

Advanced Heart Failure

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Patient LJ

- 52 y/o female with breast Ca
- 5/18 Started chemotherapy including Adriamycin
- 7/18 EF 35% (from 60%), minimally tolerated GDMT
- 8/18 Presented to RMH with decompensated HF, repeat Echo EF 20%, discharged
- Admitted soon after at CCF, low output by PAC, BB discontinued, vasodilators titrated
- 9/2018 despite close follow up: presented with nausea, fatigue, hypotension and signs of CS
- RHC on milrinone 0.125 mcg/kg/min:
 - RA 12, PA 29/18 (22), PCWP 17, CO/CI 3.2/1.7 by Fick, 2.7/1.4 by Thermo
 - SVR 1800, PVR 127, PAPI 0.92, CVP/PCWP 0.7

Thoughts:

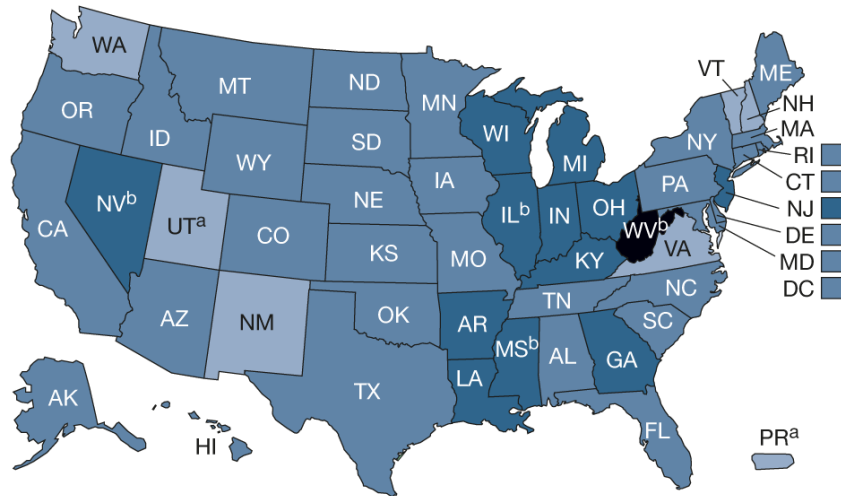
- Persistent low output despite maximally tolerated inotrope (limited by recurrent VT)
- End stage heart failure, non dischargeable (IABP and nipride are not homegoing options)
- Not a transplant candidate due to recent breast Ca (needs 5 years in remission)
- *What now?*

Proceed with DT VAD therapy:

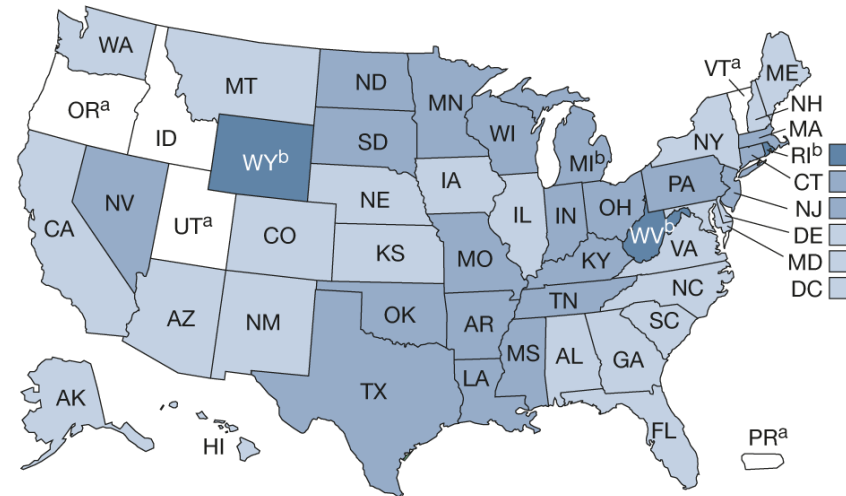
- Received HMIII that admission
- Post op course complicated by RV failure (expected, concerning preop R heart indices: PAPI 0.92, CVP/PCWP 0.7)
- Managed with dual inotropes and iNO (no VT post VAD)
- Patient at our holiday party: Ambulating, NYHA I-II, discussed how she bought her teenage son a new video game chair for Christmas

Burden of Heart Failure

1998 (Mean hospitalization rate, 2845 per 100 000 person-years)



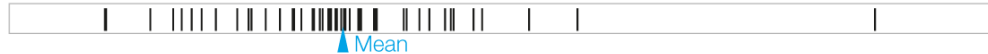
2008 (Mean hospitalization rate, 1957 per 100 000 person-years)



