

# AER 2019



**AER**

ACTUALITÉS EN RÉANIMATION

**25<sup>ème</sup> AER : 19 & 20 novembre 2020**

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# Assistances circulatoires « légères »

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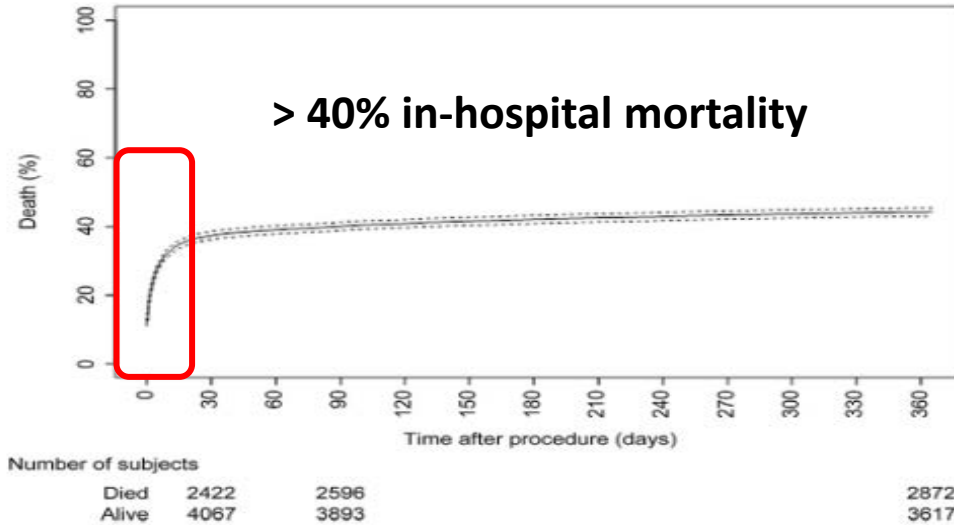
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- Grant/Research Support
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### Company

- Maquet, Abiomed, Daichii Saicko, Térumo
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- Abiomed, Abott, Thoratec, Sorin, Maquet, Pfizer, Medtronic

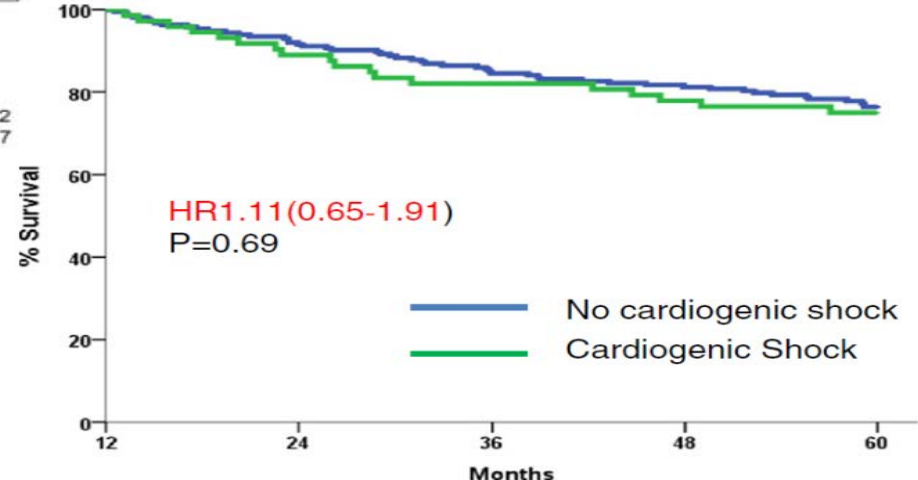
# High and early mortality: active management +++



*Kunadian et al, JACC Cardiovasc Intervention 2014*

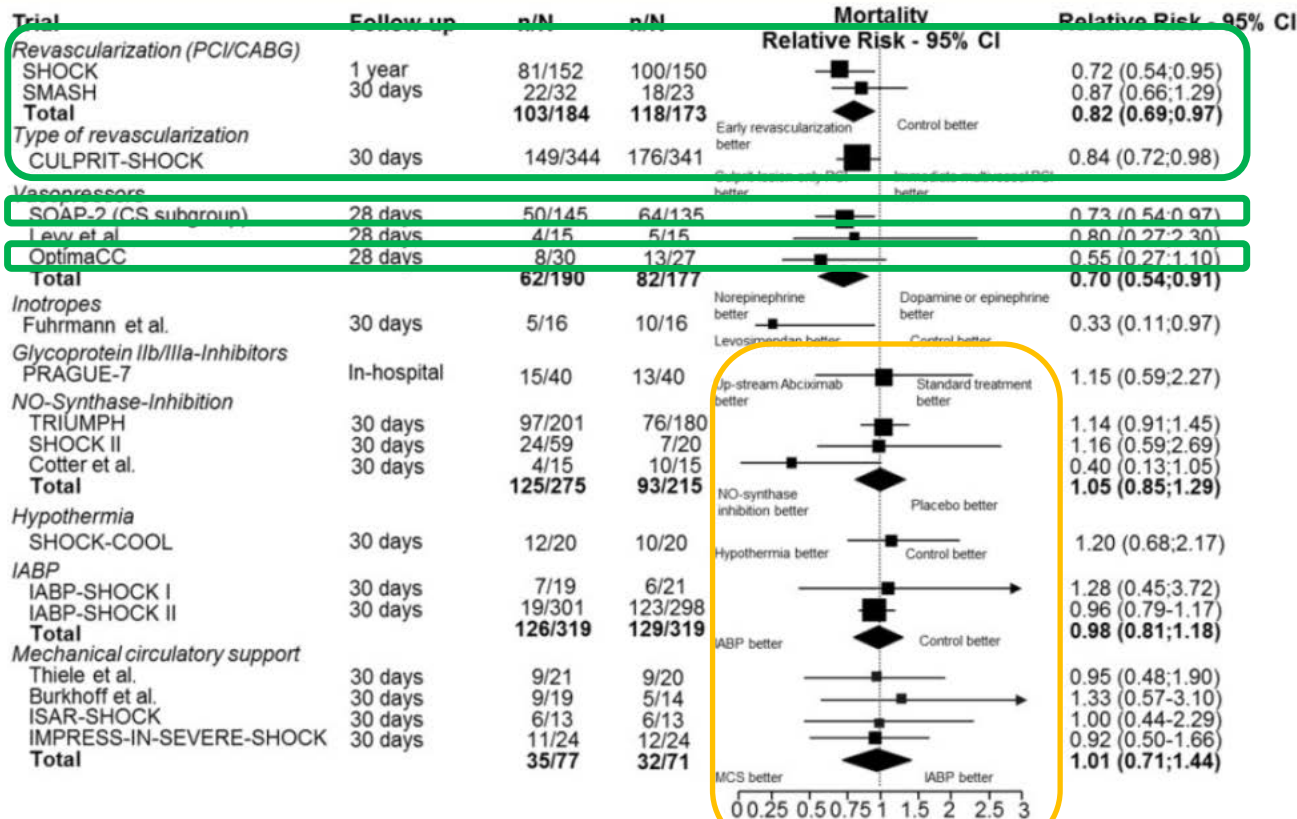
**Place of early diagnosis + stratification + treatment**

