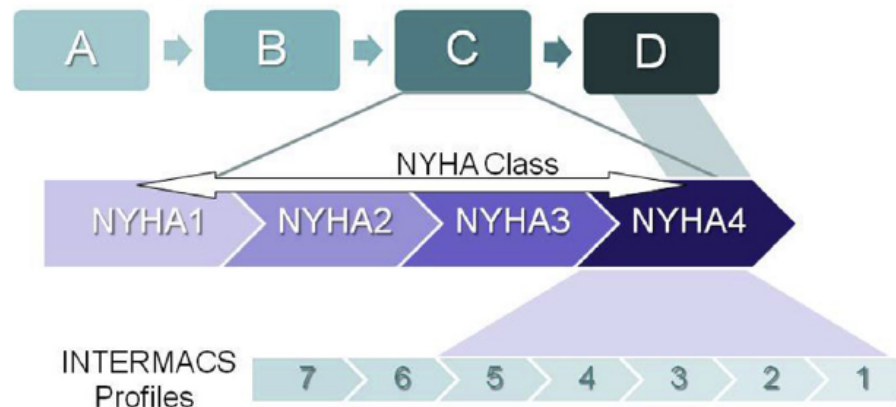


|                  |                                                                                                                                                                  |
|------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <b>Class I</b>   | No limitation of physical activity. Ordinary physical activity does not cause undue breathlessness, fatigue, or palpitations.                                    |
| <b>Class II</b>  | Slight limitation of physical activity. Comfortable at rest, but ordinary physical activity results in undue breathlessness, fatigue, or palpitations.           |
| <b>Class III</b> | Marked limitation of physical activity. Comfortable at rest, but less than ordinary physical activity results in undue breathlessness, fatigue, or palpitations. |
| <b>Class IV</b>  | Unable to carry on any physical activity without discomfort. Symptoms at rest can be present. If any physical activity is undertaken, discomfort is increased.   |

| Interagency Registry for Mechanically Assisted Circulatory Support patient profiles |                                                                                        |
|-------------------------------------------------------------------------------------|----------------------------------------------------------------------------------------|
| Patient Profile                                                                     | Official Shorthand                                                                     |
| 1                                                                                   | "Crash and burn"                                                                       |
| 2                                                                                   | "Sliding fast" on inotropes                                                            |
| 3                                                                                   | "Stable" on continuous inotropes                                                       |
| 4                                                                                   | Resting symptoms on oral therapy at home                                               |
| 5                                                                                   | "Housebound", comfortable at rest but symptoms with minimum activities of daily living |
| 6                                                                                   | "Walking wounded", activities of daily living possible by meaningful activity limited  |
| 7                                                                                   | Advanced NYHA class III                                                                |

**HFSA 2010:** presence of progressive and/or persistent severe symptoms of HF despite optimized medical, surgical and device therapy

AHA/ACC Stages



1. The criteria committee NYHA. 1994 p.253
2. Stevenson et al. JHLT 2009; 28 (6):535-41
3. Fang et al. J Card Fail 2015; 21 (6): 519
4. Chaudry et al. HF clin 2016 (12) 323-333

# Need of a new definition



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- HFPEF
- New therapies: CRT, Ivabrad, Sac/vals
- Ambulatory IV treatment
- Prognostic evidence of arrhythmias
- Increased attention to comorbidities
- Low reference

## **Advanced heart failure: a position statement of the Heart Failure Association of the European Society of Cardiology**

**Maria G. Crespo-Leiro<sup>1\*</sup>, Marco Metra<sup>2</sup>, Lars H. Lund<sup>3</sup>, Davor Milicic<sup>4</sup>, Maria Rosa Costanzo<sup>5</sup>, Gerasimos Filippatos<sup>6</sup>, Finn Gustafsson<sup>7</sup>, Steven Tsui<sup>8</sup>, Eduardo Barge-Caballero<sup>1</sup>, Nicolaas De Jonge<sup>9</sup>, Maria Frigerio<sup>10</sup>, Righab Hamdan<sup>11</sup>, Tal Hasin<sup>12</sup>, Martin Hülsmann<sup>13</sup>, Sanem Nalbantgil<sup>14</sup>, Luciano Potena<sup>15</sup>, Johann Bauersachs<sup>16</sup>, Theresa McDonagh<sup>17</sup>, Petar Seferovic<sup>18</sup>, and Frank Ruschitzka<sup>19</sup>**

Eur J Heart Fail. **2018** Nov;20(11):1505-1535.

