



Città della Salute e della Scienza

University of Turin

Division of Cardiac Surgery



AZIENDA  
OSPEDALIERO  
UNIVERSITARIA



Santa Maria  
della Misericordia  
di Udine



UNIVERSITY  
OF UDINE



BEYOND THE SLIDES 2015  
1<sup>st</sup> UDINE ECMO WORKSHOP

DECEMBER 18-19, 2015

AUDITORIUM HYPO ALPE ADRIA

TAVAGNACCO (UD)

PROMOTED BY  
CARDIOTHORACIC DEPARTMENT



SUPPORTED BY



# TIMING of ECMO IMPLANTATION

Prof. M. Rinaldi

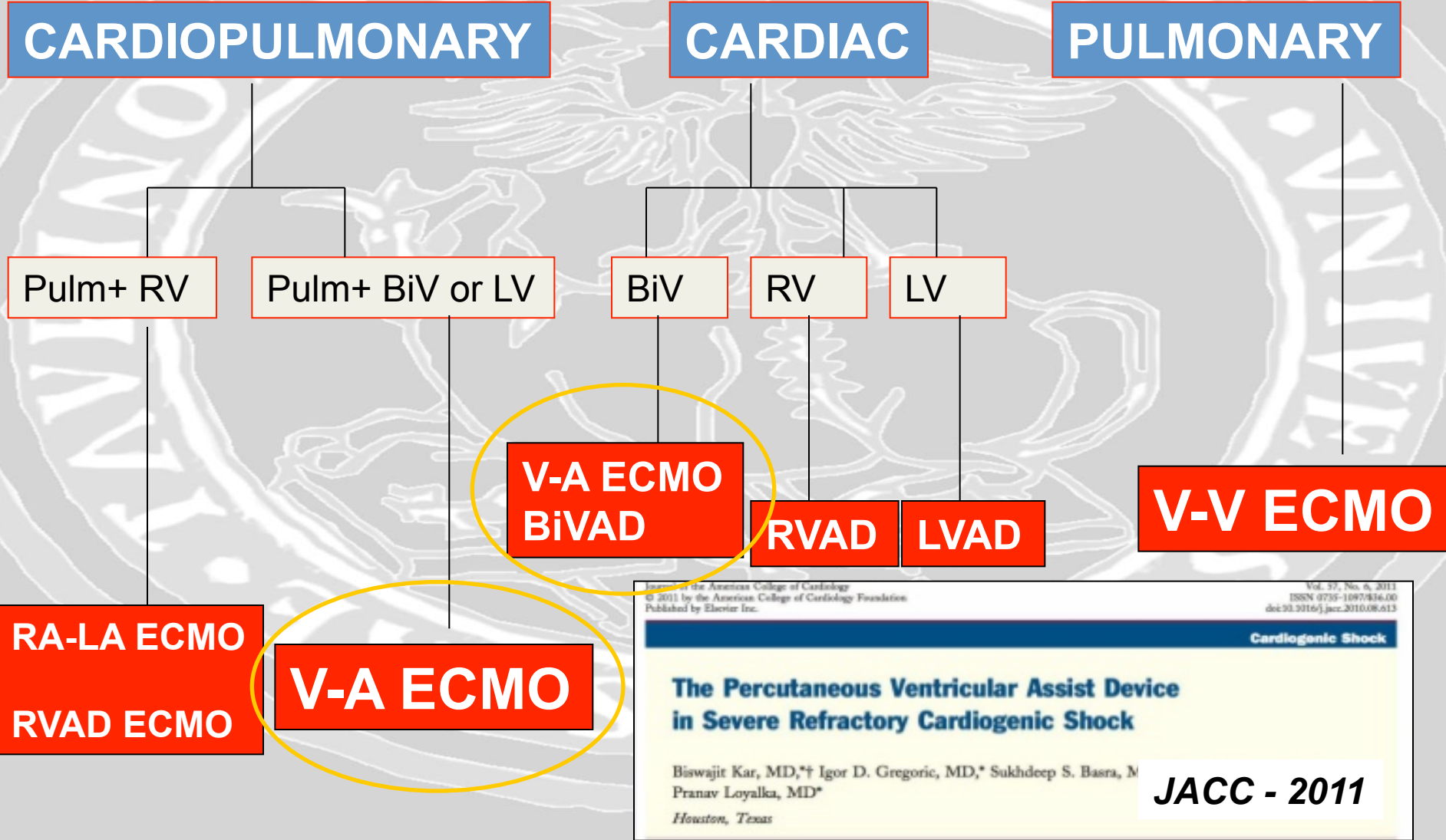
# CARDIOGENIC SHOCK

## General Considerations

- *Results of conventional therapy*
- *Risks and Benefits of the VA ECMO therapy*
  - *no randomized study data available for ECMO/ECLS use in cardiogenic shock*

# Refractory Failure and Treatment

## «The ECMO galaxy»



Journal of the American College of Cardiology  
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 Published by Elsevier Inc.

Vol. 57, No. 6, 2011  
 ISSN 0735-1097/06.00  
 doi:10.1016/j.jacc.2010.08.613

**Cardiogenic Shock**

**The Percutaneous Ventricular Assist Device  
 in Severe Refractory Cardiogenic Shock**

Biswajit Kar, MD,\*† Igor D. Gregoric, MD,\* Sukhdeep S. Basra, M  
 Pranav Loyalka, MD\*  
 Houston, Texas

**JACC - 2011**

# Timetable of VA ECMO implantation

## Precardiotomy VA ECMO

After failure of conventional therapy

## Rescue VA ECMO

In/Out of hospital

## VA ECMO for DCD

Heart failure

Cardiogenic Shock

Cardiac arrest

Death

Donor

## VA ECMO or LVAD

High risk PTCA  
High risk transcatheter valve procedure

## Postcardiotomy VA ECMO

VA ECMO or VAD in PGD after Htx

# Cardiogenic Shock: clinical definition

- **Unresponsive Hypotension**

- Prolonged MAP < 60 mmhg for > 30 min (or decrease in SBP more than 40 mmhg)
- CI < 1,8 l/min/m<sup>2</sup> or < 2,2 l/min/m<sup>2</sup> with **inotropic support**

- **High filling pressures**

- CVP > 14 mmhg
- Wedge pressure > 16 mmhg

- **Inadequate tissue perfusion**

- SVO<sub>2</sub> < 55; Lactate continuously increase (or > 3)
- Alteration in consciousness
- Urine output < 30 cc/h

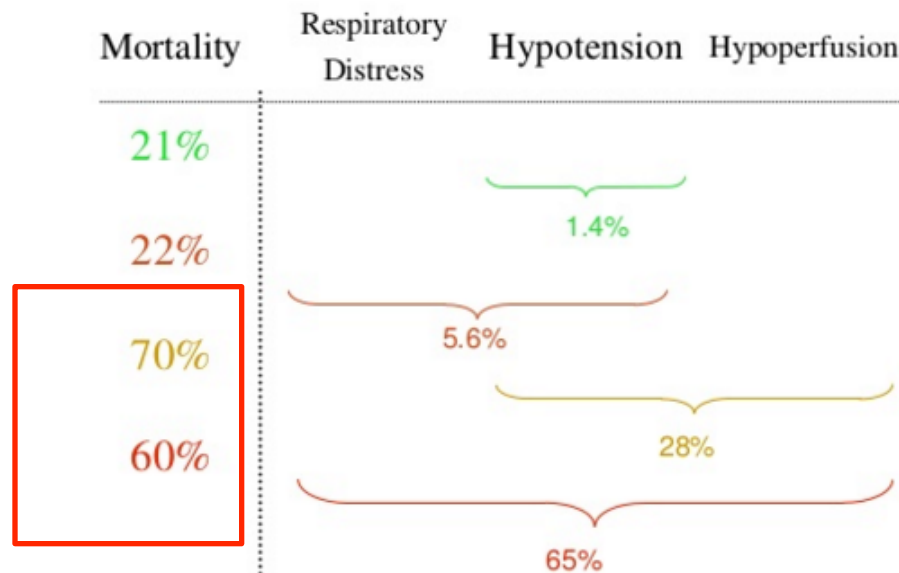
# Cardiogenic Shock Complicating Acute Myocardial Infarction—Etiologies, Management and Outcome: A Report from the SHOCK Trial Registry

Judith S. Hochman, MD, FACC,\* Christopher E. Buller, MD, FACC,† Lynn A. Sleeper, ScD,‡  
Jean Boland, MD,§ Vladimir Dzavik, MD,|| Timothy A. Sanborn, MD, FACC,¶  
Emilie Godfrey, MS, RD,\* Harvey D. White, DSc, FACC,# John Lim, BA,‡ Thierry LeJemtel, MD,\*\*  
for the SHOCK Investigators

*New York, New York; Vancouver and Edmonton, Canada; Watertown, Massachusetts; Liege, Belgium;  
Auckland, New Zealand*

SHOCK Registry JACC Sept. 2000, Supp. A

## Spectrum of Clinical Presentations



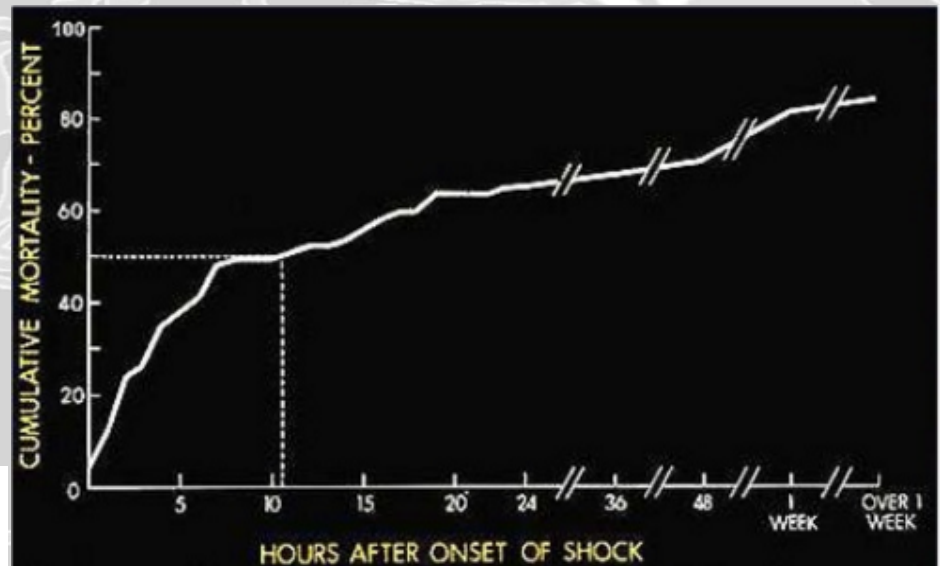
# Cardiogenic Shock survival after diagnosis

**In-hospital mortality 60- 80%**

## Long-Term Mortality of Patients With Acute Myocardial Infarction in the United States and Canada Comparison of Patients Enrolled in Global Utilization of Streptokinase and t-PA for Occluded Coronary Arteries (GUSTO)-I

Padma Karul, PhD; Paul W. Armstrong, MD; Wei-Ching Chang, PhD; C. David Naylor, MD, DPhil;  
Christopher B. Granger, MD; Kerry L. Lee, PhD; Eric D. Peterson, MD, MPH; Robert M. Califf, MD;  
Eric J. Topol, MD; Daniel B. Mark, MD, MPH

- **56% GUSTO I study**



**Figure 7.** Cumulative mortality from the time of onset of shock. Half the group are dead within 10.2 hr (thin dashed line). Overall mortality is 86 percent.

Hasdai et al. JACC 2000;36:687

- **67% Shock Trial Registry**

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## Cardiogenic Shock Complicating Acute Myocardial Infarction—Etiologies, Management and Outcome: A Report from the SHOCK Trial Registry

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for the SHOCK Investigators

New York, New York; Vancouver and Edmonton, Canada; Watertown, Massachusetts; Liege, Belgium;  
Auckland, New Zealand

PROFILE-LEVEL	Official Shorthand	General time frame for support
INTERMACS LEVEL 1	“Crash and burn”	Hours
INTERMACS LEVEL 2	“Sliding fast”	Days to week
INTERMACS LEVEL 3	Stable but Dependent	Weeks
INTERMACS LEVEL 4	“Frequent flyer”	Weeks to few months, if baseline restored
INTERMACS LEVEL 5	“Housebound”	Weeks to months
INTERMACS LEVEL 6	“Walking wounded”	Months, if nutrition and activity maintained
INTERMACS LEVEL 7	Advanced Class III	