- High mortality during the first 48h
- Prognosis seems good after



*Aissaoui et al, Crit Care Med 2014*

# Poor current evidences in CS

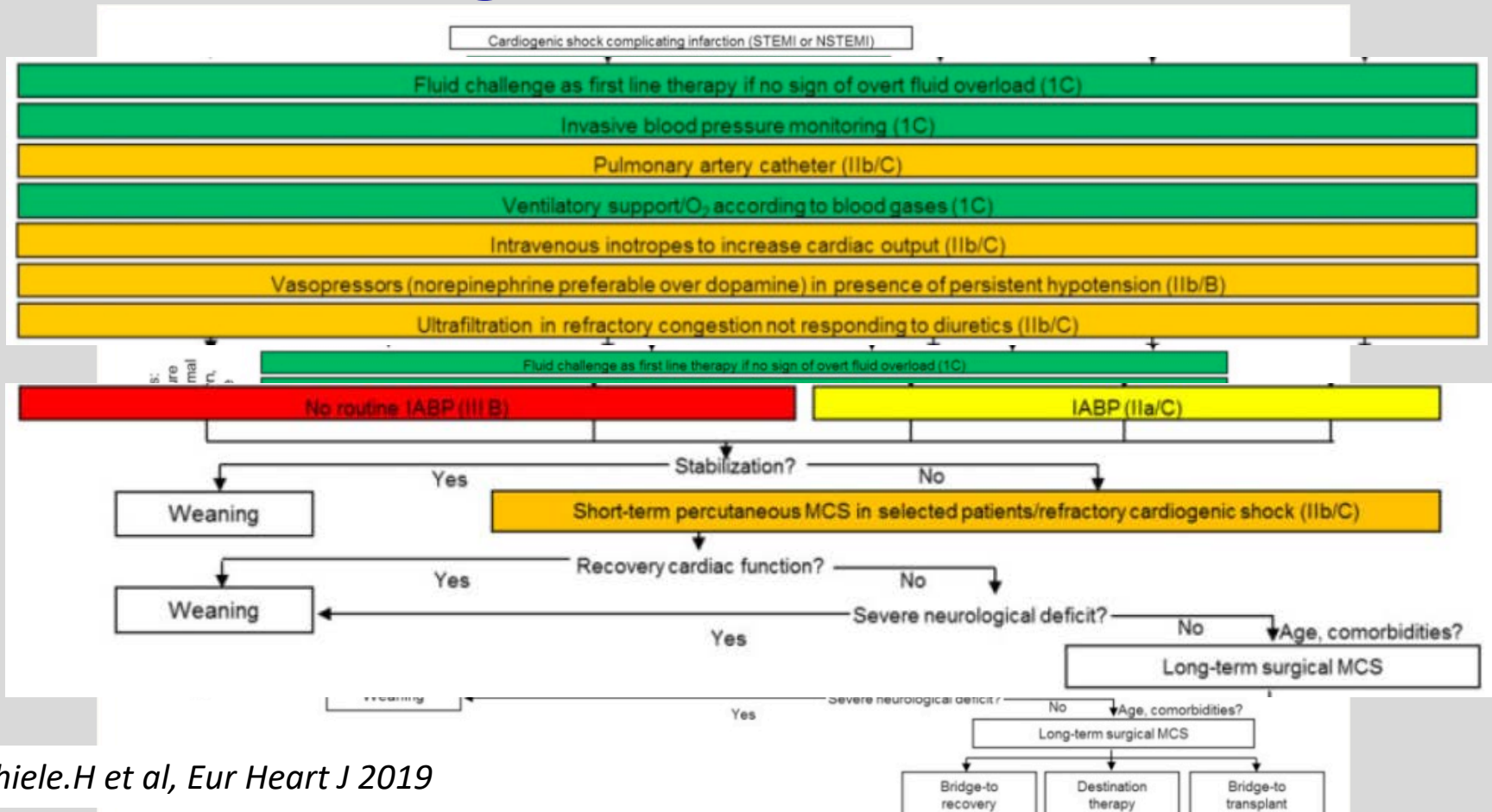


- Revascularization
- Culprit lesion only

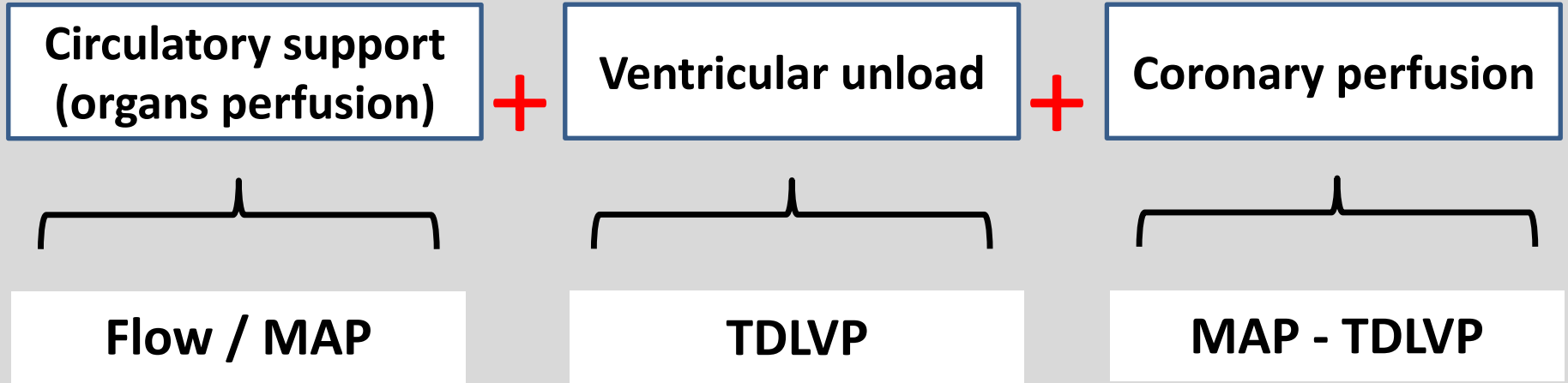
Norepinephrine

Only randomized studies were reported

# CS management: ESC Guidelines

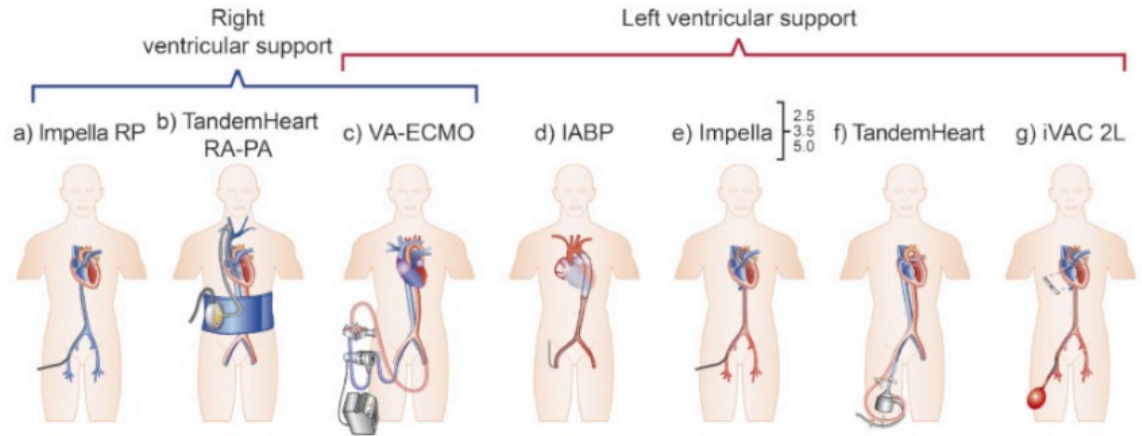


# Determinants of hemodynamic support



**Objectifs: Mettre le cœur au repos pour favoriser la récupération  
Et Eviter ou corriger la défaillance multiviscérale**

# What is a « light » circulatory support?



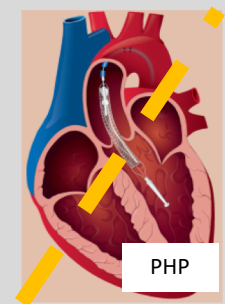
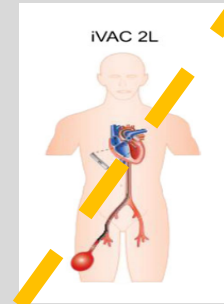
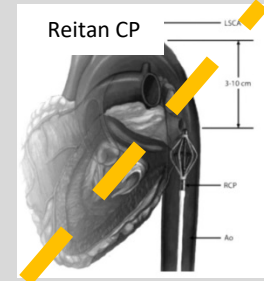
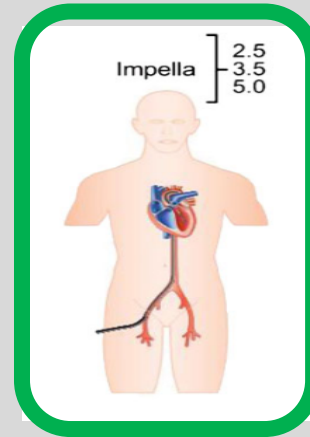
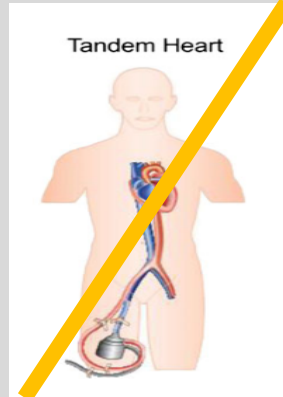
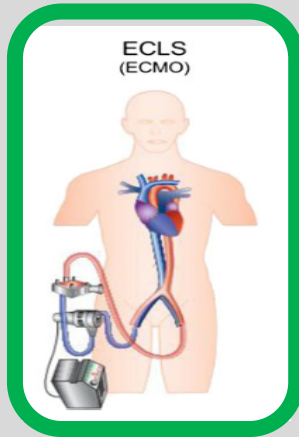
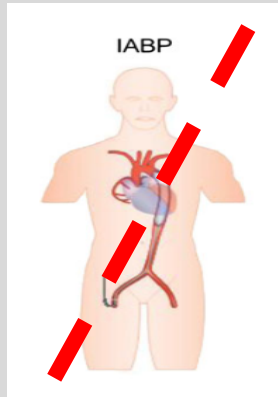
	a) Impella RP	b) TandemHeart RA-PA	c) VA-ECMO	d) IABP	e) Impella	f) TandemHeart	g) iVAC 2L
Flow:	max. 4.0 L	max. 4.0 L	max. 7.0 L		2.5-5.0 L	max. 4.0 L	max. 2.8 L
Pump speed:	33.000 rpm	max. 7.500 rpm	max. 5000 rpm		max. 51.000 rpm	max. 7.500 rpm	40 ml/beat
Cannula size:	22 F	29 F	14-19 F arterial 17-21F venous	7-8 F	12-14 F	12-19 F arterial 21F venous	17 F
Insertion/ Placement	Femoral vein	Internal jugular vein	Femoral artery Femoral vein	Femoral artery	Femoral artery	Femoral artery Femoral vein for LA access	Femoral artery
LV Unloading	-	-	-	(+)	++	++	+
RV Unloading	+	+	++	-	-	-	-

- Percutaneous ?
- W/o oxygenator?
- Flow ?
- Canula size?
- Implantation by interventional cardiologist?
- W/o complications?
- ...



# Short-term MCS

## LV support



1970

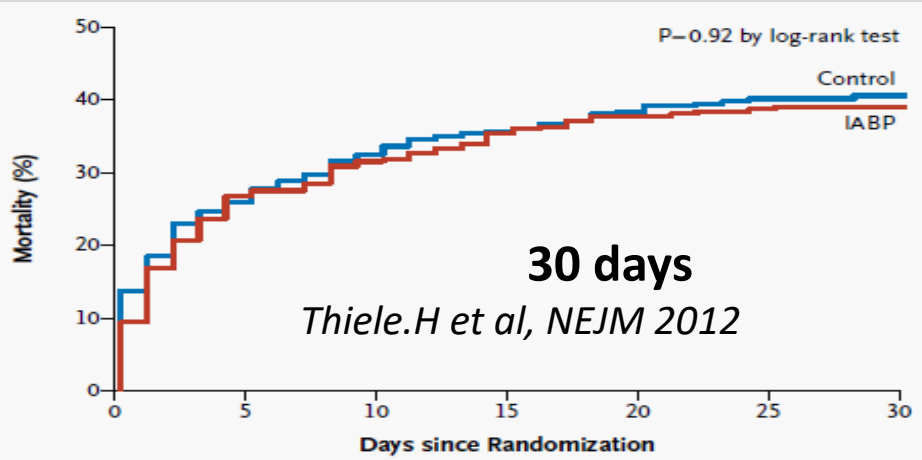
1990 2000

2004-2005

2015

Adapted from Thiele et al, EHJ 2015

# IABP place in ischemic CS?



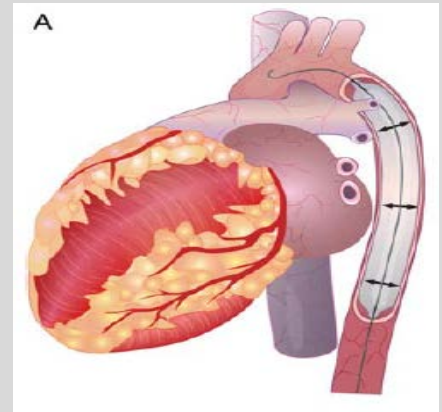
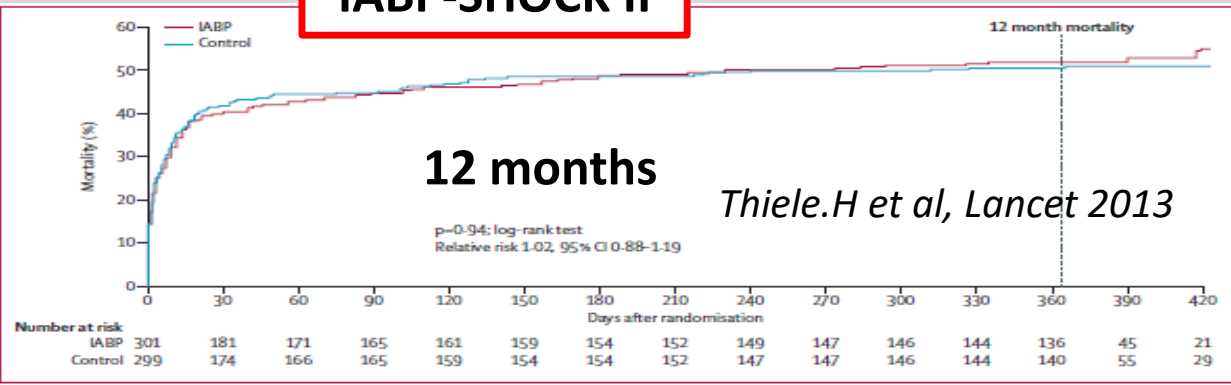
- No difference vs medical treatment
- No difference by subgroups analysis