Hospitalization rate per
100 000 person-years



Distribution 1998



2008

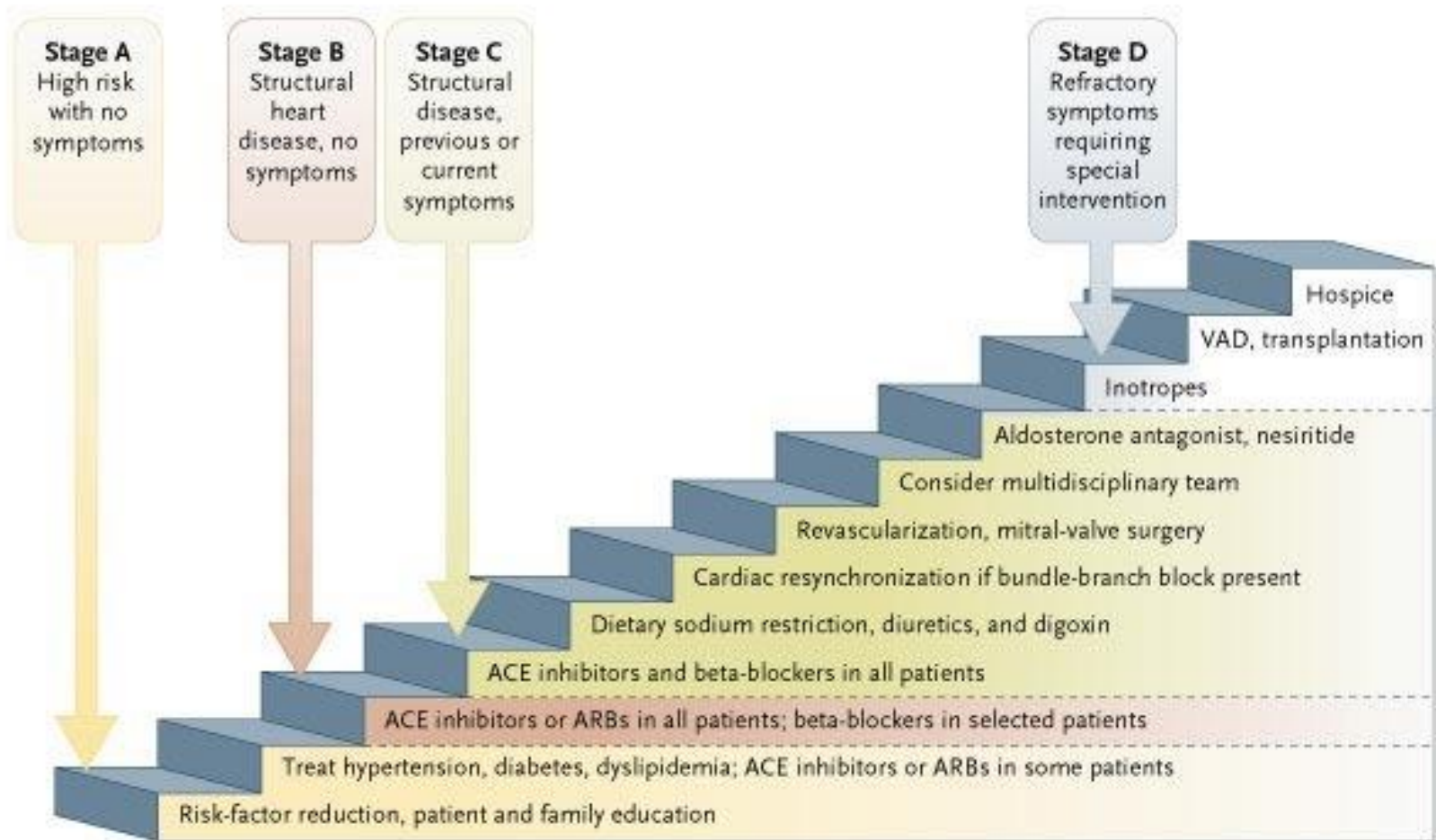


JAMA 2011

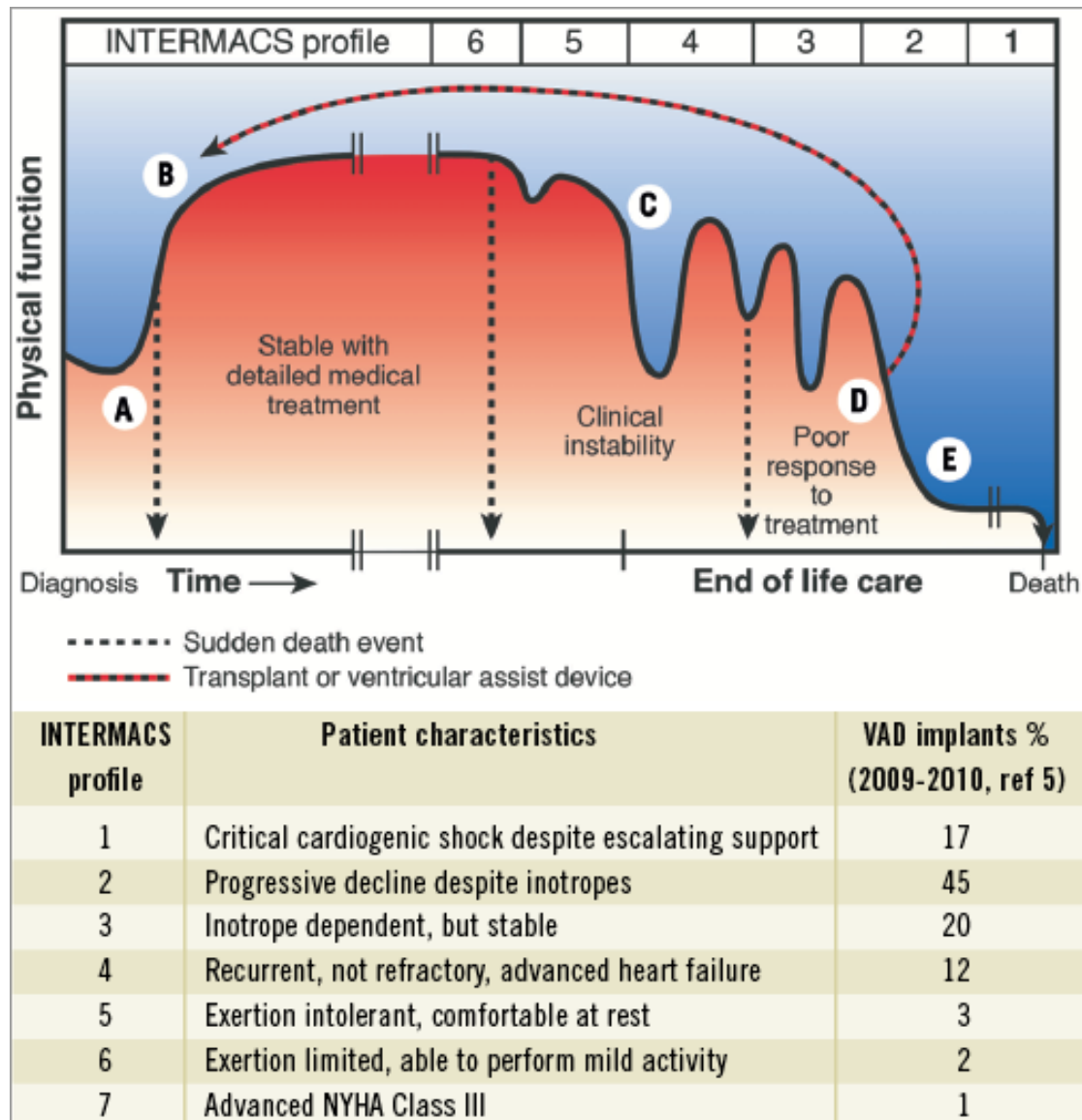
- 26 million patients with heart failure globally
- Prevalence of approximately 6 million in the US alone (2% of the population)
- Almost 4 million hospitalizations in 2004
- Cost/hospitalization of roughly \$11000
- ***Advanced Heart Failure:*** 5% is often quoted (estimates vary due to heterogeneous definitions)

1. Ponikowski et al. ESC Heart Fail 2014; 1: pp. 4-25
2. Chaudry SP, Heart Failure Clinics 2016; 12: pp. 323-3333
3. Komanduri et. al JCHIP March 2017

Recognizing Advanced Heart Failure

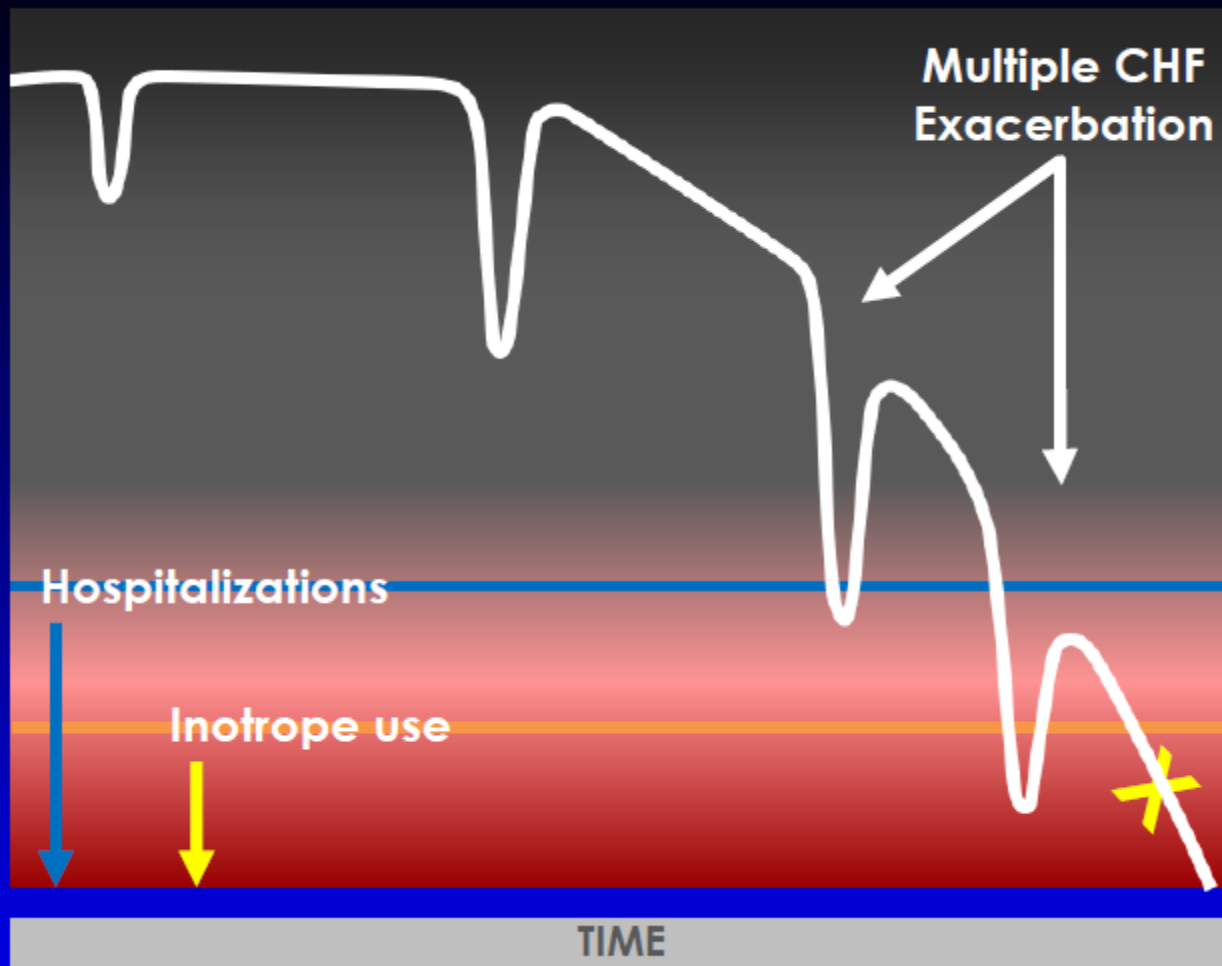


Jessup, NEJM 2003



Goodlin

BETTER
↑
Health Status
↓
WORSE



↑ Diuretics
ACE-i/BB
Intolerance

Timeframe for Definitive Interventions based on INTERMACS classifications

| AHA/ACC classification | Stage C | | | | Stage D | | |
|---------------------------------------|-------------------------------------------------------------------------------------------------|----------------------------------------------------------------|----------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------|------------------------------------------------------|---------------------------------------------|----------------------------------------------|
| NYHA classifications | Class III | | Class IIIb/IV | | Class IV | | |
| INTERMACS levels | <div><div>7</div><div>6</div><div>5</div><div>4</div><div>3</div><div>2</div><div>1</div></div> | | | | | | |
| Brief descriptions | Advanced NYHA Class III | Exertion limited/ "Walking wounded" | Exercise intolerant/ "House-bound" | Recurrent decompensation/ "Frequent flyer" | Stable but inotrope-dependent/ "Dependent stability" | Progressive decline/ "Sliding on inotropes" | Critical cardiogenic shock/ "Crash and burn" |
| Timeframe for definitive intervention | Transplantation or circulatory support <u>not</u> currently indicated | Variable, depends upon nutrition, organ function, and activity | Variable, depends upon nutrition, organ function, and activity | Elective over weeks to months as long as treatment of episodes restores stable baseline, including nutrition | Elective over a few weeks | Needed within a few days | Needed within hours |