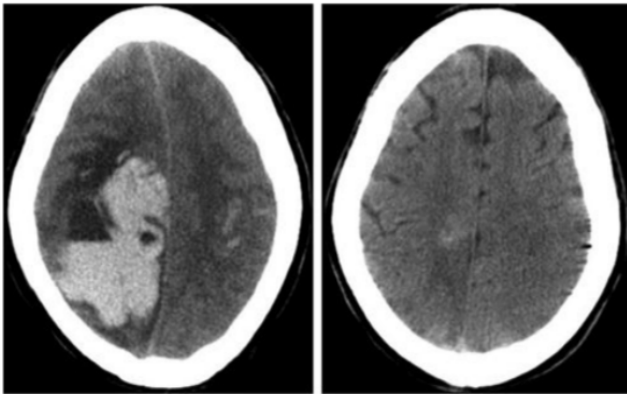


# «Science never solves problem without creating ten more...»

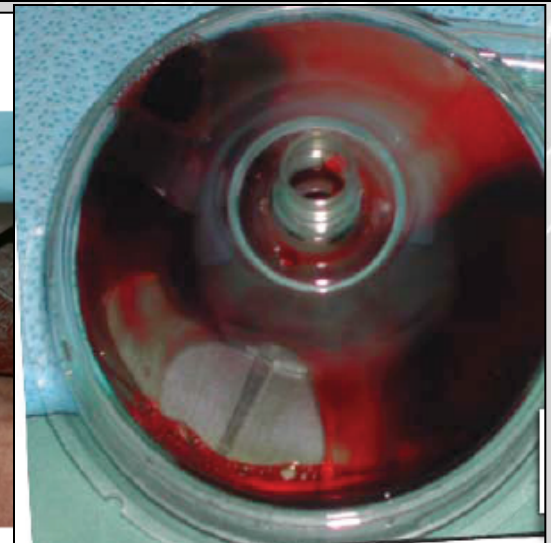
George Bernard Shaw



CNS bleeding



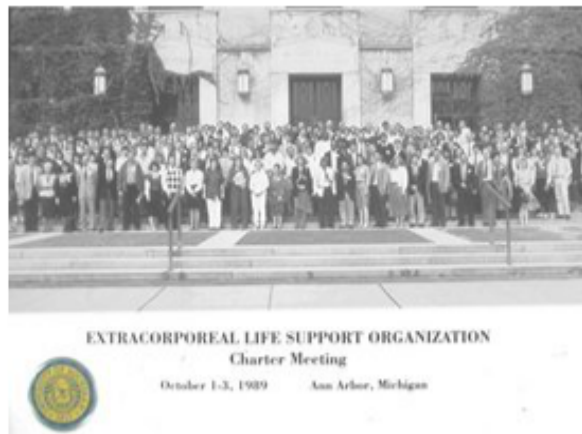
Pulmonary bleeding



# V-A ECMO critical issues

- Bleeding and thrombosis
- Aortic flow competition- *Harlequin syndrome*
- Left heart overload
- Pulmonary function impairment
- Emolysis

- Data since 1979
- Charter Meeting 1989
- **230 centers**
- International



## ELSO: What do we do?

- Registry

	Total	Surv ECLS	Surv to DC
Neonatal			
Respiratory	27,007	22,782 84%	20,093 74%
Cardiac	5,425	3,339 62%	2,206 41%
ECPR	980	626 64%	388 40%
Pediatric			
Respiratory	6,149	4,034 66%	3,496 57%
Cardiac	6,784	4,443 65%	3,388 50%
ECPR	2,071	1,123 54%	840 41%
Adult			
Respiratory	5,146	3,317 64%	2,905 56%
Cardiac	4,042	2,255 56%	1,636 40%
ECPR	1,238	476 38%	355 29%
<b>Total</b>	<b>58,842</b>	<b>42,395 72%</b>	<b>35,307 60%</b>

# ECLS Registry Report

## International Summary

January, 2014



Extracorporeal Life Support Organization  
2800 Plymouth Road  
Building 300, Room 303  
Ann Arbor, MI 48109

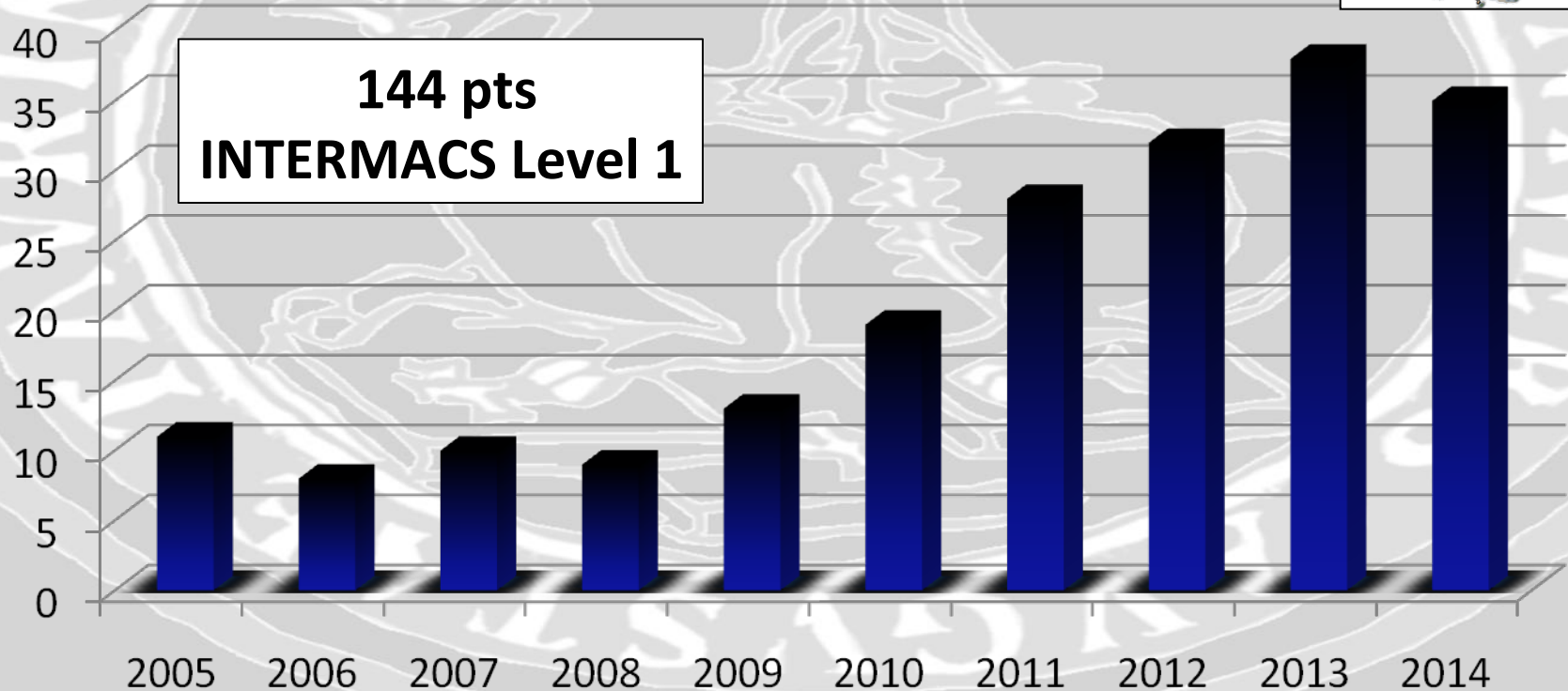
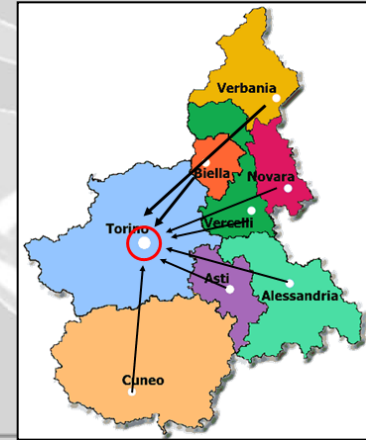
### Overall Outcomes

## Mechanical and Patient related Complications

	0-30 Days	31 Days and <1 Year	1 Year and <16 Years	>16 Years
<b>Mechanical</b>				
Oxygenator failure	7.4 (24)	8.1 (29)	9 (43)	15.1 (36)
Tubing rupture	0.3 (25)	0.6 (50)	0.7 (47)	0.2 (0)
Pump malfunction	1.6 (29)	2.1 (35)	2.1 (50)	0.7 (28)
Cannula problems	6.1 (33)	5.6 (38)	6.3 (42)	4.4 (27)
<b>Patient-related</b>				
<b>ICH</b>				
Cannula site bleeding	11.3 (23)	5.7 (29)	3.8 (21)	1.7 (7)
Surgical site bleeding	10.4 (30)	11.9 (40)	17.6 (52)	20.9 (39)
Cardiac tamponade	31.7 (30)	33 (39)	28.8 (48)	25.5 (34)
Clinical seizures	6.1 (27)	5.1 (38)	5.1 (50)	5.7 (27)
	7.3 (29)	9 (26)	4.5 (21)	2.1 (15)

# Hub-and-Spoke 2005-2014

## Short term VAD/ECMO implants



# Mechanical assistance implanted

**N= 144**

*n cases*

- |                   |     |
|-------------------|-----|
| • VA ECMO         | 113 |
| • VA ECMO to RVAD | 5   |
| • RVAD            | 13  |
| • VA ECMO to LVAD | 9   |
| • LVAD            | 4   |

# VA ECMO in Cardiogenic shock

## University of Turin

n= 113 patients INTERMACS level 1

### Results

---

Death	41/113	36%
Recovery	38/113	34%
Bridge to emergency transplant	28/113	25%
Bridge to long term LVAD	6/113	5%

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Survival to discharge	59/113	51%
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# VA ECMO Risk factors for death

## University of Turin n= 113 patients

### *Multivariate analysis for hospital death*

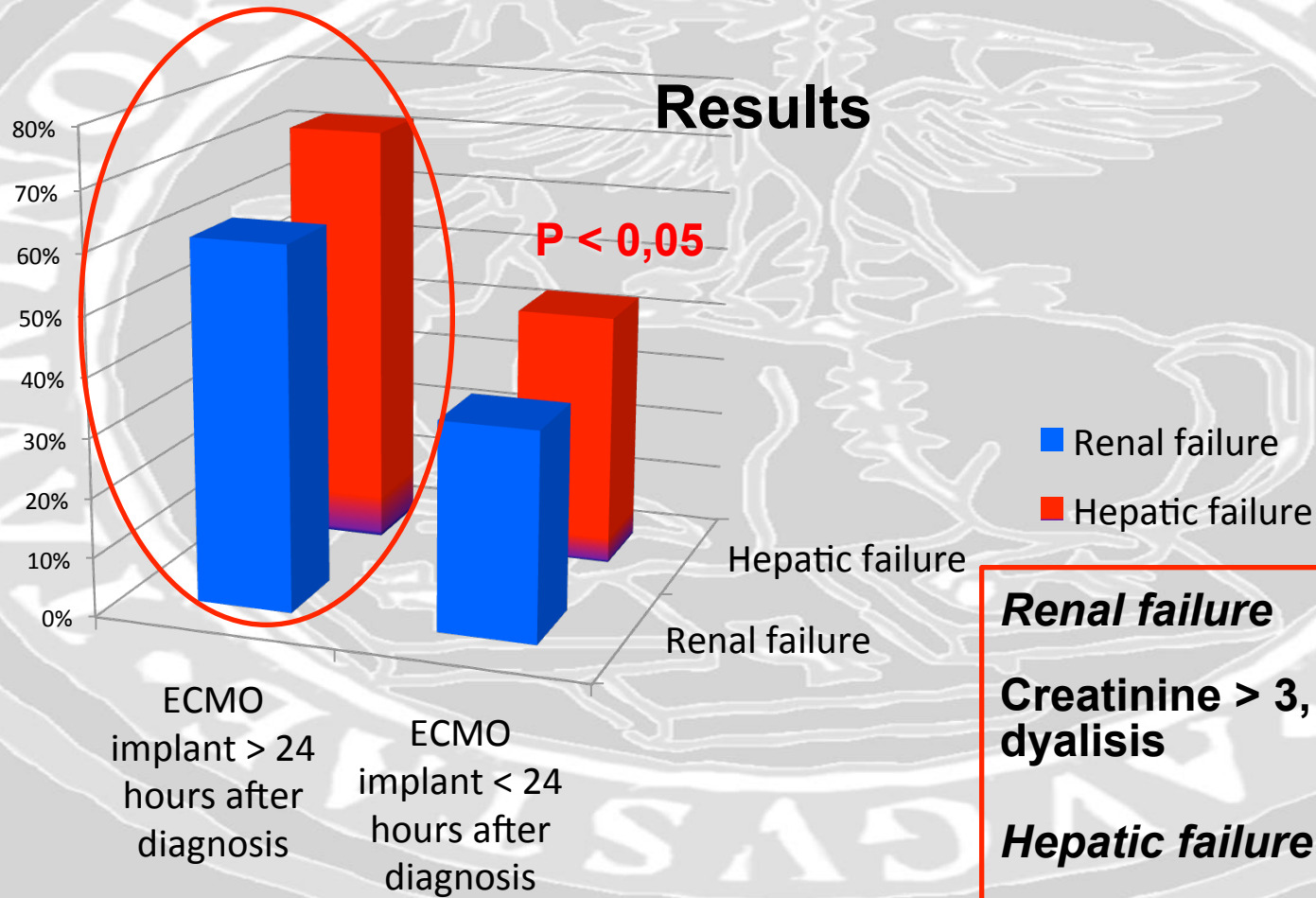
Postcardiotomic	p 0,003
No left ventricular venting	p 0,01
Central approach	p 0,01
External ECMO	p 0,22
“Unstable” ECMO	p 0,02
Previous cardiac arrest (CPR)	p 0,15
Miocardial infarction	p 0,32
Renal failure (Creatinine > 3, oligo- anuria or dialysis)	p 0,04
Hepatic failure (INR> 1,5; AST or ALT > 3 times)	p 0,03