Routine use of IABPs in patients with cardiogenic shock due to ACS is not recommended.<sup>260-262</sup>

**III** **B**

## IABP-SHOCK II



# IABP Shock 2 and after ?

French datas

CLINICAL RESEARCH

## Current indications for the intra-aortic balloon pump: The CP-GARO registry

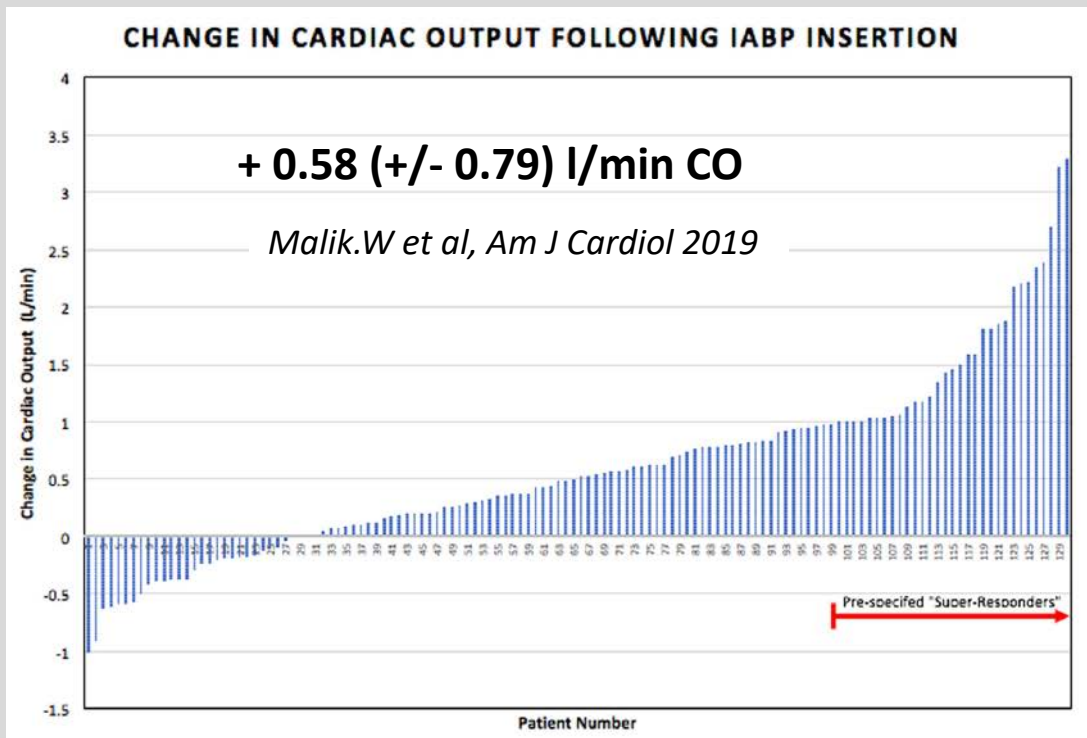
Helleu.B et al, Arch Cardiovasc Diseases 2018

172 patients in a multicentre (19 french centers) prospective registry in 2015

	Total (n = 172)	Haemodynamic indication (n = 107)	Bridge to revascularization indication (n = 34)	Coronary perfusion – related indication (n = 11)	Prophylactic indication (n = 18)
<i>Primary endpoints</i>					
In-hospital mortality	70/172 (40.7)	57/107 (53.3)	5/34 (14.7)	2/11 (18.2)	4/18 (22.2)
Cardiac mortality	51/61 (83.6)	41/50 (82.0)	4/5 (80.0)	Missing	4/4 (100)
<i>Secondary endpoints</i>					
1-year mortality	76/166 (45.8)	61/107 (57.0)	6/30 (20.0)	2/10 (20.0)	5/17 (29.4)
In-hospital stroke	6/172 (3.5)	5/107 (4.7)	0/34 (0.0)	0/11 (0.0)	0/18 (0.0)

62%

# Is there still a place for IABP in CS?: ADCHF ?



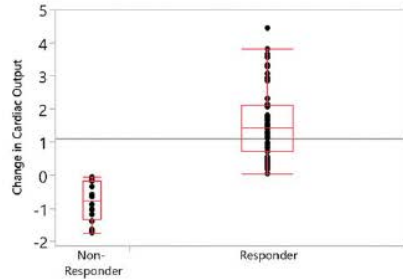
*Fried.JA et al, J Heart Lung Transplant 2018*

- **132 ADCHF patients**
- **84% 30-days survival / 78% bridge to LVAD or HTx w/o escalation to other AMCS**
- **Complication rate = 2.3%**
- **PAPi < 2 (OR 5.04 [1.86-13.03]) and ICM (OR 3.24 [1.16-9.06]) = predictive of clinical deterioration**

# Differential responses to larger volume intra-aortic balloon counterpulsation: Hemodynamic and clinical outcomes

Baran.DA et al, Catheter Cardiovasc Interv 2018

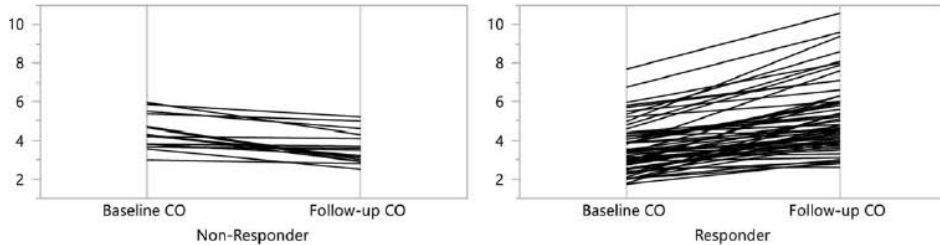
A Delta Cardiac Output: Responders vs. Non-Responders



+ 1.6 (+/- 1.1)  
l/min CO

- 76 patients
- 60% non ICM, LVEF 20%
- **18% IABP implantation at the bedside**
- 65% > +1l/min CO

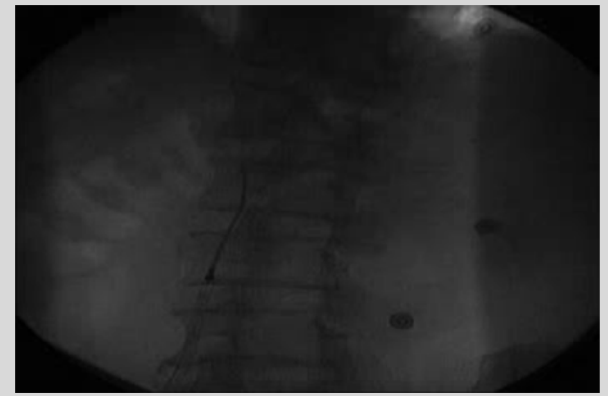
B



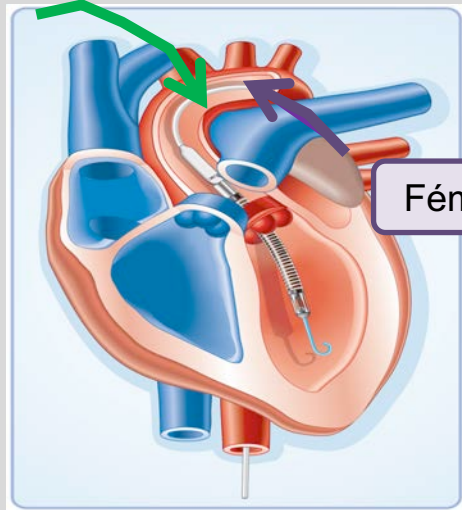
**IABP place in non ischemic CS ??**

- Low cost (400 euros)
- Large availability
- Vascular access 8Fr
- Easy to implant and to use

# Impella devices

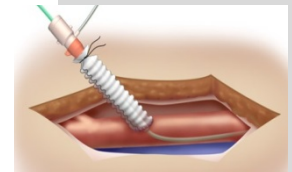


Axillaire

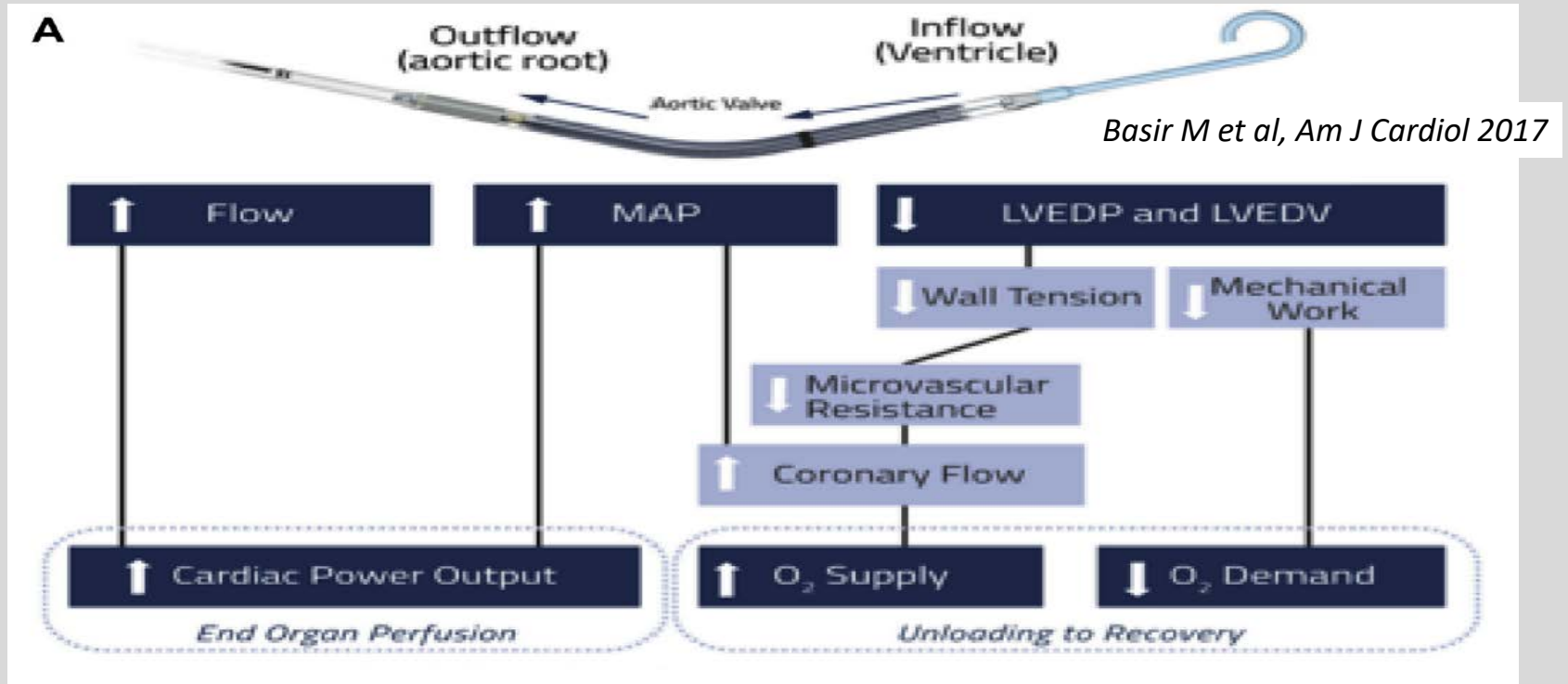


Fémoral

- Assistance mono-ventriculaire gauche
- Pompe axiale rotative (20-50000 trs/min)
- Placée à travers la Vao
- **2 types de pompes:**
  - **Abord percutané**
    - Impella 2.5 (12 Fr; 5j)
    - **Impella CP ou 3.5 (14 Fr; 5j)**
  - **Abord Chirurgical**
    - **Impella 5.0 (21 Fr; 10j) = tube en dacron**



