

# **Non pharmacological management of acute heart failure**

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# Types of LVADS used in patients with acute heart failure

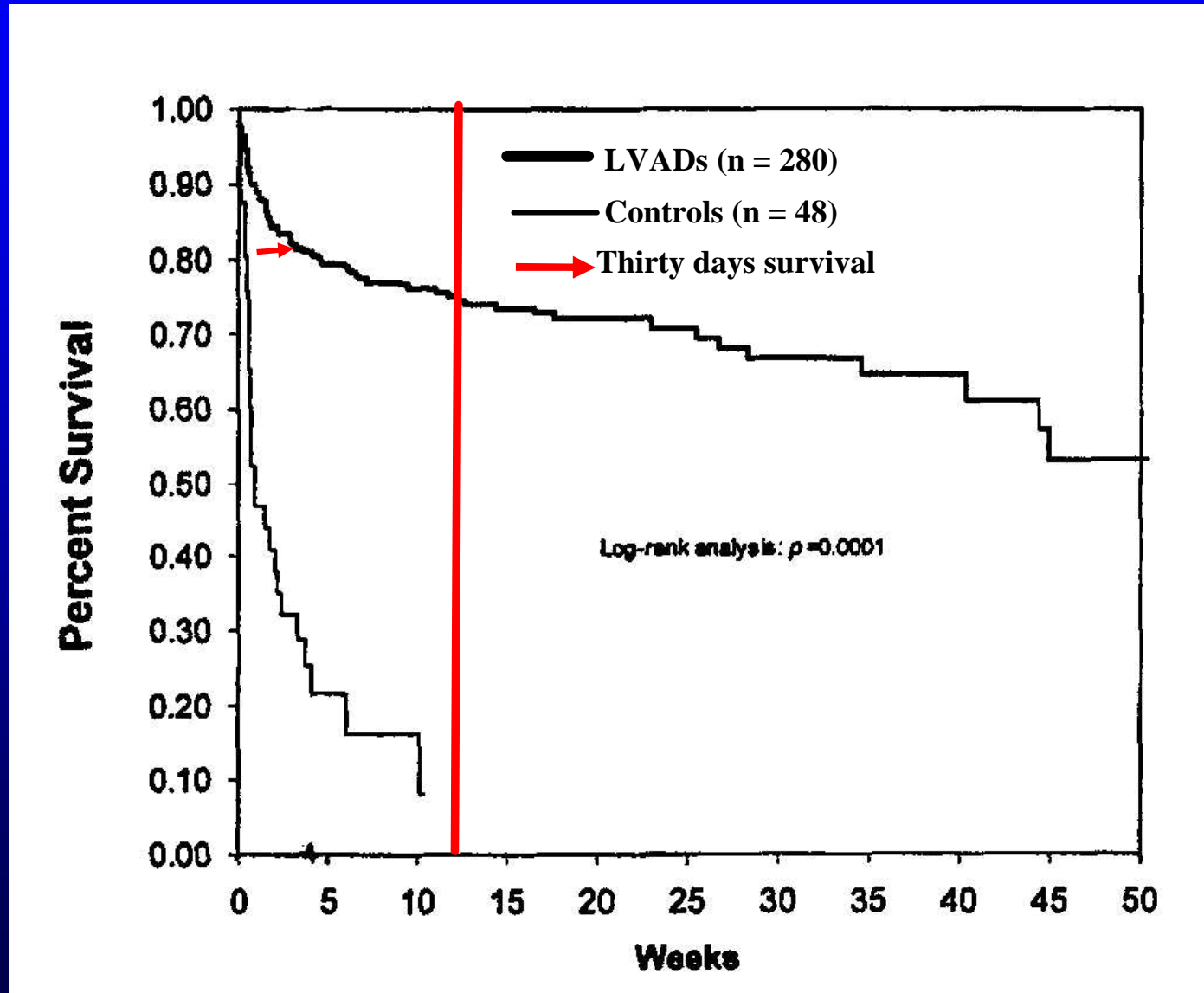
- ✓ Pulsatile extracorporeal devices:
  - **Suitable for limited-duration mechanical assistance**
- ✓ Pulsatile internal devices:
  - **Suitable for long-term mechanical assistance**

Driving systems:

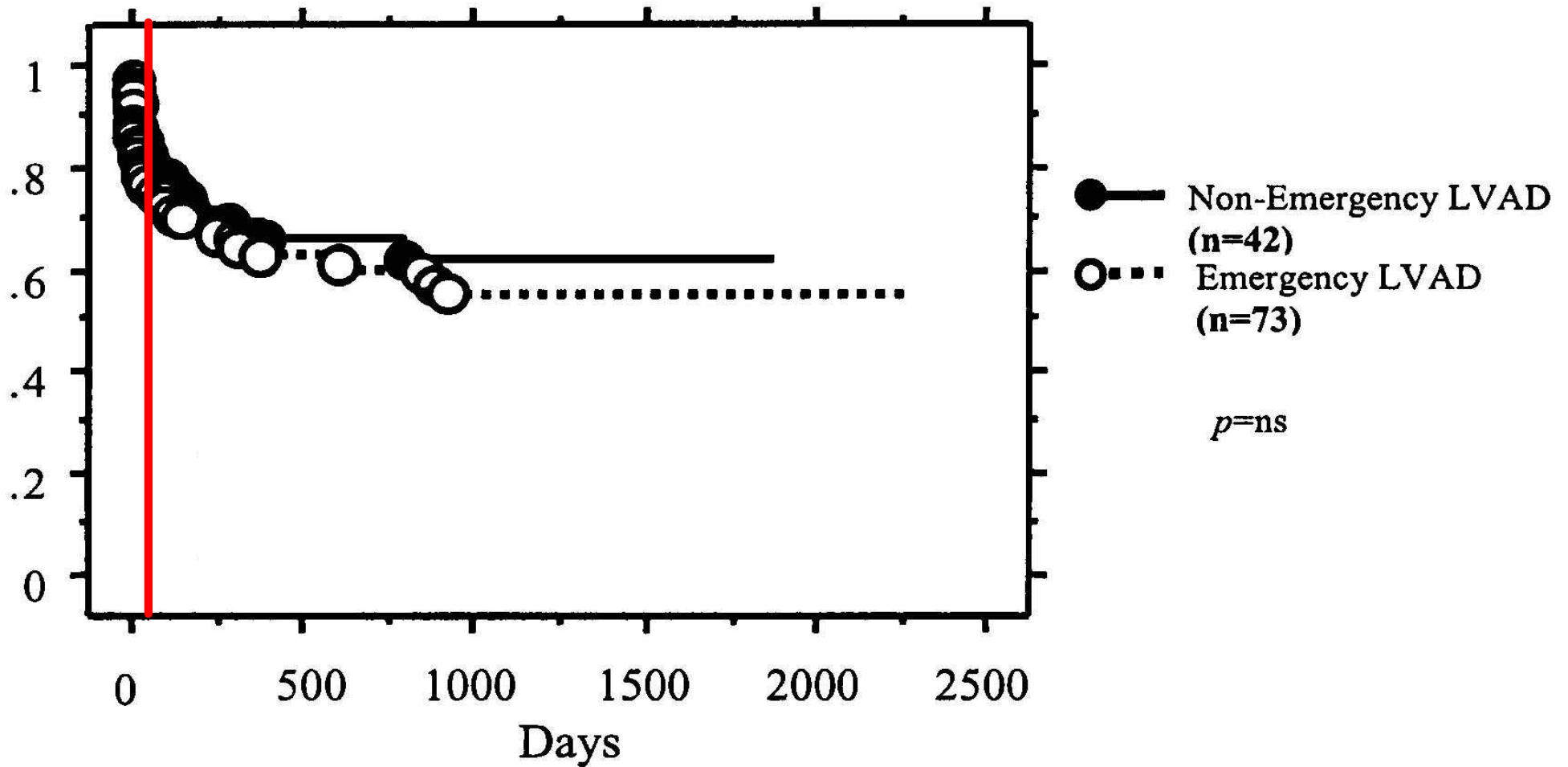
Pneumatic  $\Rightarrow$  Limiting ambulation