# VA ECMO in Cardiogenic shock

## University of Turin

n= 113 patients INTERMACS level 1



### ***Renal failure***

**Creatinine > 3, oligo- anuria or dialysis**

### ***Hepatic failure***

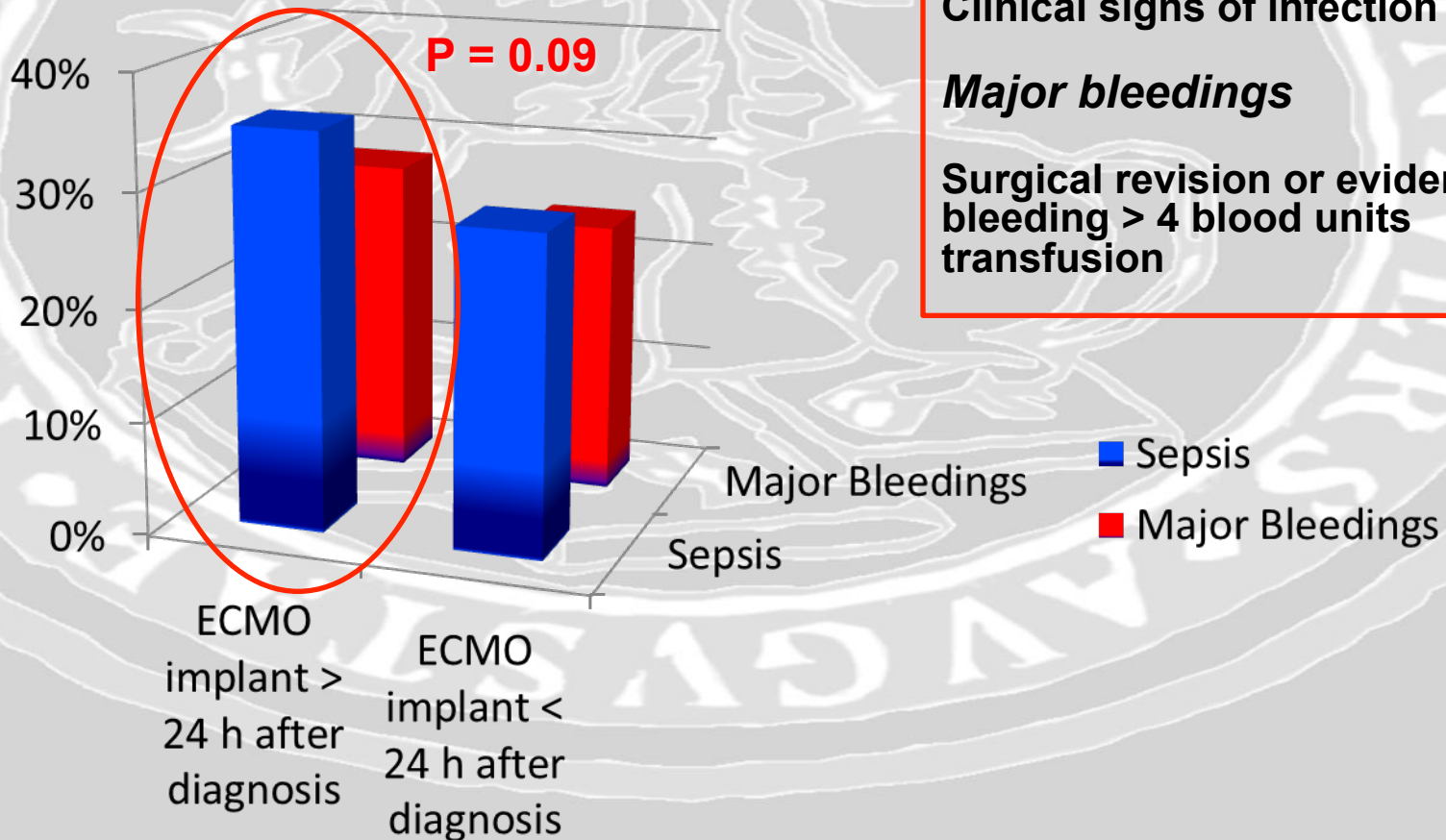
**INR > 1,5; AST or ALT > 3 times**

# VA ECMO in Cardiogenic shock

## University of Turin

n= 113 patients INTERMACS level 1

### Results



#### *Sepsis*

Positive blood cultures and Clinical signs of infection

#### *Major bleedings*

Surgical revision or evidence of bleeding > 4 blood units transfusion

# When is too early or too late?

LVEF (%)

LVEDD (mm)

>90

Cardiac Index (l/min/m<sup>2</sup>)

CVP tor  
>20

APACHE II  
>20

Inotropes (days)

Hepatic failure (bilirubin)

RVF (RVFAC)%  
<20

Ventilation (days) 0

MCS (days)

MOF (organs)