Note: This grid was based on the best interpretation of the information provided in the sources listed below
 Sources: "Heart Failure". NEJM 2003; 348:2007-18. "On the Fledgling Field of Mechanical Circulatory Support". JACC 2007; (50) 8.
 "Characteristics of Stage D heart failure: Insights from the Acute Decompensated Heart Failure National Registry Longitudinal Module (ADHERE LM)". Am J Heart 2008; 155:341-9. INTERMACS Manual of Operations version 2.2, User's Guide

ESC AHF Definition

- ***NYHA Class III-IV Symptoms***
- ***Episodes of volume overload and/or peripheral hypoperfusion***
- ***Objective evidence of severe cardiac dysfunction***
(EF<30%, Doppler Pseudonormal or Restrictive filling pattern, PCWP>16mmHg or RAP> 12 mmHg)
- ***Severely impaired functional capacity***
(Inability to exercise, 6MWD<300m, Peak VO₂<12-14 ml/kg/min)
- ***HF Hospitalizations***
(≥1 in past 6 months)
- ***Above occurring despite attempts to optimize diuretics, RAAS antagonists, BB, CRT or in the setting of intolerance to OMT***

Identifying the Advanced Heart Failure Patient:

I-NEED-HELP (also see *Table 6*)

I: IV inotropes

N: NYHA IIIB/IV or persistently elevated natriuretic peptides

E: End-organ dysfunction

E: Ejection fraction $\leq 35\%$

D: Defibrillator shocks

H: Hospitalizations >1

E: Edema despite escalating diuretics

L: Low blood pressure, high heart rate

P: Prognostic medication – progressive intolerance or down-titration of GDMT

Yancy et. al, Optimization of Heart Failure Treatment, Expert Consensus; JACC 2017

Once Stage D is Recognized: *Early Referral is Always Preferred*

-hemodynamically profound RV failure, pulmonary hypertension or profound renal insufficiency often limit advanced therapies

AHF Services

- Pre VAD and Transplant Evaluation
- Mechanical Circulatory Support (temporary, percutaneous, extracorporeal, and durable)
- Cardiogenic Shock Team
- CPET (Cardiopulmonary exercise testing) for risk stratification
 - Peak VO₂ < 14 (or <12 on a beta blocker) have 20% mortality risk at 1 year
- Endomyocardial biopsy
- CardioMEMS
- Clinical trials
- Palliative care services/Hospice transition
- Genetic testing
- AVAILABILITY – Staffed to allow q2week (between physician and APP) follow ups for sick patients
- Easy integration with additional advanced programs (CMR/PET, structural heart, VT ablation)

Our Team:

- 3 AHF Physicians
- 1 VAD Surgeon
- 4 AHF NPs
- 2 VAD coordinators
- HF Pharmacist
- HF coordinator
- HF Clinic RN team
- Social worker
- Dietician
- Palliative Care
- Finance



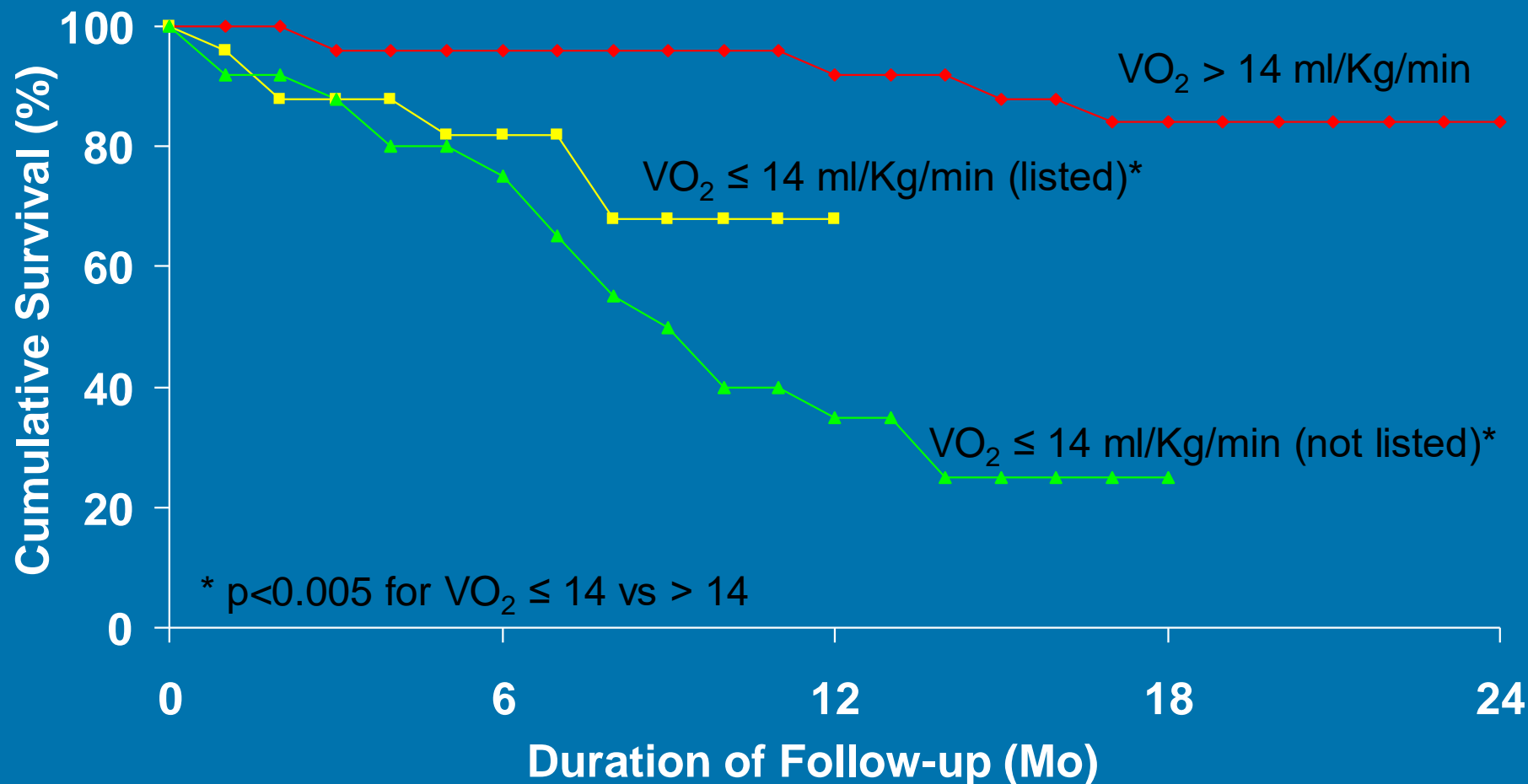
TABLE 1. Functional Impairment During Incremental Treadmill Testing in Heart Failure: The Weber Classification