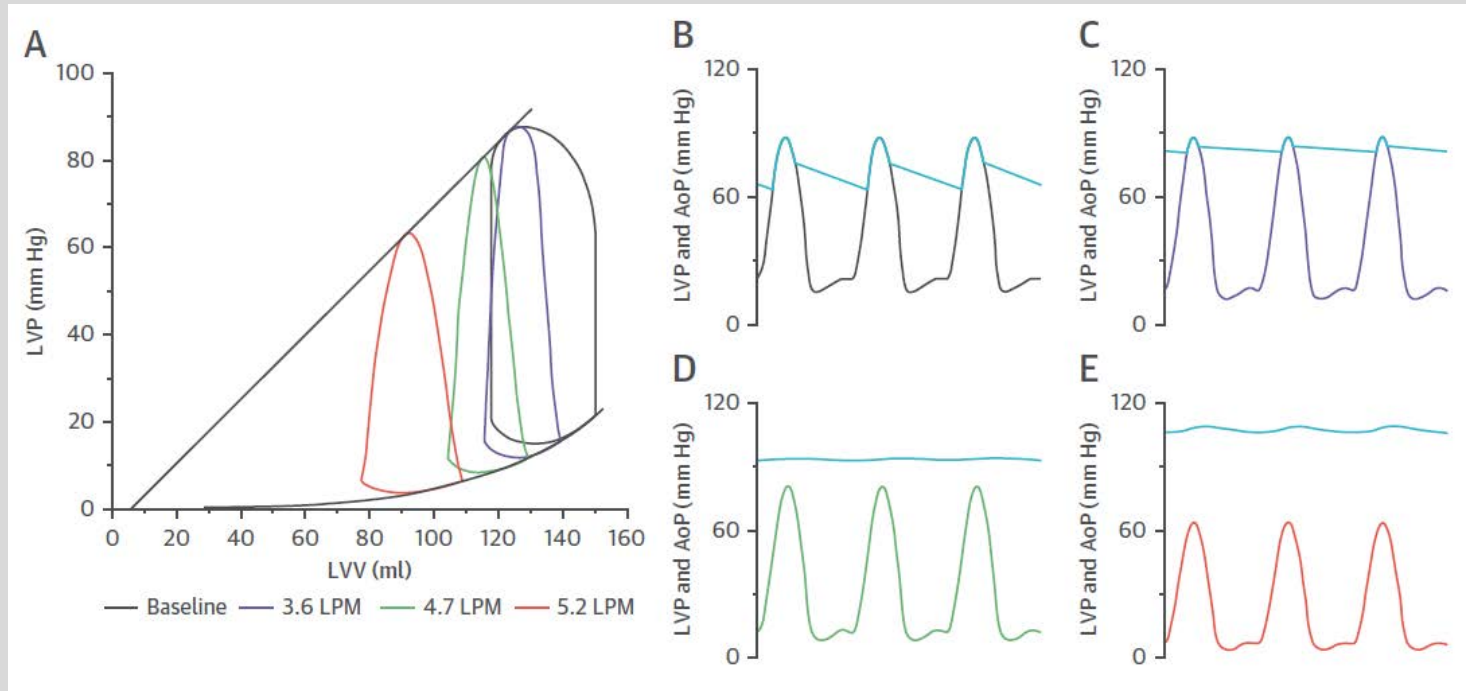
# Hemodynamic and clinical effects of Impella



Sauren LD, Artif organs 2007; Meyns B JACC 2003; Rimmelink M Catheter Cardiovasc Interv 2007; Agel RA, J Nucl Cardiol 2009; Lam K Clin Res Cardiol 2009



# Impella = acute LV assist device



*Uriel.N, JACC 2018 / Burkhoff.D et al, JACC 2015*

- **LV unloading** // No oxygenation and decarboxylation
- **Gradual LV- Ao pressure gradient decoupling**= Impella support



# Associated complications

- Usual complications 4-35%
- More than IABP

*Cheng.M et al, EHJ 2009*

Expected complications

Expected benefits

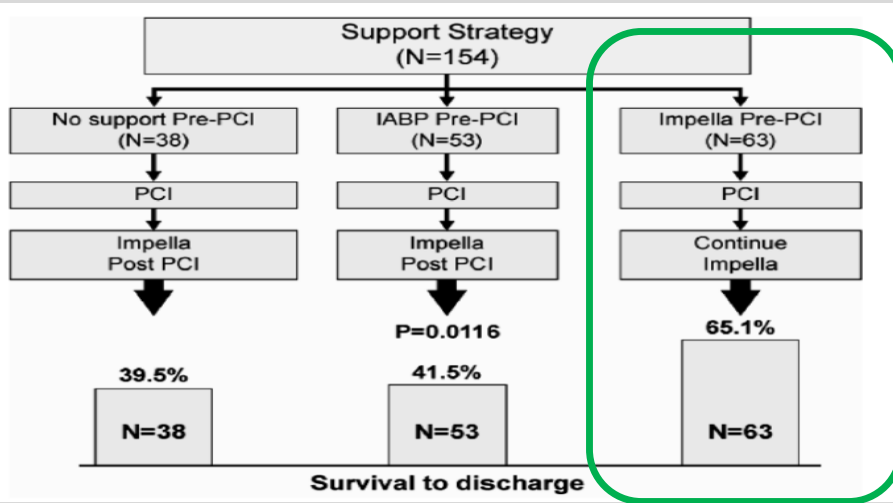


*Jensen PB et al, Eur Heart J ACC 2018*

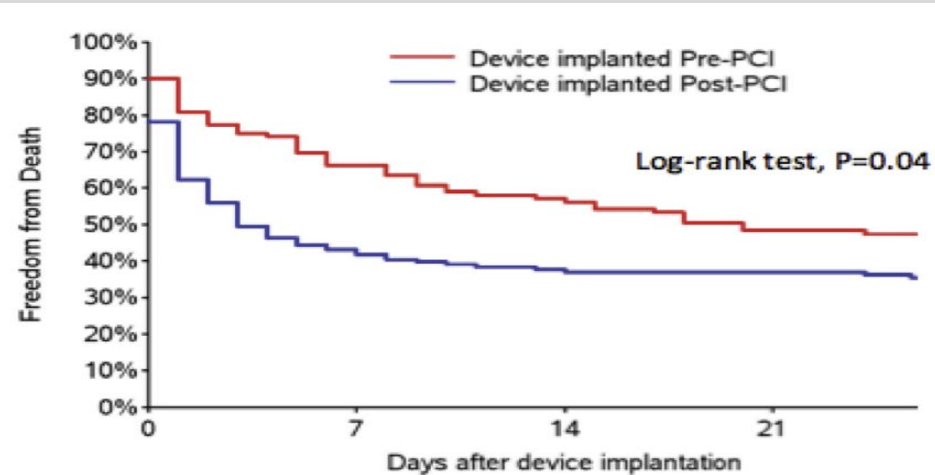
	MI and shock (n=79)	AHF and shock (n=16)
Impella CP	73 (92%)	15 (94%)
Impella 5.0	7 (9%)	1 (6%)
Impella RP	2 (3%)	2 (13%)
Pre-admission cardiac arrest (%)	29 (37%)	6 (38%)
BMI, kg/m <sup>2</sup>	29±5	27±6
Lactate at placement, mmol/L	7.6±6.0	8.6±4.9
pH at placement	7.19±0.18	7.14±0.18
Systolic BP at placement, mmHg	78±17	75±13
One or more suction events (%)	27 (34%)	6 (38%)
Echo guided reposition (%)	46 (58%)	10 (63%)
≥3 Repositions	18 (23%)	4 (25%)
Haemolysis (%)	8 (10%)	3 (19%)
Limb ischaemia		
Limb ischaemia without intervention (%)	3 (4%)	1 (6%)
Limb ischaemia with intervention (%)	4 (5%)	0 (0%)
Limb ischaemia with amputation (%)	1 (1%)	0 (0%)
Bleeding*		
Minor (%)	23 (29%)	6 (38%)
Moderate (%)	15 (19%)	2 (13%)
Severe (%)	4 (5%)	2 (13%)
Significant groin bleeding (%)	10 (13%)	2 (13%)
Confirmed heparin-induced thrombocytopenia (%)	0 (0%)	0 (0%)
Significant aortic regurgitation (%)	2 (3%)	1 (6%)
Stroke during support (%)	0 (0%)	0 (0%)
Impella pump stop (%)	2 (3%)	1 (6%)

➔ What timing for implantation and what type of support ?