**Electrical  $\Rightarrow$  Ambulatory with almost total independence**

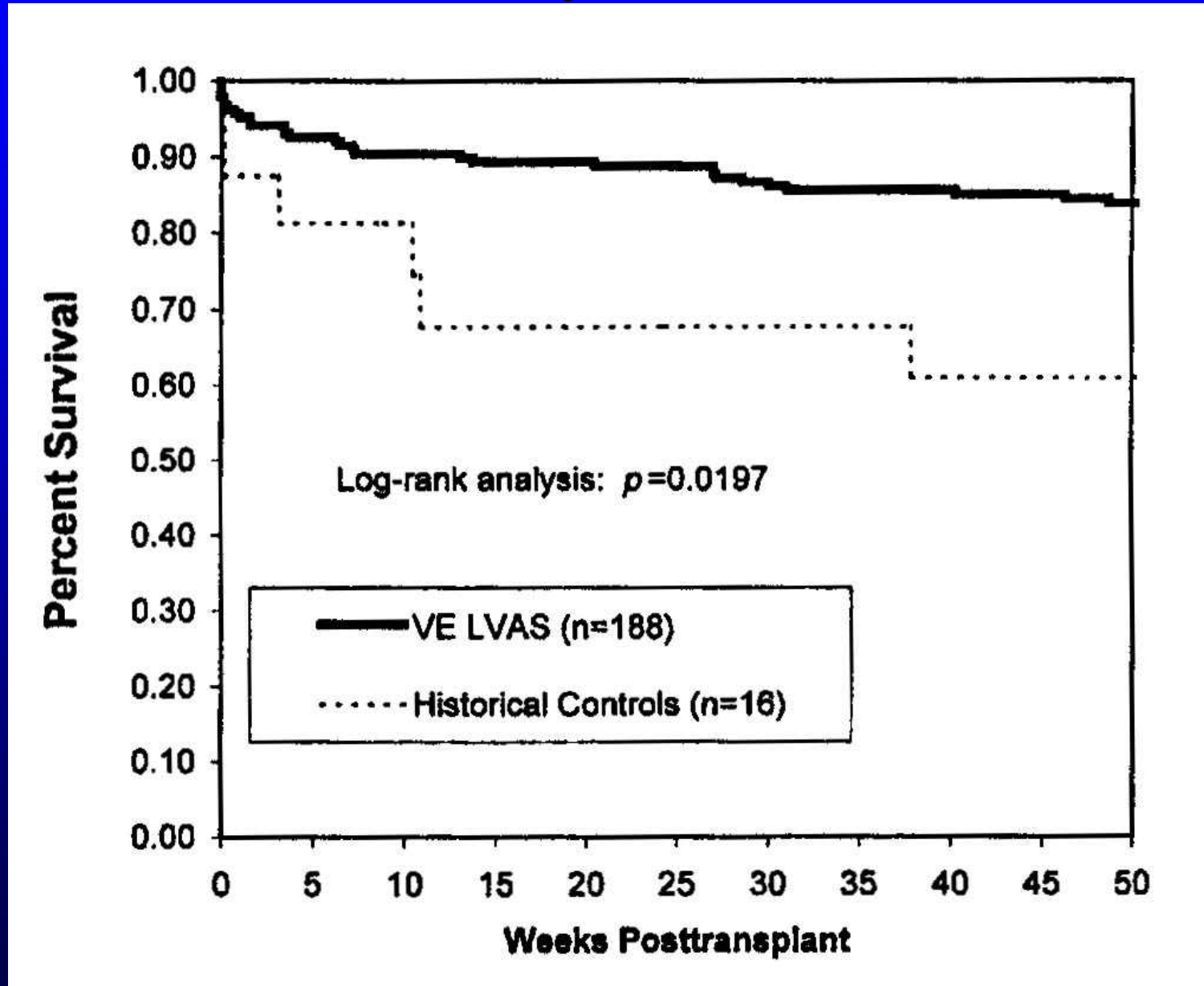
# Survival to transplantation of patients supported by LVAD versus control patients (device unavailability or family refusal)



# Survival (irrespective of receiving a transplant) of the non-emergency and emergency LVAD recipients



# Post-transplantation survival for LVAD-treated patients versus control (device unavailability or family refusal)



# Survival to transplantation of patients assisted by LVADs

**Survival to transplant (%)**

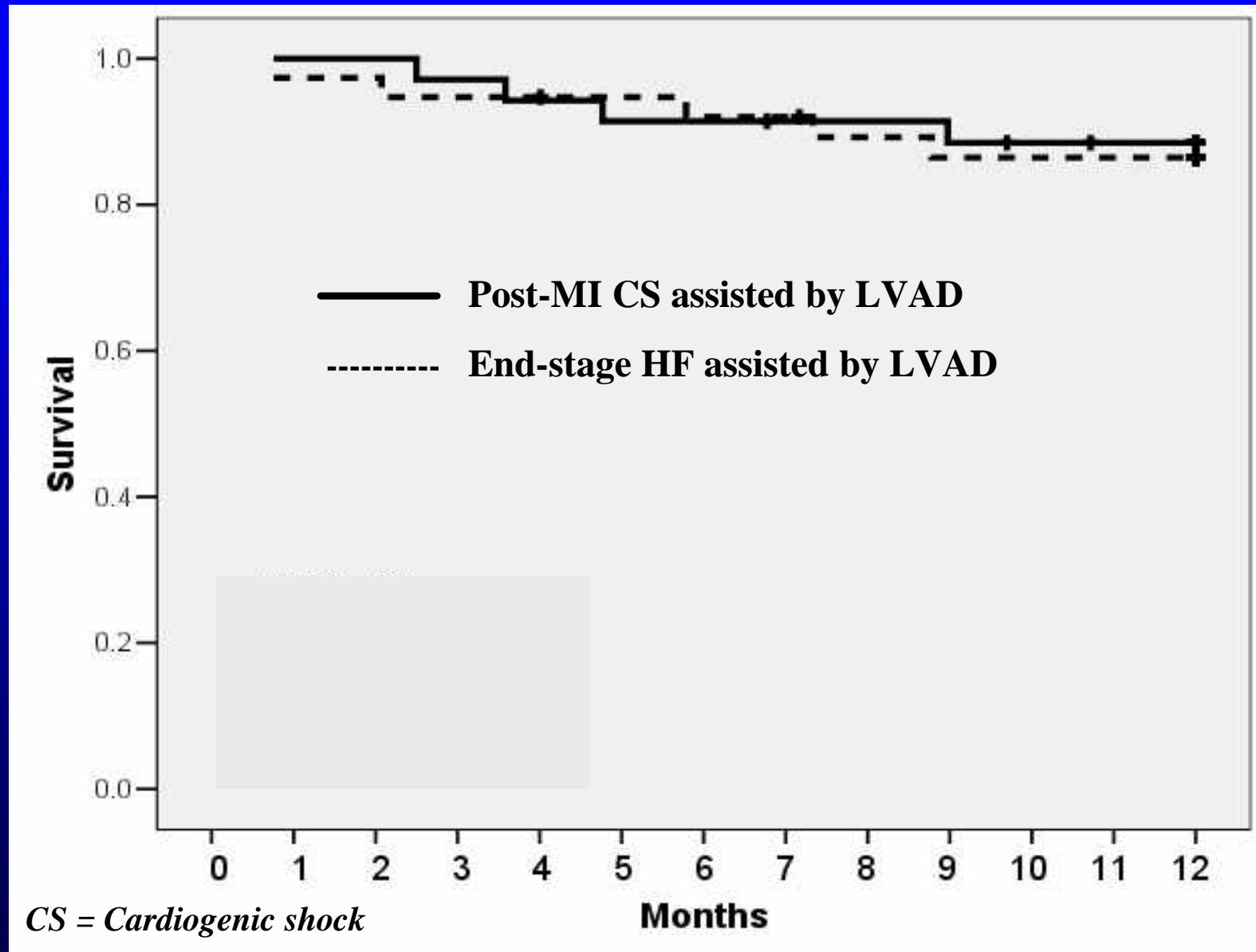
<b>Device</b>	<b>No. of Patients</b>	<b>1992</b>	<b>1995</b>	<b>2002</b>
<b>Thoratec PVAD</b>	<b>1333</b>	<b>65</b>	<b>63</b>	<b>67</b>
<b>IP</b>	<b>1301</b>	<b>64</b>	<b>66</b>	<b>63</b>
<b>VE</b>	<b>1905</b>	<b>-</b>	<b>-</b>	<b>65</b>
<b>Novacor</b>	<b>1600</b>	<b>63</b>	<b>65</b>	<b>-</b>