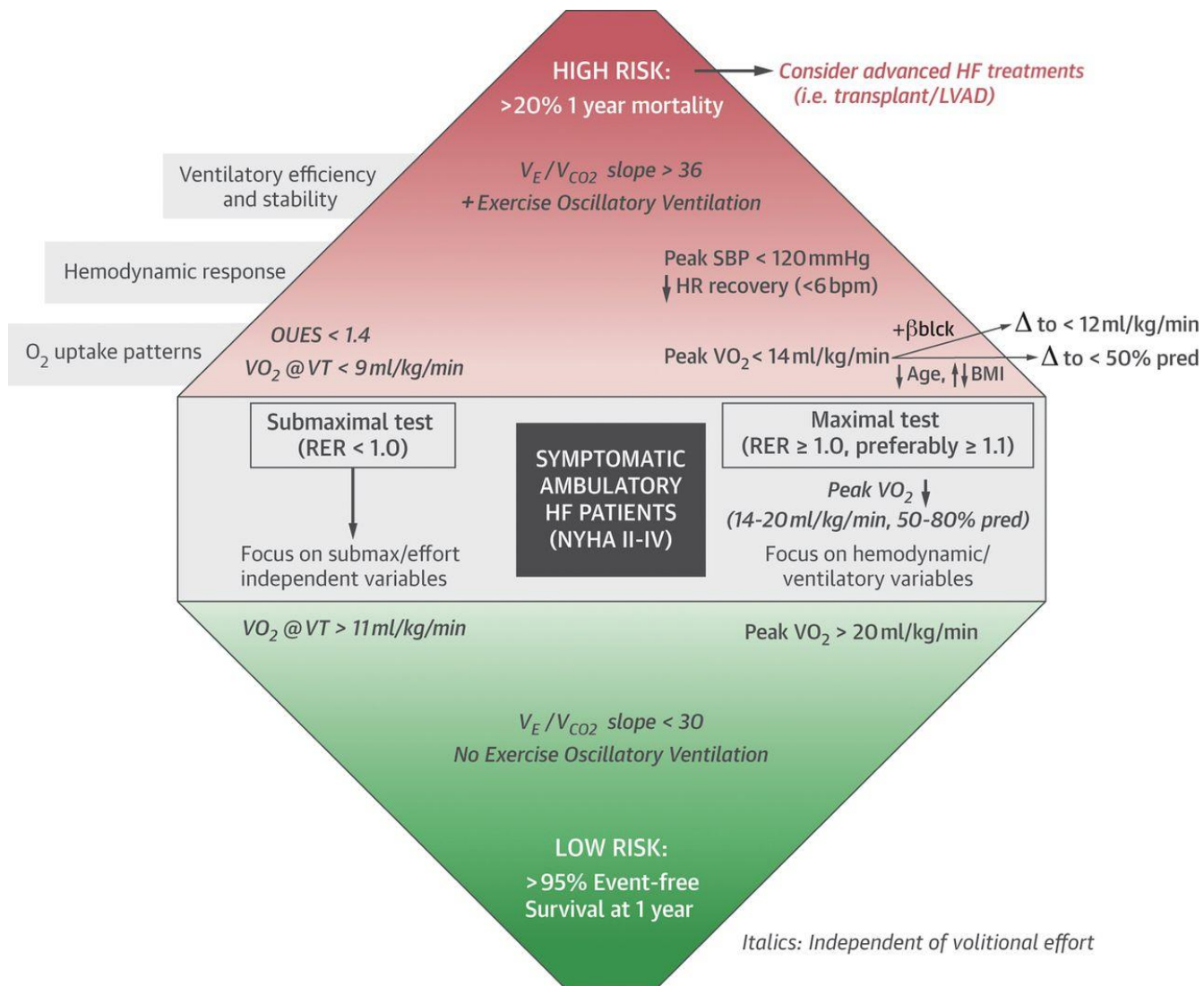
| Class | Severity | Peak $\dot{V}O_2$ $\text{mL} \cdot \text{kg}^{-1} \cdot \text{min}^{-1}$ | VT | CI max, $\text{L} \cdot \text{min}^{-1} \cdot \text{m}^{-2}$ |
|-------|--------------------|-----------------------------------------------------------------------------|-------|-----------------------------------------------------------------|
| A | Mild to none | >20 | >14 | >8 |
| B | Mild to moderate | 16–20 | 11–14 | 6–8 |
| C | Moderate to severe | 10–16 | 8–11 | 4–6 |
| D | Severe | 6–10 | 5–8 | 2–4 |
| E | Very severe | <6 | <4 | <2 |

VT indicates ventilatory threshold; CI max, maximum cardiac index.

Adapted with permission from Weber et al, "Determination of aerobic capacity and the severity of chronic cardiac and circulation failure." (*Circulation*. 1987;76[suppl VI]:VI-40–VI-45.)