35 30 25 20 15 <10

65 70 75 80 85  
*reversible* *irreversible*



2,4 2,2 2 1,8 1,6 1,4

10 12 14 16 18

<10 <10 11-15 11-15 16-20

0 0 1 2-4 5-10 >10

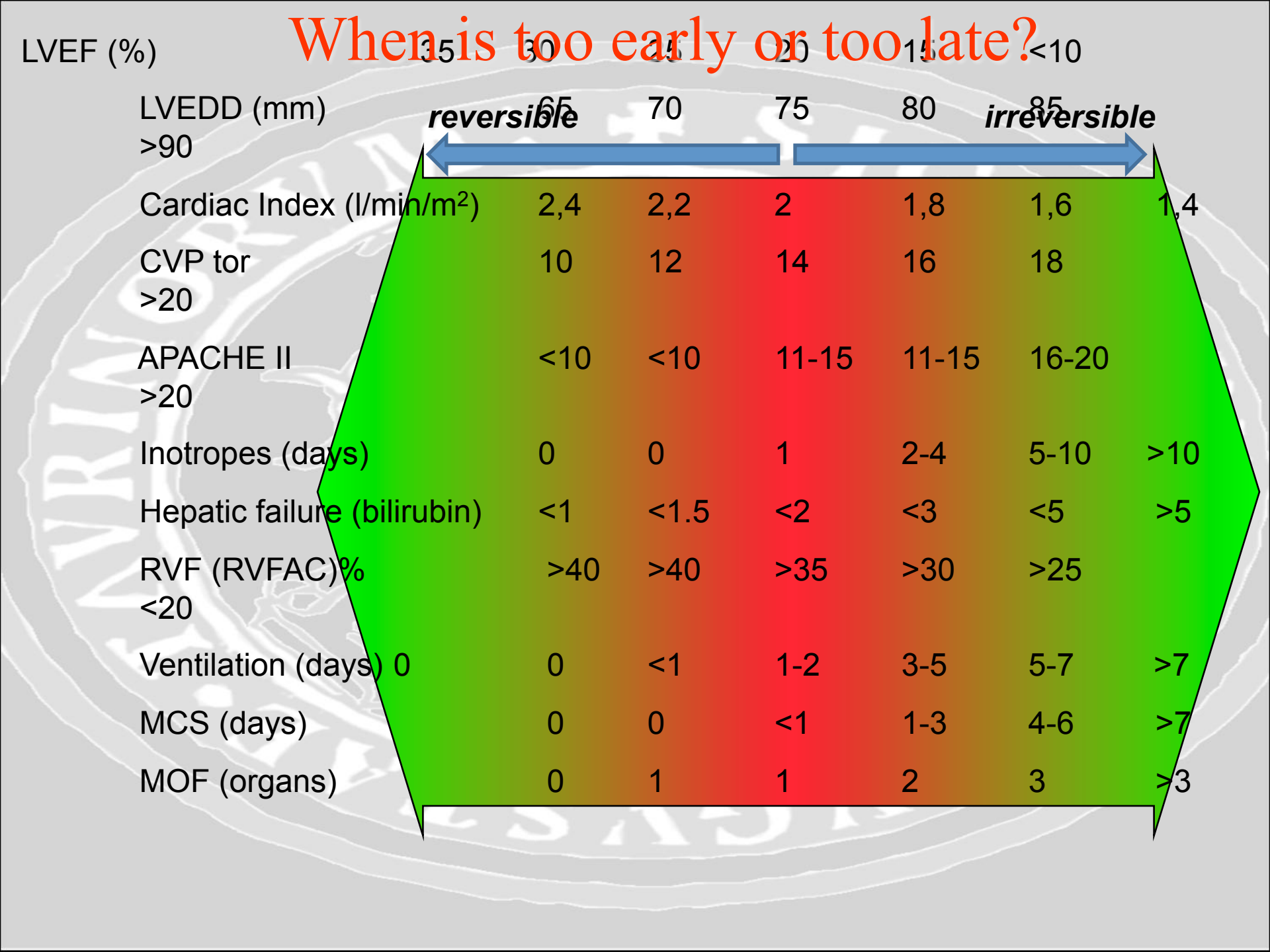
<1 <1.5 <2 <3 <5 >5

>40 >40 >35 >30 >25

0 0 <1 1-2 3-5 5-7 >7

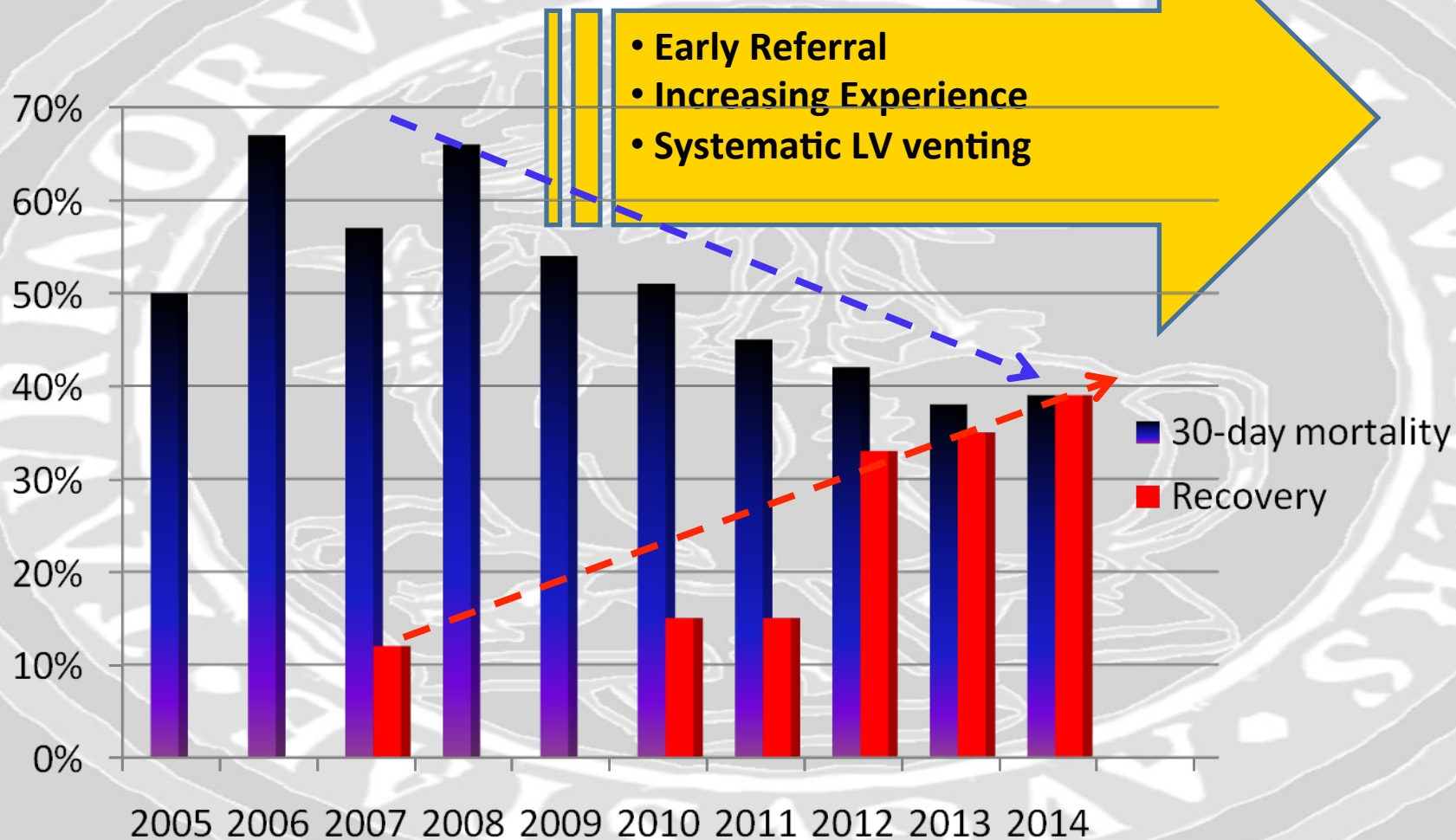
0 0 <1 1-3 4-6 >7

0 1 1 2 3 >3



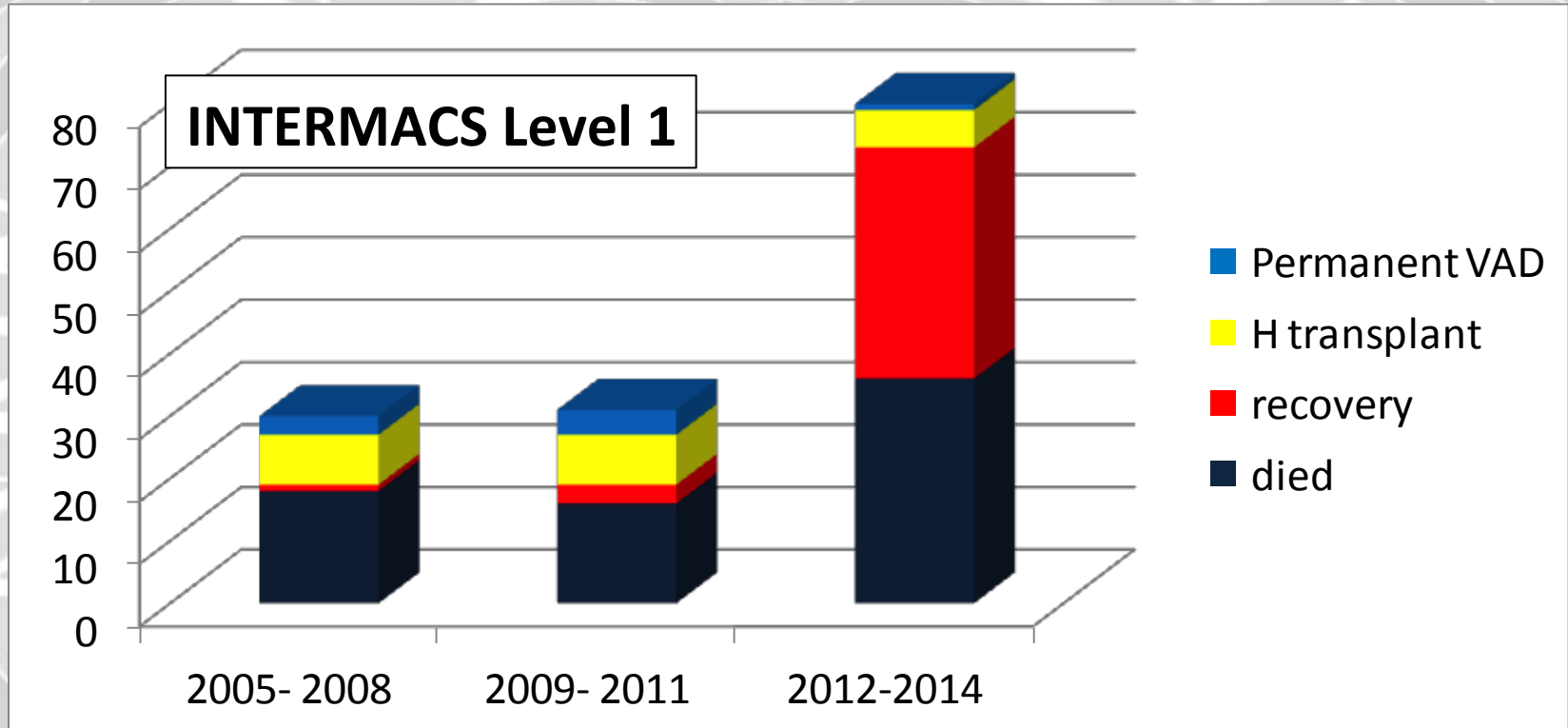
# VA ECMO changing outcomes

University of Turin n= 113 patients



**INTERMACS Level 1**

# VADs-ECMO outcomes



**30-day  
mortality  
60%**

**30-day  
mortality  
52%**

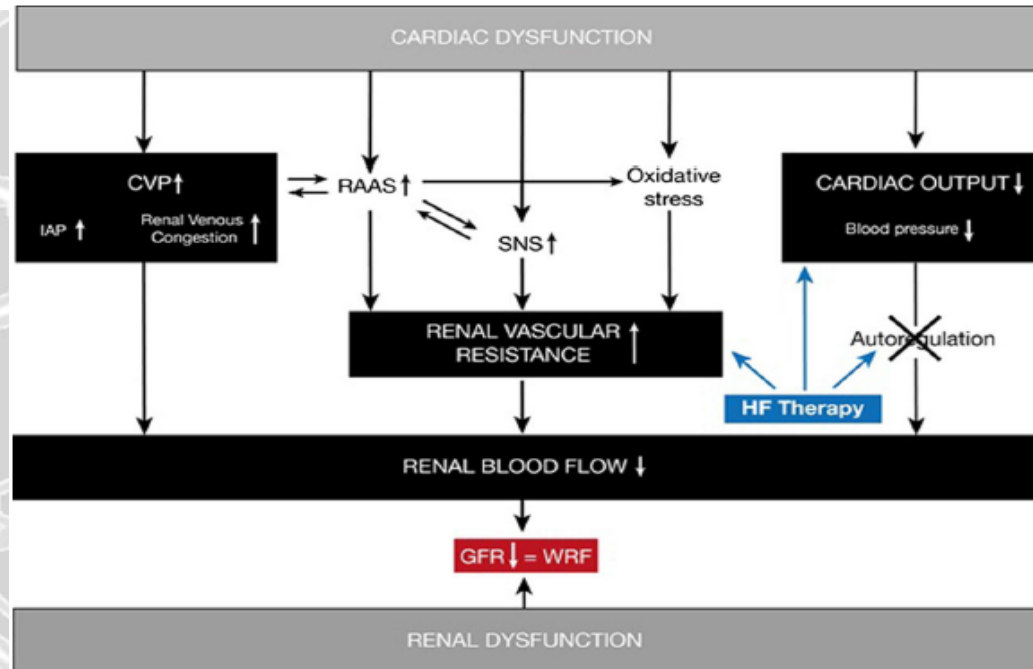
**30-day  
mortality  
44%**

The background of the slide features a large, faint, light-gray watermark of the seal of the University of Göttingen. The seal is circular and contains a central figure of a griffin or eagle with its wings spread, perched on a globe. The Latin text "SIGILLUM UNIVERSITATIS GÖTTINGENSIS" is visible around the perimeter of the seal.

***When the Heart kills  
the Liver and the Kidney***

## The cardiorenal syndrome in heart failure: cardiac? renal? syndrome?

Filippos Triposkiadis · Randall C. Starling ·  
 Harisios Boudoulas · Gregory Giamouzis ·  
 Javed Butler



**Table 1** Classification of acute kidney injury (AKI)

Stage	Creatinine criteria	Urine output (UO) criteria
Risk or stage 1	Creatinine increase $\geq 0.3$ mg/dl or creatinine $\geq 150$ and $< 200\%$ than baseline	UO $< 0.5$ ml/kg/h for 6 h
Injury or stage 2	Creatinine $\geq 200\%$ and $< 300\%$ than baseline	UO $< 0.5$ ml/kg/h for 12 h
Failure or stage 3	Creatinine $\geq 300\%$ than baseline, or $\geq 4.0$ mg/dl and increase $\geq 0.5$ mg/dl	UO $< 0.3$ ml/kg/h for 24 h or anuria for 12 h

STATE-OF-THE-ART PAPER

# Cardiohepatic Interactions in Heart Failure

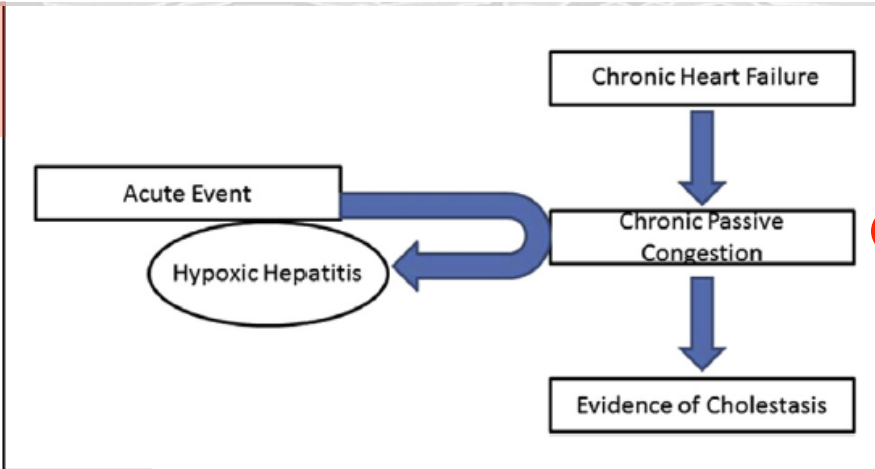
## An Overview and Clinical Implications

Marc D. Samsky, MD, Chetan B. Patel, MD, Tracy A. DeWald, PHARM.D, Alastair D. Smith, MB, CHB,  
G. Michael Felker, MD, MHS, Joseph G. Rogers, MD, Adrian F. Hernandez, MD, MHS

Durham, North Carolina

Hypoxic Hepatitis  
Reversible damage

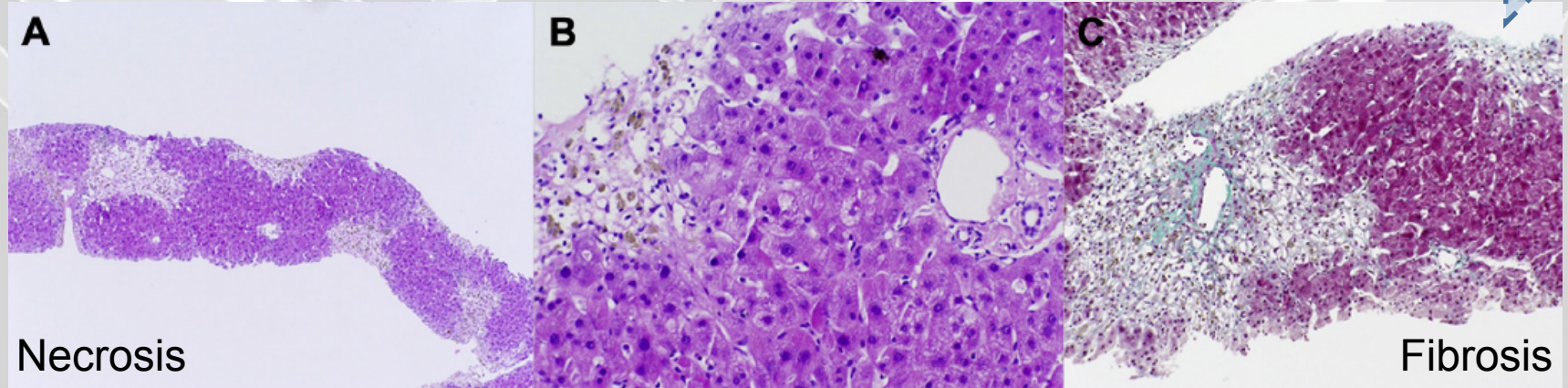
AST, ALT, LDH, INR



Chronic Hepatitis  
Irreversible damage

Bilirubin, Albumin, INR

Time hours days weeks





## Cardiogenic Hepatic Injury-Renal Impairment

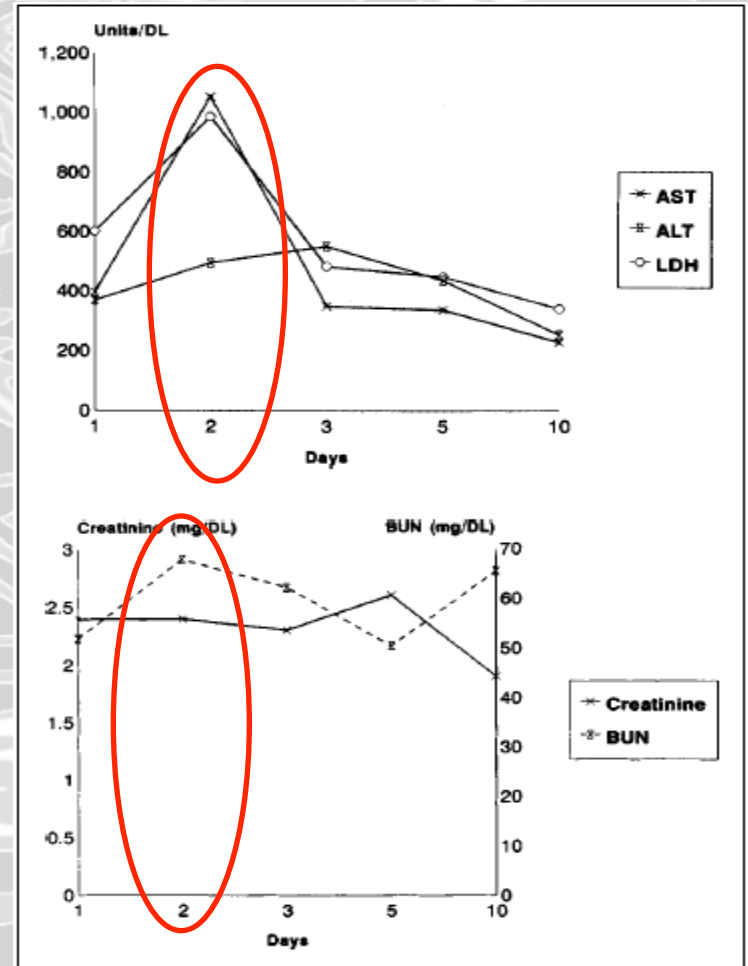
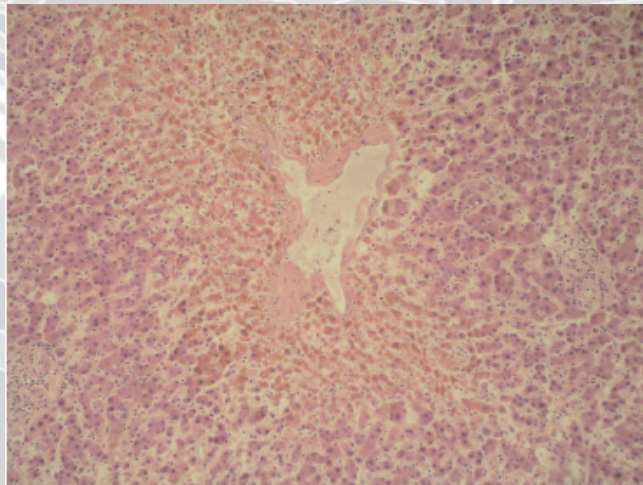
J. E. Naschitz<sup>1</sup>, N. Elias<sup>1</sup>, E. Zuckerman<sup>1</sup>, E. Sabo<sup>2</sup>, E. G. Abinader<sup>3</sup>, D. Yeshurun<sup>1</sup>