# Timing of implantation ?



Before PCI in case of ischemic CS



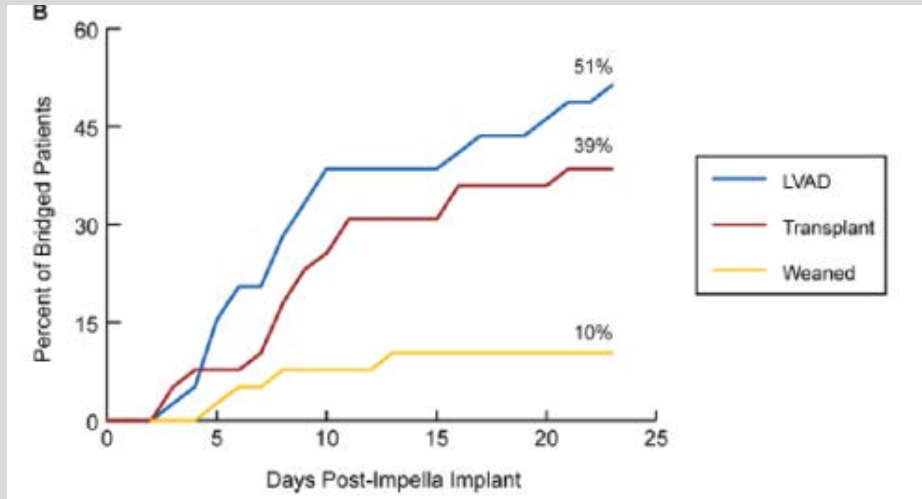
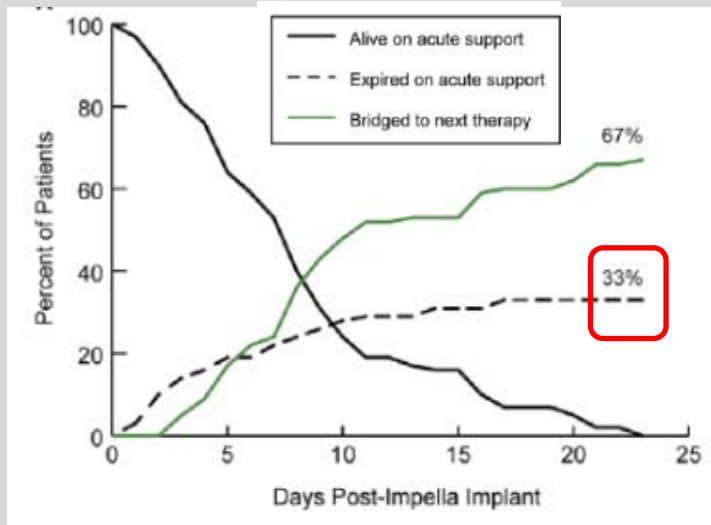
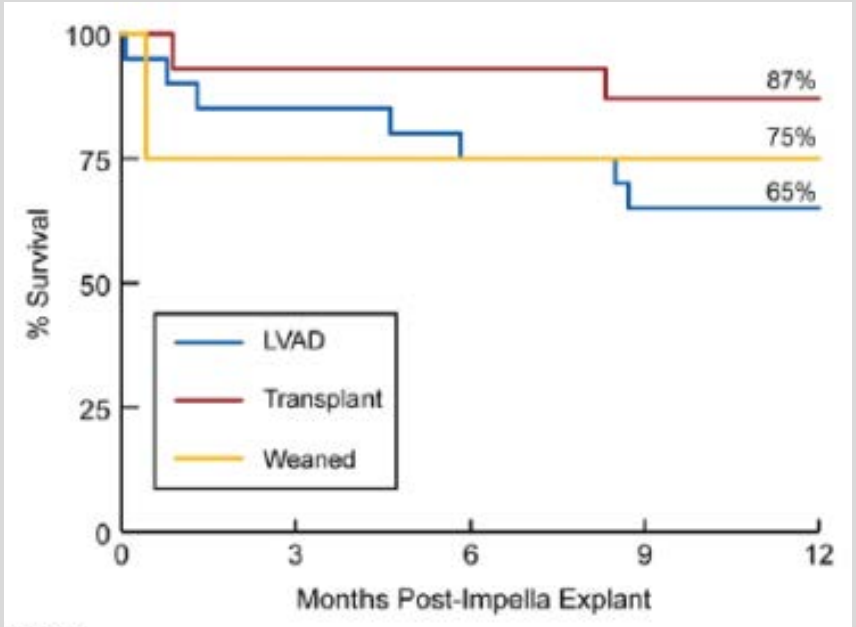
O'Neill WW, J Interv Cardiol 2014

- cVAD registry = 287 patients with ischemic CS
- **Laying time = 17 min**

Basir M et al, Am J Cardiol 2017

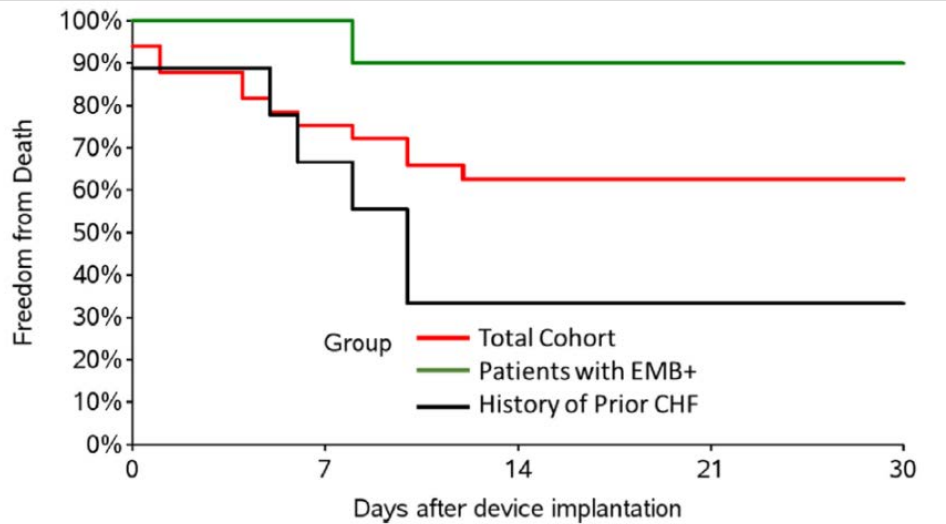
# Use of a percutaneous temporary circulatory support device as a bridge to decision during acute decompensation of advanced heart failure

Hall.SA et al, J Heart Lung Transplant 2018



# The Impella Microaxial Flow Catheter Is Safe and Effective for Treatment of Myocarditis Complicated by Cardiogenic Shock: An Analysis From the Global cVAD Registry

*Annamalai.SK et al, J Card Fail 2018*

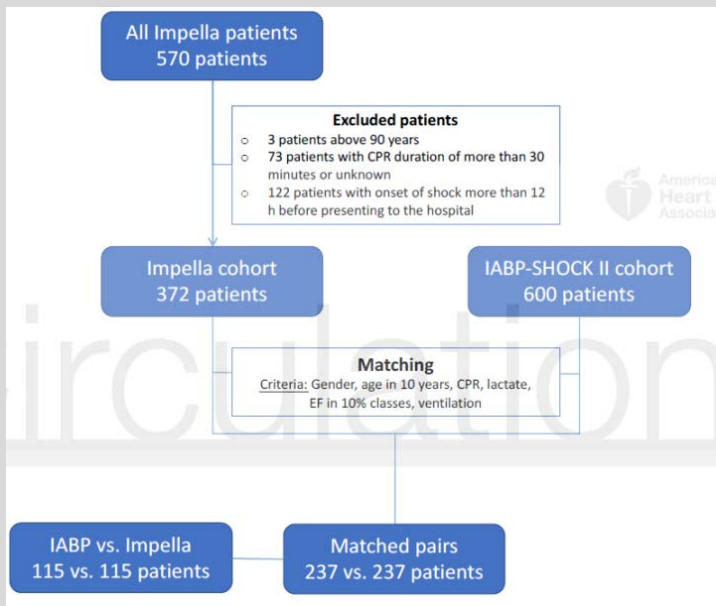


- 34 patients /cVAD registry / 88 sites US and Europe
- Mean age 42 yo
- 26% with previous congestive HF
- Mean LVEF 18 +/- 10%
- 85% inotropes and vasopressors, 22% IABP
- Impella 2.5 (41%), CP (35%) and 5.0 (24%)

In-hospital survival 62%  
LVEF at discharge 37%

Stroke 5.88%, 21% Transfusion, Limb ischemia 9%, Hemolysis 12%, RRT 33%

# But

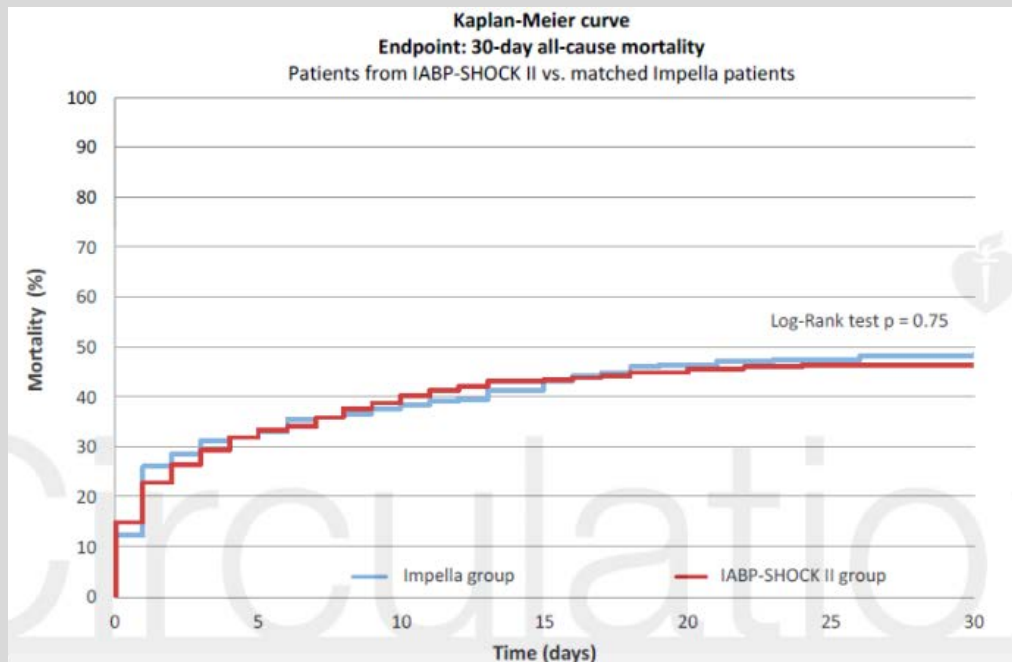


- 237 patients EUROSHOCK registry matched with 237 patients from IABP Shock 2 study

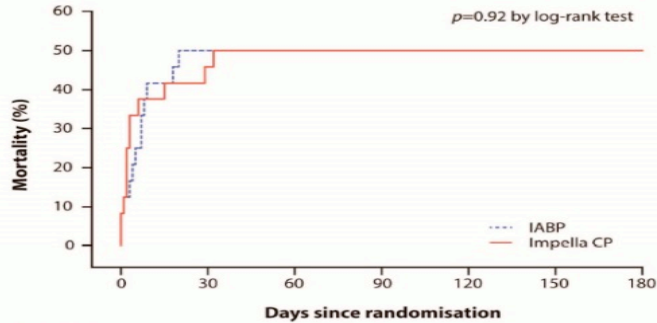
## Impella Support for Acute Myocardial Infarction Complicated by Cardiogenic Shock: A Matched-Pair IABP-SHOCK II Trial 30-Day

### Mortality Analysis

Schrage.B et al, *Circulation* 2018



# IMPRESS study: gaps and limits



Number at risk		0	30	60	90	120	150	180
IABP	24	12	11	11	11	11	11	11
Impella CP	24	13	12	12	12	12	12	12

## Very severe patients:

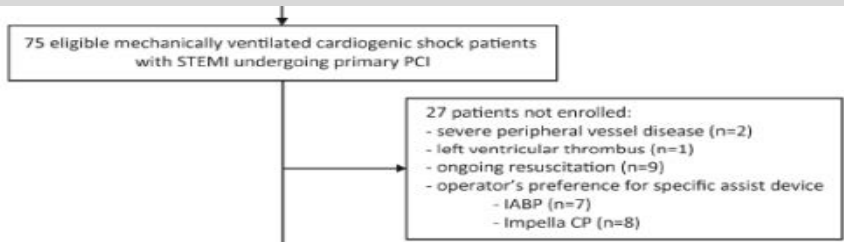
- 100 vs 83% prior CA
- 100% ETI and MV
- 100% vs 92% under inotrops
- Traumatic lesion = 21% vs 8%
- Lactates 7.5 vs 8.9mmol/l
- pH 7.14 vs 7.17