Prognostic Value of peak VO_2

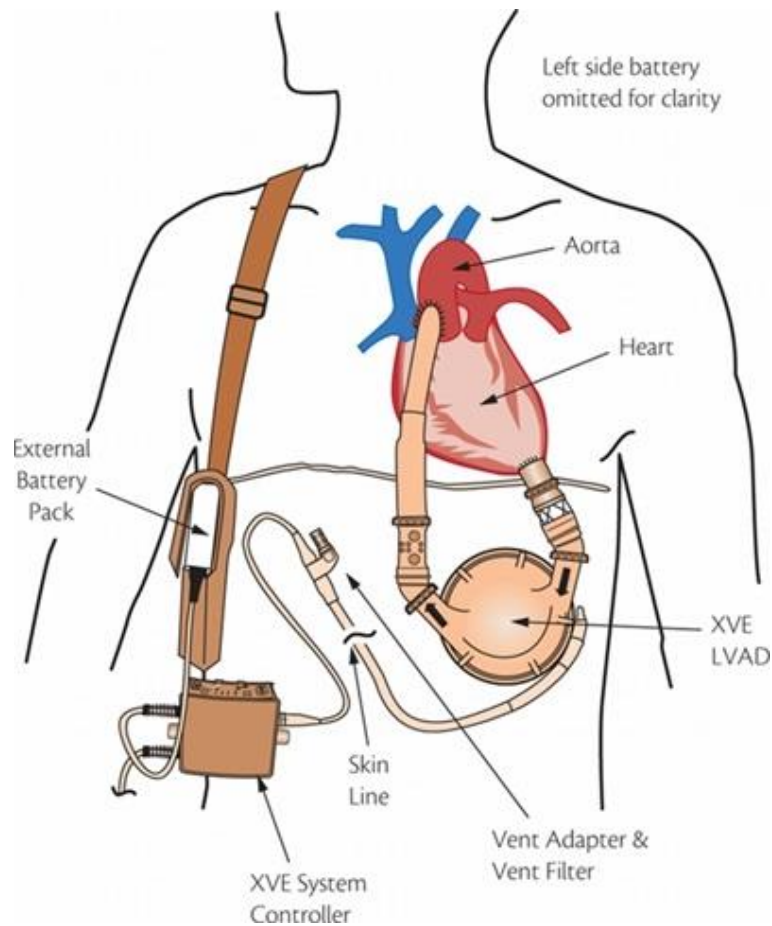




JACC HF 2018

VAD Therapy

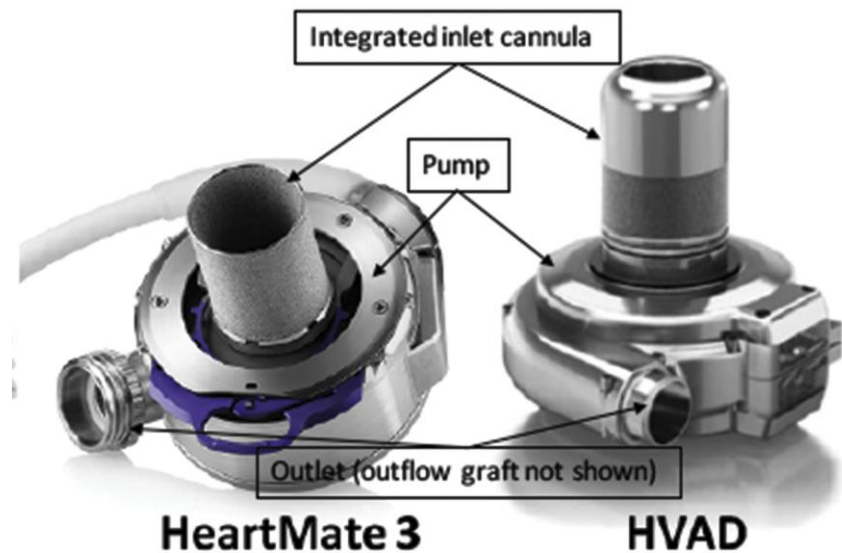
1st Generation VAD: Heartmate HVE

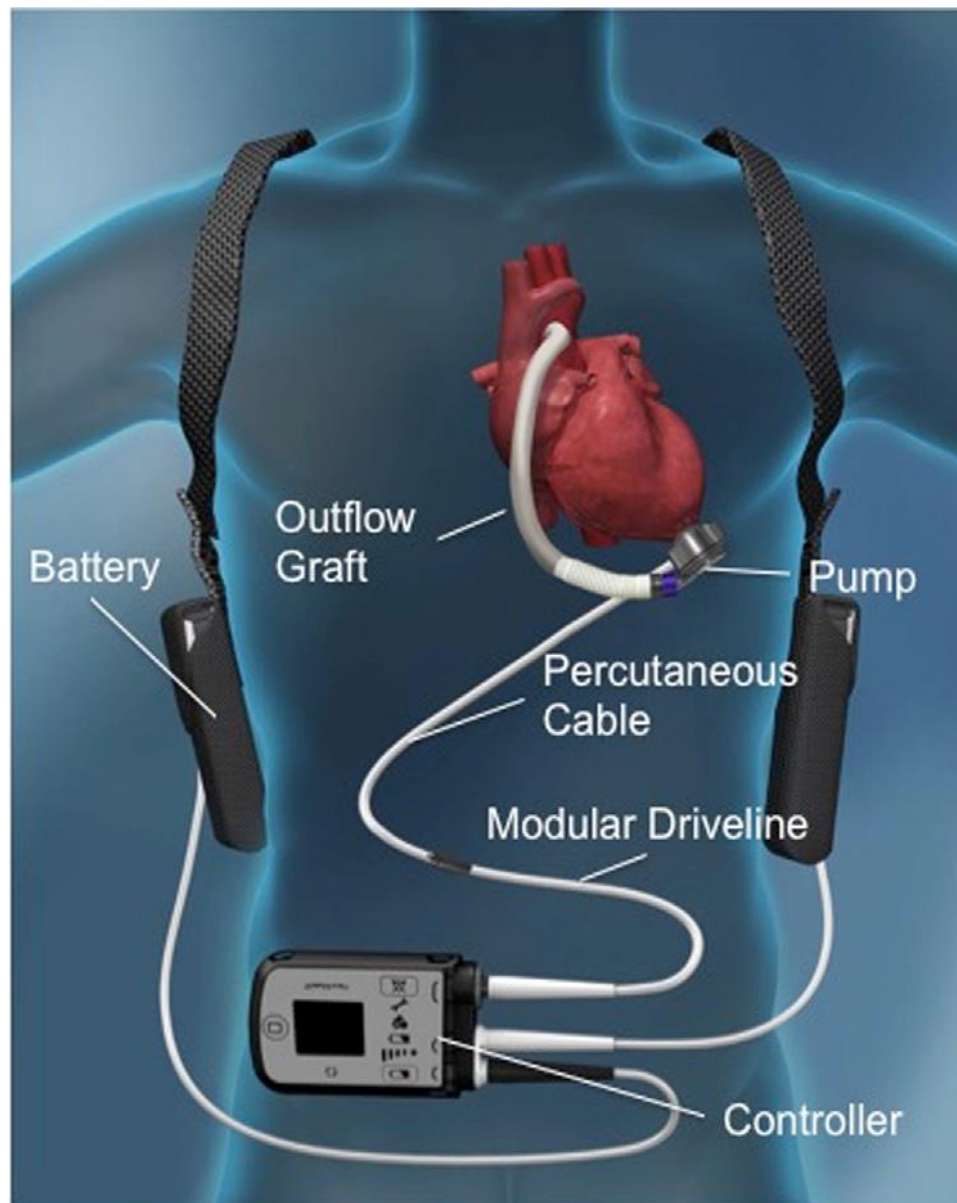


2nd Generation: Heartmate II

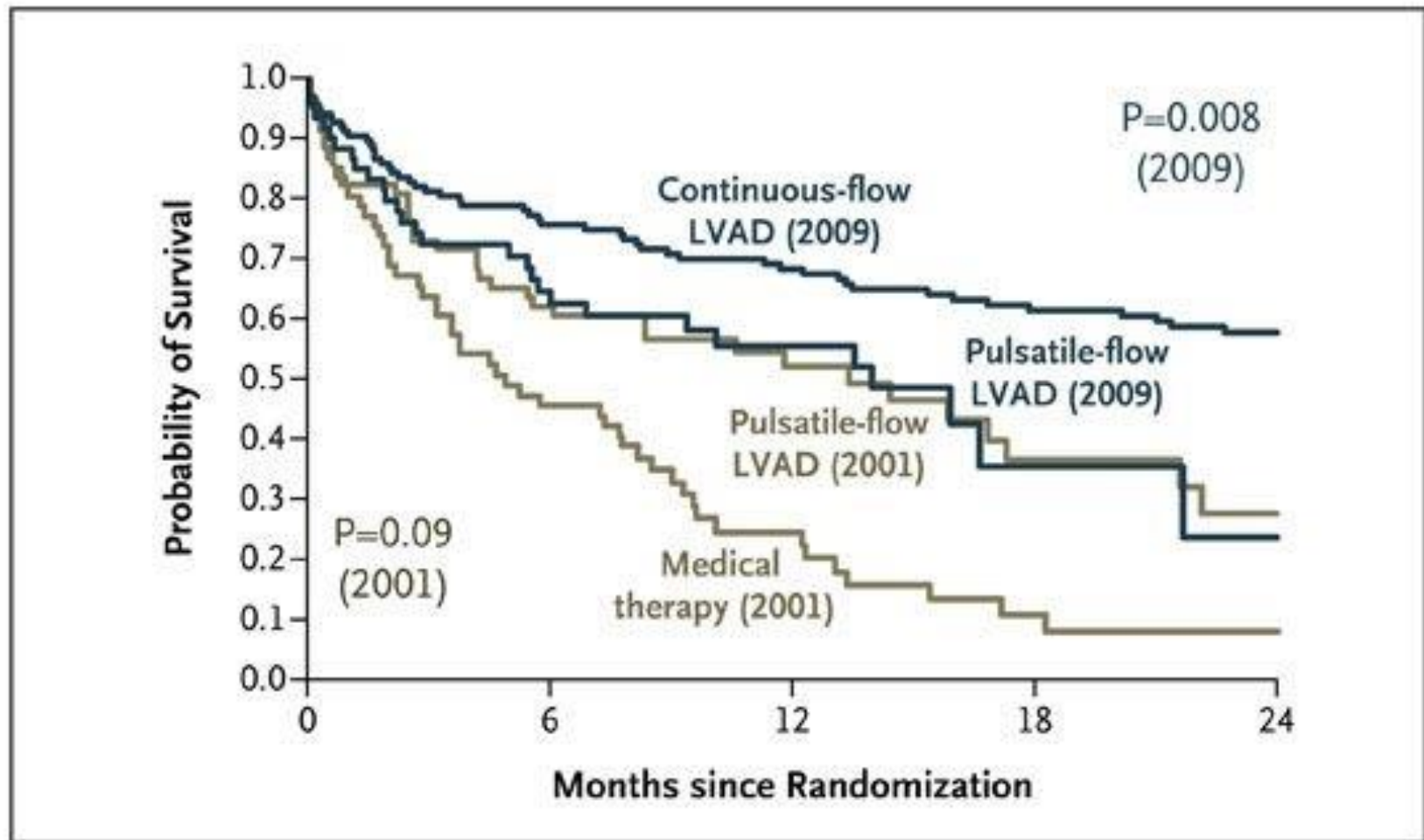


3rd Generation: HVAD and HMIII



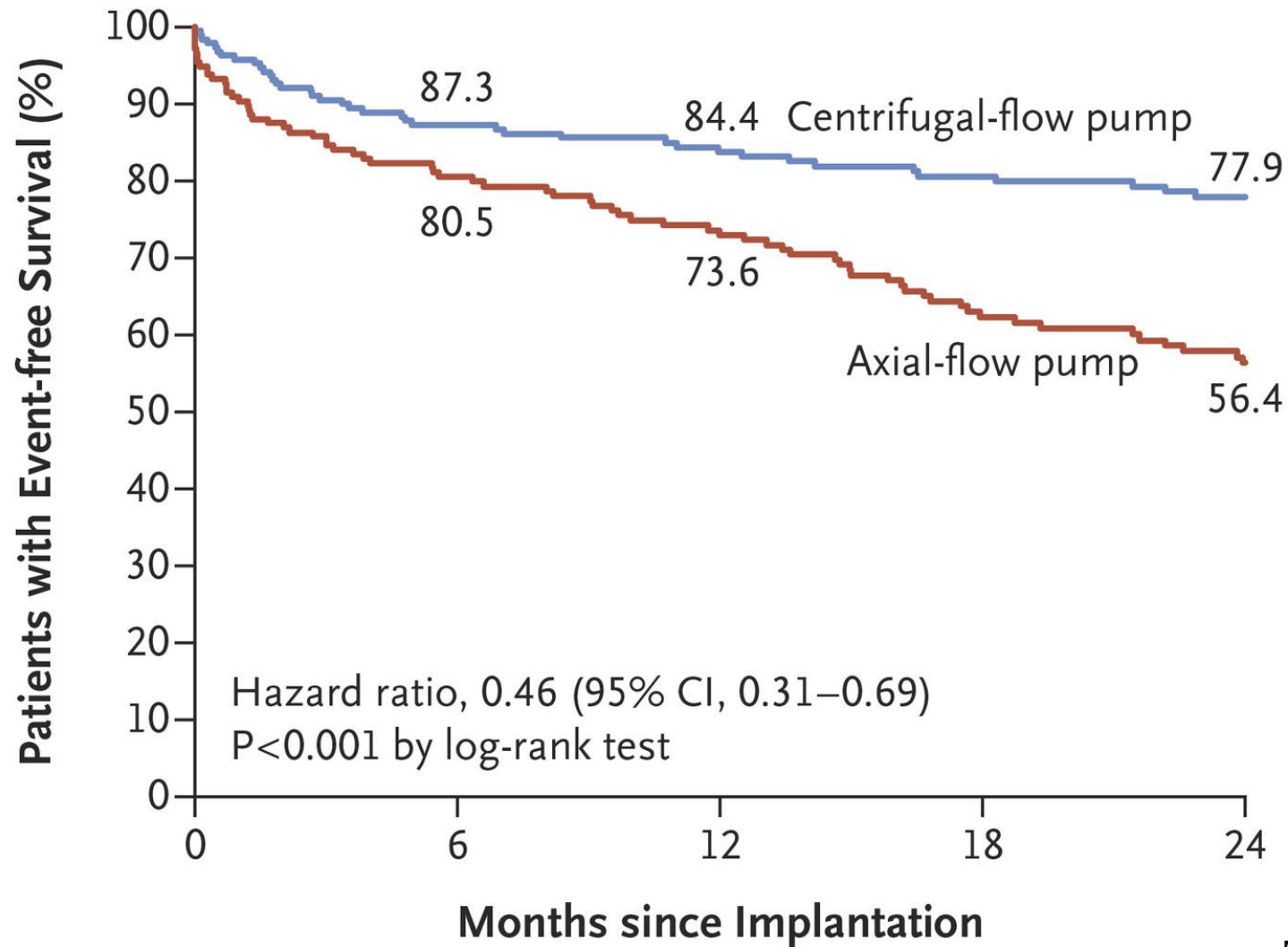


OMT vs CF LVAD



Fang NEJM 2009

HMII vs HMIII



