

# **Extracorporeal Circulation and Myocardial Protection**

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# **Extracorporeal Circulation**

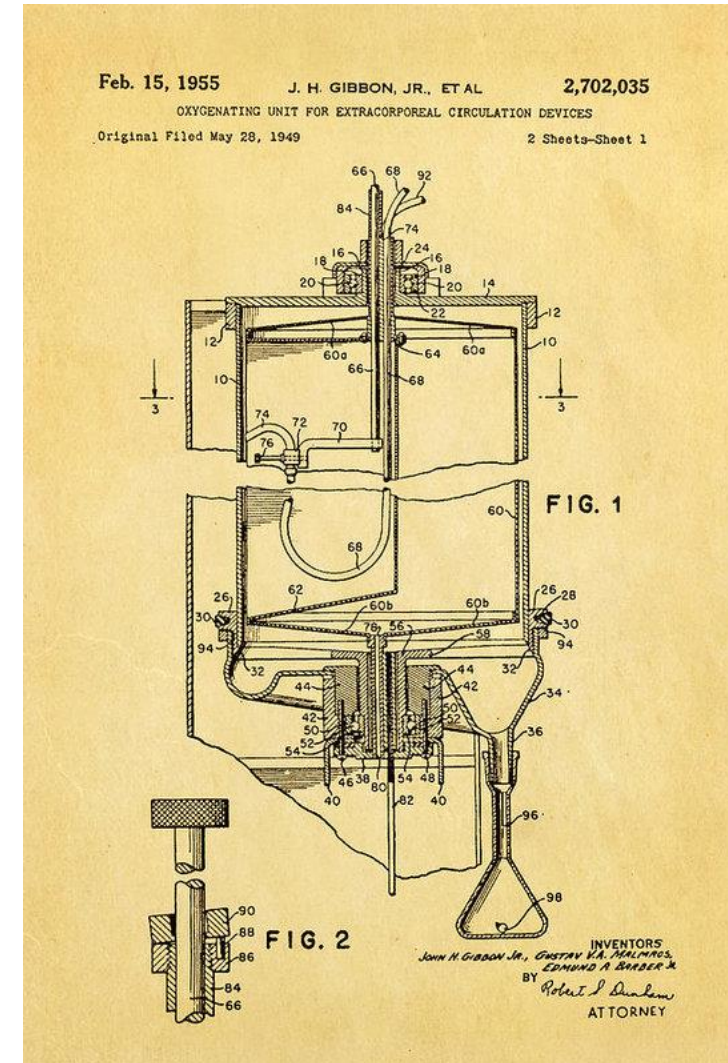


# 1<sup>st</sup> Cardiopulmonary Bypass

- **John Gibbon** - IBM engineer

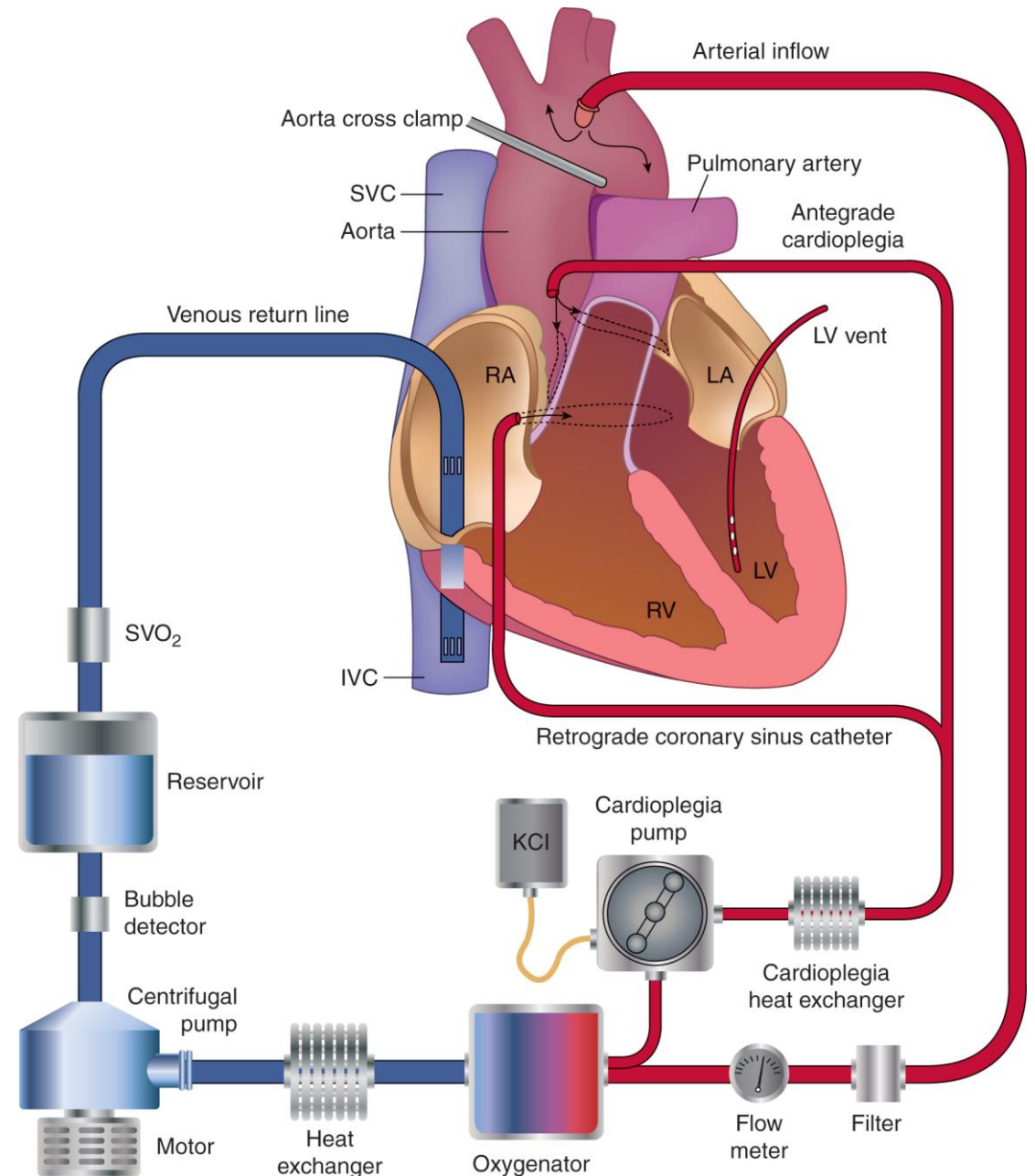


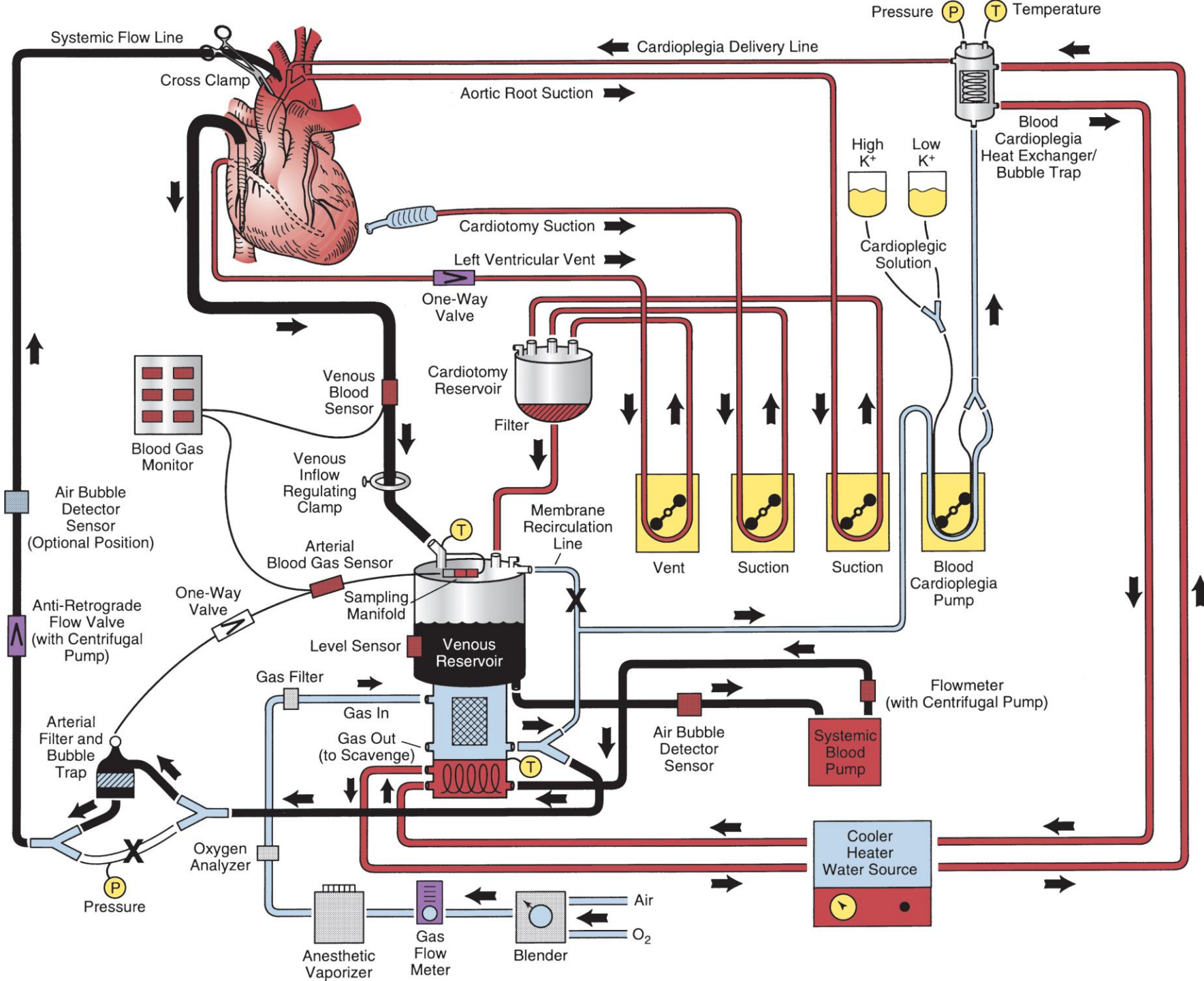
One of the most important biomedical inventions