**N= 25 in cardiogenic shock in medical therapy**

**CAHIRI** is potentially reversible

**CAHIRI** (ischemic hepatic and renal failure)  
Peak of necrosis within 48 hours

Liver autopsy:  
Central necrosis



# Short term ECMO/VAD - Indications

## SCENARIO

*Cardiology – Cardiac Surgery*

Precardiotomy

**Acute Cardiogenic Shock**

Postcardiotomy

Low cardiac output

- **high dose inotropic drugs**

10 units:

10 gamma/kg/min dobutamine

or

0,1 gamma /kg/min adrenaline

**Inotropic  
drugs**

**IABP**

**Short-term  
ECMO/VAD**

Persistent Low cardiac output

**Check signs of renal**

**and hepatic injury**

Whithin 6 hours

# TIMING: ECMO implant before Liver and Kidney Irreversible ischemic Injury

## Hepatic function cut-off

- AST, ALT < 3 times than normal value
- LDH < 1,5 times than normal value
- INR < 1,5 times than normal value
- Bilirubin and Albumin are usually preserved*

## Kidney function cut-off

- Creatinine < 2 times than normal value and BUN < 60
- UO > 0,5 ml/kg/h
- GF > 40 ml/min/m<sup>2</sup>

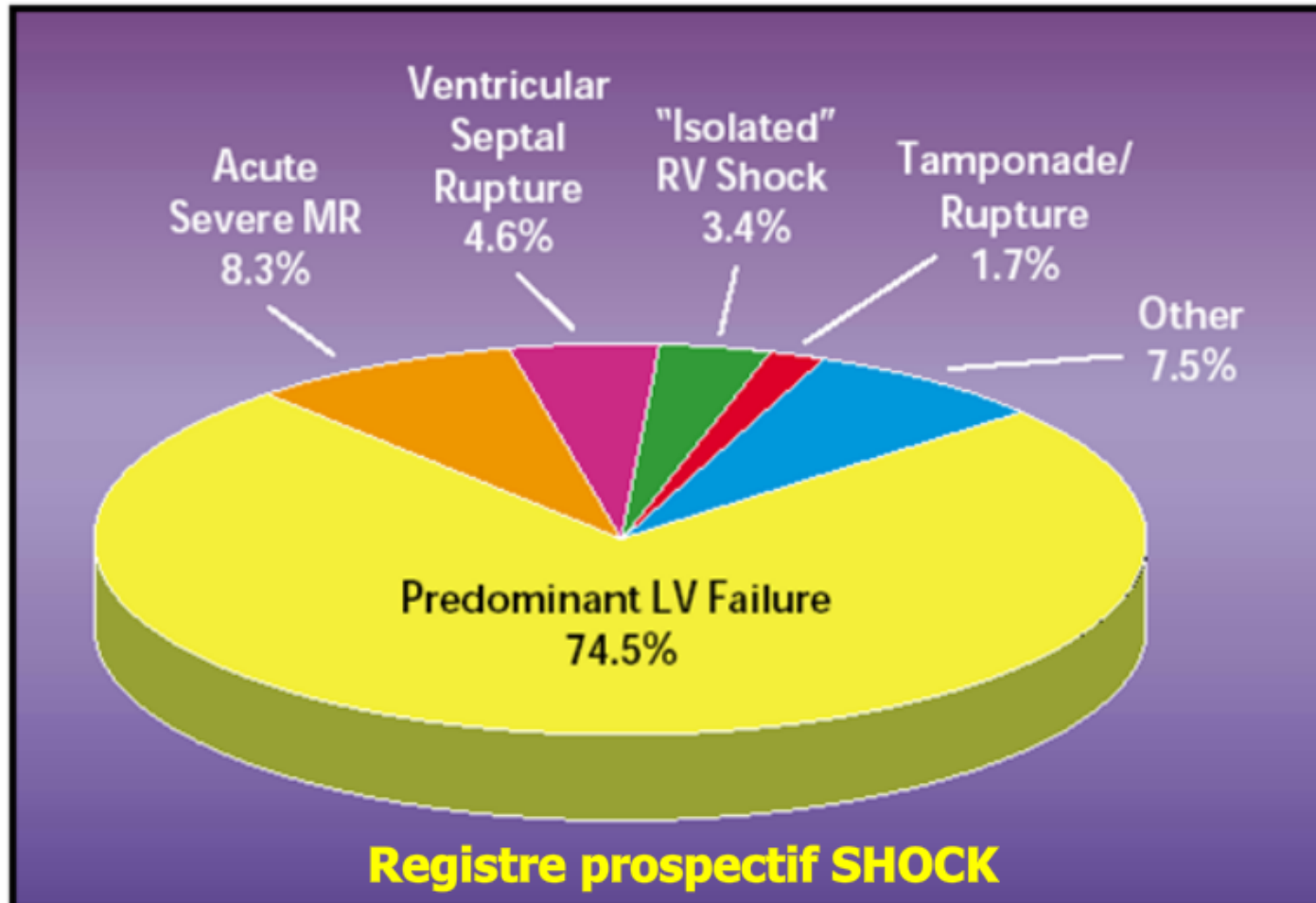
# CARDIOGENIC SHOCK

## *Basic Approach*

- **RAPID REPERFUSION** of the myocardium
- **SUPPORT THE HEMODYNAMIC SYSTEM** assuring adequate fluid volume, maintaining adequate cardiac output with drugs or **mechanical assistance**, correcting gas-exchange abnormalities and providing adequate oxygenation
- It is not clear the chance of **RECOVERY**

# Post-AMI cardiogenic shock

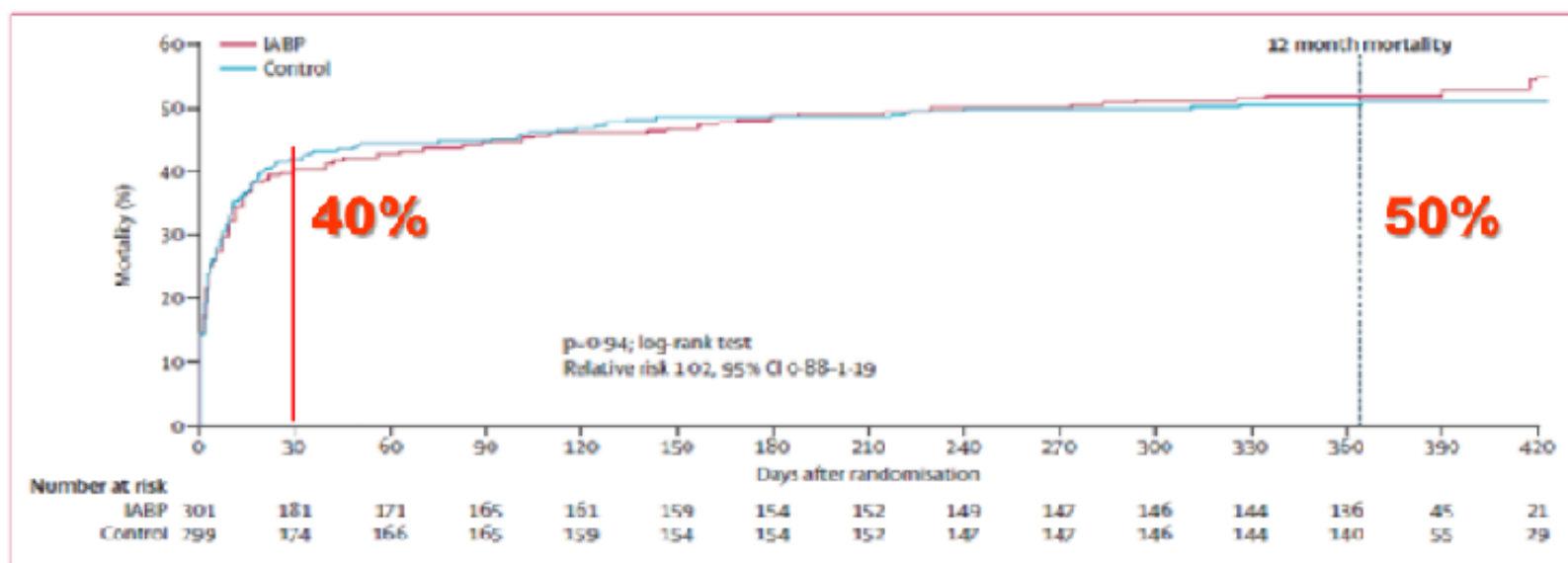
*Hochman, Circ, 1995*





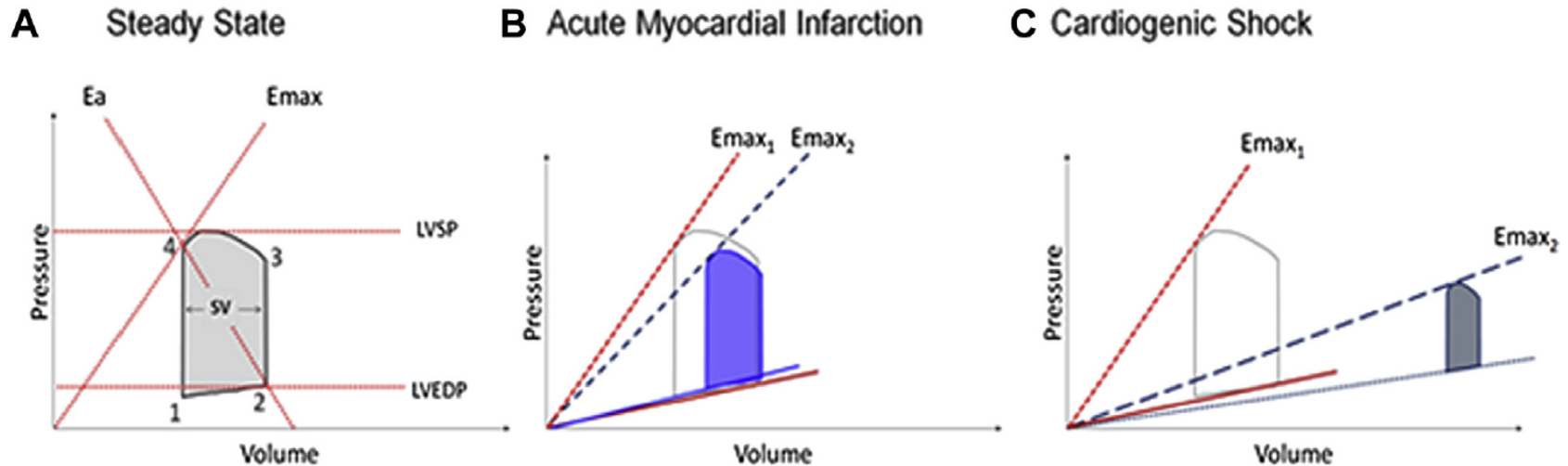
## Intra-aortic balloon counterpulsation in acute myocardial infarction complicated by cardiogenic shock (IABP-SHOCK II): final 12 month results of a randomised, open-label trial

www.thelancet.com Published online September 3, 2013



**2015 SCAI/ACC/HFSA/STS Clinical Expert Consensus Statement on the Use of Percutaneous Mechanical Circulatory Support Devices in Cardiovascular Care (Endorsed by the American Heart Association, the Cardiological Society of India, and Sociedad Latino Americana de Cardiologia Intervencion; Affirmation of Value by the Canadian Association of Interventional Cardiology—Association Canadienne de Cardiologie d'intervention)\***

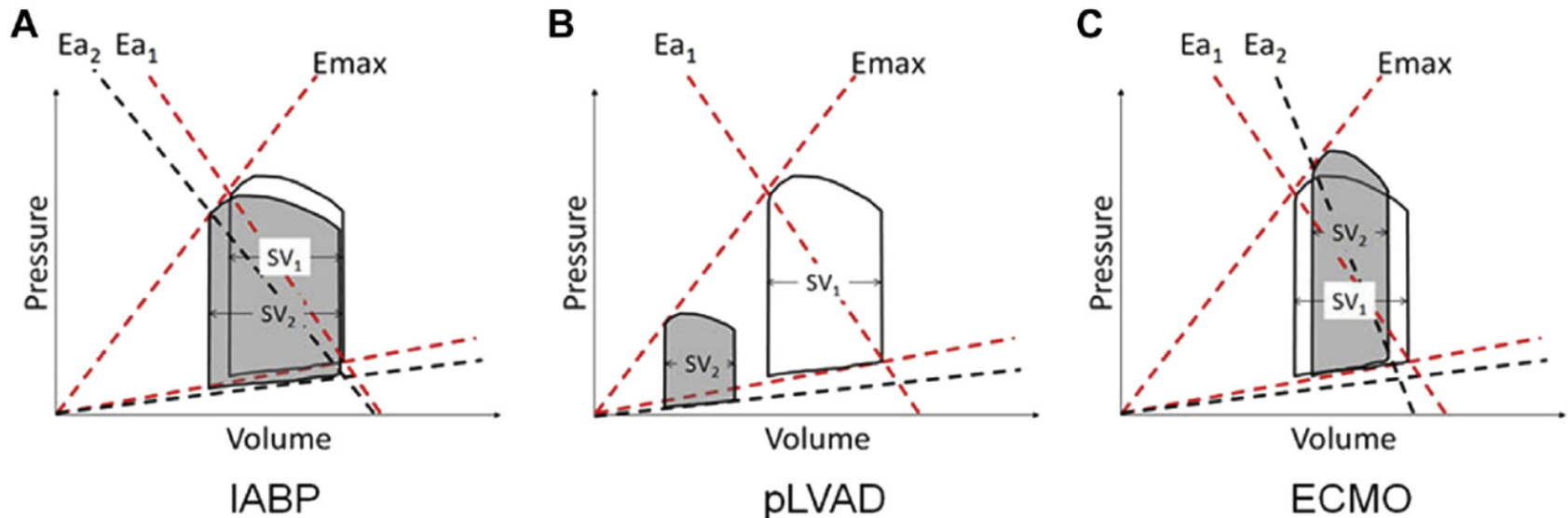
Journal of Cardiac Failure Vol. 21 No. 6 2015