*IP = implantable pneumatic, PVAD = paracorporeal ventricular assist device,*

*VE = vented electric*

*Ann Thorac Surg 2003;75:S66-71*

# Post-transplantation survival of patients assisted by LVAD either because of intractable post-MI CS or end-stage HF



# Complications in candidates for Tx assisted (n=280) by LVAD

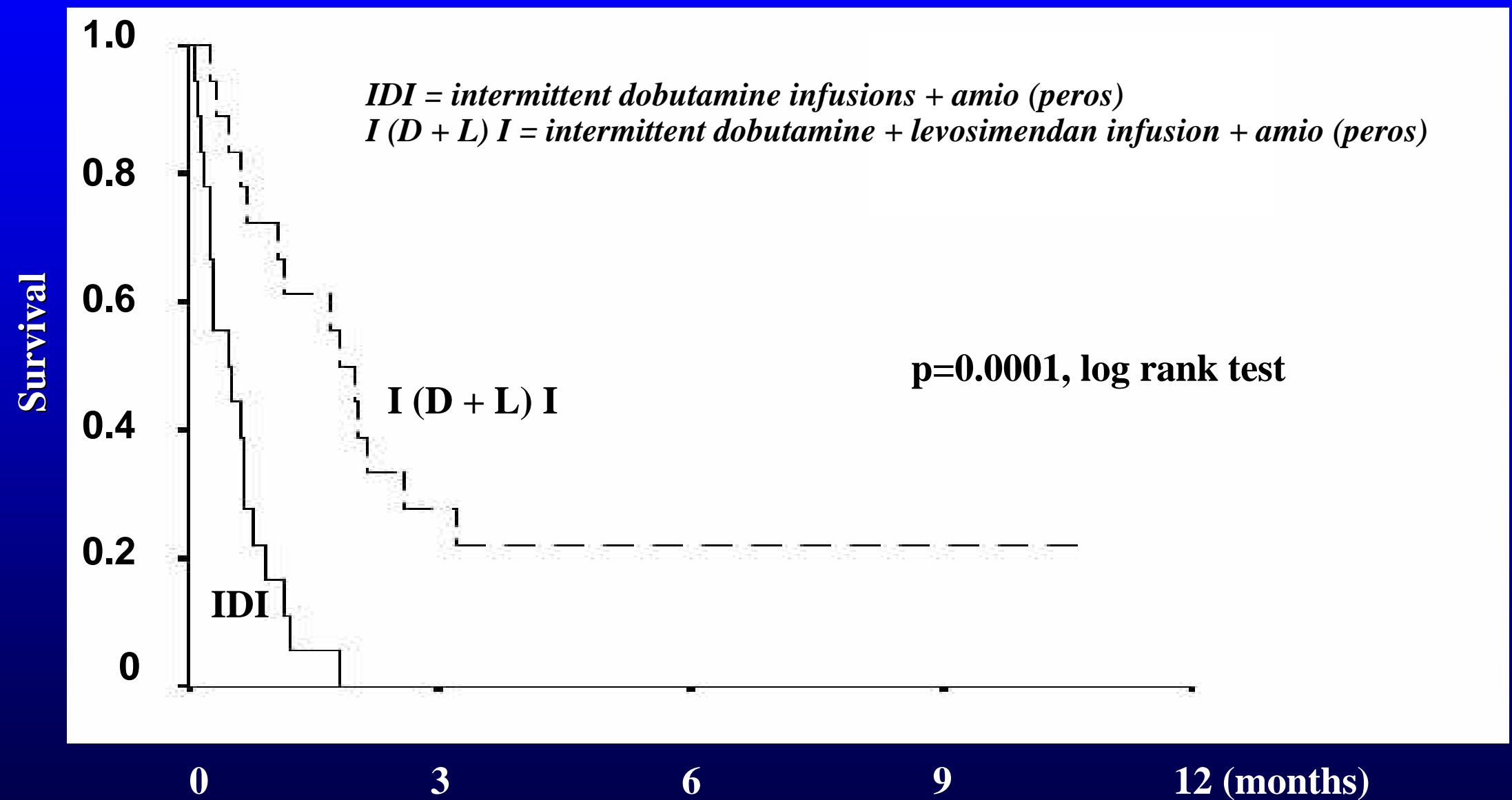
<b>Complication</b>	<b>% of all assisted LVAD patients</b>
<b>Renal dysfunction</b>	<b>56</b>
<b>Bleeding</b>	<b>48</b>
<b>Infection</b>	<b>45</b>
<b>Neurologic dysfunction</b>	<b>27</b>
<b>Thromboembolic events</b>	<b>12</b>
<b>Right heart failure</b>	<b>11</b>
<b>Mechanical failure</b>	<b>1</b>

# Risk factors in LVAD recipients for poor survival to transplantation

<b>Variable</b>	<b>Hazard ratio</b>	<b>P value</b>
Age (range, 11-72 y)	1.03	.0163
Prior heart surgery	1.69	.0366
Baseline creatinine (range, 0.4-8.9mg/dL)	1.38	.0005
Baseline total bilirubin (range, 0.1-23.0 mg/dL)	1.08	.0043

*J Thorac Cardiovasc Surg 2001;122:1186-1195*

# Long term survival of acute decompensated end-stage HF intractable to initial 24 hrs aggressive treatment including inotropes



# Benefits and risks of LVAD implantation as a bridge to transplantation

## Benefits

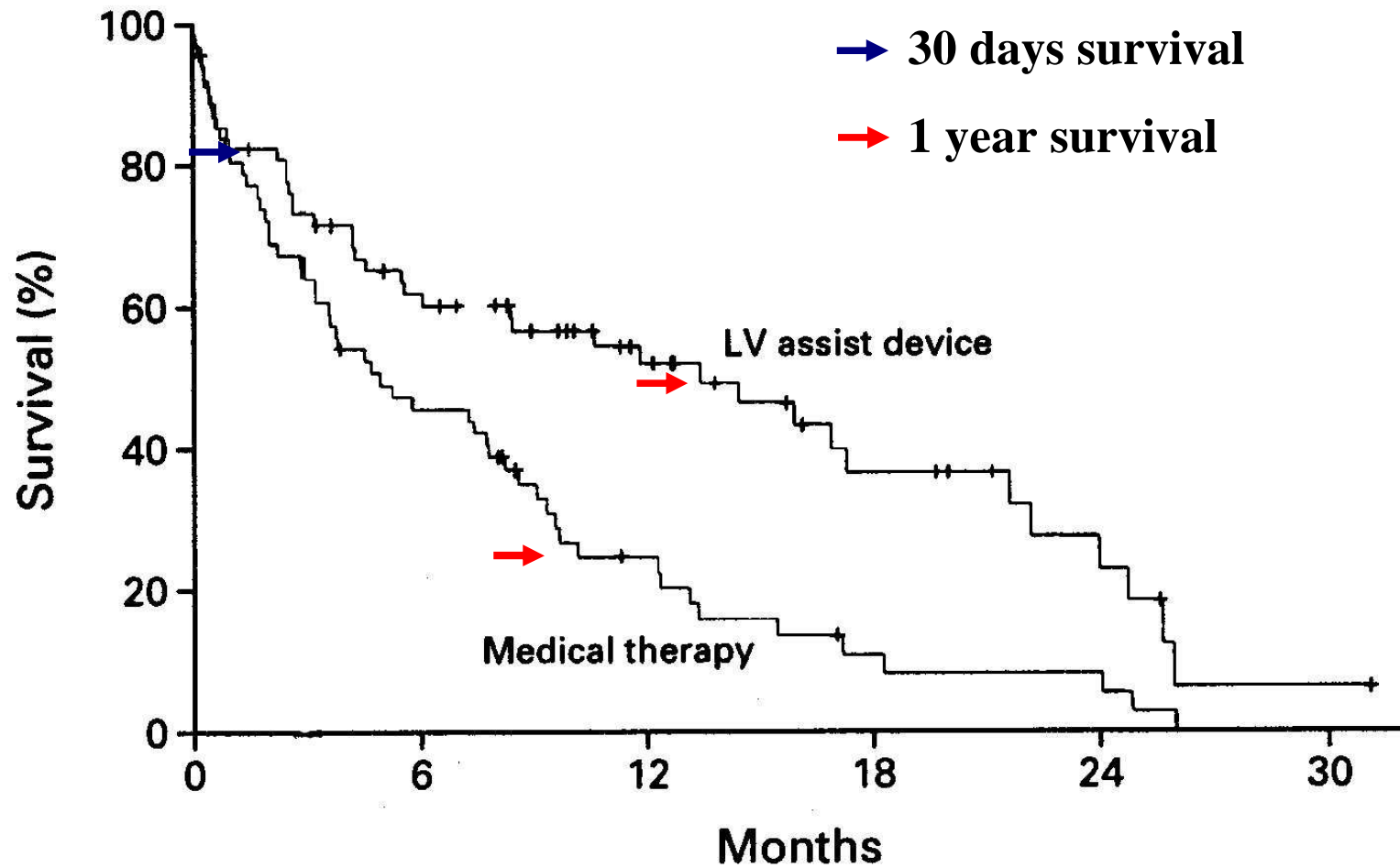
- ✓ Increased survival to transplantation
- ✓ Increased post-transplantation survival
- ✓ Improved quality of life

## Risks

- ✓ High perioperative mortality rate
- ✓ High frequency of complications
- ✓ Unpredictable outcome for an individual patient

*(Compared to inotrope dependent patients)*

# Survival of end-stage HF supported by LVAD versus medically treated patients



No. AT RISK

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

# Survival benefit in end – stage HF randomized studies

<b>Study (Therapy)</b>	<b>Control vs therapy 1 year survival (%)</b>	<b>Relative Benefit, %</b>	<b>Absolute benefit No, Patients/100</b>
<b>SOLVD (ACEI)</b> <i>N Engl J Med 1991;325:293-302</i>	<b>86 vs 89</b>	<b>3.5</b>	<b>3</b>
<b>CONSENSUS (ACEI)</b> <i>N Engl J Med 1987;316:1429-35</i>	<b>38 vs 55</b>	<b>45</b>	<b>17</b>
<b>COPERNICUS (β- blocker)</b> <i>N Engl J Med 2001;344:1651-8</i>	<b>81.5 vs 89</b>	<b>9</b>	<b>7.5</b>
<b>RALES (spironolactone)</b> <i>N Engl J Med 2002;346:1845-53</i>	<b>75 vs 83</b>	<b>11</b>	<b>8</b>
<b>REMATCH – (LVAD)</b> <i>N Engl J Med 2001;345:1435-43</i>	<b>25 vs 52</b>	<b>108</b>	<b>27</b>