# Perfusion Systems

- Venous and arterial cannulation
- Venous reservoir
- Oxygenators
- Heat exchangers
- Pumps
- Filters and bubble traps
- Tubing and connectors
- Heparin-coated circuits
- Cardiotomy reservoir and field suction
- Venting
- Cardioplegia delivery systems
- Hemoconcentrators (Hemofiltration/ultrafiltration)
- Monitors and safety devices

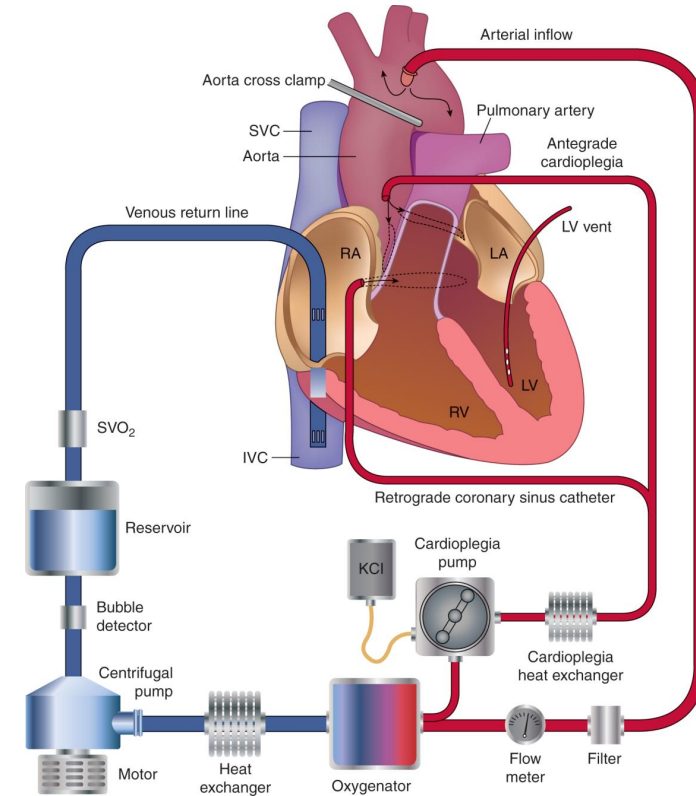




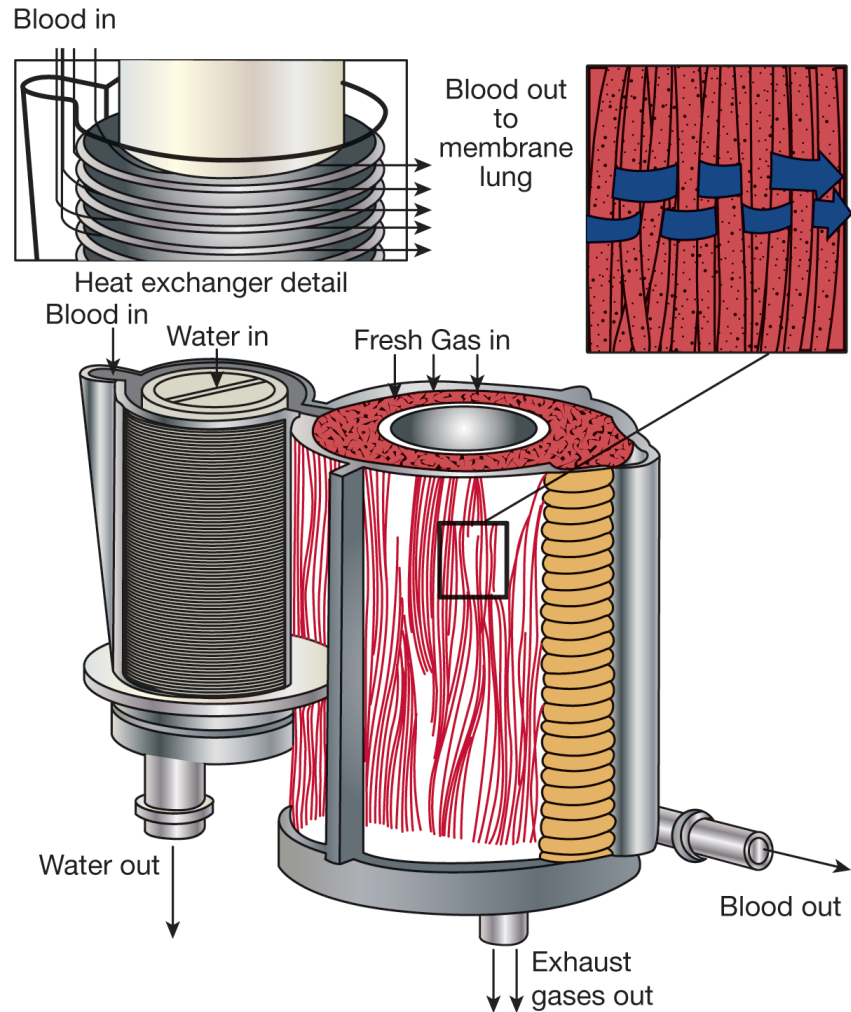


# Venous Reservoir

- 1~3L of blood
- Store excess blood
- Bubble trap
- Access for drugs, fluids, or blood
- To provide time for the perfusionist to act if venous drainage is sharply reduced or stopped



# Oxygenators



## Recommendations for selection of an oxygenator

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref <sup>c</sup>
Microporous membrane oxygenators are recommended as the first choice for use in CPB.	I	B	[78]
Polymethylpentene membrane oxygenators are not recommended when volatile anaesthetics are used during the procedure.	III	B	[84, 86]

<sup>a</sup>Class of recommendation.

<sup>b</sup>Level of evidence.

<sup>c</sup>References.

CPB: cardiopulmonary bypass.