## Non optimal management

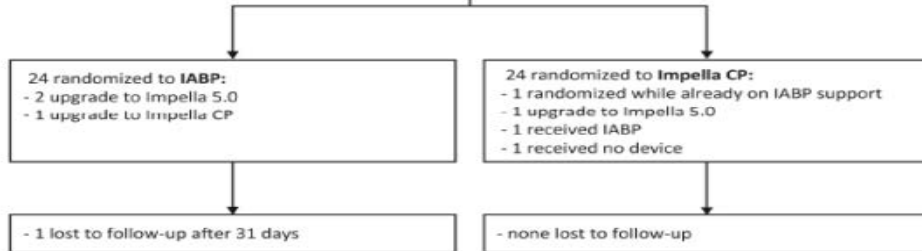
- 21 vs 13% before PCI
- Time under support 49 vs 48h

## Crossover:

- 4.2 vs 12.5%



*Ouweneel.DM et al, JACC 2016*



**Conclusion: For post CA CS during ACS, Impella CP is not better than IABP**



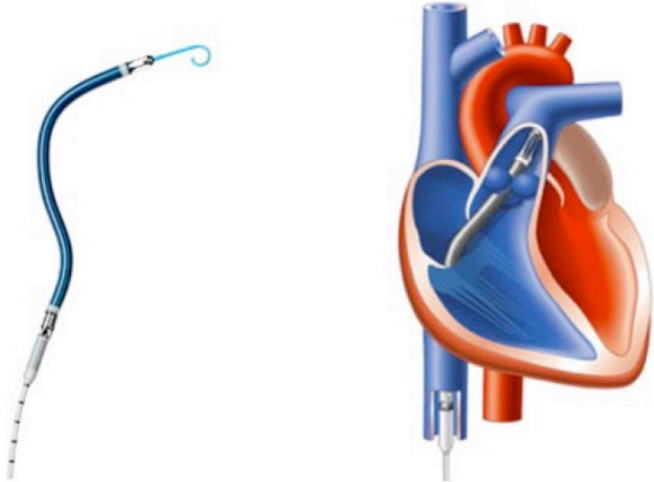
# Lack of randomized datas!

Prospective studies to evaluate the efficacy of Impella in ischemic cardiogenic shock

Table 2 Trials performed using the Impella<sup>®a</sup> device; most were stopped because of poor inclusion rate.

Study	Registry number	Condition	Patients required (n)	Patients enrolled (n)	Duration (months)	Status	Reason for discontinuation
FRENCH TRIAL (2006)	NCT00314847	AMI CS	200	19	52	Discontinued	Low enrolment
ISAR-SHOCK (2006)	NCT00417378	AMI CS	26	26	19	Completed	N/A
IMPRESS in STEMI trial (2007)	NTR1079 <sup>b</sup>	STEMI pre-CS	130	21	22	Discontinued	Low enrolment
RECOVER I FDA (2008)	NCT00596726	PCCS	Up to 20	17	28	Completed	N/A
RECOVER II FDA (2009)	NCT00972270	AMI CS	384	1	18	Discontinued	Low enrolment
RELIEF I (2010)	NCT01185691	ADHF	20	1	33	Discontinued	Low enrolment
DANSHOCK (2012)	NCT01633502	AMI CS	360	~50	40	Enrolling	N/A

# What about RV support?



*Cheung AW et al J Heart Lung Transplant 2014*

## Impella RP

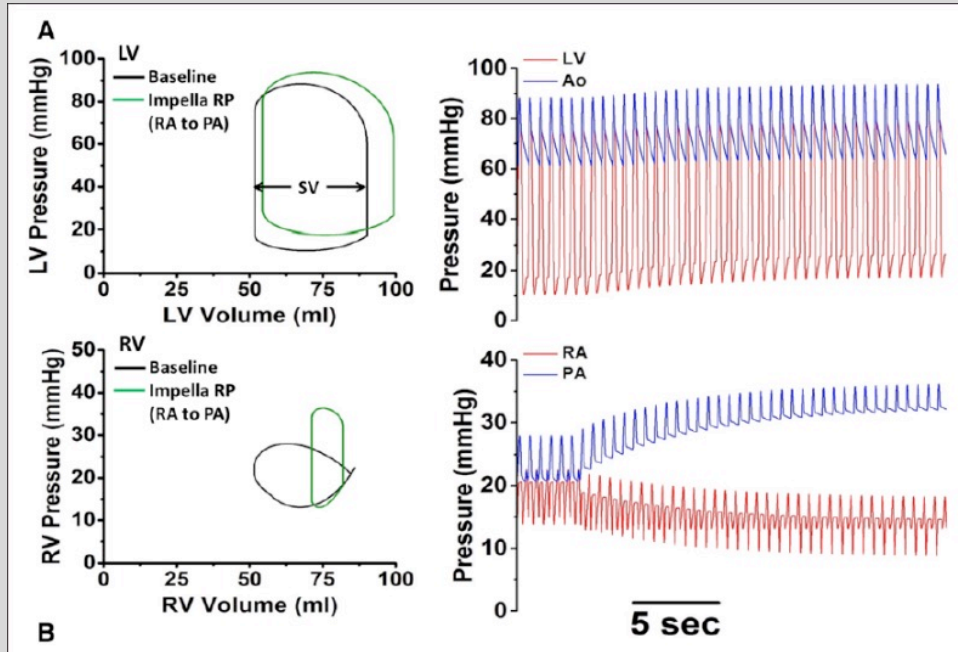


- **Percutaneous 22F microaxial pump munted on a 11 F catheter**
- Aspirate on the IVC and expels it into the PA
- **33000 rpm = 4.0L/min**
- ACT 160-180
- **Up to 14 days**

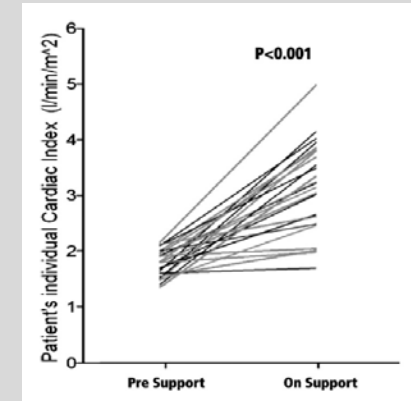
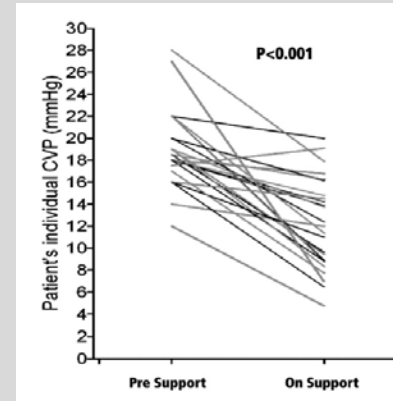
*Anderson et al, J Heart Lung Transplant 2015*



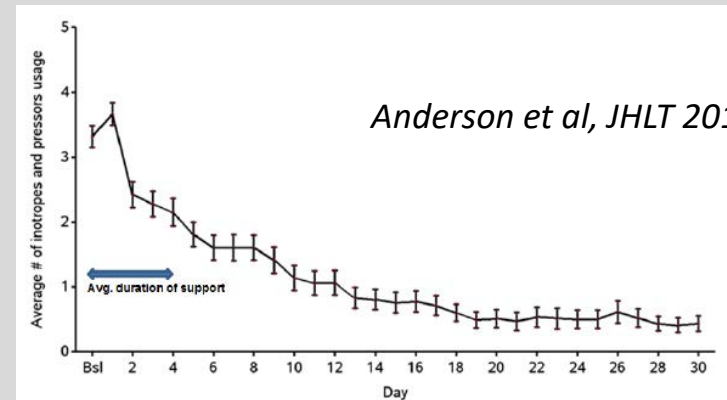
# Impella RP : Hemodynamic effects



*Kapur NK et al, Circulation 2017*

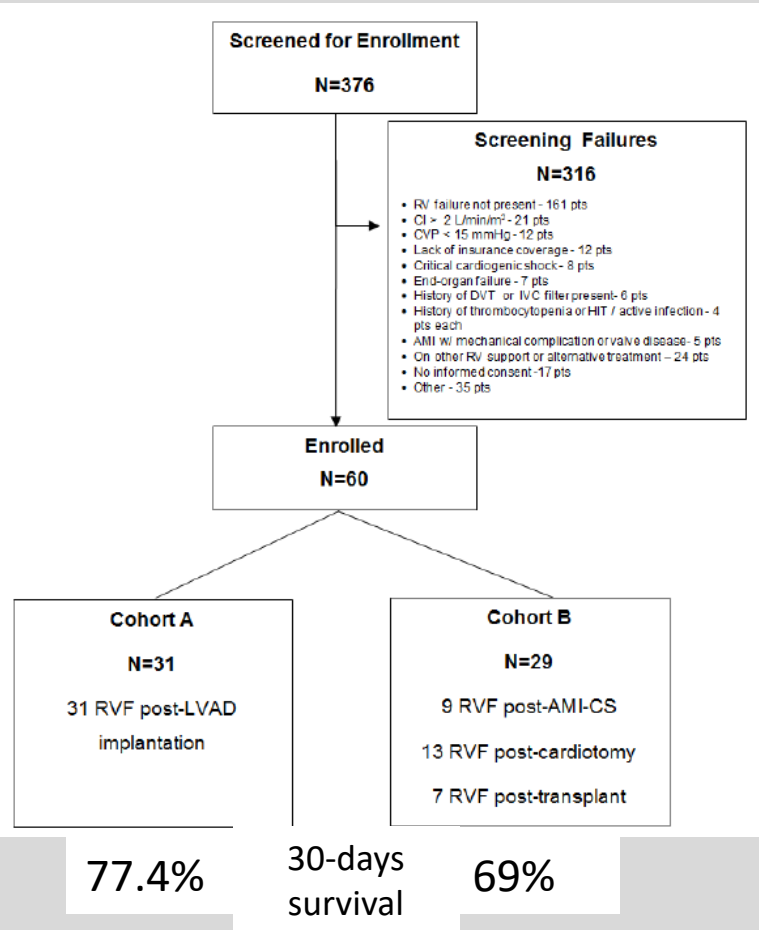


*Anderson et al, J Heart Lung Transplant 2015*



*Anderson et al, JHLT 2018*

# Impella RP: Clinical results



Safety Endpoints	All Patients (N=60 patients)	Cohort A (N=31patients)	Cohort B (N=29 patients)	P-value
Death	26.7% (16/60)	22.6% (7/31)	31.0% (9/29)	0.563
Major Bleeding	48.3% (29/60)	54.8% (17/31)	41.4% (12/29)	0.316
Device access site bleeding	1.7% (1/60)	-	3.4% (1/29)	0.297
Postoperative bleeding	43.3% (26/60)	54.8% (17/31)	31.4% (9/29)	0.063
Transfusion with no overt bleeding	1.7% (1/60)	=	3.4% (1/29)	0.329
Other	1.7% (1/60)	-	3.4% (1/29)	0.329
Hemolysis	21.7% (13/60)	25.8% (8/31)	17.2% (5/29)	0.421
Pulmonary Embolism	0.0% (0/60)	0.0% (0/31)	0.0% (0/29)	--
Tricuspid and Pulmonary Valve Dysfunction*	2.9% (1/34)	2.9% (1/34)	2.9% (1/34)	2.9% (1/34)

*\*based on echocardiographic core lab analysis*

## Increased Rate of Mortality in Patients Receiving Abiomed Impella RP System - Letter to Health Care Providers

May 21, 2019 UPDATE: The FDA issued an updated Letter to Health Care Providers to provide the most recent, interim post-approval study results for Abiomed's Impella RP System.