# Usual causes of death in patients assisted by LVAD in the *REMATCH* study

**Cause of death**

**LVAD group**

**% of the total deaths**

**Sepsis**

**42**

**LVAD Failure**

**17**

**Non-cardiac**

**12**

**Cerebrovascular disease**

**10**

**Pulmonary embolism**

**5**

**Perioperative bleeding**

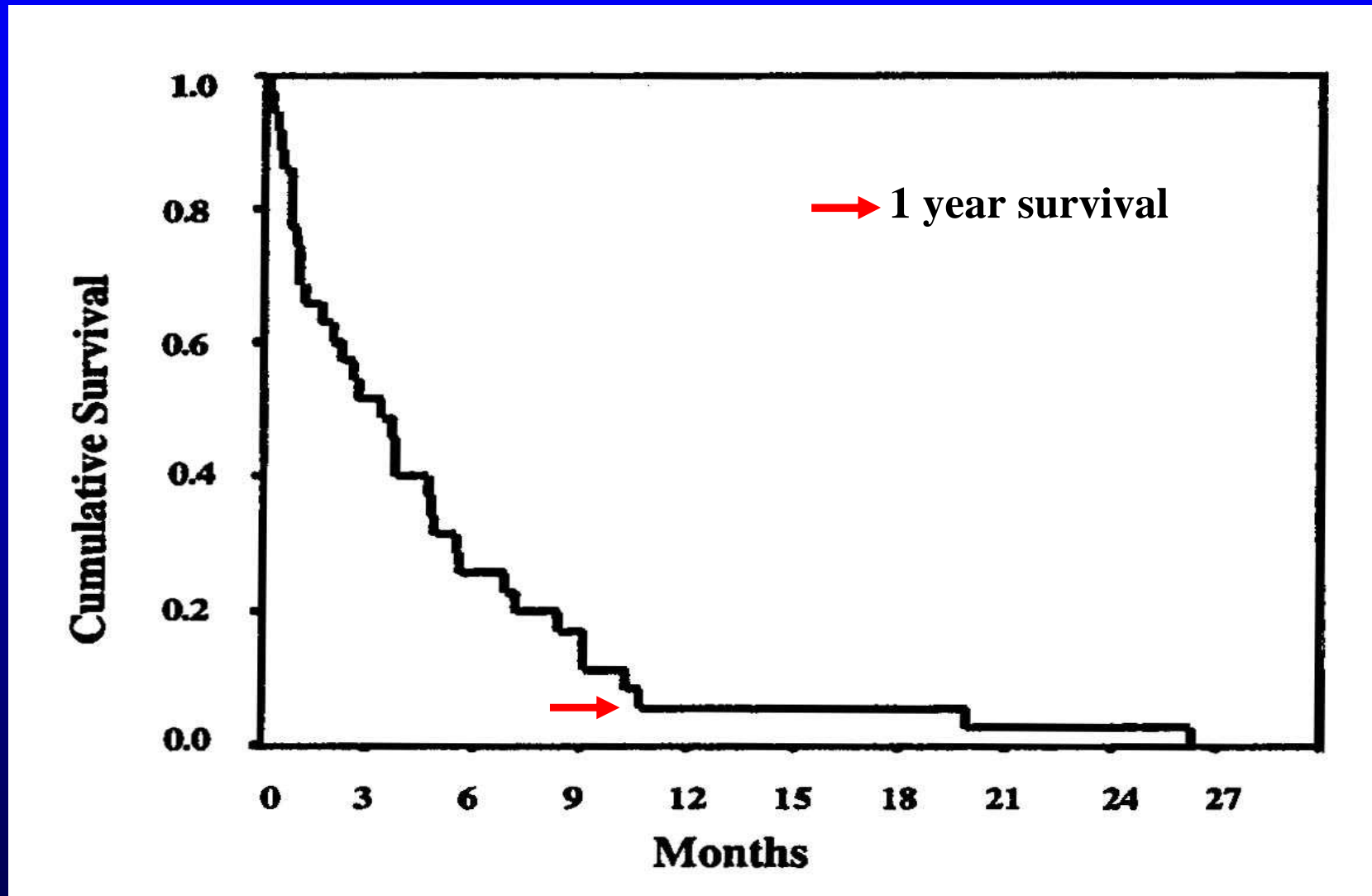
**3**

**Total**

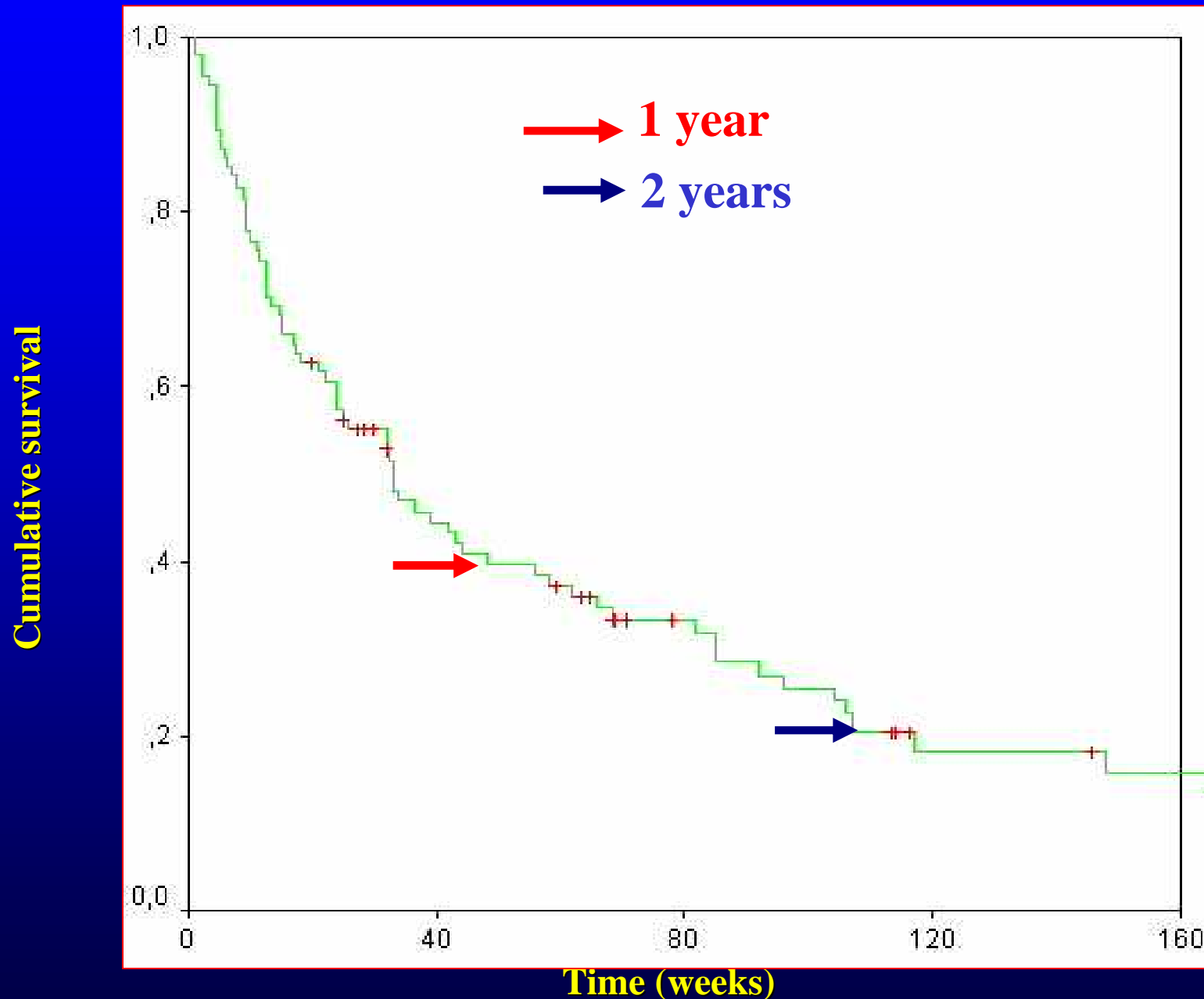
**89**

**In the medical therapy group, 92% of the total deaths were due to LV dysfunction**

# Survival of patients with persistent decompensated HF dependent on inotrope infusion

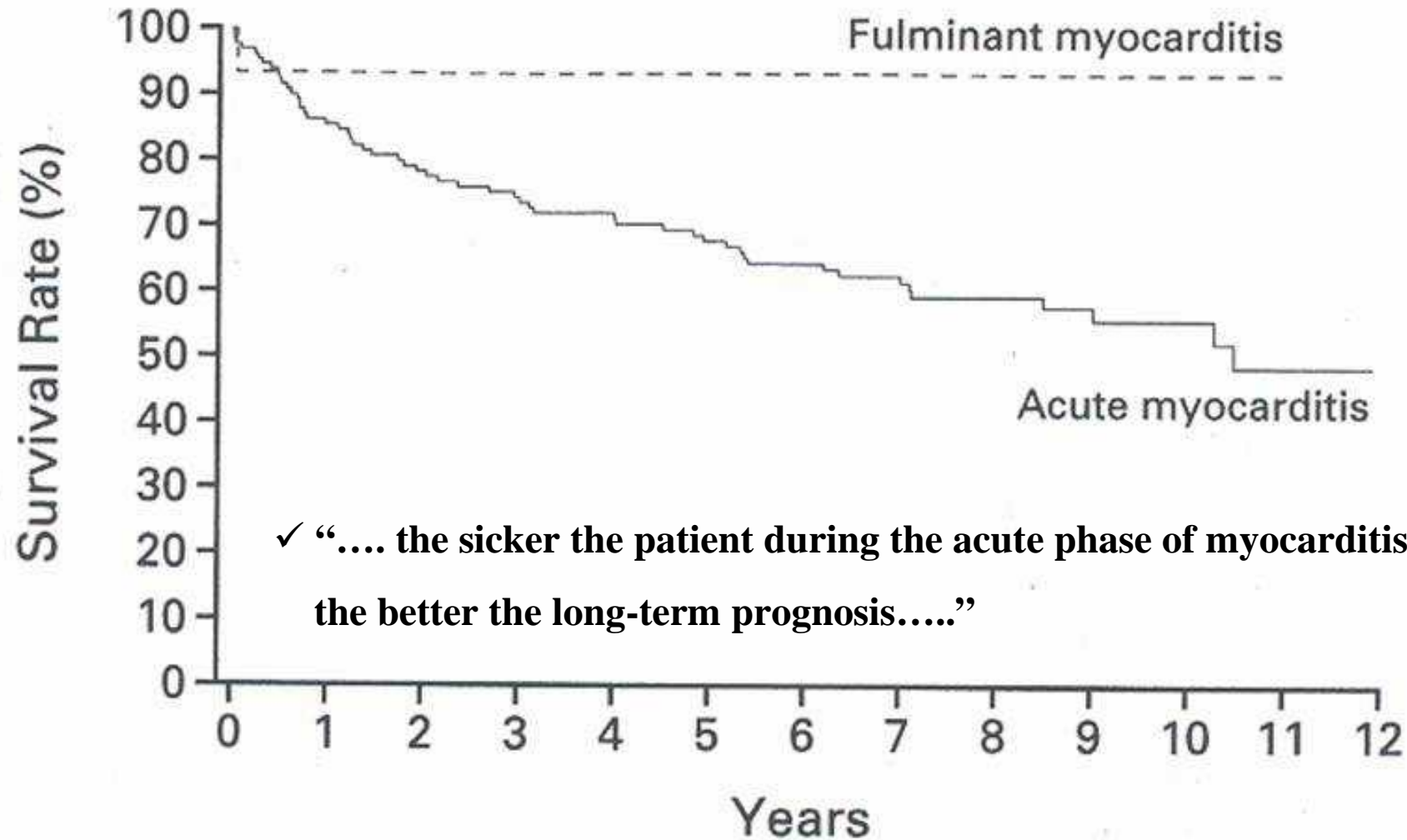


# Survival of patients (n=101) with acute decompensated end stage HF clinically responded to initial 24-72 aggressive treatment including inotropes



*J. Nanas, Dept Clin Therapeutics, University of Athens, Athens, Greece (Unpublished Data)*

# Transplantation-free survival in patients with fulminant and acute myocarditis



# Brief History and Hemodynamics of patients with fulminant myocarditis complicated by CS

*Dept of Clin Therapeutics, Univ of Athens*

**Patient #1**

**Patient #2**

<b>Sex</b>	<b>F</b>	<b>F</b>
<b>Age</b>	<b>48</b>	<b>43</b>
<b>HR (bpm)</b>	<b>125</b>	<b>120</b>
<b>BP (mmHg)</b>	<b>90/60</b>	<b>100/80</b>
<b>LVEF</b>	<b>23%</b>	<b>36%</b>
<b>EMB</b>	<b>AM</b>	<b>AM</b>
<b>RAP (mmHg)</b>	<b>9</b>	<b>13</b>
<b>PCWP (mmHg)</b>	<b>15</b>	<b>25</b>
<b>CI (L/min/m<sup>2</sup>)</b>	<b>2.0</b>	<b>2.1</b>
<b>Treatment</b>	<b>IABP + I + MV</b>	<b>IABP + I ⇒ LVAD</b>
<b>Outcome</b>	<b>Alive</b>	<b>Death</b>

*AM = Acute myocarditis, I = Inotrope, EMB = Endomyocardial biopsy, MV = Mechanical ventilation, CS = Cardiogenic shock*