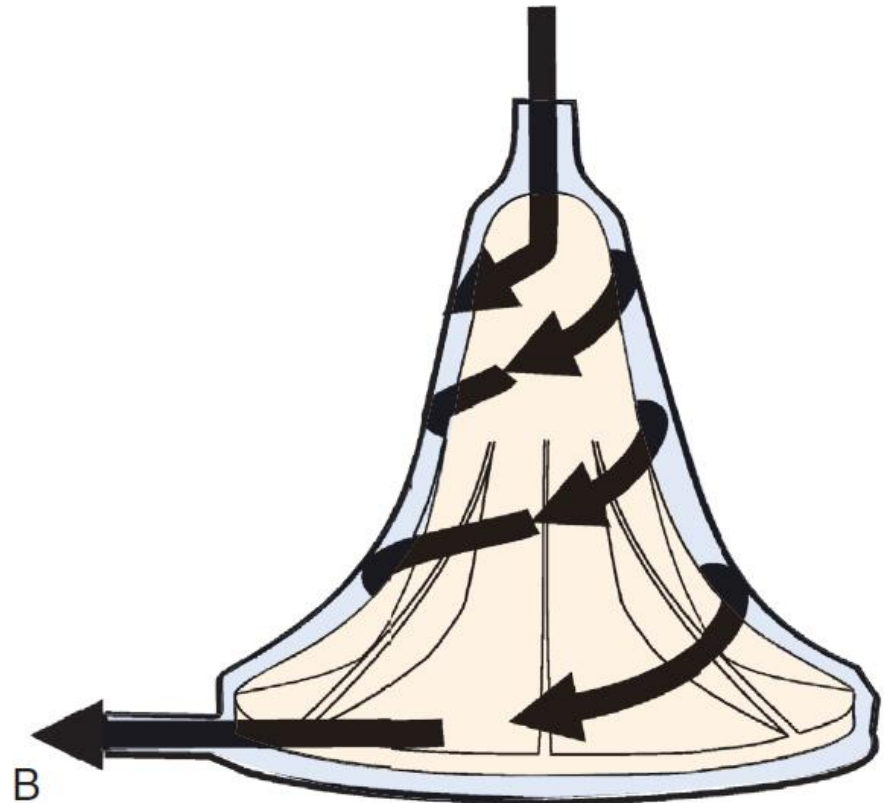
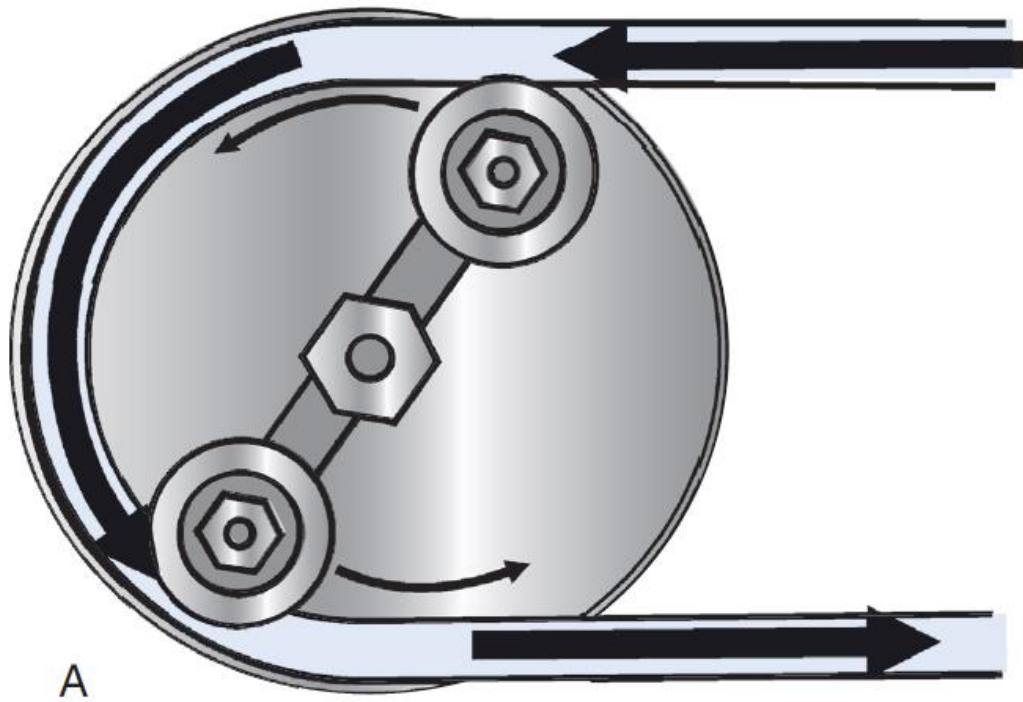
# Heat Exchangers



- Heated temperature  $< 40\text{ }^{\circ}\text{C}$
- Temperature gradient between body and circuit  $< 10\text{ }^{\circ}\text{C}$



# Types of Pump



# Roller vs Centrifugal Pumps

	Roller pump	Centrifugal pump
Description	Nearly occlusive Afterload independent	Nonocclusive Afterload sensitive
Advantages	Low prime volume Low cost No potential for backflow Shallow sine-wave pulse	Portable, position insensitive Safe positive and negative pressure Adapts to venous return Superior for right or left heart bypass Preferred for long-term bypass Protects against massive air embolism
Disadvantages	Excessive positive and negative pressure Spallation Tubing rupture Potential for massive air embolism Necessary occlusion adjustments Requires close supervision	Large priming volume Requires flowmeter Potential passive backward flow Higher cost

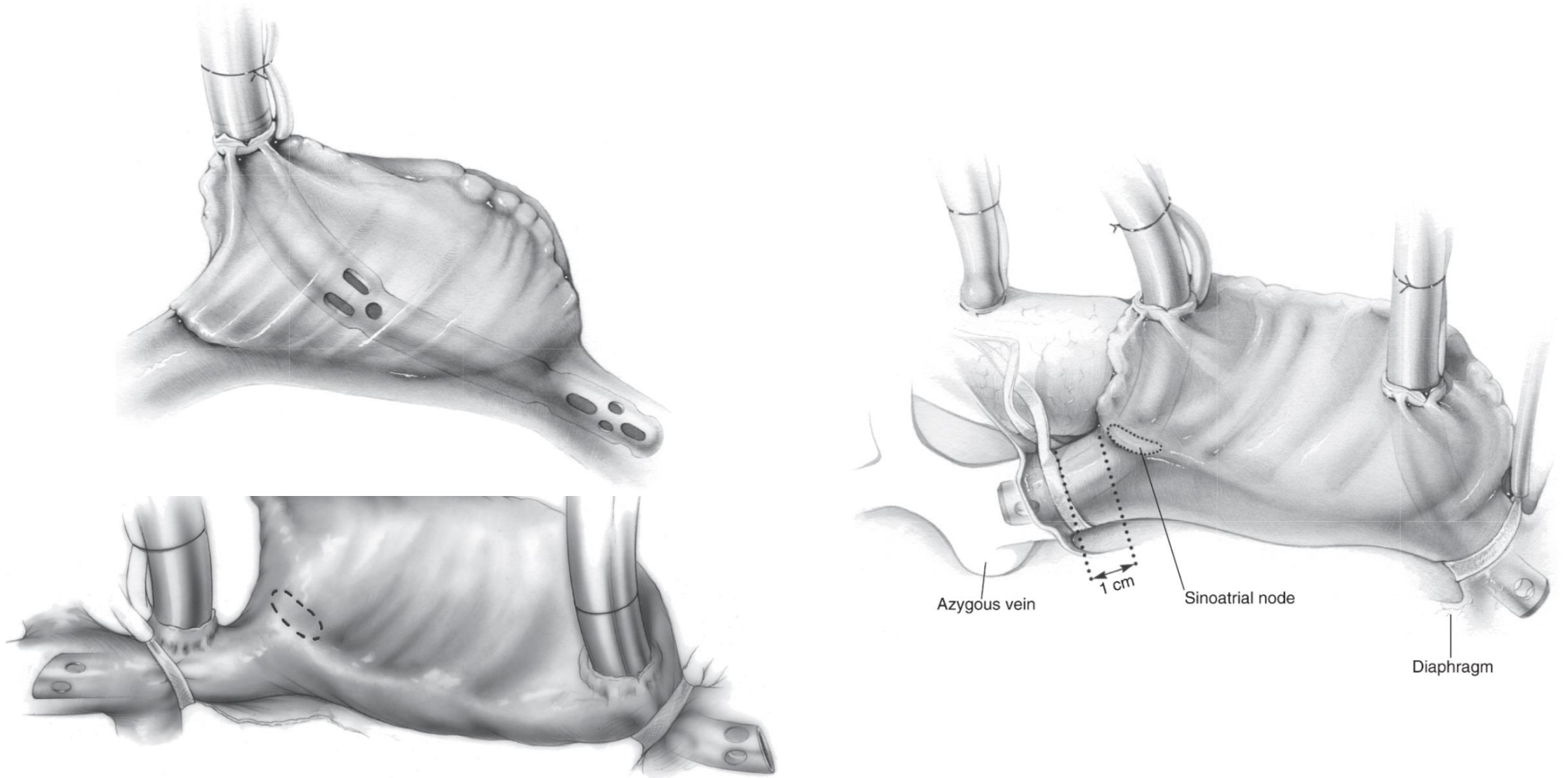
# Centrifugal Pump and Roller Pump in Adult Cardiac Surgery: A Meta-Analysis of Randomized Controlled Trials