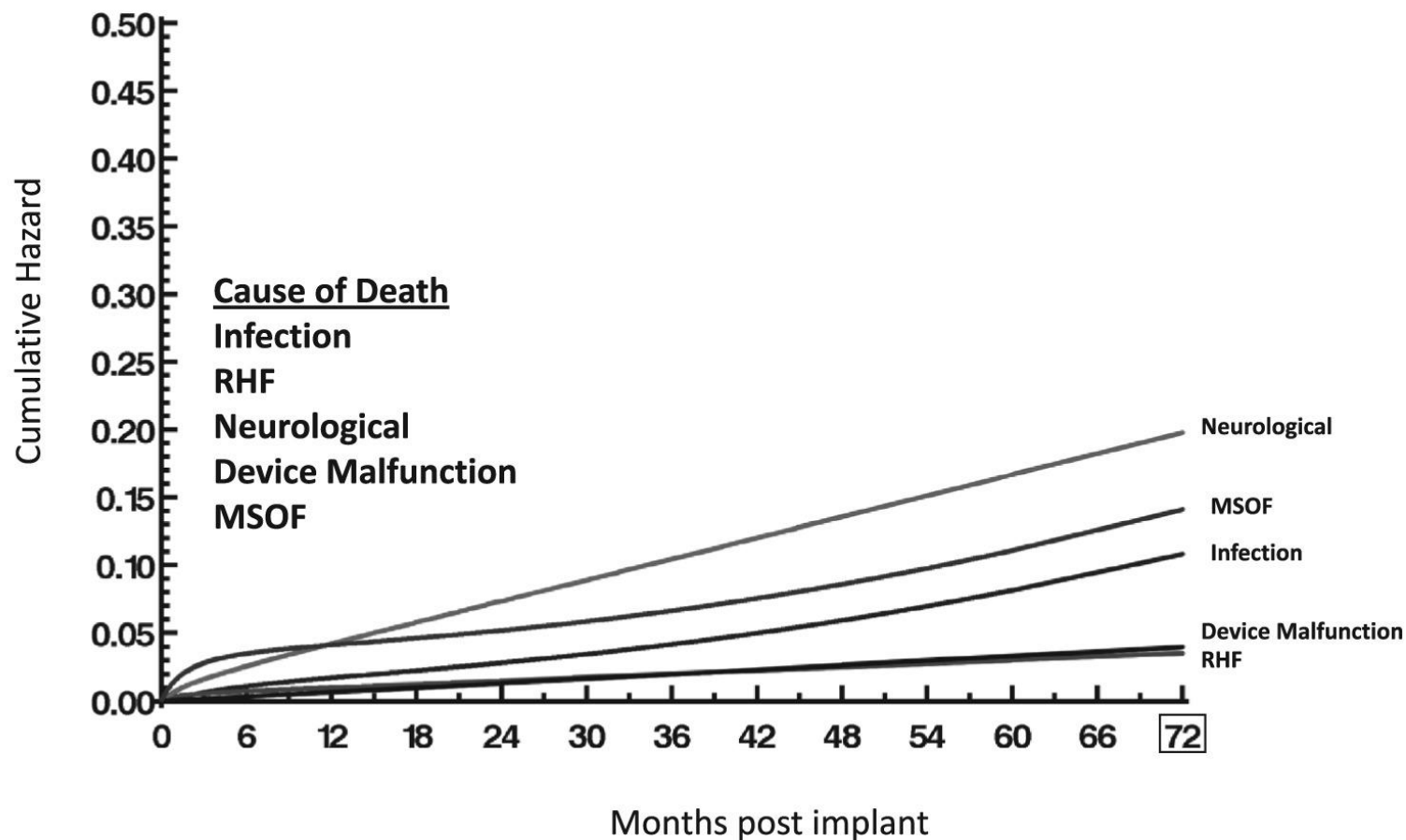
Mehra NEJM 2018

Adverse Effects – SIGNIFICANT AND MANY:

- GI Bleed
- Stroke
- Infection (including chronic driveline)
- Death
- RV failure – early and delayed
- Pump failure/pump thrombosis
- Costs

Intermacs **Continuous Flow LVAD/BiVAD Implants: 2008 – 2016, n=17633**

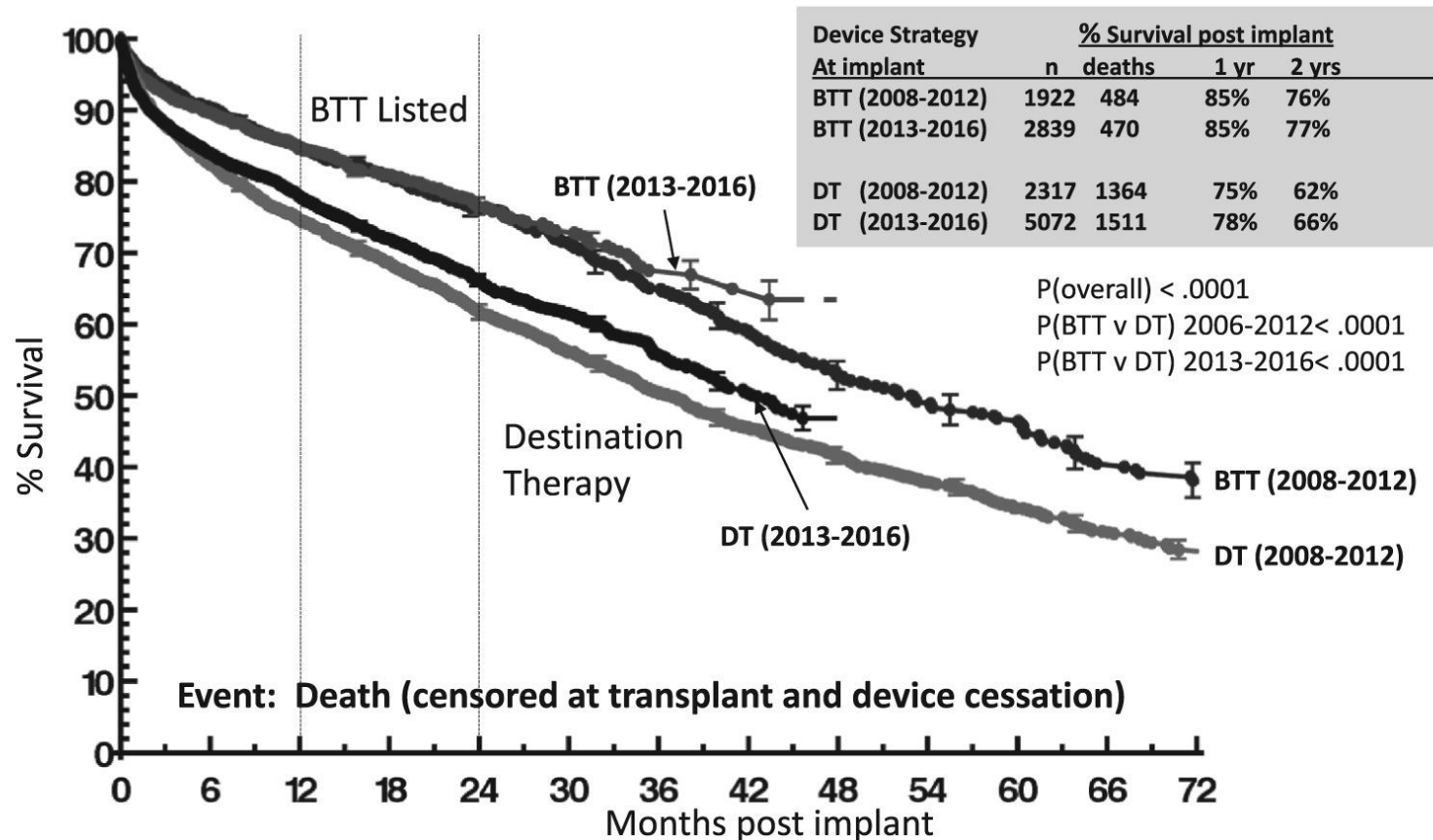
Cumulative Death Rate (Hazard) for selected causes