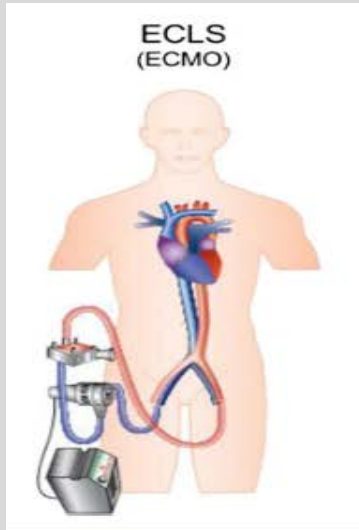
*Khalid.N et al, Cardiovasc Revasc Med 2019*

*Anderson et al, J Heart Lung Transplant 2018*

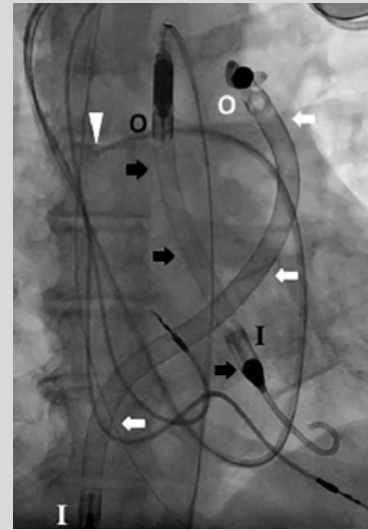
# Short –term MCS

Biventricular support

+/- respiratory support



VA ECMO



BiPELLA

# Percutaneous versus surgical femoro-femoral veno-arterial ECMO: a propensity score matched study

Daniel.P et al, Int Care Med 2018

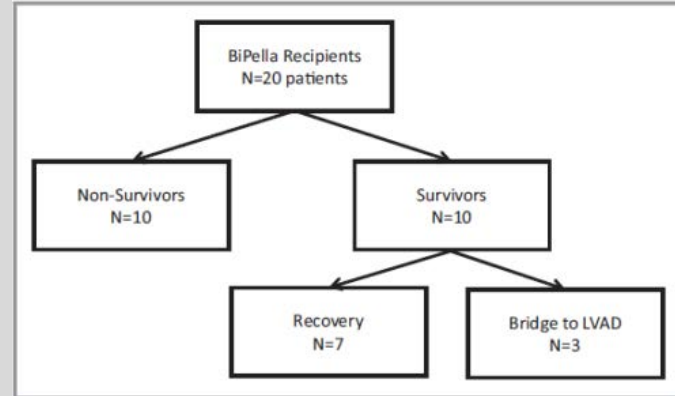
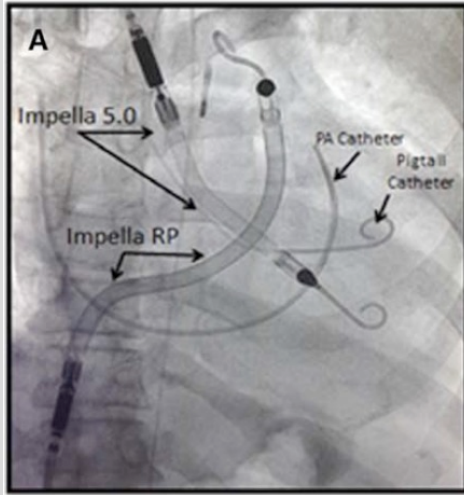
	Surgical group n = 266 (%)	Percutaneous group n = 266 (%)	p value
30-Day overall survival	150 (56.3)	170 (63.8)	0.034
Cannulation site infection	74 (27.8)	44 (16.5)	0.001
Infection requiring surgical revision <sup>a</sup>	40 (15.0)	14 (5.3)	< 0.001
Vascular complications at cannulation <sup>b</sup>	7 (2.6)	10 (3.8)	0.663
Limb ischemia	33 (12.4)	23 (8.6)	0.347
Cannula relocation or removal	25 (9.4)	15 (5.6)	0.258
Limb fasciotomy	10 (3.8)	6 (2.3)	0.310
Amputation	2 (0.8)	2 (0.8)	1.000
Vascular complications after cannula removal	9 (3.4)	39 (14.7)	< 0.001
Surgical revision for persistent bleeding early after decannulation	4 (1.5)	25 (9.4)	< 0.001
Surgical revision in the days after decannulation <sup>c</sup>	5 (1.9)	14 (5.3)	0.035
Lower limb sensory-motor deficit	6 (2.3)	7 (2.6)	0.779

## Is it time to implant in cathlab ?

De Waha S et al, Eurointervention 2016

- Larger availability
- ↘ « time to support » ?
- Need specialized ICCU +++

# Double Impella: BiPELLA???



- 14 Fr arterial + 11 Fr venous catheter
- **BiPella**=
  - No respiratory support
  - LV and RV unloading / possible evaluation of RV function
  - Possible bridge to LVAD with initial RV support

*Kapur et al, Circ Heart Fail 2015*

20 patients / **50% mortality**

Major complications: acute limb ischemia (5%); hemolysis (30%); TIMI major bleeding (35%)

**Right flow 3.2 L/min and LV flow 3.5 L/min**

↗ CI and ↘ RAP, sPAP, PCWP

Kuchibhotla S et al, J Am Heart Assoc 2017



# What recommendations for (cardio)-circulatory support in CS ?

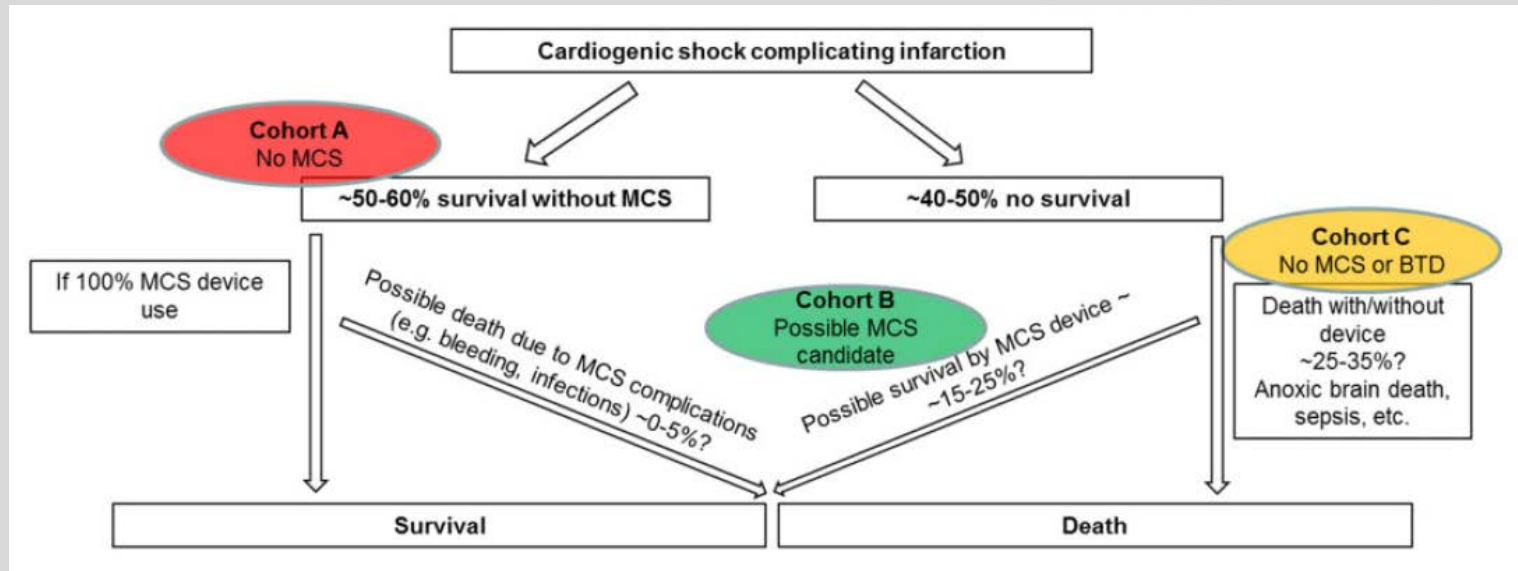
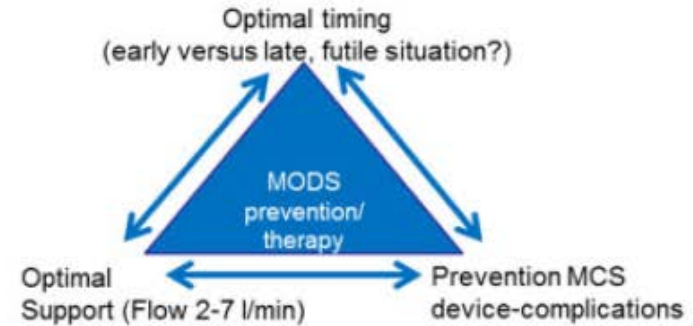


Assistance	ESC/EACTS guidelines (2012 et 2016)	ACCF/AHA/SCAI guidelines (2013)	Experts SRLF (2015)	HFA ESC/SEM /SAEM consensus (2015)
<b>CPLA</b>	Classe IIIa (level of evidence B)	Classe IIa C	Ne doit pas être utilisée (accord faible)	Non recommandé
<b>Impella</b>	Classe IIb C ou IIa C	Classe IIaB	Impella 5.0 si expertise (accord faible)	Peut être utilisée
<b>ECLS</b>		Néant	ECLS est préférée (accord fort)	Peut être utilisée

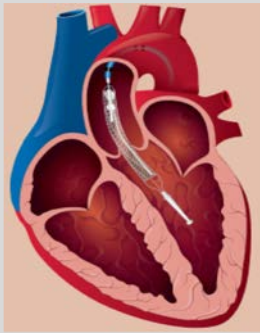
**Des niveaux de recommandations de faible niveau mais qui se précisent ...**

# Considerations on use in MCS in CS patients

**MCS = Only Flow !!**

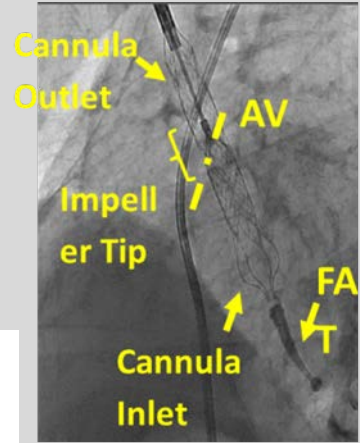






# PHP = percutaneous heart pump (Thoratec®)

- 14 Fr sheath (femoral artery > 5 mm)
- Expansible to 24Fr
- 16000 – 20500 rpm / > 4.0L/min support



	Design	Condition	n	Trial n <sup>o</sup> *	Status
HeartMate PHP CE Mark Clinical Investigation Plan	Prospective registry		50	NCT02156609	Complete
Coronary InterventionS in High-Risk PatiEnts Using a Novel Percutaneous Left Ventricular Support Device (SHIELDS-II)	Randomized trial 2:1 PHP vs. Impella LP 2.5	High risk PCI	425	NCT02468778	<b>Stop</b>
Thoratec Corporation HeartMate PHP cardiogenic Shock Trial	Prospective registry	Cardiogenic shock	25	NCT02279979	<b>5 patients</b>

## SHIELD 1 trial

- ↗ CO = 1.2-2.2L/min
- ↗ MAP and CPO
- **No change in PCWP**

Dudek.D et al, Am J Cardiol 2018

## BUT

Trial **enrollment was stopped in 01.2017** due to mechanical issues ...