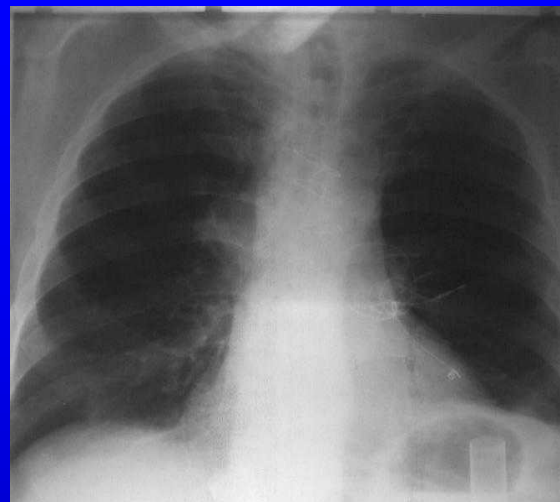
1st



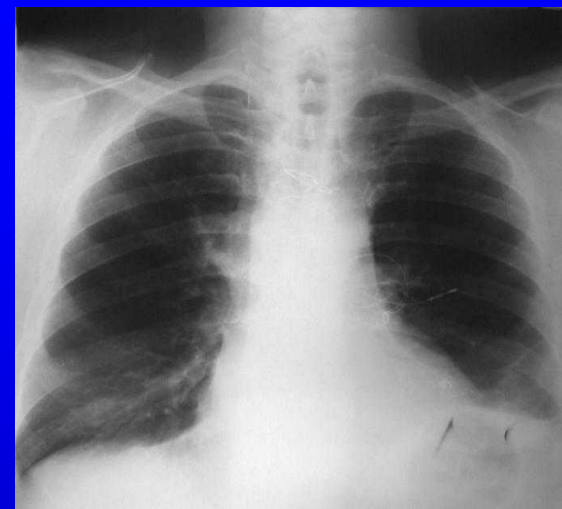
2nd



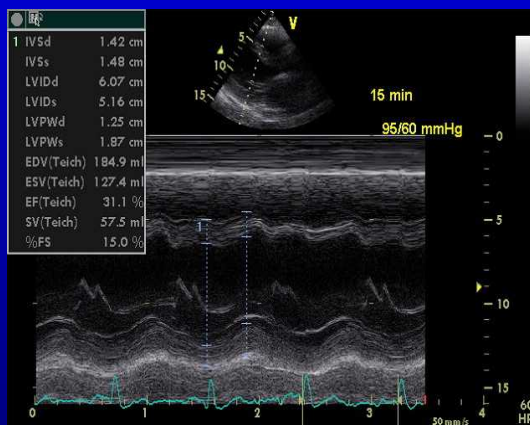
3rd



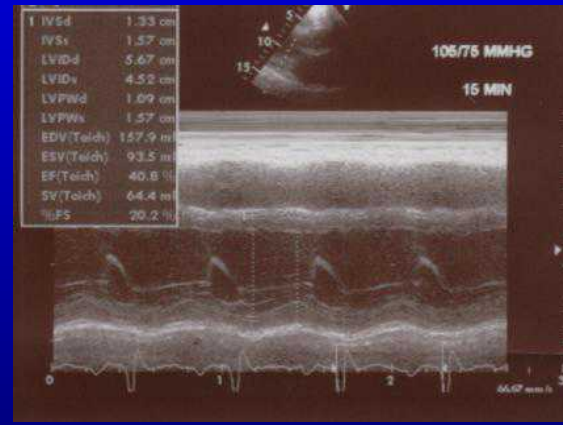
4th



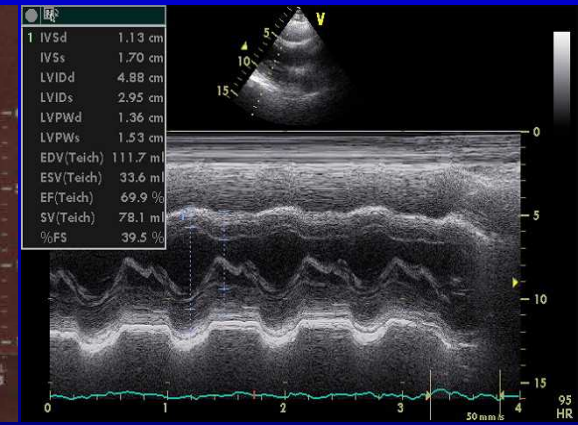
Pre-implantation