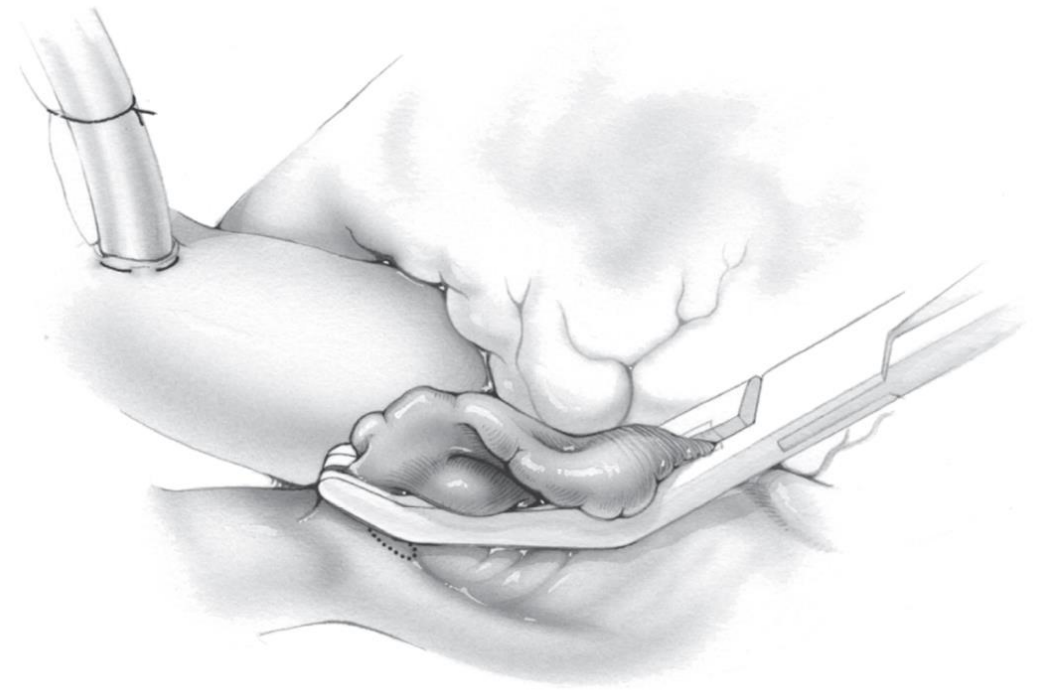
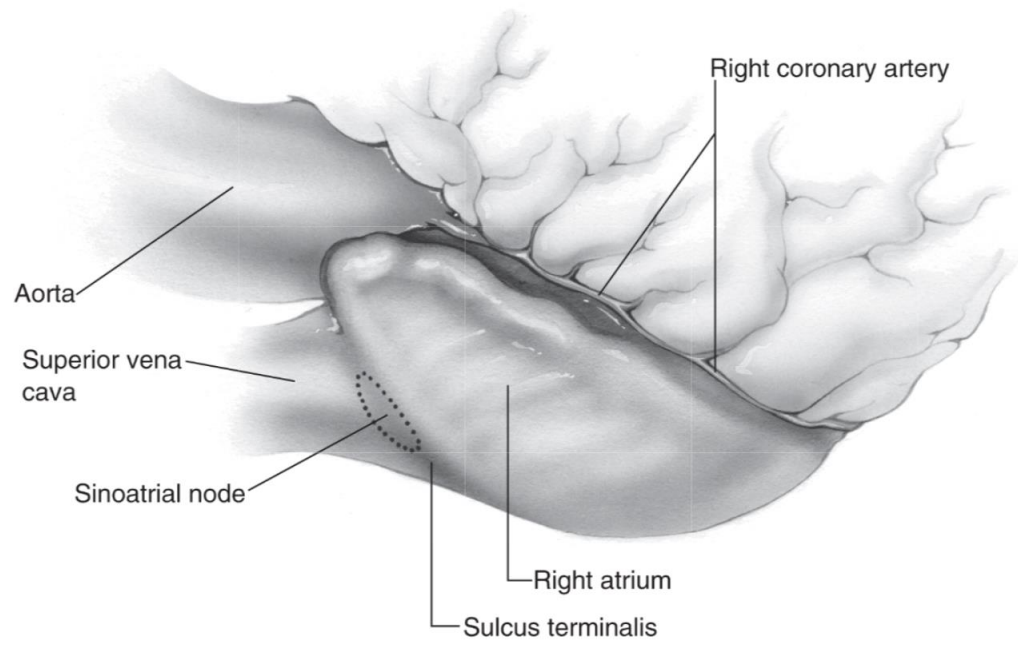
Richard Saczkowski, Michelle Maklin, Thierry Mesana, Munir Boodhwani, and Marc Ruel

*Department of Cardiac Surgery, Royal Columbian Hospital, New Westminster, British Columbia, Canada*

- 18 randomized controlled trials with 1868 patients
- Predominantly isolated CABG
- **No significant difference** for hematological variables, postoperative blood loss, transfusions, neurological outcomes, or mortality

# Venous Cannulation





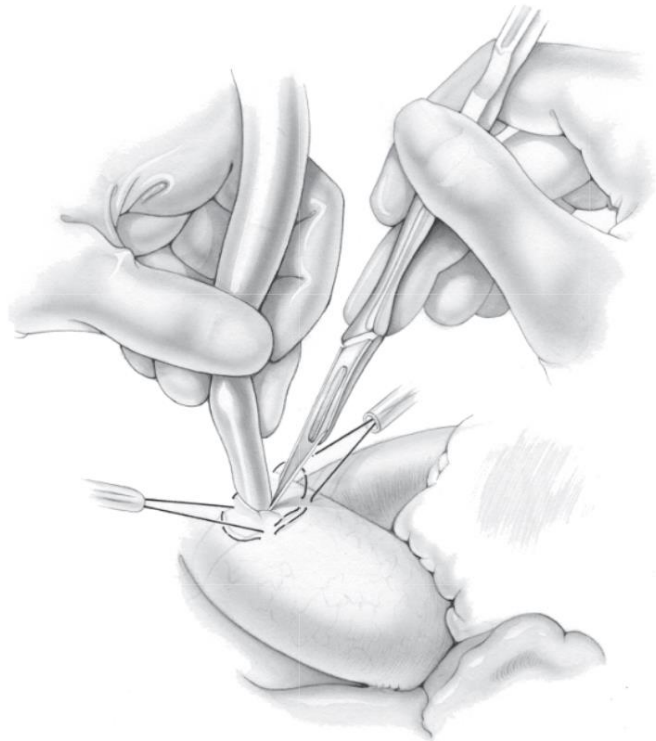


# Causes of Low Venous Return

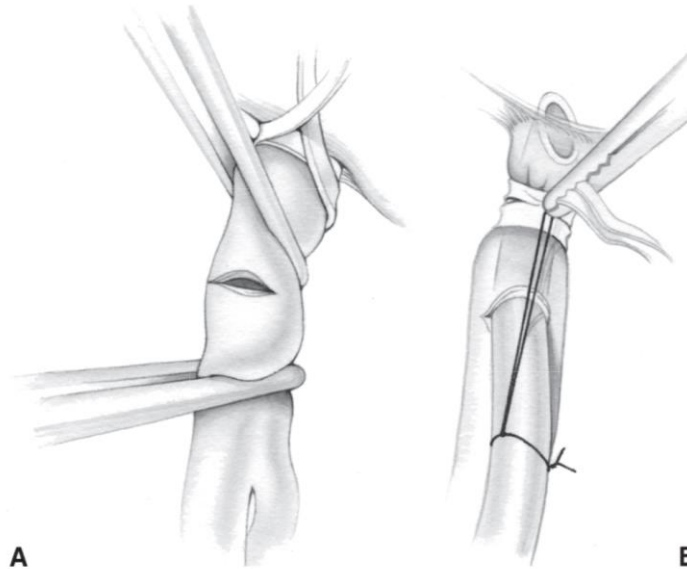
- Low venous pressure
- Hypovolemia
- Drug- or anesthetic-induced venous dilatation
- Inadequate height between the heart and the reservoir
- Inadequate cannula size
- Cannula obstruction or kinking, air-lock

# Arterial Cannulation

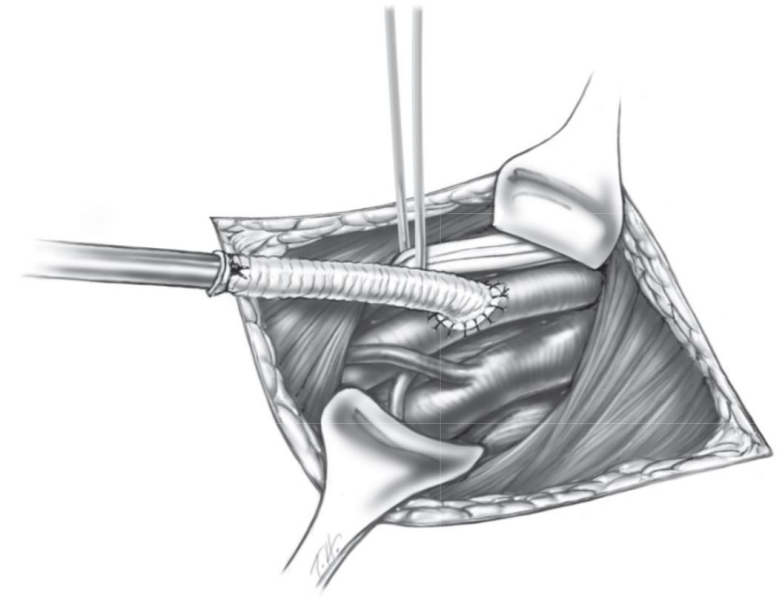
- Optimal arterial BP during cannulation
  - MAP 70~80mmHg, **sBP 100~120mmHg**