**2015 SCAI/ACC/HFSA/STS Clinical Expert Consensus Statement on the Use of Percutaneous Mechanical Circulatory Support Devices in Cardiovascular Care (Endorsed by the American Heart Association, the Cardiological Society of India, and Sociedad Latino Americana de Cardiologia Intervencion; Affirmation of Value by the Canadian Association of Interventional Cardiology—Association Canadienne de Cardiologie d'intervention)\***

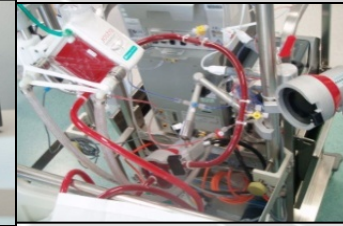
Journal of Cardiac Failure Vol. 21 No. 6 2015

**Left ventricular overload on ECMO**





# Currently available percutaneous MCS



	iVAC 2L <sup>®</sup>	TandemHeart <sup>™</sup>	Impella <sup>®</sup> 5.0	Impella <sup>®</sup> 2.5	Impella <sup>®</sup> CP	ECLS (multiple systems)
Catheter size (F)	11 (expandable)	–	9	9	9	
Cannula size (F)	17	21 venous 12–19 arterial	21	12		17–21 venous 16–19 arterial
Flow (L/min)	Max 2.8	Max. 4.0	Max. 5.0	Max. 2.5	3.7–4.0	Max. 7.0
Pump speed (rpm)	Pulsatile, 40 mL/beat	Max. 7500	Max. 33 000	Max. 51 000	Max. 51 000	Max. 5000
Insertion/placement	Percutaneous (femoral artery)	Percutaneous (femoral artery + vein for left atrium)	Peripheral surgical (femoral artery)	Percutaneous (femoral artery)	Percutaneous (femoral artery)	Percutaneous (femoral artery + vein)
LV unloading	+	++	++	+	+	–
Anticoagulation	+	+	+	+	+	+
Recommended duration of use	–21 days	–14 days	10 days	10 days	10 days	–7 days
CE-certification	+	+	+	+	+	+
FDA	–	+	+	+	+	+
Relative costs	++	+++++	++++	+++	++++	+(+)

Reproduced from Ref. [57] with permission

# MCS Strategy

**Acute setting**

**Chronic setting**

**Short term VAD**

**Long term VAD**

*?unknown patients?*

**BRIDGE to LIFE**

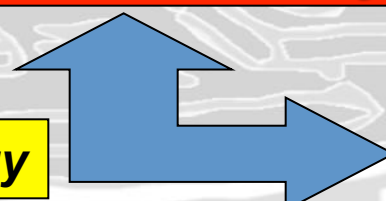
*!known patients!*

**Bridge to Decision**

**Bridge to Recovery**

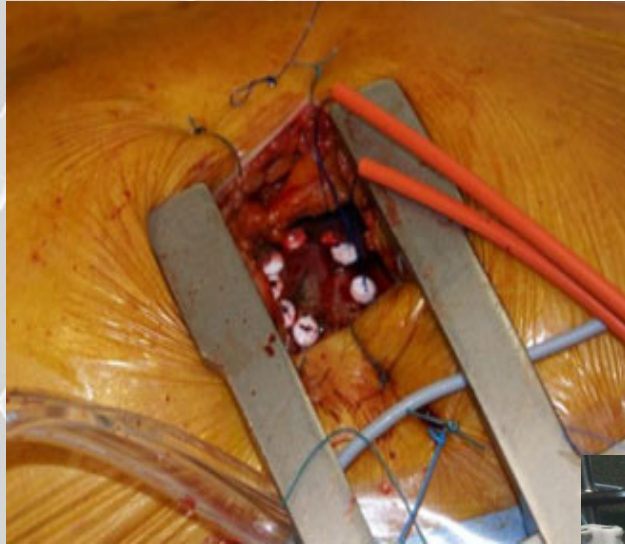
**Dynamic strategy**

**Emergency Htx  
Permanent VAD**

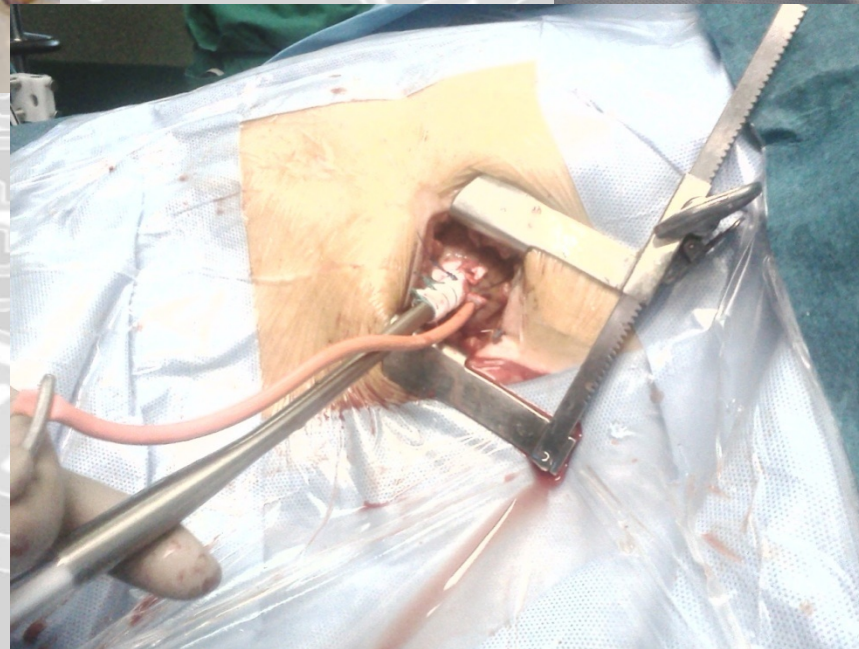
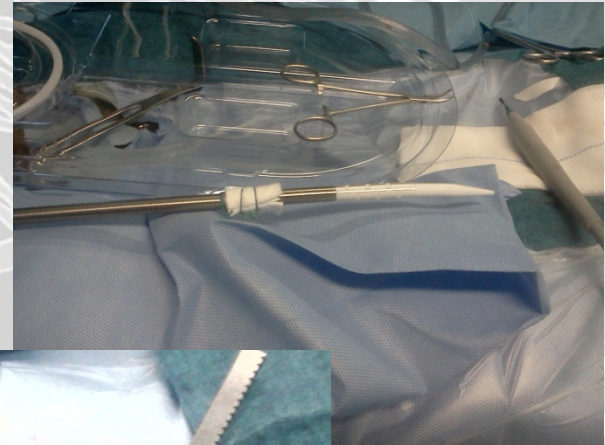


# Transapical Left Ventricular Venting

**Anterolateral Mini left thoracotomy**



**Medtronic High flow cannula 21 Fr**



# Postcardiotomy Heart Failure

## *the IABP Score for VA ECMO indication*

### Prognosis After the Implantation of an Intra-Aortic Balloon Pump in Cardiac Surgery Calculated With a New Score

Harald Hausmann, MD; Evgenij V. Potapov, MD; Andreas Koster, MD; Thomas Krabatsch, MD; Julia Stein; Ruhi Yeter, MD; Marian Kukucka, MD; Ralf Sodian, MD; Hermann Kuppe, MD, PhD; Roland Hetzer, MD, PhD

**TABLE 3. Results of Multivariate Analysis**

Parameter	Odds Ratio (OR)	Range (OR)	<i>P</i>
Adrenaline dose >0.5 $\mu\text{g/kg/BW/min}$	6.6	2.3–18.8	0.0005
Diuresis <100 ml	2.5	1.2–5.4	0.026
SVO <sub>2</sub> <60%	2.5	1.0–6.3	0.048
LAP >15 mm Hg	3.0	1.1–8.5	0.036

Multivariate analysis showed 4 parameters as statistically significant for the prediction of survival or death 1 hour after IABP implantation in patients with cardiac low-output syndrome in cardiac surgery. All other parameters were not significant.

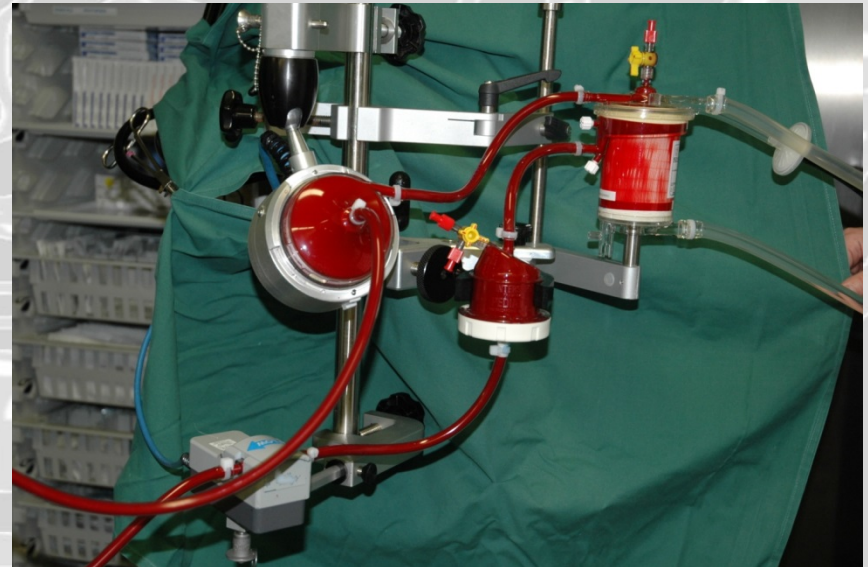
# Short term biventricular support *postcardiotomy setting*

ASAIO Journal 2009

## Levitronix CentriMag to Berlin Heart Excor: A “Bridge to Bridge” Solution in Refractory Cardiogenic Shock

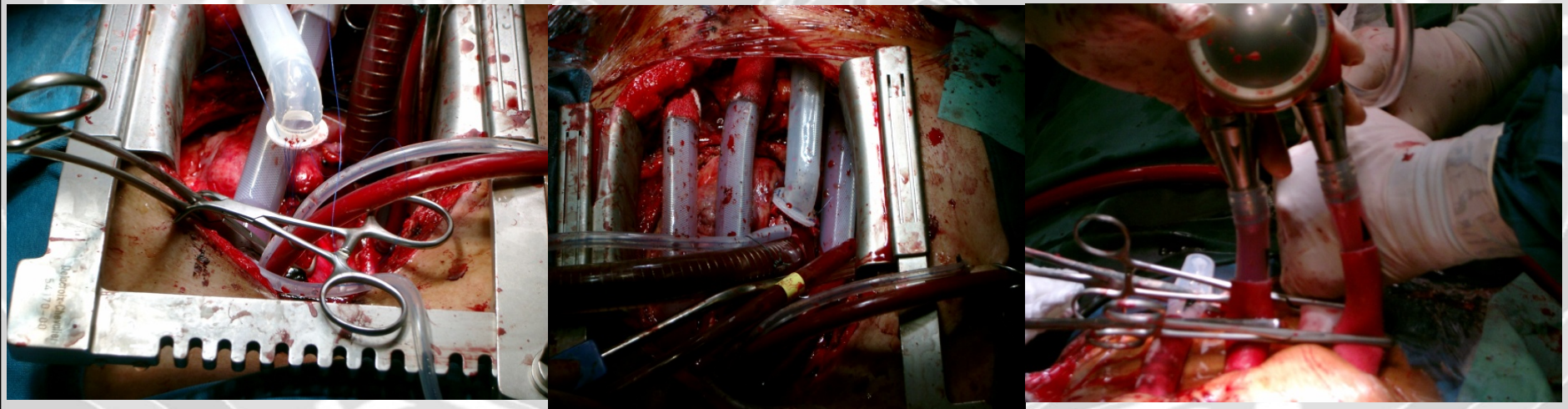
ANTONIO LOFORTE, EVGENIJ POTAPOV, THOMAS KRABATSCH, MICHELE MUSCI, YUGUO WENG,  
MIRALEM PASIC, AND ROLAND HETZER