8 weeks post implantation,  
pre-clenbuterol initiation,  
LVAD off 15 min



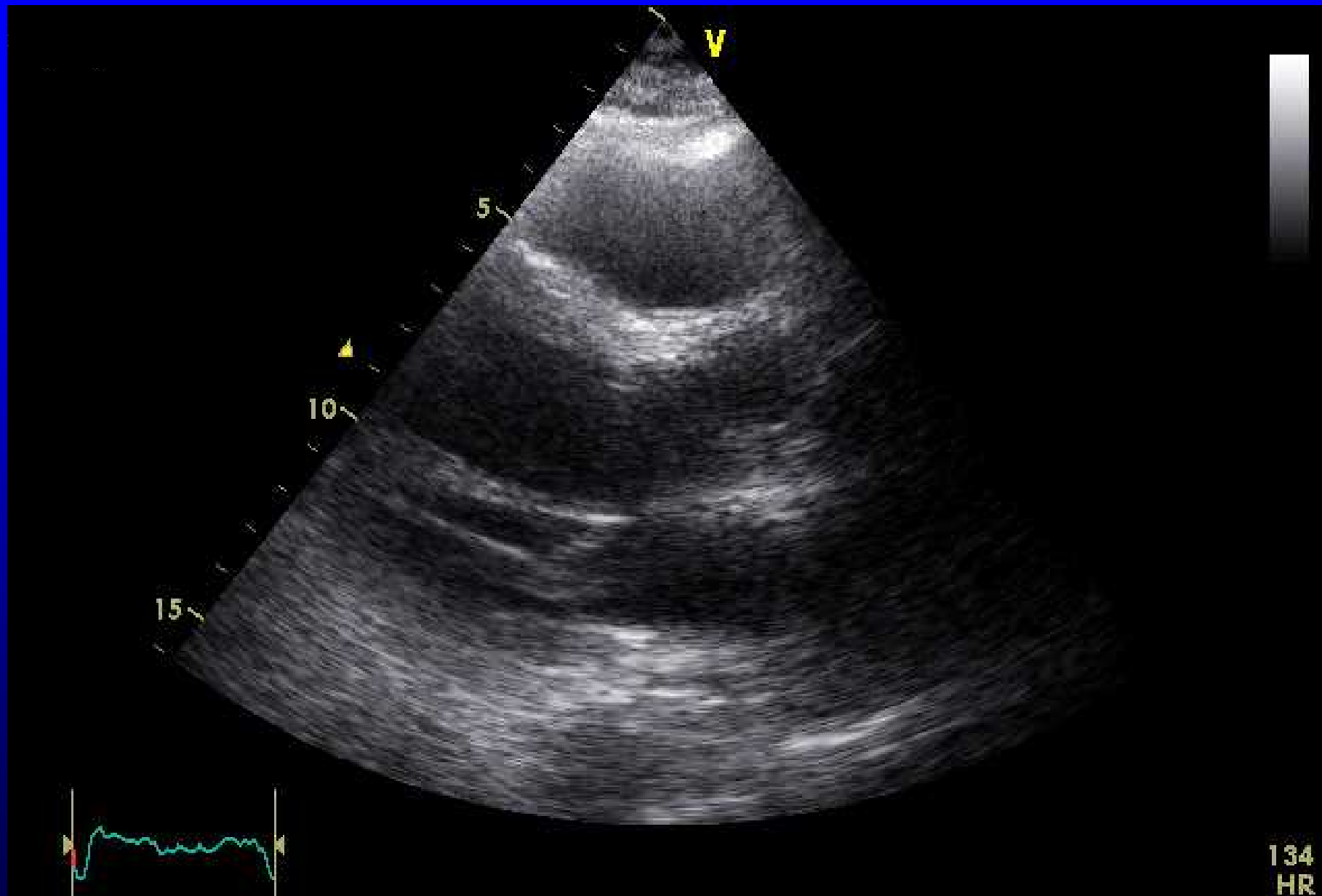
14 weeks post implantation,  
6 weeks on clenbuterol,  
LVAD off 15 min

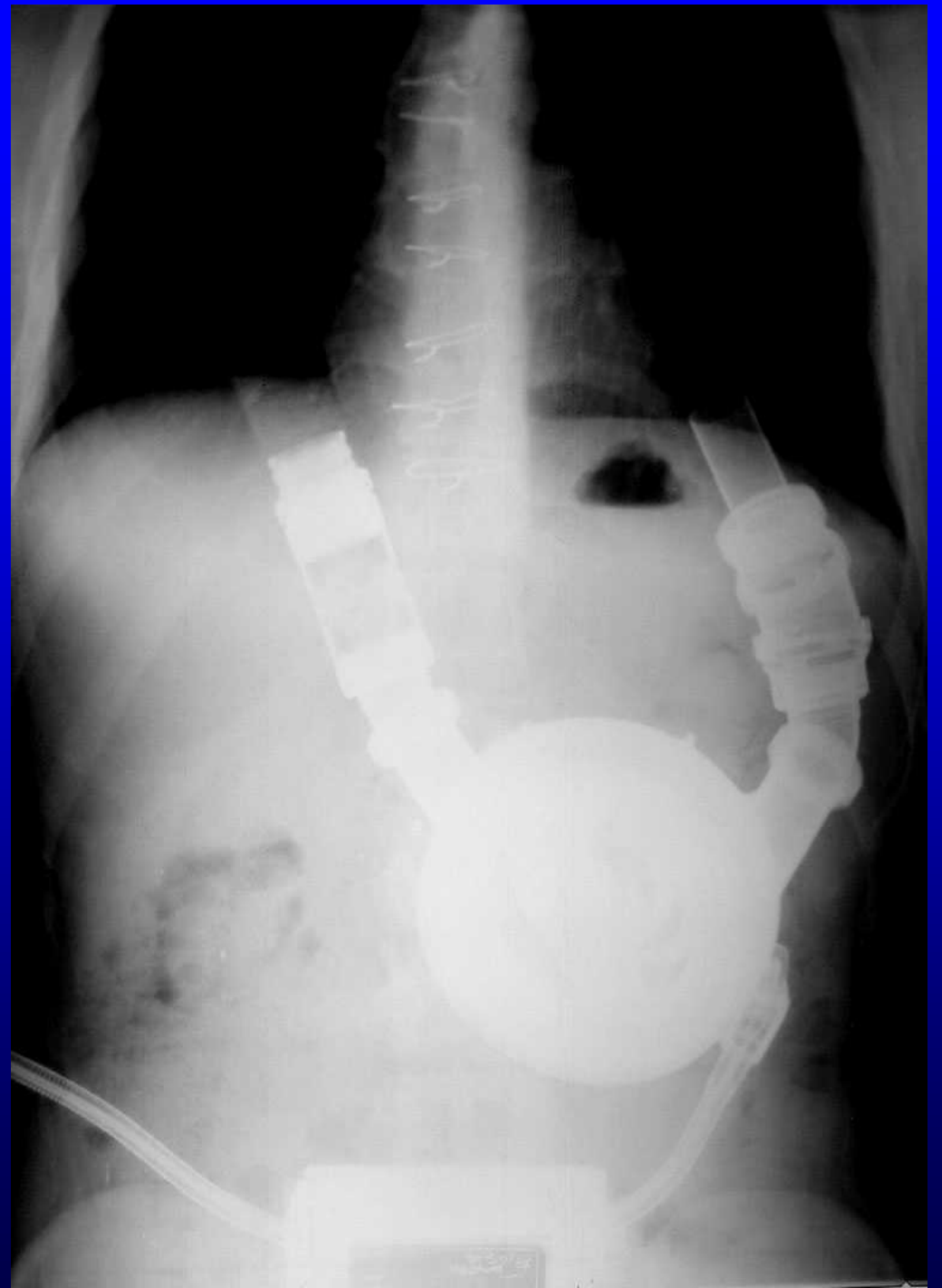
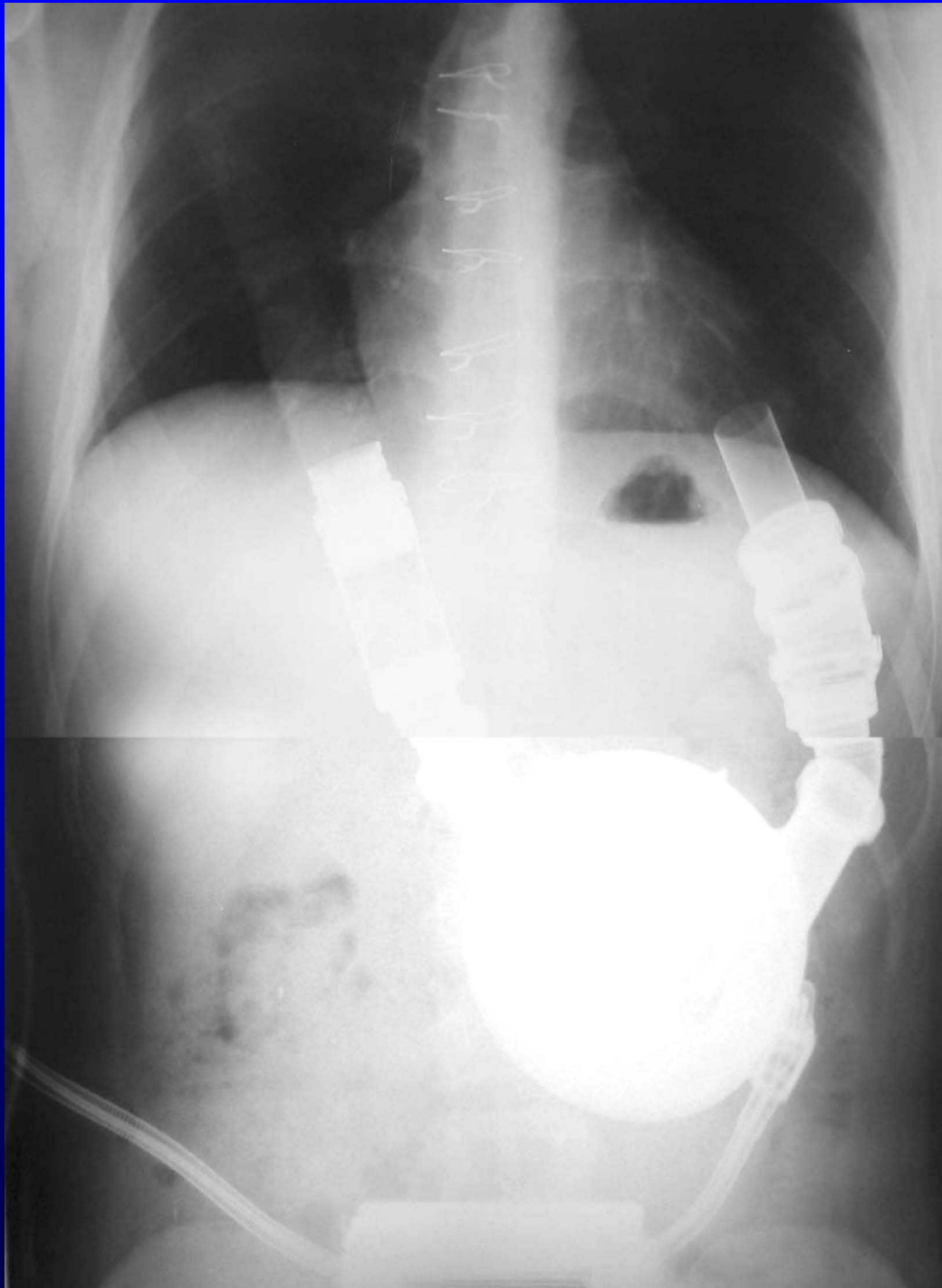