Ascending aorta

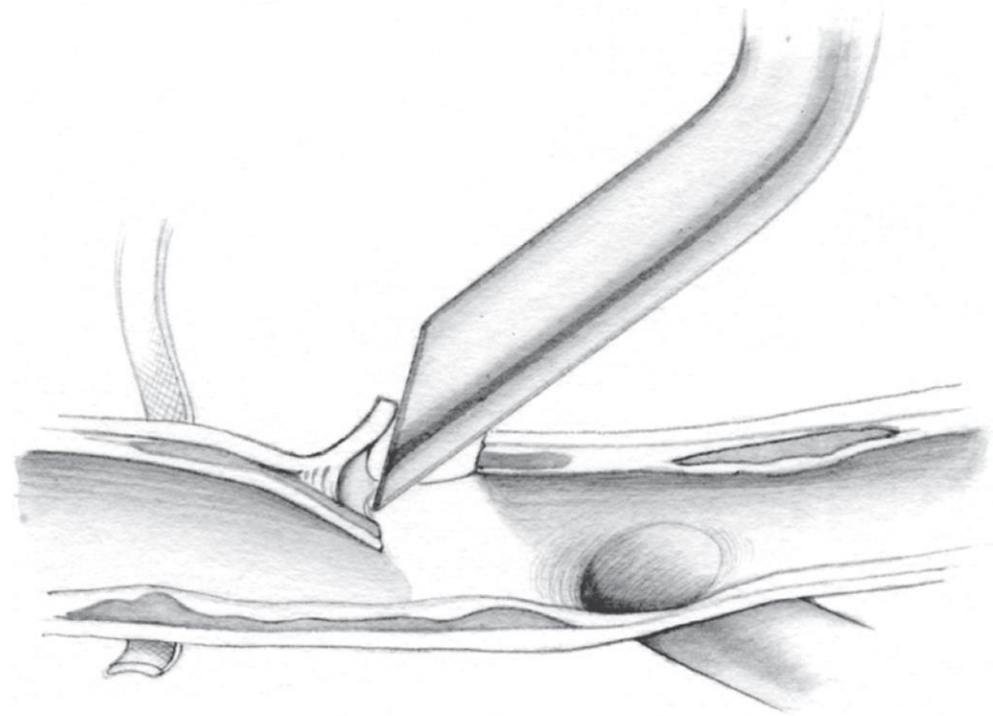
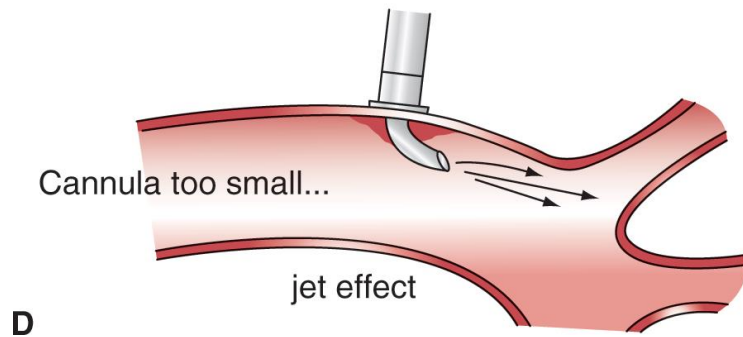
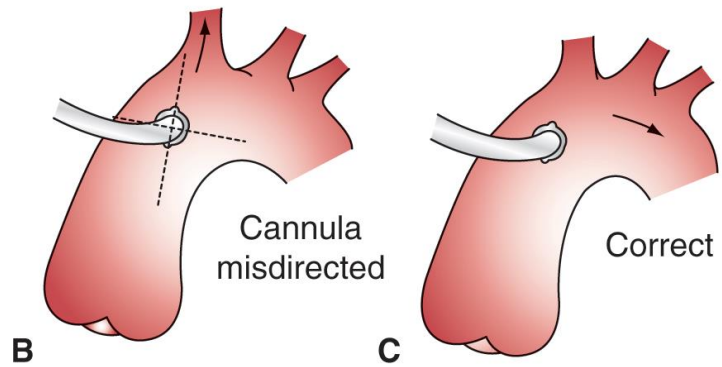
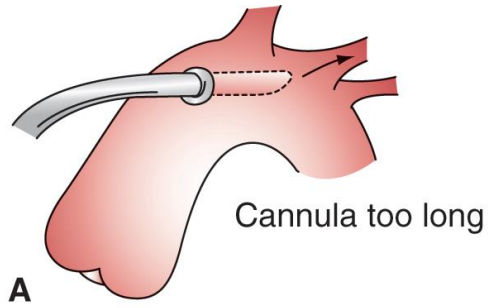


Femoral artery

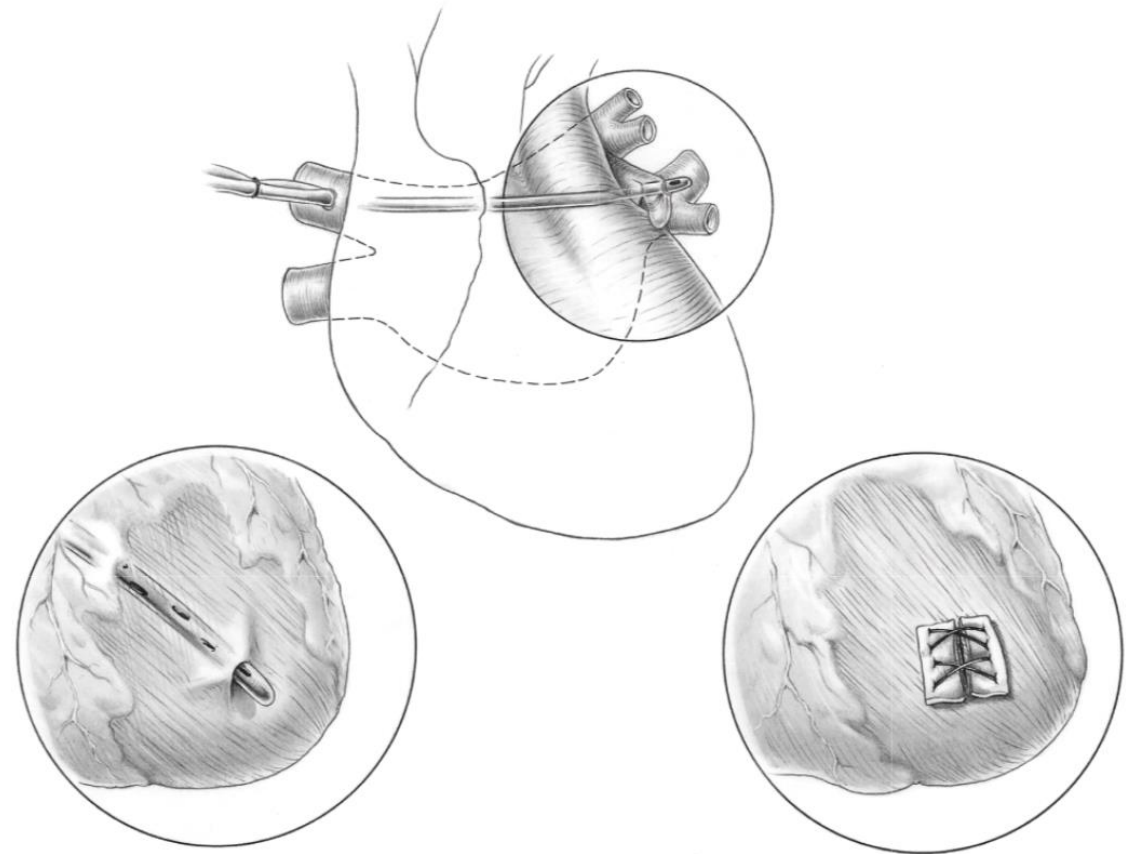
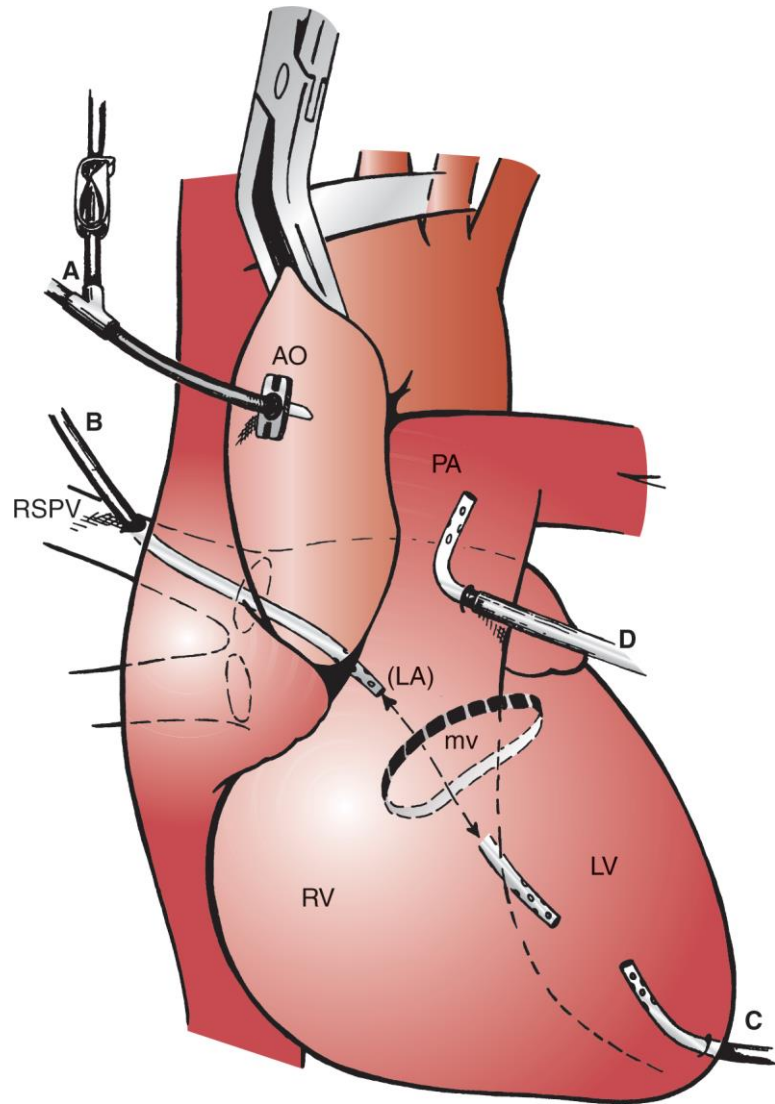