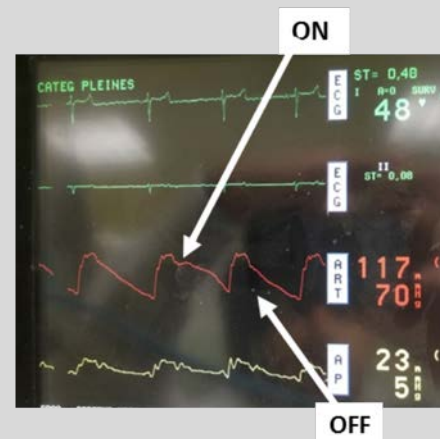
Van Mieghem et al, Eurointervention 2016

Maly J et al, JHLT 2017



# iVAC 2.0L (Terumo®)

- 13.5 / 17 Fr
- Same consol as IABP
- Same principle as IABP but with **real unloading of LV**
- Synchronization with AP or EKG
- Up to **2L/min** for 24h (pulsatile)

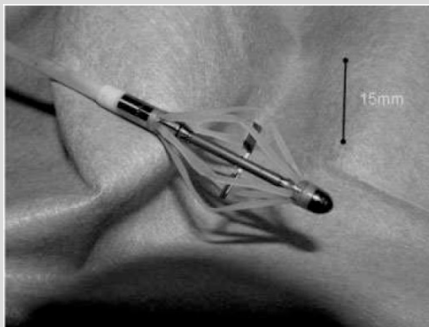


*Van Mieghem et al, Eurointervention 2015*

	Baseline	During support	p-value
MAP	66 [52-82]	83 [71-102]***	<0.001
HR	71 [51-100]	72 [56-98]	ns
MPAP	22 [12-44]	17 [10-39]*	<0.05
Wedge	12 [3-25]	9 [5-21]	ns
CO	3.7 [2.3-6.9]	5.0 [3.1-8.1]***	<0.001
CI	1.8 [1.3-3.6]	2.5 [1.7-4.3]***	<0.001
CPI	0.29 [0.16-0.51]	0.48 [0.27-0.87]***	<0.001
iVAC output, L/min	-	1.4 [1.2-2.0]	-
SvO <sub>2</sub>	64 [33-72]	67 [40-76]**	<0.05

- 14 high risk PCI patients
- 1 iliac arterial dissection during sheath insertion
- Total support time: 67 (23-147) min

*Den Uil CA et al, Eurointervention 2017*

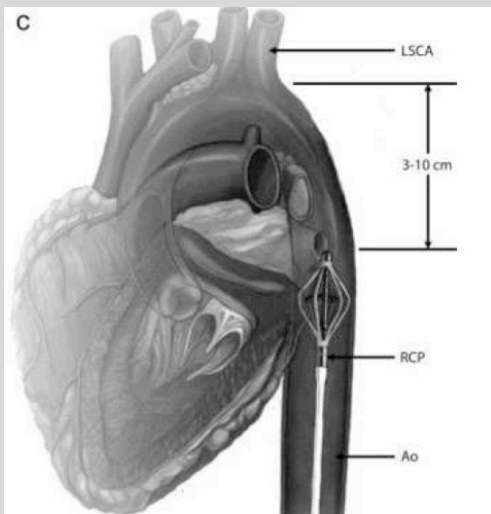


Smith EJ et al, Cath Card Int 2009

# Reitan catheter pump

- **14 fr** femoral sheath/ descending aorta > 22mm
- **Continuous non phasic** pump: 8000 – 13000 rpm/min  
=>  $\searrow$  LV afterload and  $\nearrow$  organ perfusion (lower part of the body+++)
- **Not contraindicated in case of LV thrombus or aortic regurgitation**

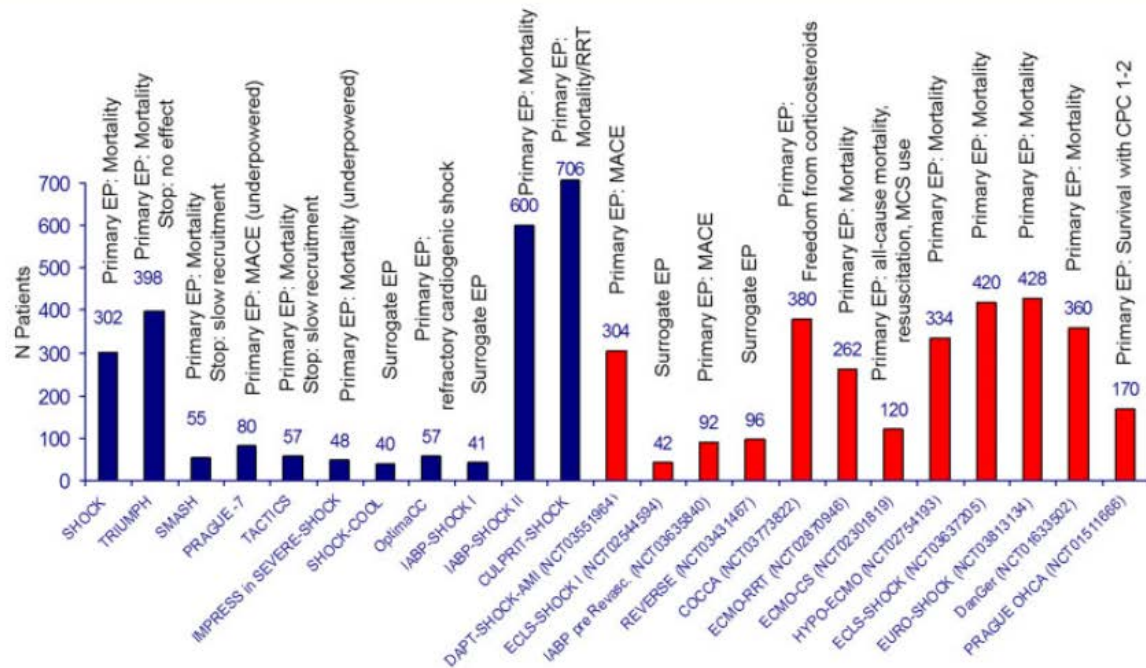
Reitan O et al, ASAIO 1999 / Reitan O et al, ASAIO 2003



- 20 ADCHF patients in 4 european centers
- LVEF 20%; CI 1.79L/min/m<sup>2</sup>; eGFR 37.8ml/min; inotrops 45%
- Support time 18.3h
- $\nearrow$  CI (**0.57 l/min/m<sup>2</sup> / + 31%**),  $\searrow$  CVP (- 6mmHg)
- Trends  $\searrow$  PCWP et PAPm
- 20% 30-days mortality; **10% vascular complications**

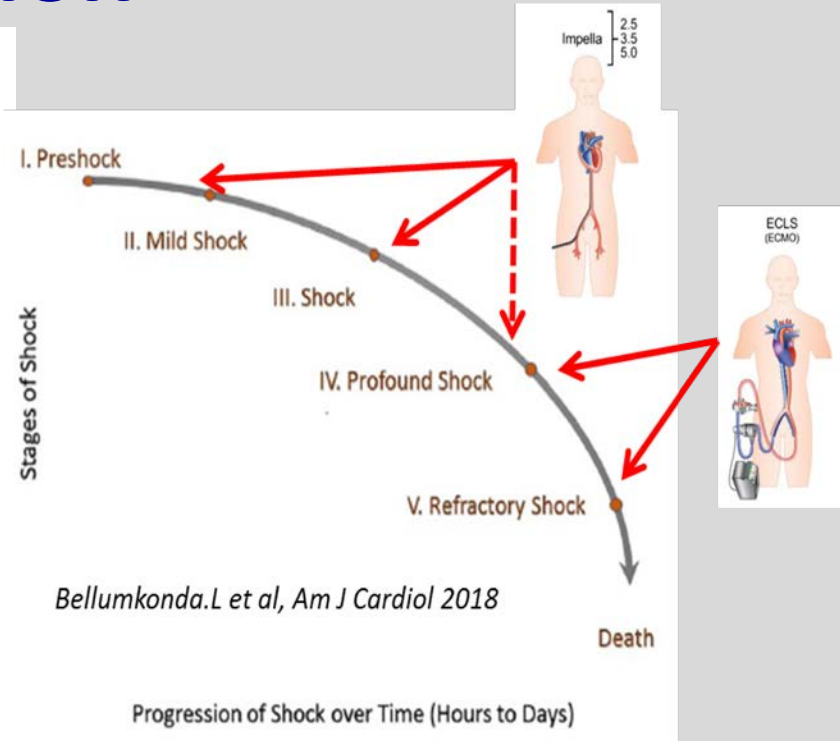
Keeble.TR et al, Int J Cardiol 2019

# Perspectives in CS trials



# Conclusion

- **CS is a continuum**
- Several possible supports = No solid data !!
  - Left: **Impella CP**, VA-ECMO vs Impella 5.0, VA-ECMO
  - Right: Impella RP, **VA-ECMO**, RA-PA ECMO
  - Biventricular: **VA-ECMO**, BiPella (??)
- **Type and Timing of implantation**
  - **Initial severity of CS** (MOF, RV function, respiratory status)
  - **Available device**
  - **Local experiences and capacities**
  - **Ischemic CS: pre-PCI?**
- Non exclusive and evolutive
  - **Monitoring** (TTE, PAC?)
  - Association / upgrading



Randomized data during CS are needed ...

Lyon, AER 22.11.2019

# Assistances circulatoires « légères »

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