6 months  
post- explantation

**Harefield** (Prof M Yacoub) — **Athens** (Dr S Theodoropoulos/Prof J Nanas) **Recovery Program (HAPR)**

# Post-explantation





# Problems that need to be solved

- High peri-operative mortality rate even in experienced surgical teams
- High frequency of complications in specialized centers
- Large number of potential candidates > 150000 (Europe + USA)
- Need for more than 3000 centers (Europe + USA)

## *New Directions*

- Simple device for its implantation and follow up

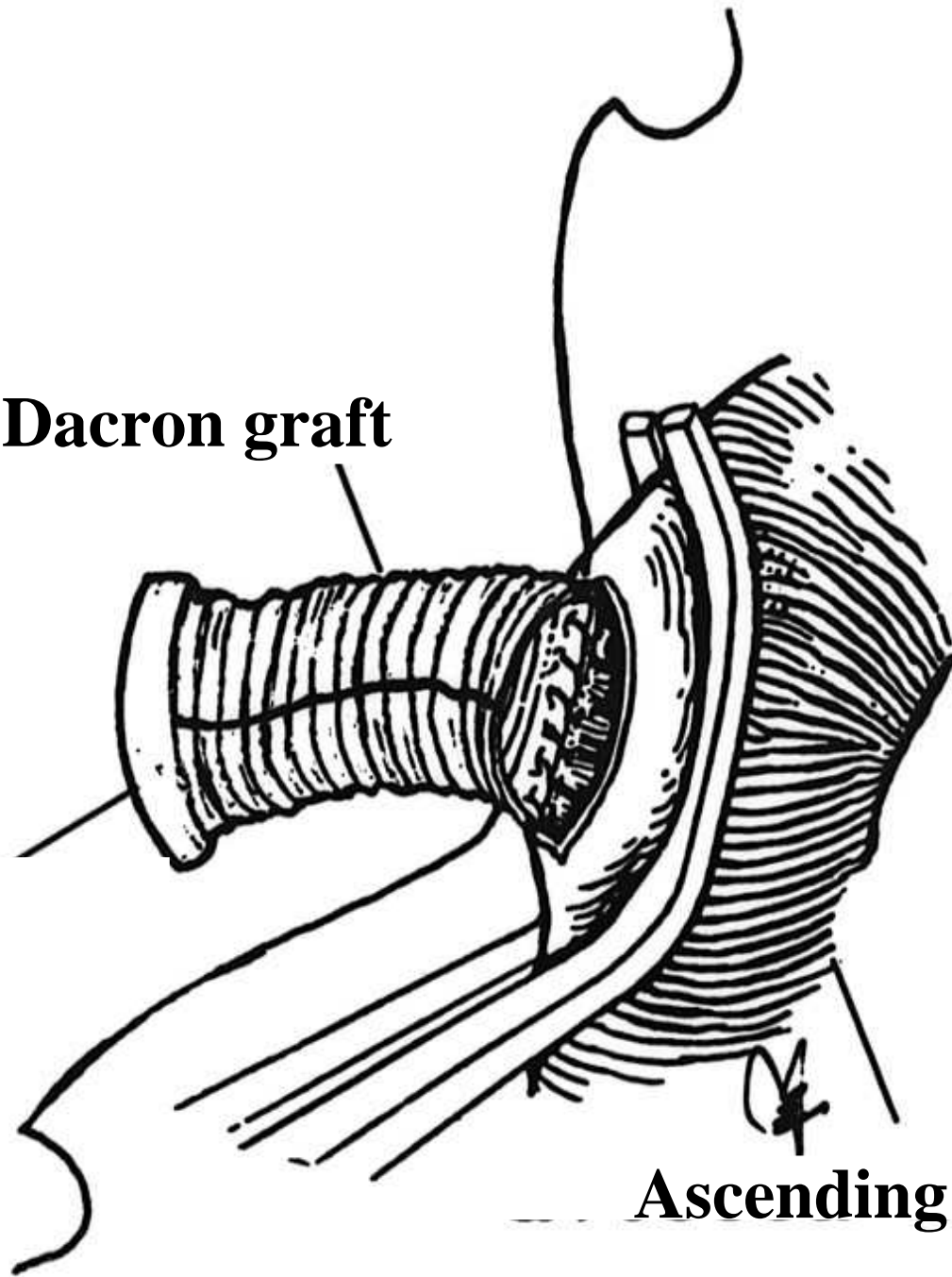
# LVADs under clinical investigation

- **Axial flow LVADs (various types) > 300 pts**
- **Counterpulsation LVADs**
  - **Cardio VAD (KCV) 8 pts**
  - **Sunshine Heart 2 pts**
  - **Paraaortic CD 5 pts**