Kirklin JHLT 2017

Interm@cs Continuous Flow LVAD/BiVAD Implants: 2008 – 2016, n=17633

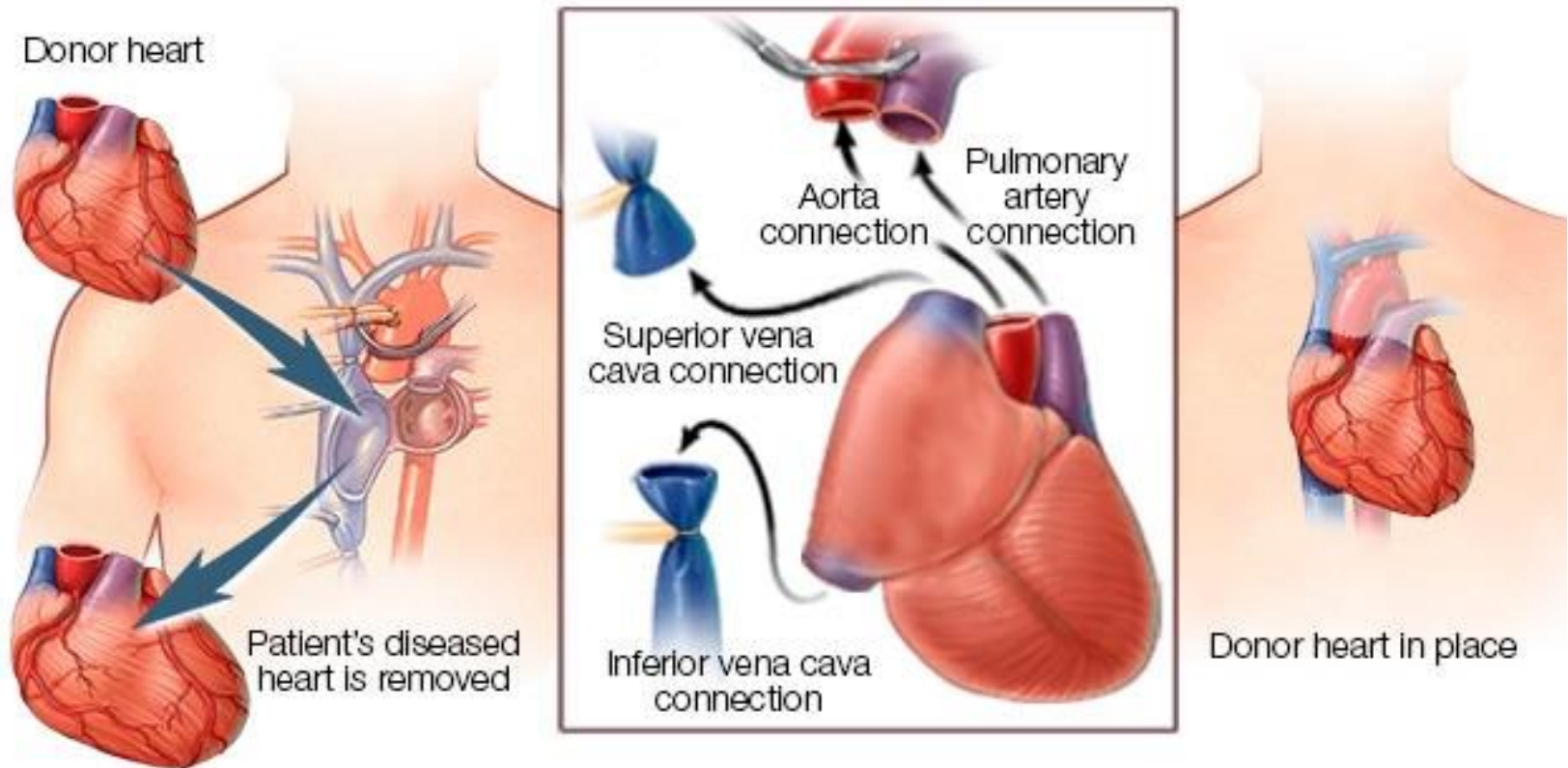
Bridge to Transplant Listed and Destination Therapy by Era (n=12150)