Axillary artery

# Aortic Cannulation Problems



# Sites for Venting LV



# Determinants of Safe Perfusion



- Blood flow rate



- Arterial pressure



- Hematocrit



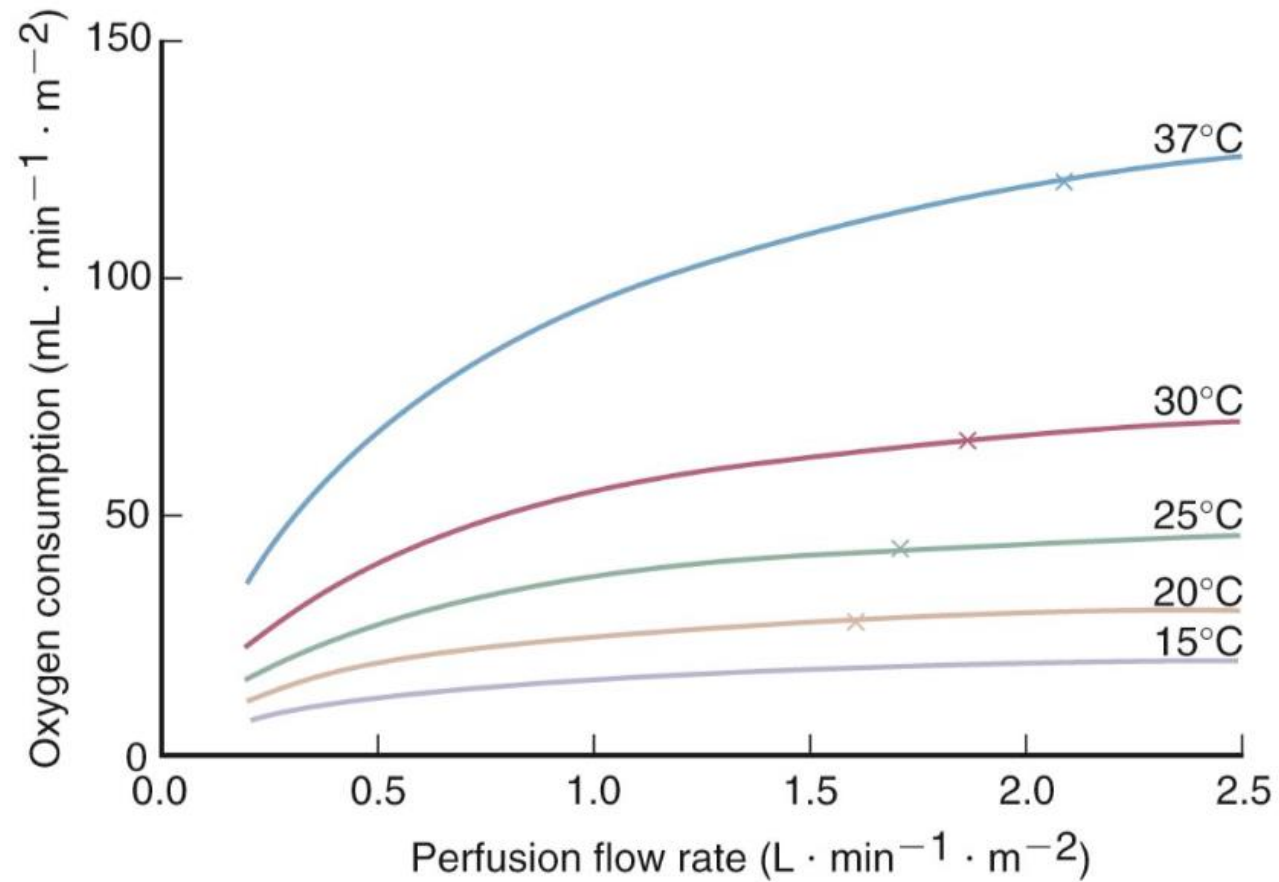
- Temperature



- Acid/Base management



# Blood Flow Rate



**Accepted flow rate : 2.2~2.5L/min/m<sup>2</sup> at 35~37°C, Hct 25%**

# Arterial Pressure



- Mean arterial pressure  
→ near **70mmHg**



- Older patients with vascular disease or hypertension  
→ **70~80mmHg**

# Pressure Monitoring

## Recommendations for control of mean arterial blood pressure during cardiopulmonary bypass

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref <sup>c</sup>
It is recommended to adjust the MAP during CPB with the use of arterial vasodilators (if MAP >80 mmHg) or vasoconstrictors (if MAP <50 mmHg), after checking and adjusting the depth of anaesthesia and assuming sufficiently targeted pump flow.	I	A	[186, 187]
The use of vasopressors to force the MAP during CPB at values higher than 80 mmHg is not recommended.	III	B	[186, 191, 199]
It is recommended that vasoplegic syndrome during CPB be treated with $\alpha$ 1 adrenergic agonist vasopressors.	I	C	
In patients with vasoplegic syndrome refractory to $\alpha$ 1-adrenergic agonist vasopressors, alternative drugs (vasopressin, terlipressin or methylene blue) should be used, alone or in combination with $\alpha$ 1-agonists.	IIa	B	[194, 196, 197]
Hydroxocobalamin may be used to treat vasoplegic syndrome during CPB.	IIb	C	

<sup>a</sup>Class of recommendation.

<sup>b</sup>Level of evidence.

<sup>c</sup>References.

CPB: cardiopulmonary bypass; MAP: mean arterial pressure.

# Hematocrit

- Ideal hematocrit : controversial, but **usually 20~25%** during CPB
- Low hematocrit
  - Reduce blood viscosity and hemolysis
  - Reduce oxygen-carrying capacity
  - Increase cerebral blood flow
- Viscosity remains stable when percent Hct and blood temperature are equal (ex. Same viscosity : 37°C 37% Hct = 20°C 20% Hct)

# Temperature

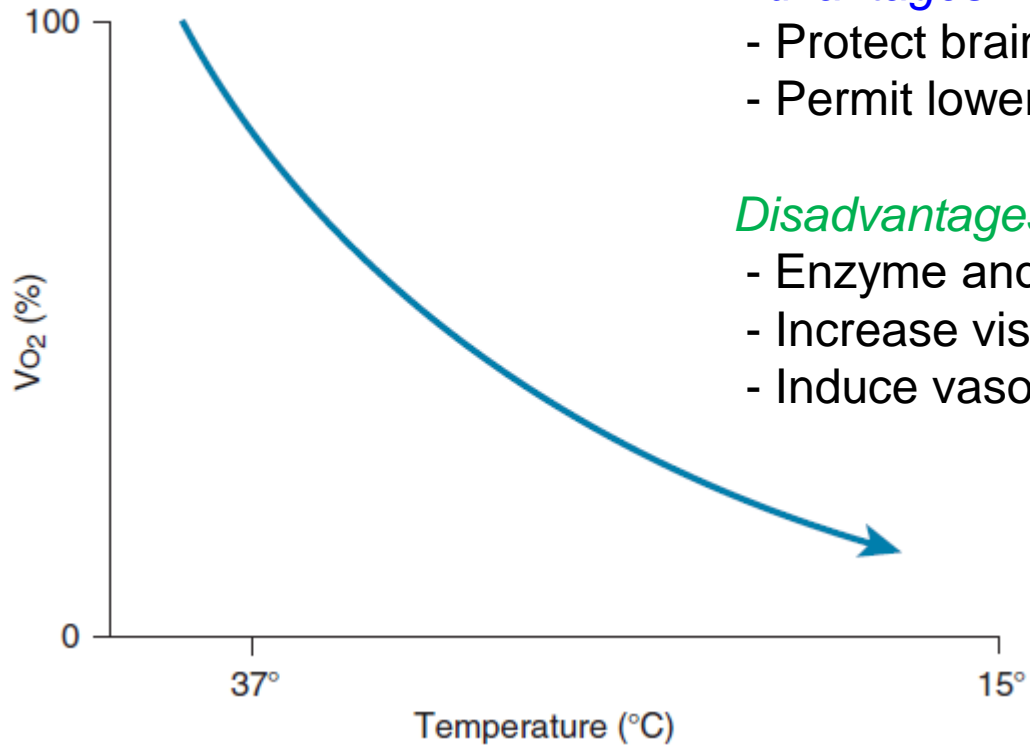
## Hypothermia

### Advantages

- Protect brain
- Permit lower flow perfusion and lower hematocrit

### Disadvantages

- Enzyme and organ dysfunction (bleeding $\uparrow$ , SVR $\uparrow$ , delayed cardiac recovery)
- Increase viscosity
- Induce vasoconstriction