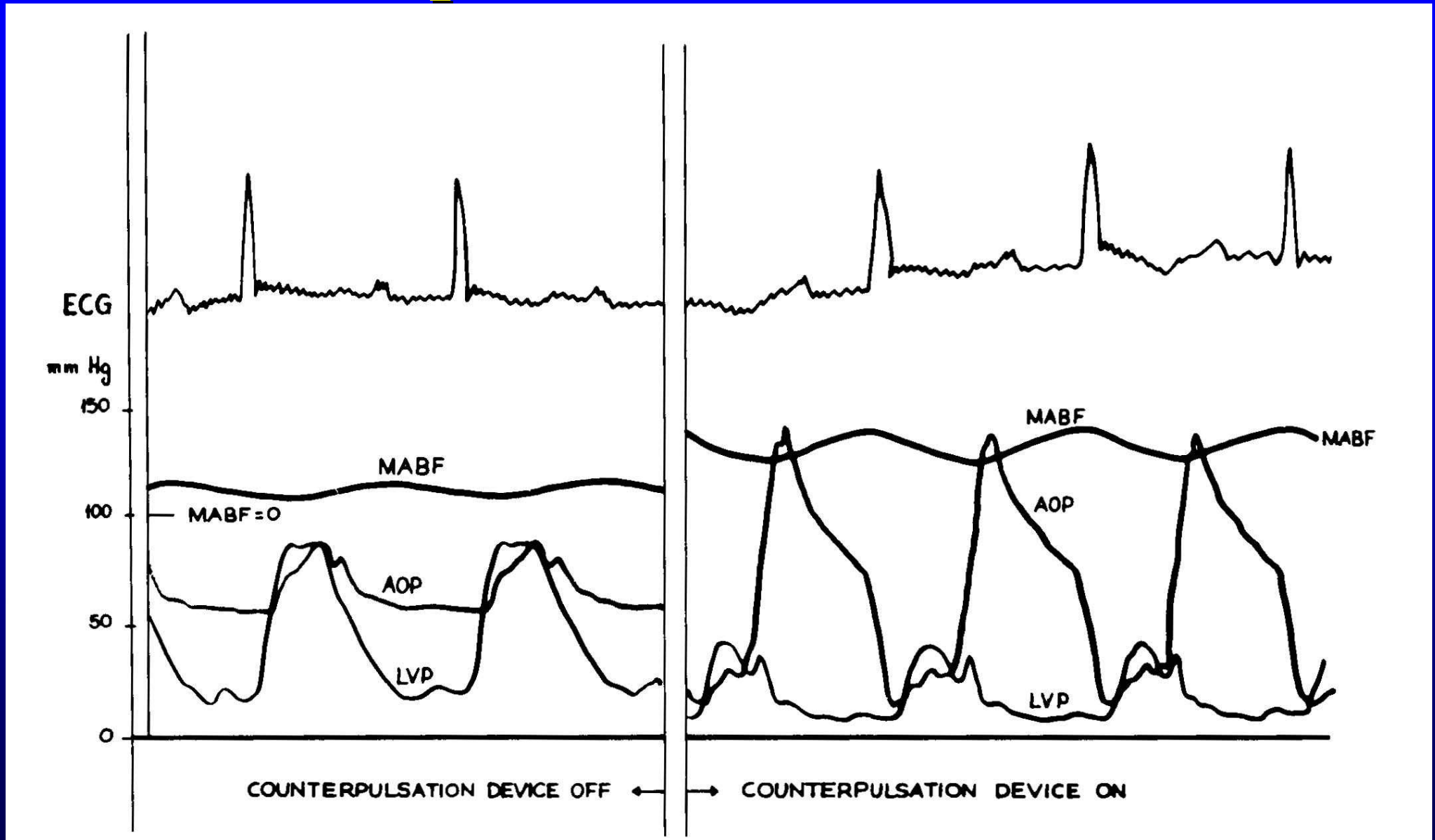
**Connector**

**Dacron graft**



**Ascending aorta**

# Hemodynamic effects of the PACD in experimental acute HF

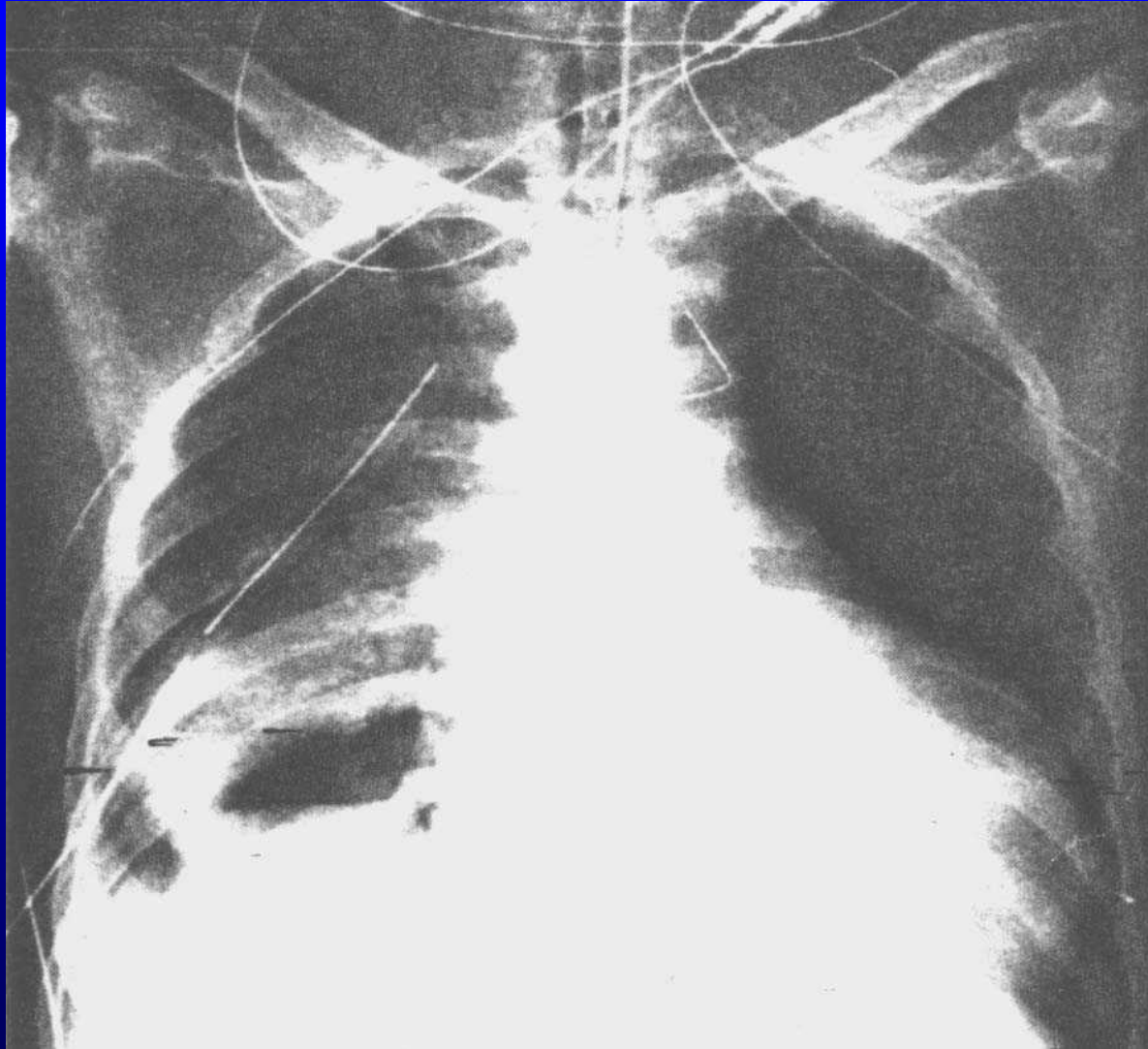


# Clinical and laboratory pre-implantation data of a patient assisted by the PACD for 54 days

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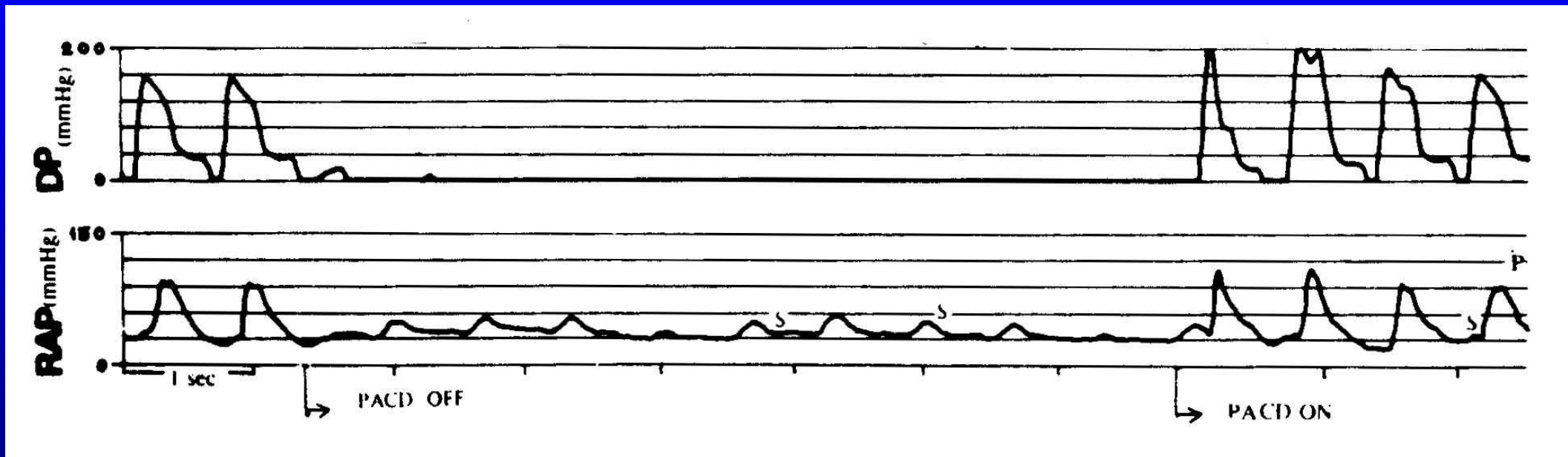
Age (yr)	33
Diagnosis	IDC
LVEF (%)	6
<b>RAP (mmHg)</b>	<b>17</b>
<b>PCWP (mmHg)</b>	<b>35</b>
BP (mm Hg)	90/60
<b>CI (L/min/m<sup>2</sup>)</b>	<b>1.3</b>
<b>Na (mEq/L)</b>	<b>120</b>
<b>Cr (mg/dl)</b>	<b>3.6</b>

# After the PACD implantation



*Nanas J et al, J Thorac Cardiovasc Surg 1996;111:55-61*

# Recordings of the driving and radial arterial pressures of a patient assisted by the PACD for 54 days



*Nanas J et al, J Thorac Cardiovasc Surg 1996;111:55-61*

## **Conclusion (1/3)**

# **Management of potential candidates for LVAD among patients with acute decompensated end-stage HF or de-novo acute HF**

- ✓ **Exclude reversible causes**
- ✓ **Tailor medical therapy to exclude adequate response**
- ✓ **Determine candidacy for assistance by LVAD as:**
  - 1. Bridge to transplantation**
  - 2. Bridge to recovery**
  - 3. Destination therapy**

## **Conclusion (2/3)**

# **Therapeutic targets according to etiologies of acute heart failure in potential candidates for LVAD assistance**

### **✓ Bridge to Transplantation**

- Acute decompensation of end-stage HF in patients listed for Tx**
- Post-MI Cardiogenic Shock**

### **✓ Destination**

- Persistent decompensation of CHF in patients ineligible for Tx**

### **✓ Bridge to Recovery**

- Fulminant myocarditis**
- Acute decompensation of end-stage HF due to IDC**

## **Conclusion (3/3)**

**The new LVADs require a far less traumatic procedure for their implantation and are likely to be beneficial for patients with less severe HF.**