**centrifugal pump ± oxygenator (ECMO)**



## Levitronix CentriMag to Berlin Heart Excor: A “Bridge to Bridge” Solution in Refractory Cardiogenic Shock

ANTONIO LOFORTE, EVGENIJ POTAPOV, THOMAS KRABATSCH, MICHELE MUSCI, YUGUO WENG,  
MIRALEM PASIC, AND ROLAND HETZER



# V-A ECMO out of hospital



# Out-of-Hospital Resuscitative ECMO



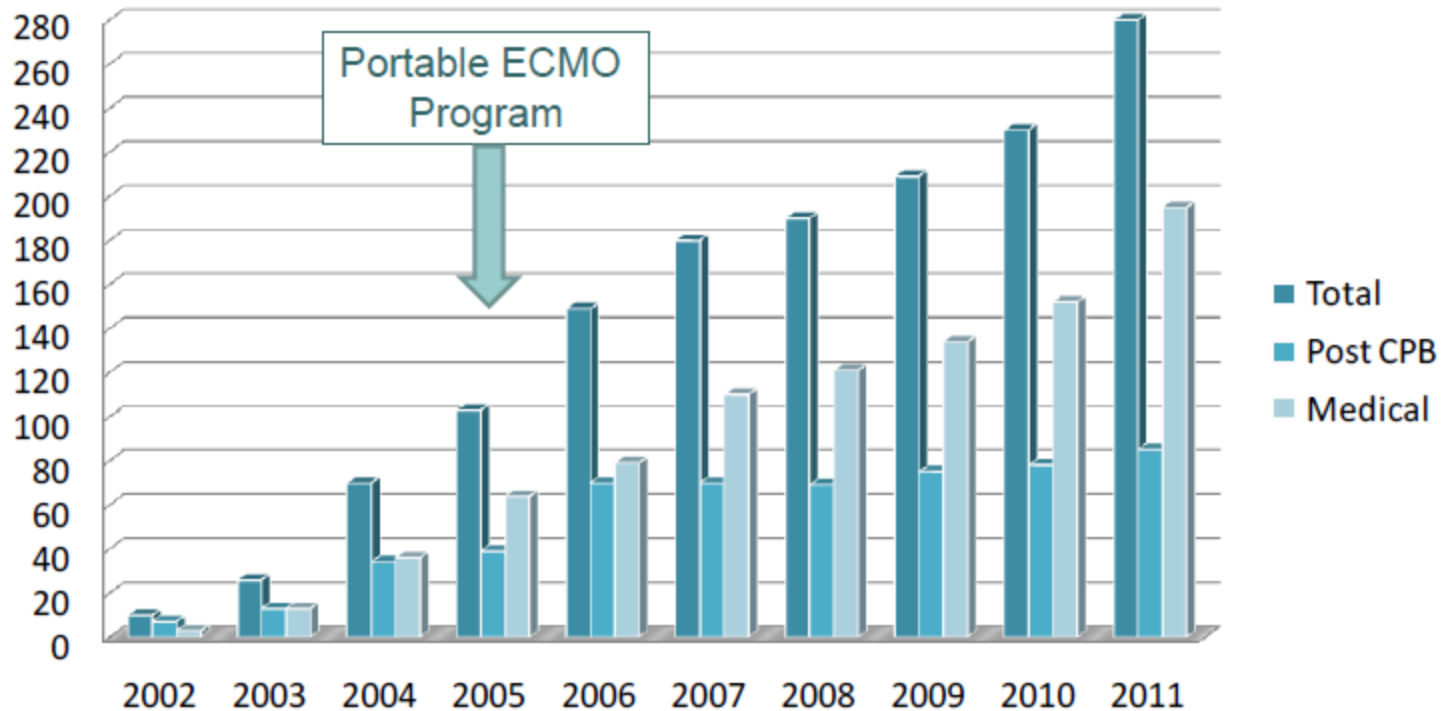
**CARDIO HELP- Maquet**







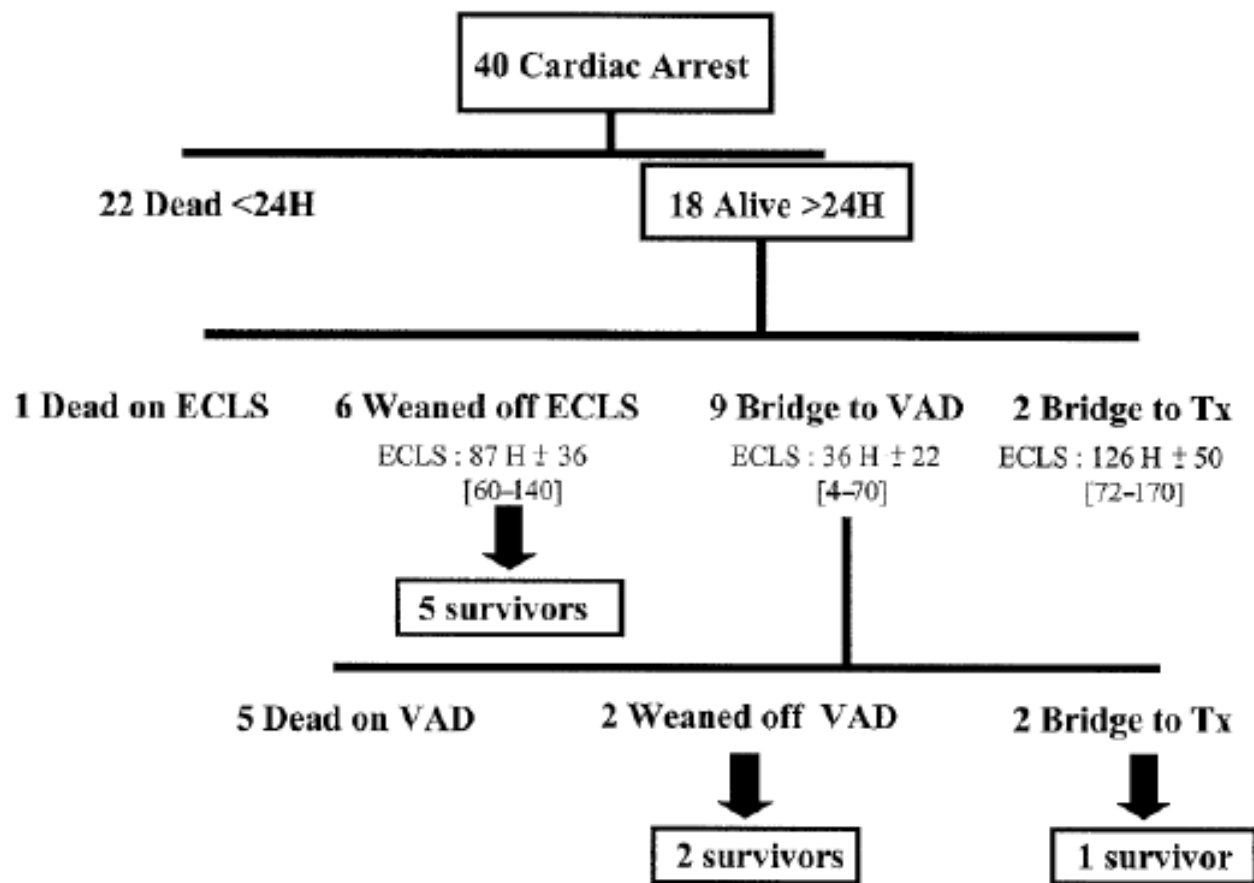
## ECMO program at La Pitié, Paris



# Back from Irreversibility: Extracorporeal Life Support for Prolonged Cardiac Arrest

Massimo Massetti, MD, Marine Tasle, MD, Olivier Le Page, MD, Ronan Deredec, MD,  
Gerard Babatasi, MD, Dimitrios Buklas, MD, Sylvain Thuaudet, MD,  
Pierre Charbonneau, MD, Martial Hamon, MD, Gilles Grollier, MD,  
Jean Louis Gerard, MD, and André Khayat, MD

Ann Thorac Surg 2005;79:178-84



# Extracorporeal life support following out-of-hospital refractory cardiac arrest



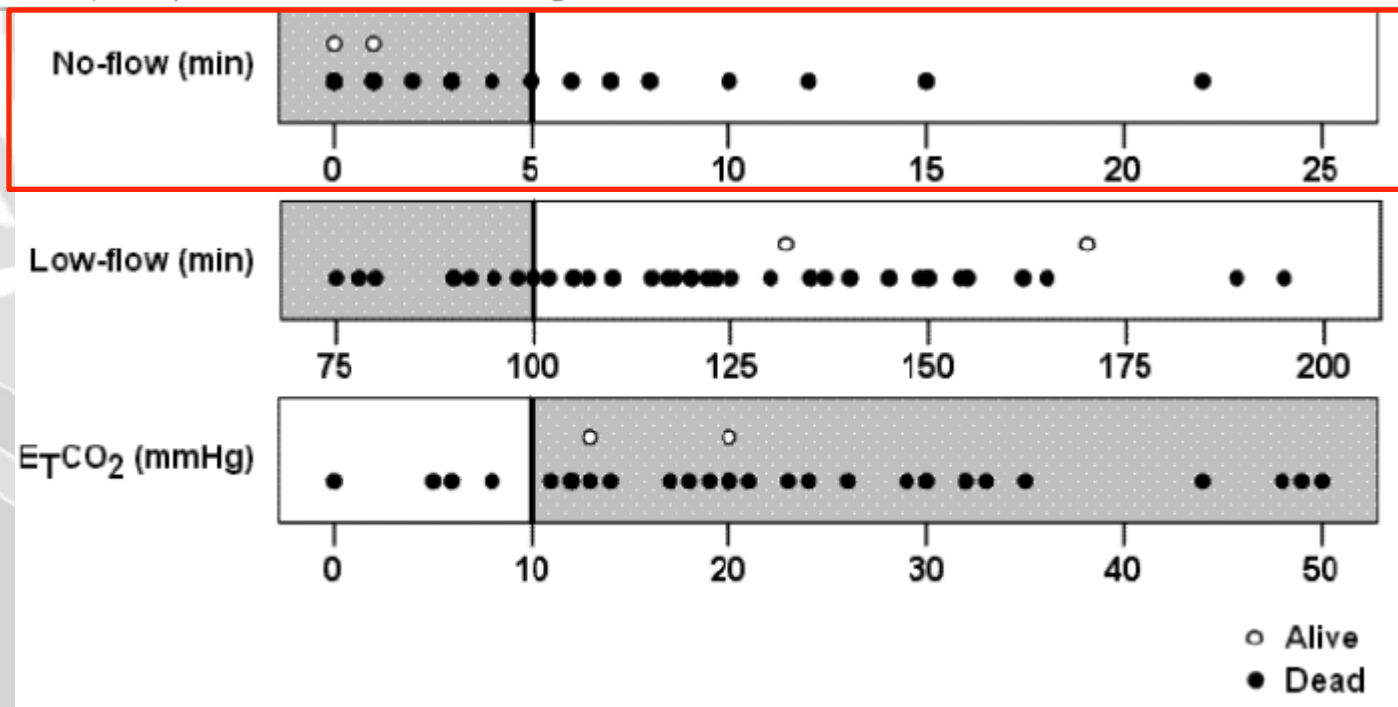
Morgan Le Guen<sup>1</sup>, Armelle Nicolas-Robin<sup>1</sup>, Serge Carreira<sup>1</sup>, Mathieu Raux<sup>1</sup>, Pascal Leprince<sup>2</sup>, Bruno Riou<sup>3\*</sup>, Olivier Langeron<sup>1</sup>

*Critical Care* 2011, 15:R29

## Abstract

**Introduction:** Extracorporeal life support (ECLS) has recently shown encouraging results in the resuscitation of in-hospital (IH) refractory cardiac arrest. We assessed the use of ECLS following out-of-hospital (OH) refractory cardiac arrest.

**Methods:** We evaluated 51 consecutive patients who experienced witnessed OH refractory cardiac arrest and received automated chest compression and ECLS upon arrival in the hospital. Patients with preexisting severe hypothermia who experienced IH cardiac arrest were excluded. A femorofemoral ECLS was set up on admission to the hospital by a mobile cardiothoracic surgical team.