O<sub>2</sub> consumption decreases by 50% for every 10°C drop in temperature



# Acid/Base Management



- $1^{\circ}\text{C} \downarrow \rightarrow \text{pH } 0.015 \text{ units } \uparrow$
- Alkalosis and hypocarbia  $\rightarrow$  cerebral blood flow  $\downarrow$
- **pH-stat strategy**
  - Addition of  $\text{CO}_2$  during hypothermia
  - Preferable in children
- **$\alpha$ -stat strategy**
  - No active correction of pH with hypothermia
  - Preferable in adult

# Heparin and Protamine

- Heparin dose : 300U/kg
- Target activated clotting time (ACT) : 400~480 sec
- Protamine dose : 1mg/100U of heparin
- The most common problem related to protamine
  - Heparin rebound
  - Hypotension
  - Anaphylactoid reactions
  - Pulmonary vasoconstriction
  - Direct antiplatelet effect

## Recommendations for periprocedural anticoagulation management

Recommendations	Class <sup>a</sup>	Level <sup>b</sup>	Ref <sup>c</sup>
<b>Heparin management</b>			
ACT above 480 s during CPB should be considered in CPB with uncoated equipment and cardiotomy suction. The required target ACT is dependent on the type of equipment used.	Ila	C	
Individualized heparin and protamine management should be considered to reduce postoperative coagulation abnormalities and bleeding complications in cardiac surgery with CPB.	Ila	B	[165, 166, 169]
In the absence of individual heparin dosing tools, it is recommended that ACT tests be performed at regular intervals based on institutional protocols, and heparin doses have to be given accordingly.	I	C	
<b>Protamine management</b>			
Protamine overdosing should be avoided in order to reduce postoperative coagulation abnormalities and bleeding complications in cardiac surgery with CPB.	Ila	B	[172]
<b>Alternative anticoagulation</b>			
In patients with contraindications to heparin and/or protamine usage and in need of an operation requiring CPB, anticoagulation with bivalirudin should be considered.	Ila	B	[174, 176]
In patients with contraindications to heparin and/or protamine usage, in need of an operation requiring CPB and significant renal dysfunction, anticoagulation with argatroban may be considered.	IIb	C	

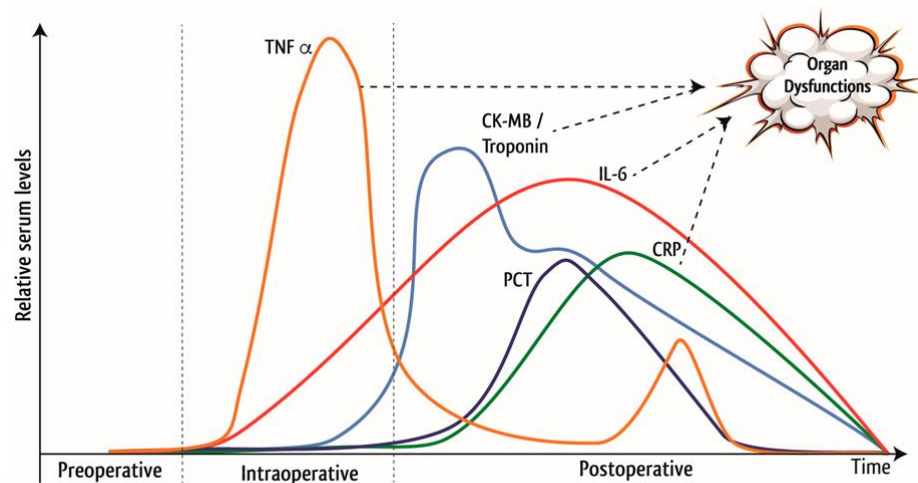
<sup>a</sup>Class of recommendation.

<sup>b</sup>Level of evidence.

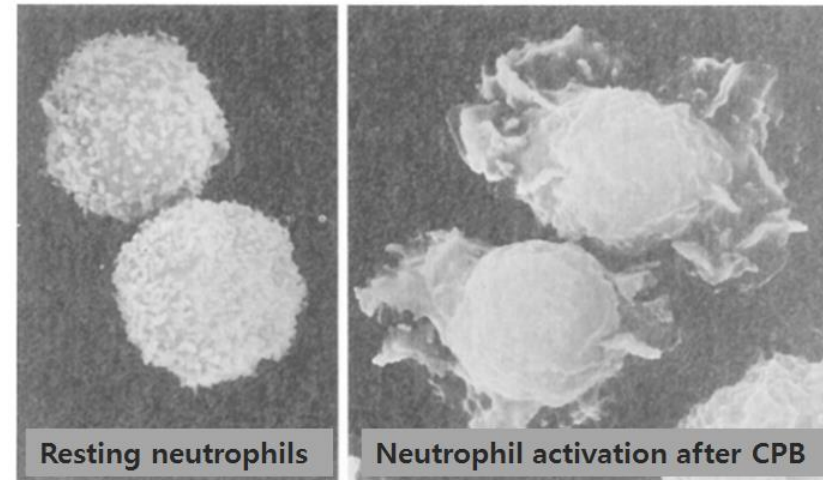
<sup>c</sup>References.

ACT: activated clotting time; CPB: cardiopulmonary bypass.

# Inflammatory Responses to CPB



Aileen H., et al. Nutrients 2018



Baggiolini M. Nature 1998; 392:565.

**Activation of inflammatory mediator and alteration of immune function**

# Coagulopathy Related to CPB








- Hemodilution
- Hypothermia
- Hemolysis
- Heparinization
- Activation of the coagulation system

# Endocrine and Electrolytes Responses to CPB

- Antidiuretic hormone (vasopressin)↑
- Adrenocorticotropin↑
- T3 and T4 responses to TSH↓
- Adrenal responses↑
- Hyperglycemia and hypoinsulinemia
- Hypomagnesemia

# Organ Damage Related to CPB

- Neurologic injury
  - Stroke
  - Delirium
  - Cognitive decline
- Lung injury
  - Atelectasis
  - ARDS
- Renal injury