# Updated HFA-ESC criteria for defining advanced heart failure



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## Symptoms

## Cardiac dysfunction

## Decompensation

## Impaired exercise capacity

All the following criteria must be present despite optimal guideline-directed treatment:

1. Severe and persistent symptoms of heart failure [NYHA class III (advanced) or IV].
2. Severe cardiac dysfunction defined by a reduced LVEF  $\leq 30\%$ , isolated RV failure (e.g. ARVC) or non-operable severe valve abnormalities or congenital abnormalities or persistently high (or increasing) BNP or NT-proBNP values and data of severe diastolic dysfunction or LV structural abnormalities according to the ESC definition of HFpEF and HFmrEF.
3. Episodes of pulmonary or systemic congestion requiring high-dose intravenous diuretics (or diuretic combinations) or episodes of low output requiring inotropes or vasoactive drugs or malignant arrhythmias causing  $>1$  unplanned visit or hospitalization in the last 12 months.
4. Severe impairment of exercise capacity with inability to exercise or low 6MWT ( $<300$  m) or pVO<sub>2</sub> ( $<12-14$  mL/kg/min), estimated to be of cardiac origin.

In addition to the above, extra-cardiac organ dysfunction due to heart failure (e.g. cardiac cachexia, liver, or kidney dysfunction) or type 2 pulmonary hypertension may be present, but are not required.

Criteria 1 and 4 can be met in patients who have cardiac dysfunction (as described in criterion #2), but who also have substantial limitation due to other conditions (e.g. severe pulmonary disease, non-cardiac cirrhosis, or most commonly by renal disease with mixed aetiology). These patients still have limited quality of life and survival due to advanced disease and warrant the same intensity of evaluation as someone in whom the only disease is cardiac, but the therapeutic options for these patients are usually more limited

Crespo-Leiro et al, Eur J Heart Fail. 2018 Nov;20(11):1505-1535.

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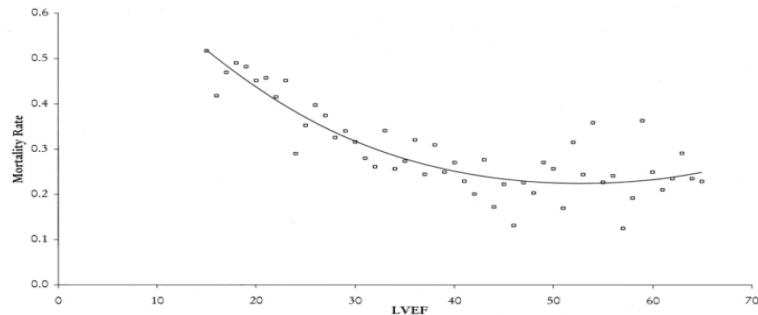
2. Severe cardiac dysfunction defined by
- a reduced LVEF  $\leq 30\%$  and/or
  - isolated RV failure (e.g. ARVC) or
  - non-operable severe valve abnormalities or
  - congenital abnormalities or
  - persistently high (or increasing) BNP or NT-proBNP values and data of severe diastolic dysfunction or LV structural abnormalities according to the ESC definition of HFpEF and HFmrEF.