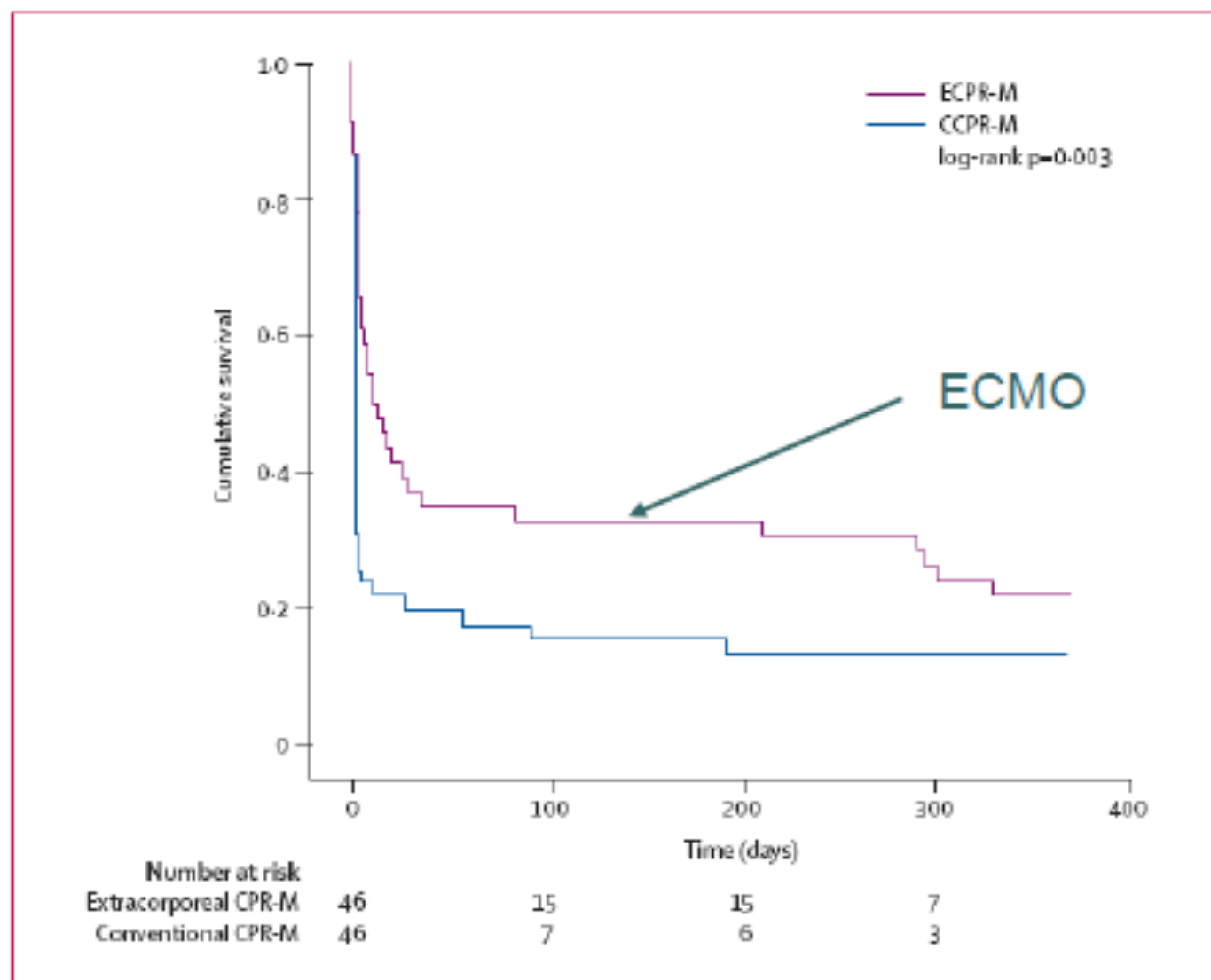
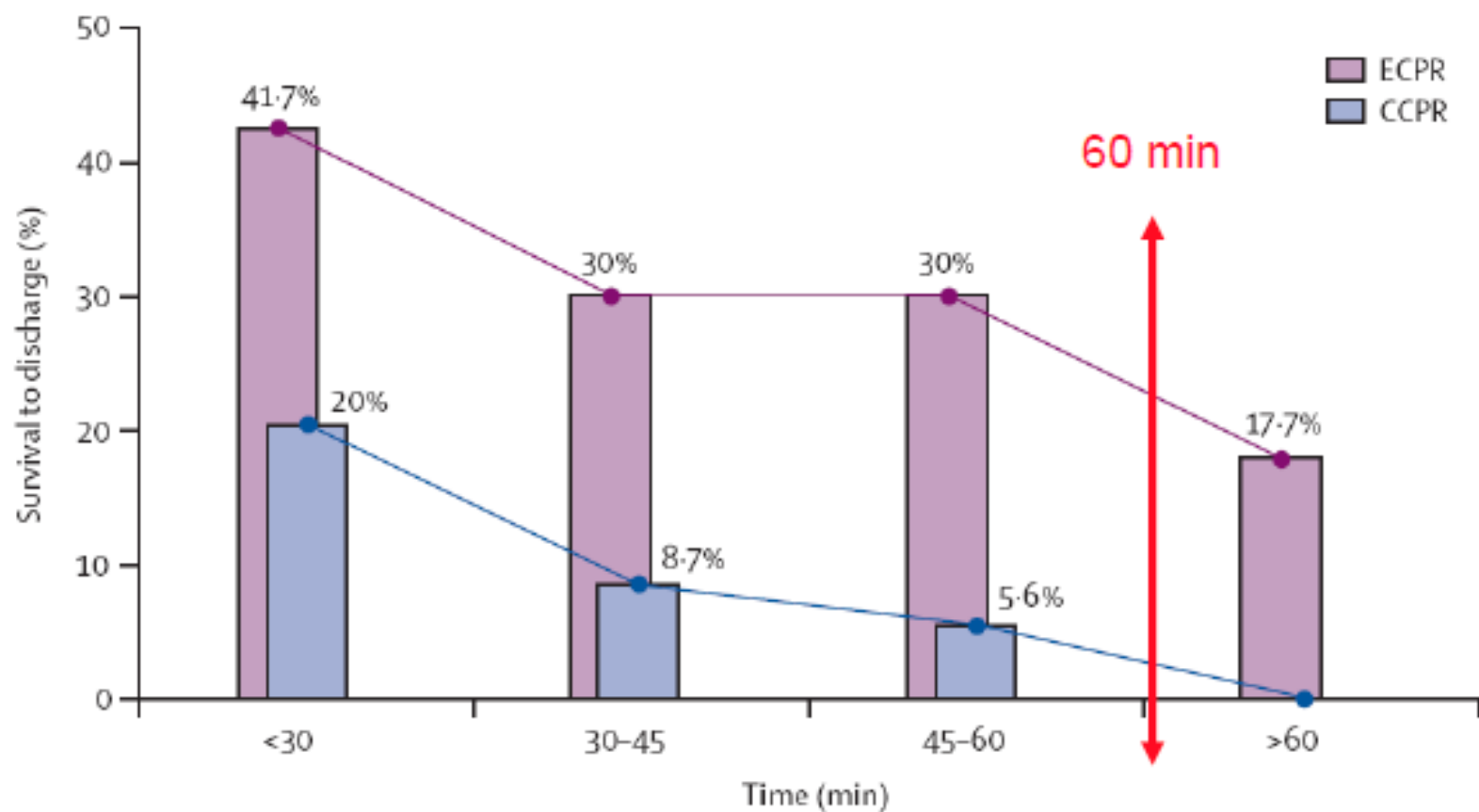


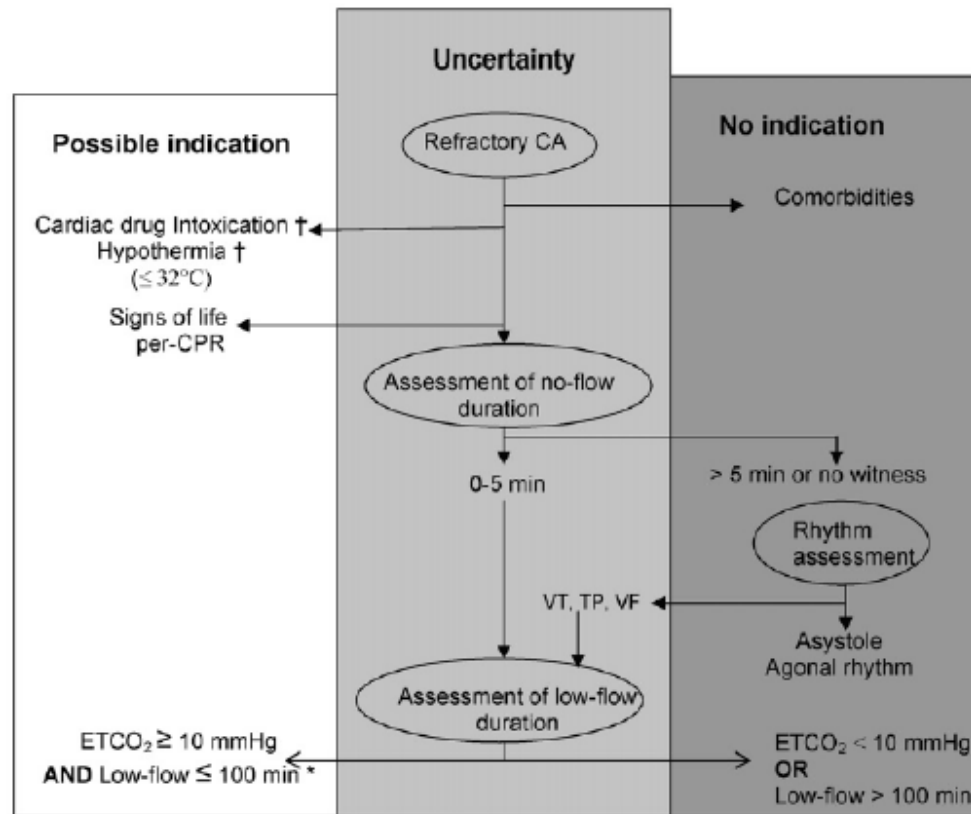
Figure 3: Kaplan-Meier plot of the survival curves in the extracorporeal CPR-M and conventional CPR-M groups for 1 year



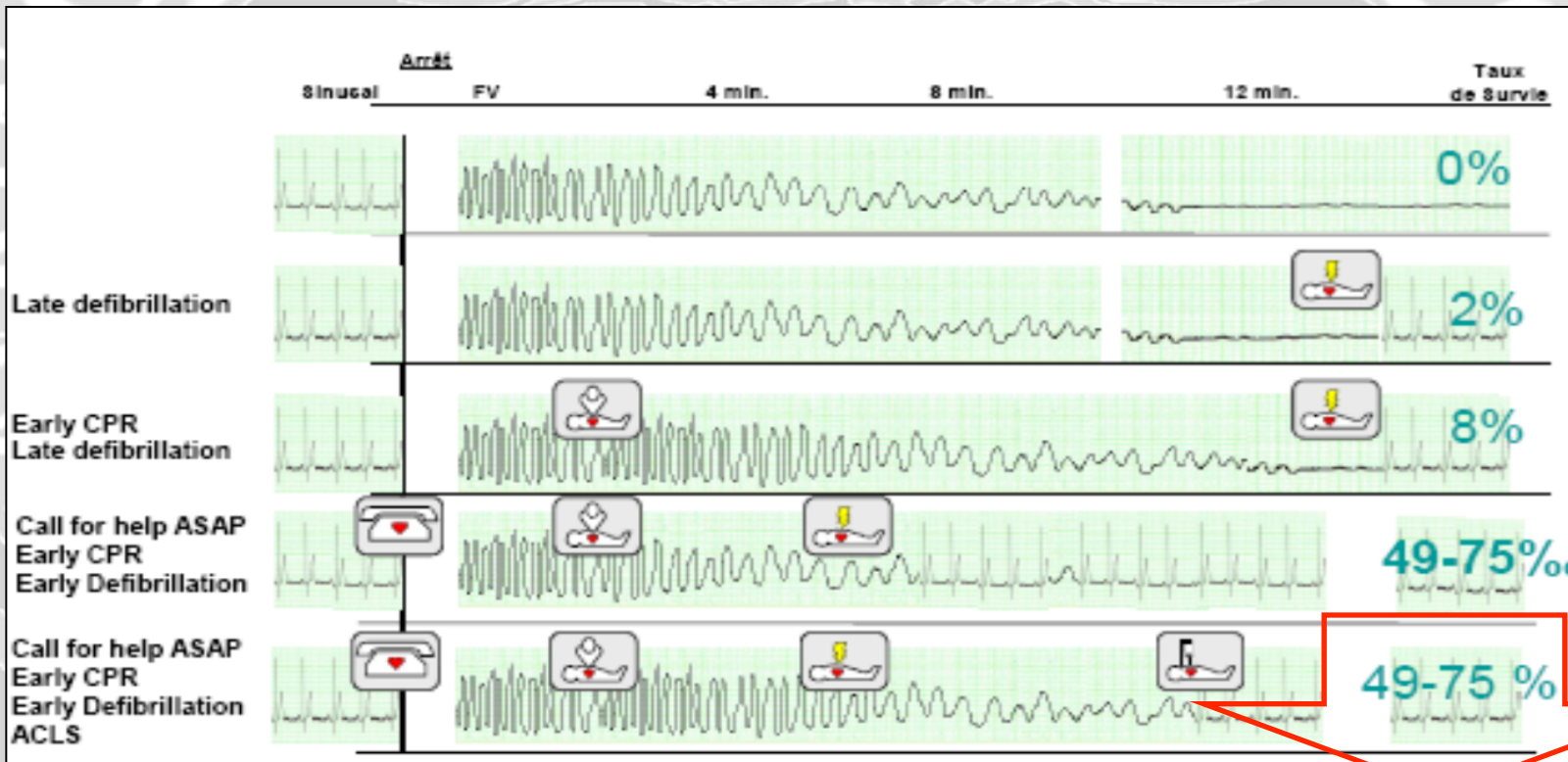
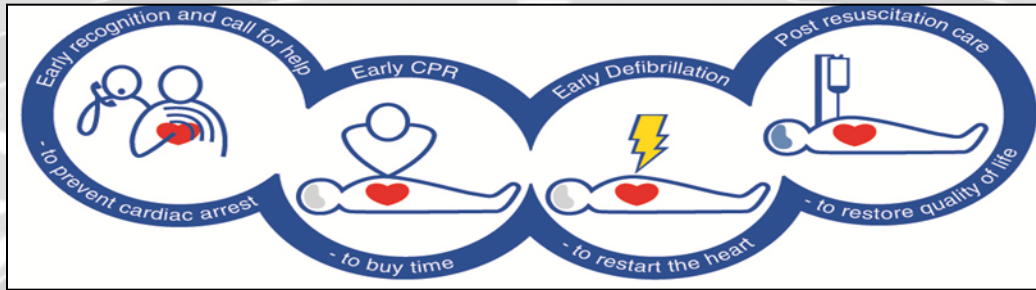
**Figure 1: Relation between CPR duration and the survival rate to discharge**  
ECPR=extracorporeal CPR. CCPR=conventional CPR.

# Guidelines for indications for the use of extracorporeal life support in refractory cardiac arrest<sup>☆</sup>

Annales Françaises d'Anesthésie et de Réanimation 28 (2009) 187-190



# Survival Chain



L.Becker, A.H.A. datas

**V-A ECMO**





# CONCLUSIONS

- Timing of ECMO implantation is strictly dependent from renal and hepatic function
- The earlier the better: within 6-12 hs after inotropes and IABP if renal and hepatic dysfunction progresses
- Systematic LV apical venting
- In out-of-hospital: only no-flow, low-flow times and ETCO<sub>2</sub> drive the timing and indication