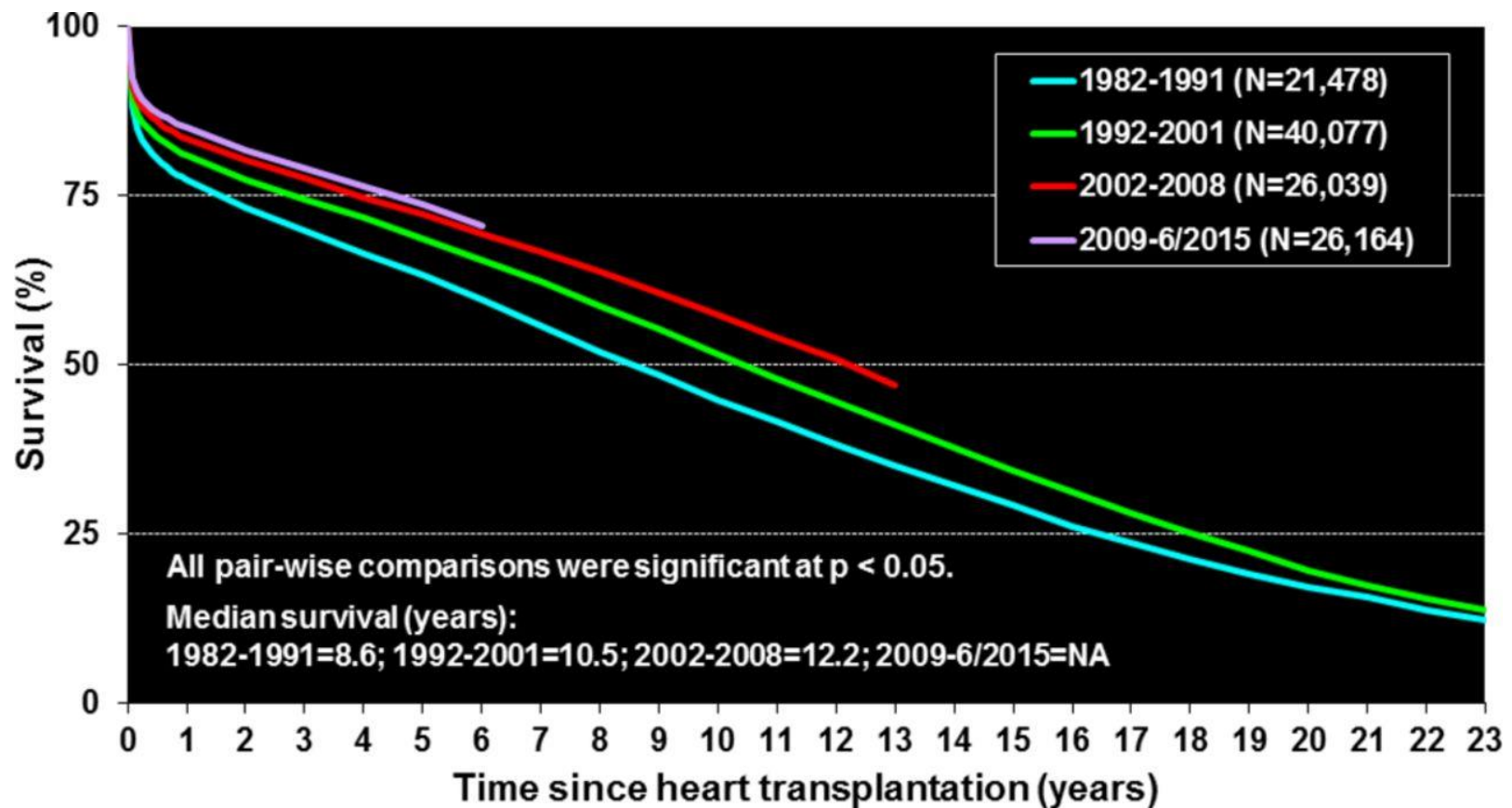
Kirklin JHLT 2017

Cardiac Transplantation

Heart transplant procedure



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BMJ

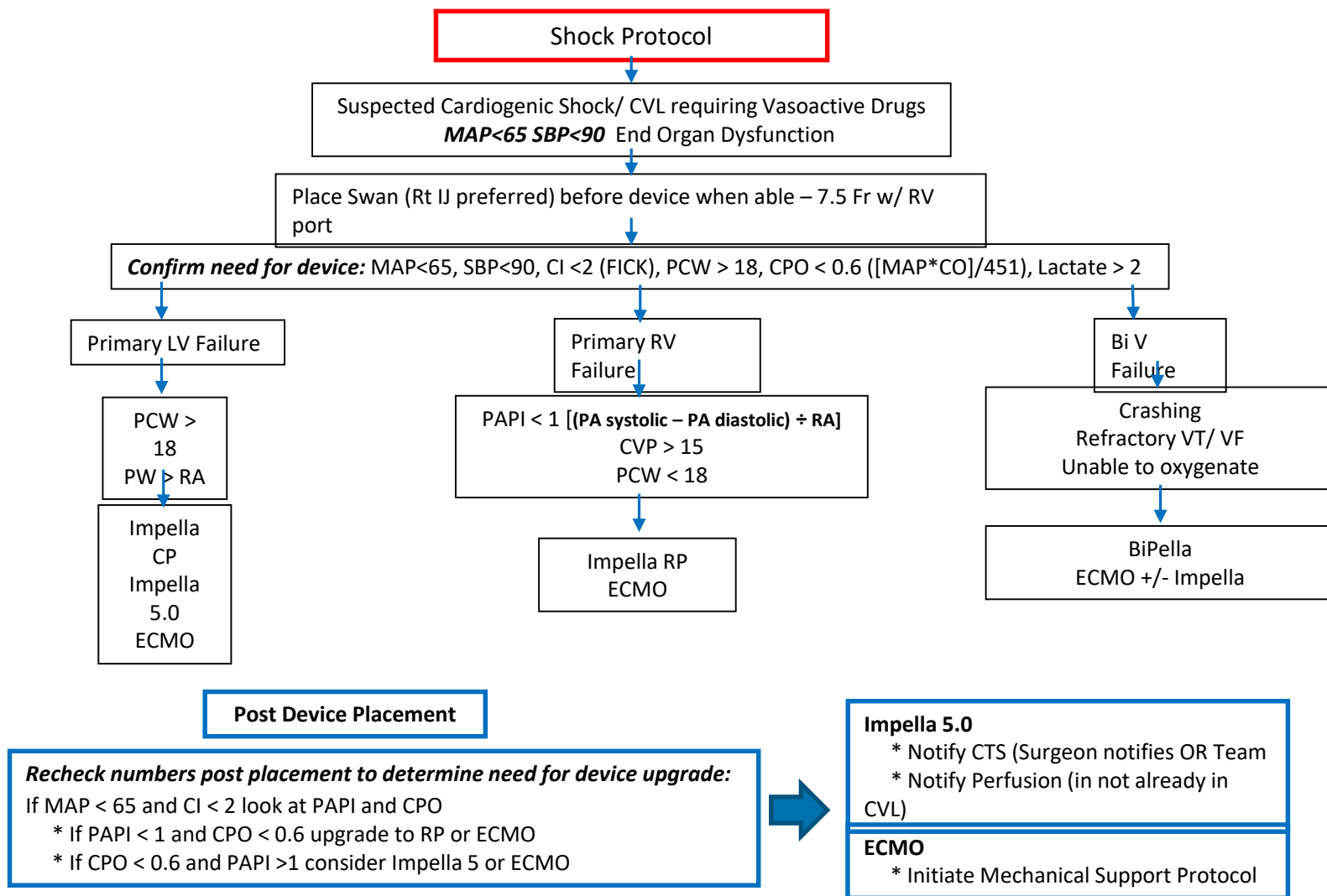
Complications

- Immunosuppression – infection, malignancy, renal failure
- Post op including stroke
- Cardiac allograft vasculopathy
- Cellular and antibody mediated rejection

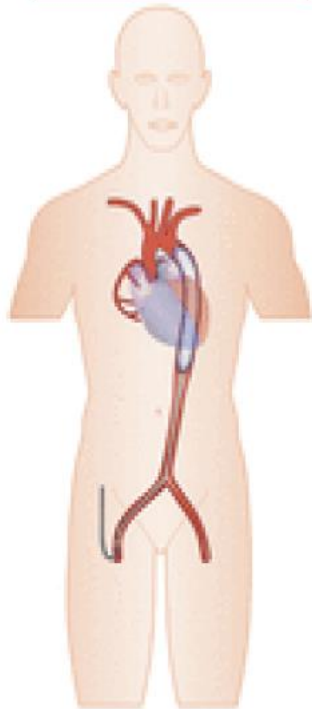
Shock Program/Shock Team

Shock Team

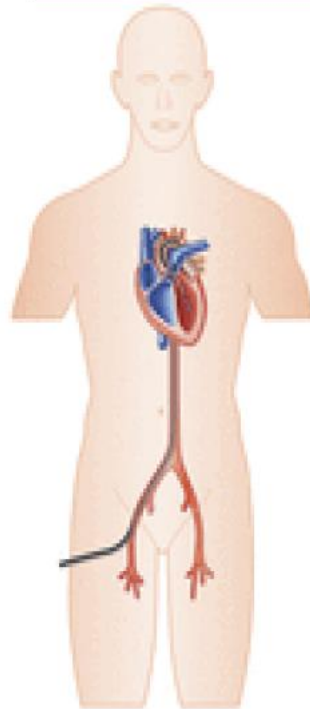
- Multidisciplinary, involving combination of AHF physician, interventionalist, and CT surgery
- Structuralized approach to Cardiogenic Shock
- Temporary support devices available:
 - IABP
 - Impella CP and RP
 - ECMO
 - Centrimag



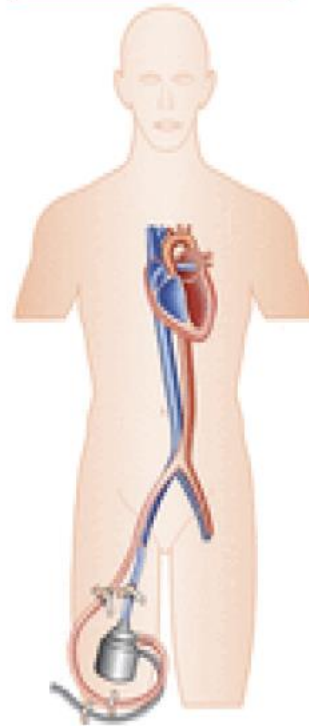
IABP



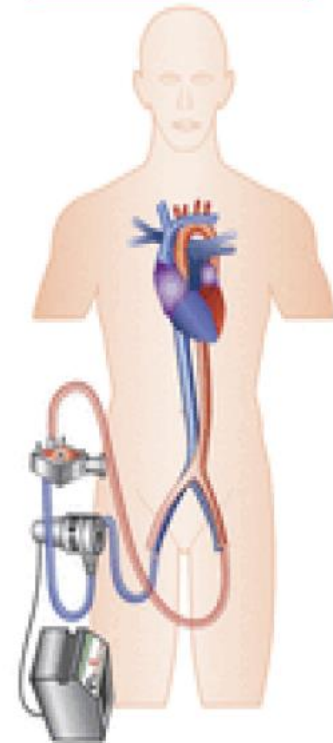
Impella



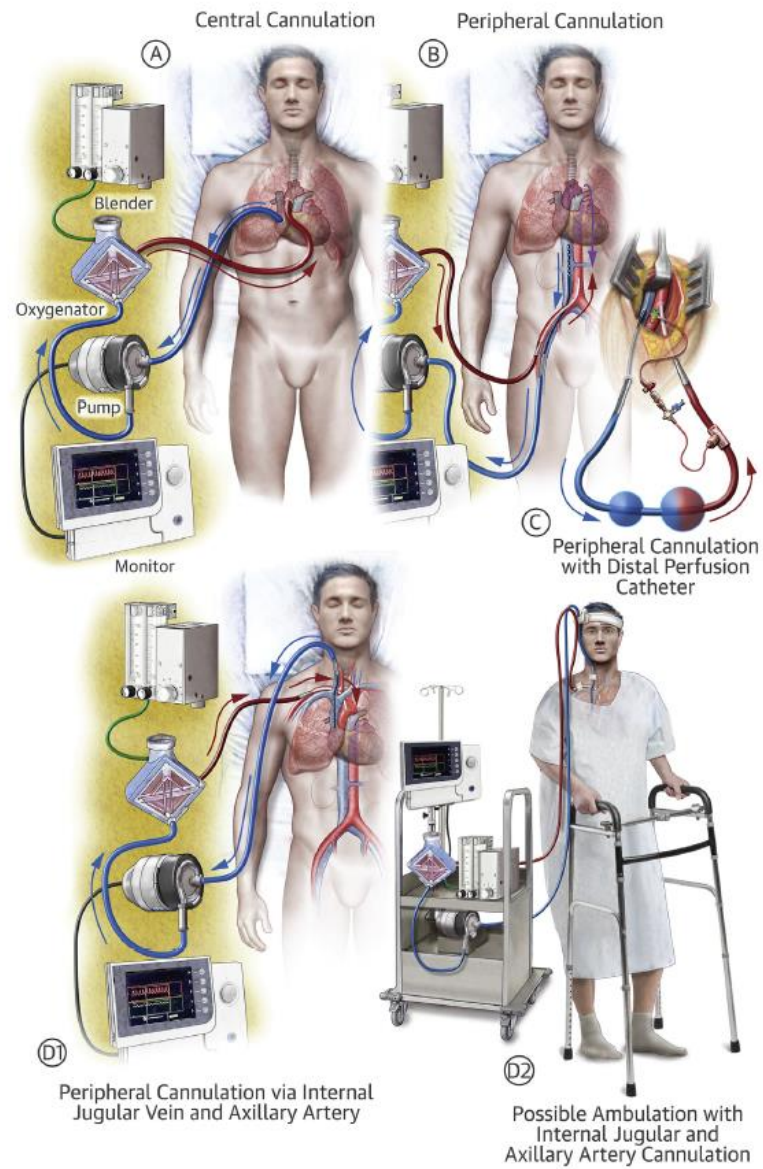
Tandem Heart



ECMO



Werdan 2013, Eur Hrt J



Keebler, M.E. et al. J Am Coll Cardiol HF. 2018;6(6):503-16.

Palliative Options for the non Advanced Therapies Candidate:

- IV diuretics in the office
- Scheduled intermittent thiazide (metolazone)
- Continuous lasix infusion pumps
- Palliative inotrope home infusion therapy
- Digoxin
- Hospice care

Questions?