# EF and mortality

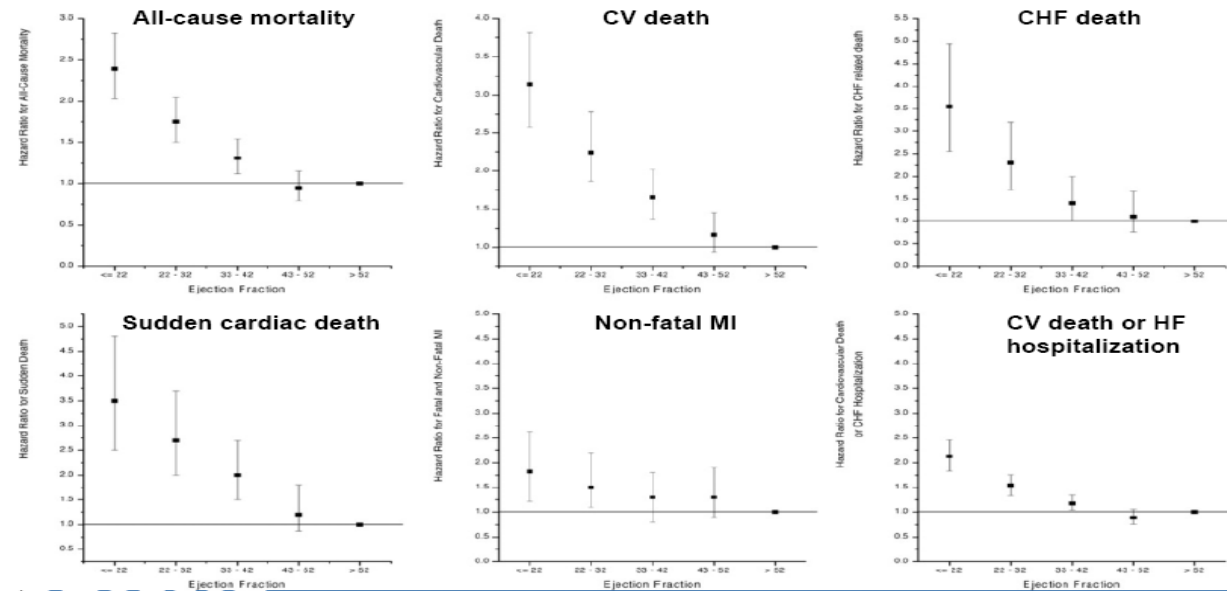


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**The association of LVEF and mortality in stable outpatients with heart failure: DIG trial**



## CHARM: Hazard ratios based on LVEF



Curtis et al. JACC 2003; 42:736-742  
Solomon et al. Circulation 2005; 112:3738-44

# Updated HFA-ESC criteria for defining advanced heart failure



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2. Severe cardiac dysfunction defined by
- a reduced LVEF  $\leq 30\%$  and/or
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# Updated HFA-ESC criteria for defining advanced heart failure



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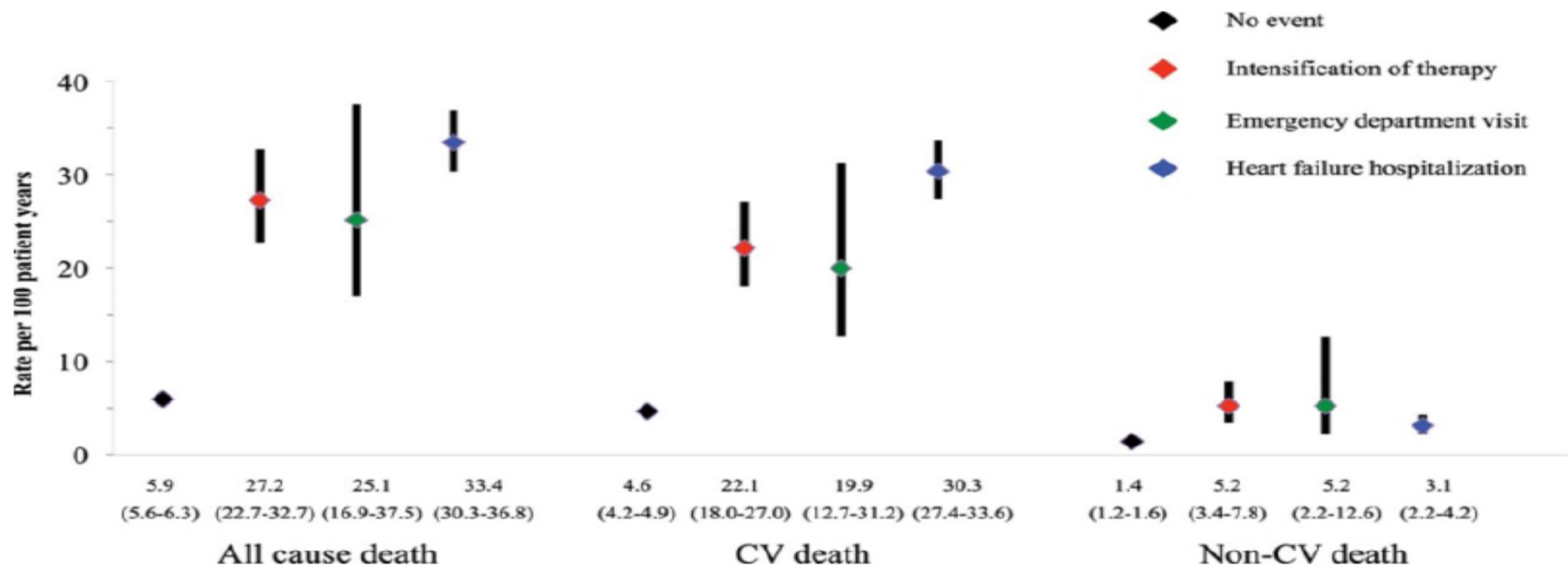
3.

Episodes of pulmonary or systemic congestion requiring high-dose intravenous diuretics (or diuretic combinations) or episodes of low output requiring inotropes or vasoactive drugs or malignant arrhythmias causing >1 unplanned visit or hospitalization in the last 12 months.

# Mortality in patients with decompensation events (PARADIGM)



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Okumura et al. Circulation 2016; 133: 2254-2262

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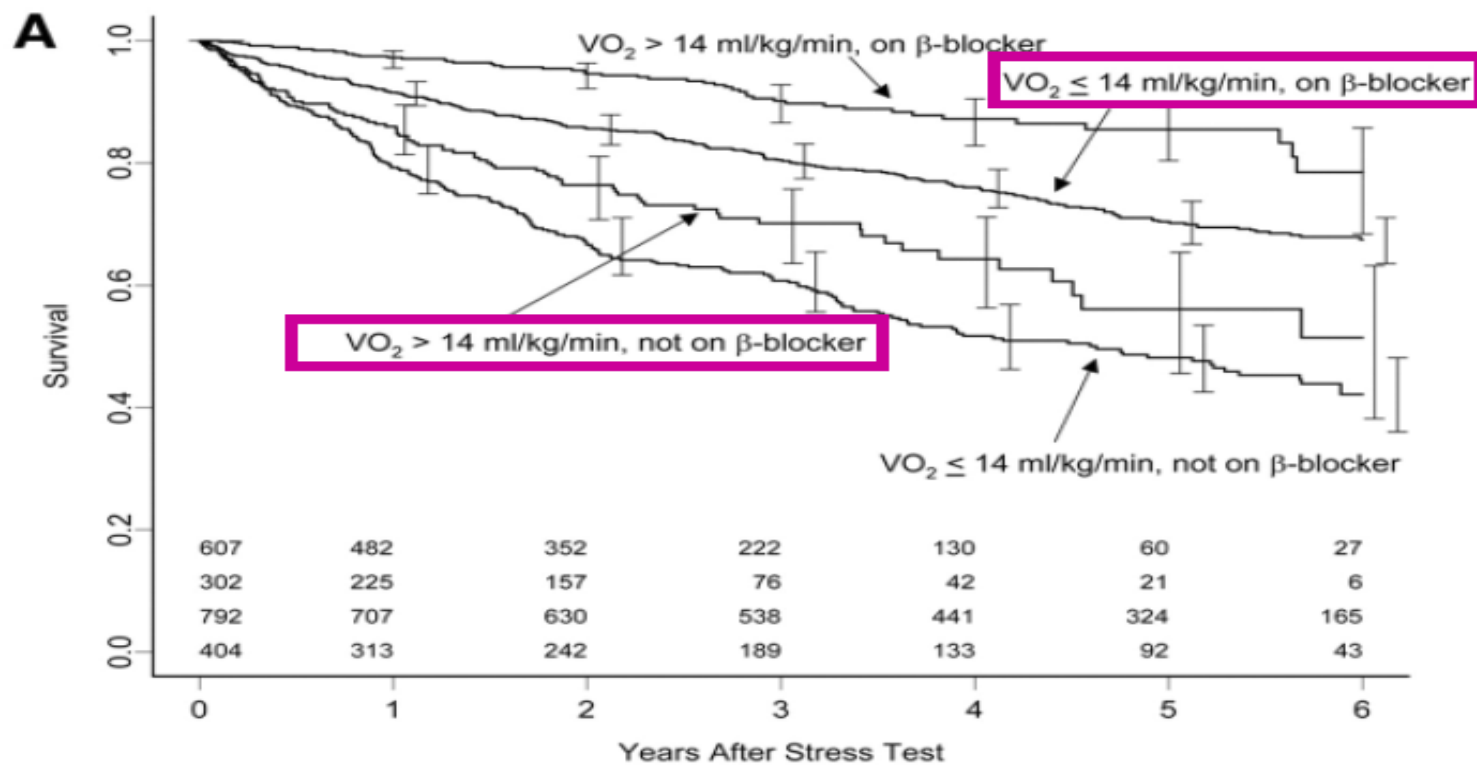
3.

Severe impairment of exercise capacity with inability to exercise or low 6MWT ( $<300$  m) or  $pVO_2$  ( $<12$ – $14$  mL/kg/min), estimated to be of cardiac origin.

# Survival and peak oxygen uptake



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O'Neill et al. Circulation 2005; 111:2313-2318

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In addition to the above, extra-cardiac organ dysfunction due to heart failure (e.g. cardiac cachexia, liver, or kidney dysfunction) or type 2 pulmonary hypertension may be present, but are not required.

# Updated HFA-ESC criteria for defining advanced heart failure



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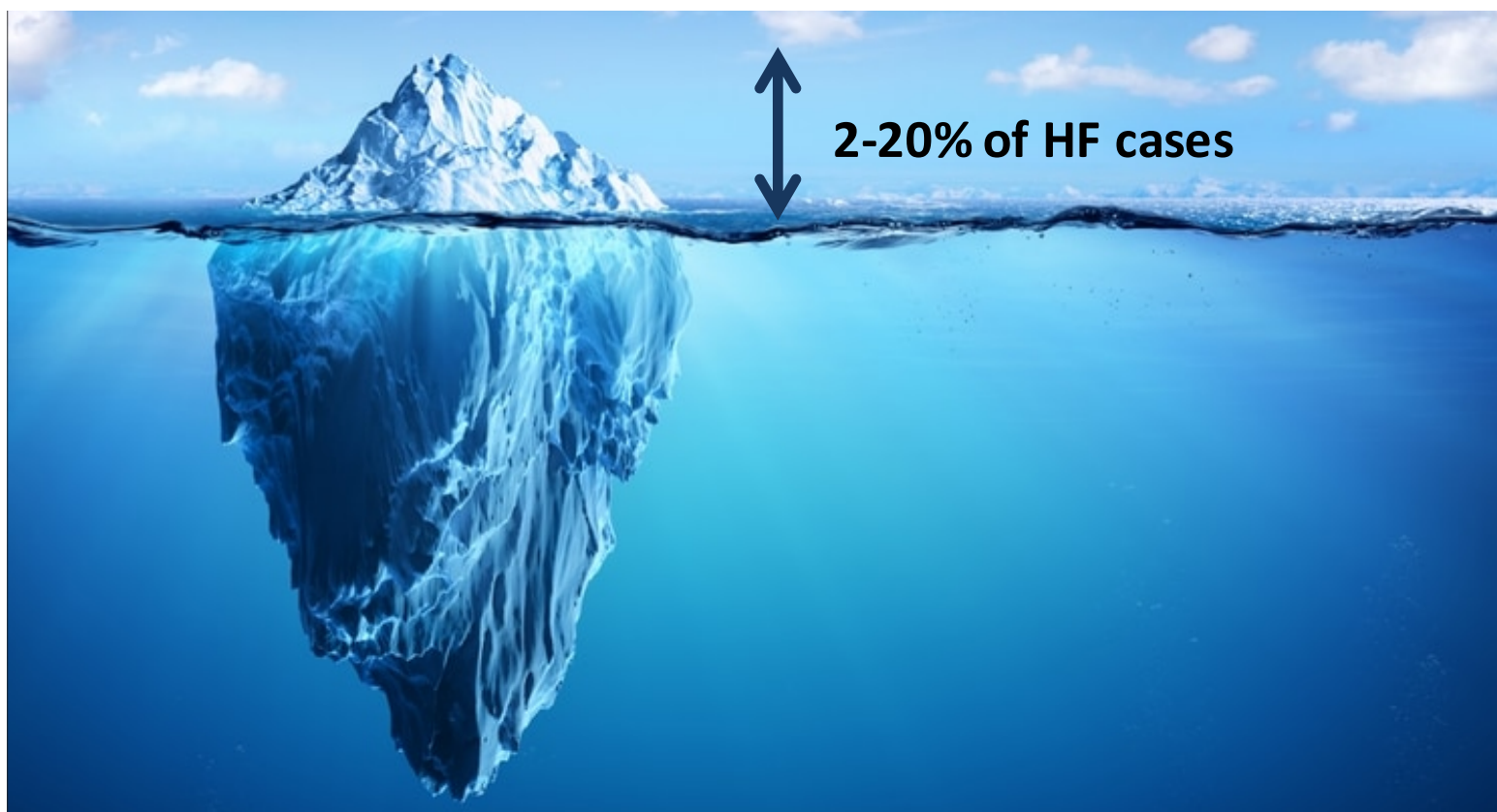
(Symptoms) (Exercise intolerance)  
Criteria 1 and 4 can be met in patients who have cardiac dysfunction (as described in criterion #2), but who also have substantial limitation due to other conditions (e.g. severe pulmonary disease, non-cardiac cirrhosis, or most commonly by renal disease with mixed aetiology). These patients still have limited quality of life and survival due to advanced disease and warrant the same intensity of evaluation as someone in whom the only disease is cardiac, but the therapeutic options for these patients are usually more limited

# What is the prevalence of AHF?



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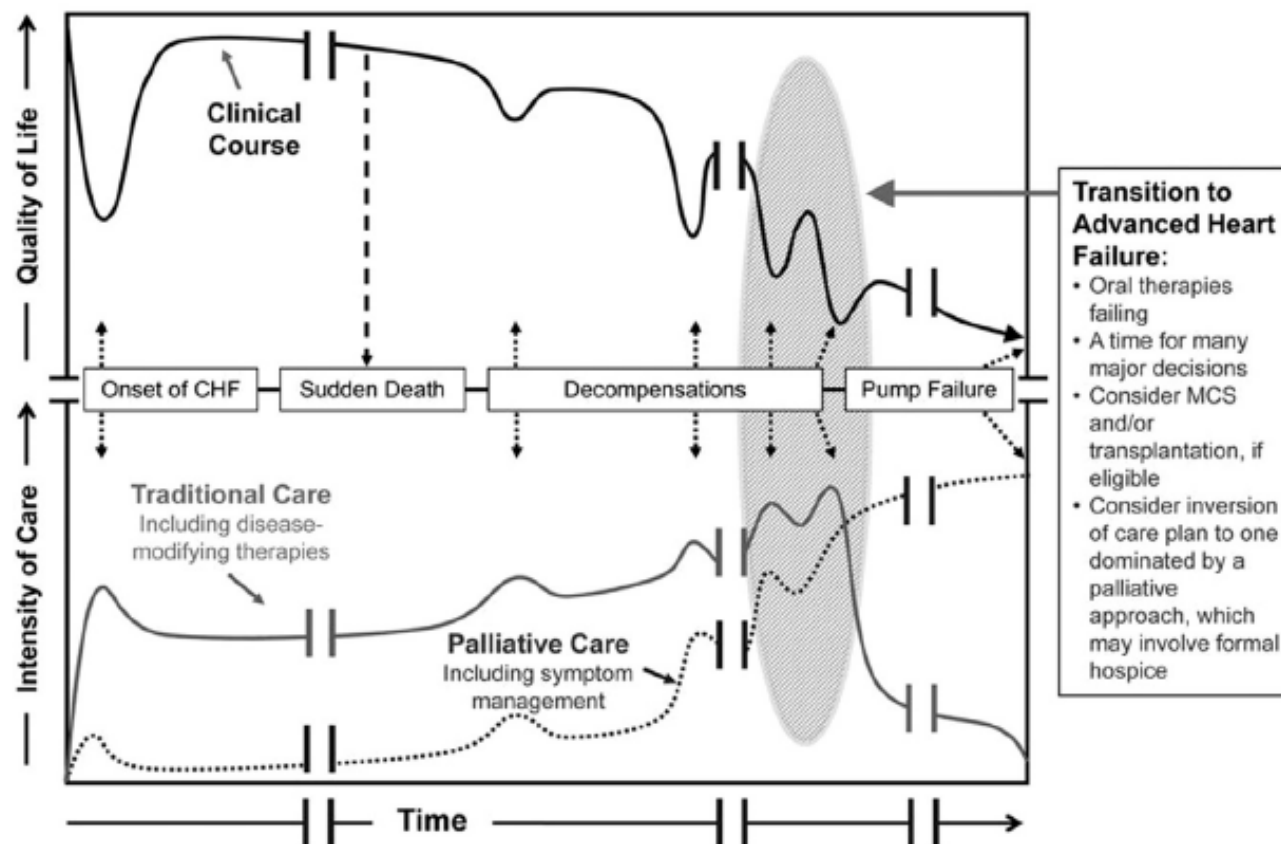


# Why do we need to identify it?



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Allen et al. Circulation 2012; 125 (5) 1930

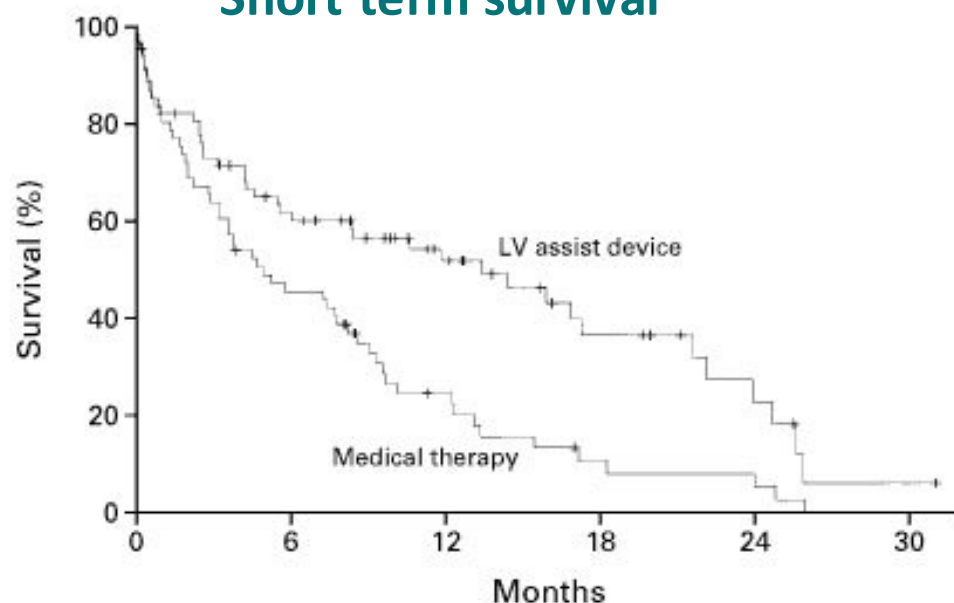
# Why do we need to identify it?



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## Short term survival



No. AT RISK

|                  |    |    |    |    |   |   |
|------------------|----|----|----|----|---|---|
| LV assist device | 68 | 38 | 22 | 11 | 5 | 1 |
| Medical therapy  | 61 | 27 | 11 | 4  | 3 | 0 |

## Low reference

Europe 750 million

HF 15 million

Advanced HF ~1 million

Excluding age >75 and  
comorbidity

**LVAD candidates**  
**>> 100,000**

Rose et al. N Engl Med 2001 Nov 15;345(20):1435-43  
Crespo-Leiro et al, Eur J Heart Fail. 2018 Nov;20(11):1505-1535

# Advanced HF needs organization

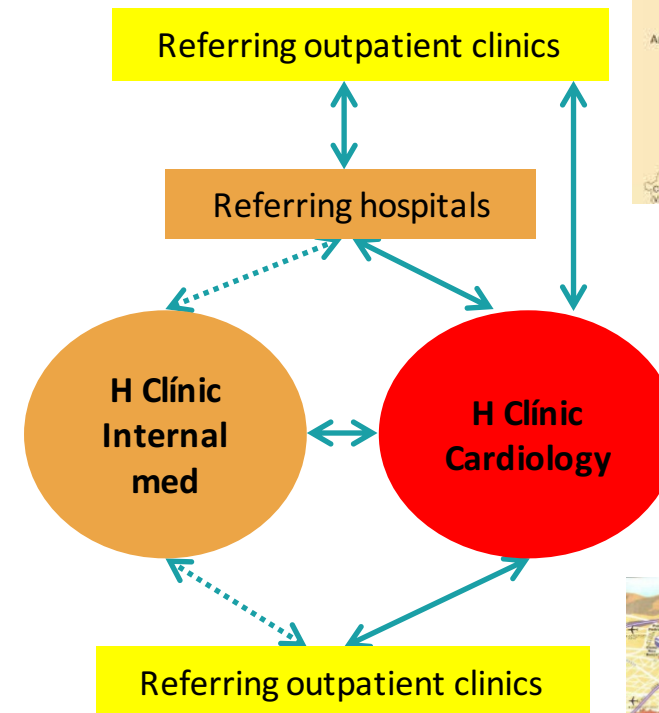


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**Community HF units**

**Specialized HF units**

**Advanced HF units**



- Hospitalization
- Day care clinic
- Specialized outpatient clinics
- Home hospital
- Home day care



Crespo-Leiro et al, Eur J Heart Fail. 2018 Nov;20(11):1505-1535

# Advanced heart failure management



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**Time to decide...**

# Advanced heart failure: needs



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## Multidisciplinary team

HF Cardiologist  
HF Nurse

Other cardiologists  
Internal Medicine  
Cardiac surgeon  
Social Workers  
Dietist  
Psychologist  
Nephrologist  
Pneumologist  
Rehabilitation

## Patient



## Tailored treatment

Optimal treatment (HF drugs,  
iron, inotropes, surgery)

Cardiac rehabilitation

Devices (ICD- CRT, mitra-clip,  
cardioMEMs, REDUCE-LAP, barostim, tricu-  
clip, cardiomodulation... )

Short-term VAD

Long-term LVAD

Heart transplantation

Palliative care

# To take home...



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- HF is a highly prevalent and deadly disease
- Advanced heart failure is present in patients with refractory symptoms despite optimal therapy
- Advanced HF needs to be identified at any level of patient care
- Organization in patient care promotes good referral to an advanced HF unit when needed
- Wide range of therapies to be provided by a multi-disciplinary team: from heart transplant to palliative care



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# Muito obrigada!

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