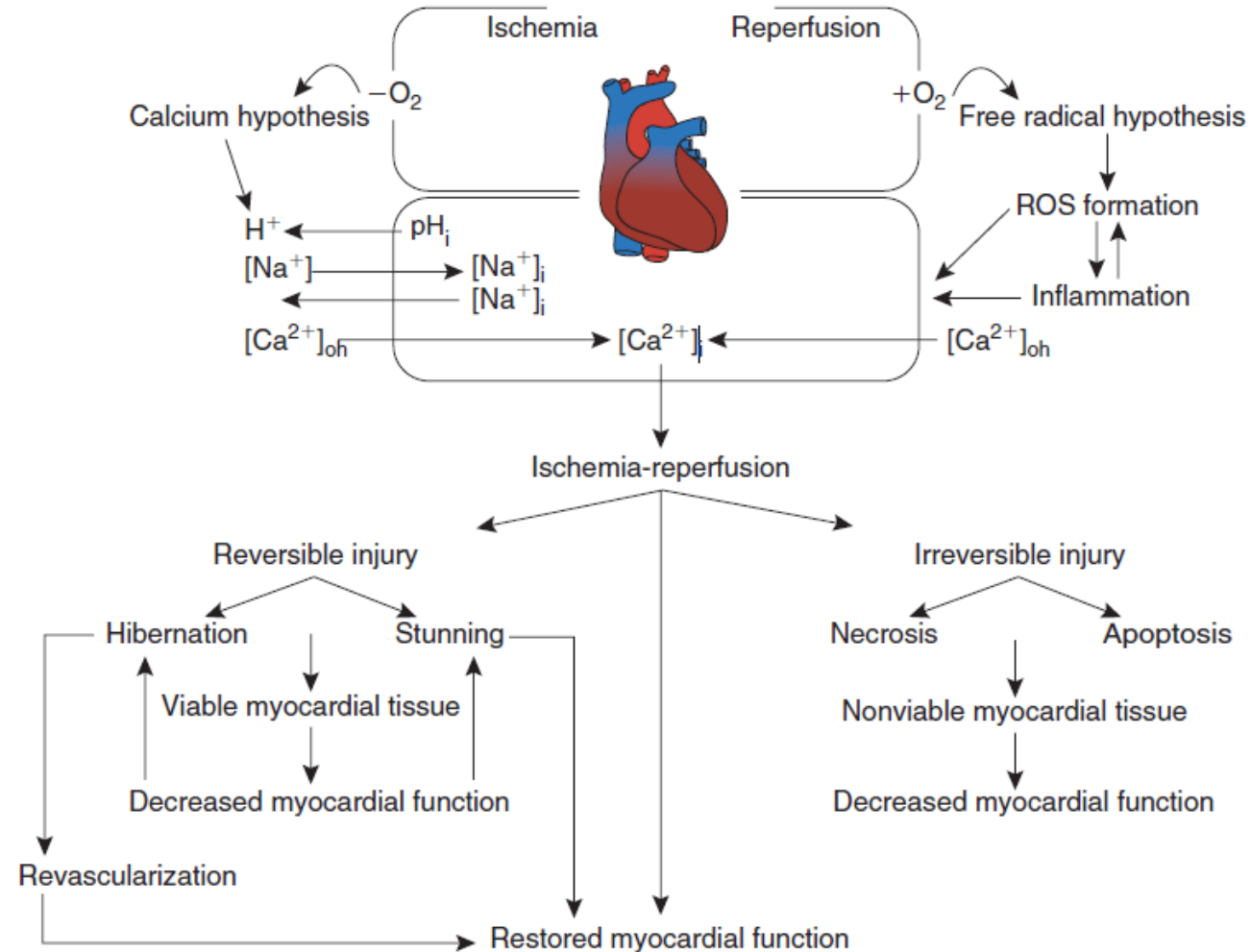
	Mediators of injury	Ensuing organ damage
	<ul style="list-style-type: none"> <li>⚡ Vasodilation through histamine and bradykinin</li> <li>⚡ Vasomotor paresis through cortisol deficiency</li> <li>⚡ Capillary fluid leak</li> </ul>	 <ul style="list-style-type: none"> <li>⊖ Systemic hypotension</li> <li>⊖ General edema</li> </ul>
	<ul style="list-style-type: none"> <li>⚡ Increased permeability of blood-brain barrier</li> <li>⚡ Cerebral edema</li> </ul>	 <ul style="list-style-type: none"> <li>⊖ Disturbed thermoregulation</li> <li>⊖ Dysregulation of the autonomic nervous system</li> <li>⊖ Disrupted hypothalamic-pituitary-adrenal axis</li> <li>⊖ Cognitive dysfunction and delirium</li> </ul>
	<ul style="list-style-type: none"> <li>⚡ Inhibition of cardiomyocyte contraction</li> <li>⚡ Decreased ventricular compliance</li> <li>⚡ Impaired systolic function</li> </ul>	 <ul style="list-style-type: none"> <li>⊖ Low cardiac output</li> </ul>
	<ul style="list-style-type: none"> <li>⚡ Pulmonary edema</li> <li>⚡ Impaired surfactant production</li> <li>⚡ Decreased lung compliance</li> <li>⚡ Pulmonary vascular dysfunction</li> </ul>	 <ul style="list-style-type: none"> <li>⊖ ARDS</li> <li>⊖ Ventilation-perfusion mismatch</li> <li>⊖ Hypoxemia</li> </ul>
	<ul style="list-style-type: none"> <li>⚡ Tubular injury and edema</li> <li>⚡ Reduced glomerular filtration rate and creatinine clearance</li> </ul>	 <ul style="list-style-type: none"> <li>⊖ Acute kidney injury</li> </ul>
	<ul style="list-style-type: none"> <li>⚡ Increased intestinal permeability</li> </ul>	 <ul style="list-style-type: none"> <li>⊖ Bacterial translocation and endotoxemia</li> </ul>



# Myocardial Protection



# Ischemia-Reperfusion Injury



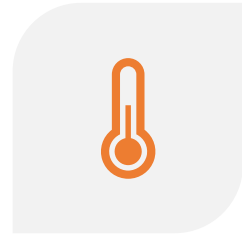
# Cardioplegia

- By *Will Sealy*, 1958, Duke University
  - Cardio- ; heart
  - Plegia ; paralysis, cessation of motion
- Maintaining the myocardium in a state of '***reversible injury***' for longer periods

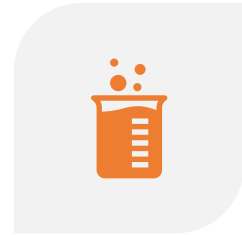
# Principles for Myocardial Protection



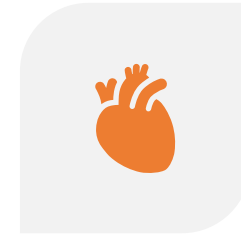
RAPID INDUCTION  
OF ARREST



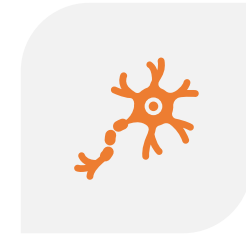
MILD OR MODERATE  
HYPOTHERMIA



APPROPRIATE  
BUFFERING OF THE  
CARDIOPLEGIC  
SOLUTION



AVOIDANCE OF  
MYOCARDIAL  
EDEMA



AVOIDANCE OF  
SUBSTRATE  
DEPLETION

# Types of Cardioplegia



Crystalloid cardioplegia



Blood cardioplegia

# Rationale for Using Blood for Cardioplegia

- Provides an oxygenated environment
- Limits hemodilution
- Affords an excellent buffering capacity and osmotic properties
- Allows for physiologic electrolytes and pH
- Offers endogenous antioxidants and free-radical scavengers
- Less complex to prepare

# Components of Cardioplegic Solutions

	Na	K	Mg	Ca	Buffer	pH	Osm	Other components
Intracellular crystalloid CP								
Bretschneider's no. 3	12.0	10.0	4.0	0	Histidine	7.4	320	Procaine, mannitol
Bretschneider's HTK	15.0	9.0	4.0	0	Histidine	7.3	310	Ketoglutarate, typtophan, mannitol
Roe's	27.0	20.0	3.0	0	THAM	7.6	347	Glucose
Extracellular crystalloid CP								
del Nido solution	140	5	0.75	0	Bicarbonate	7.4	375	Lidocaine, mannitol
St. Thomas no. 1	144.0	20.0	32.0	4.8	None	5.5	285	Procaine
St. Thomas no. 2 (Plegisol)	110.0	16.0	32.0	1.2	Bicarbonate	7.8	324	None
Tyer's	138.0	25.0	3.0	1.0	Bicarbonate	7.8	275	Acetate, gluconate
Blood CP								
Cold induction	118.0	18.0	1.6	0.3-0.5	±THAM	7.6-7.8	320-340	Glucose, oxygen
Warm induction	122.0	25.0	1.6	0.15-0.25	±THAM	7.5-7.6	340-360	Glucose, oxygen, glutamate, aspartate

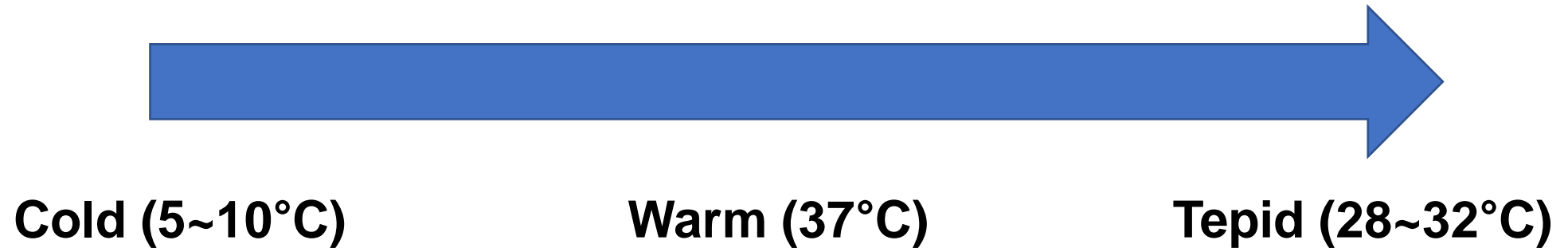
\* Intracellular type : Absent or low concentrations of sodium and calcium

\* Extracellular type : Higher concentrations of sodium, calcium, and magnisium

\* THAM : tris (hydroxymethyl) aminomethane

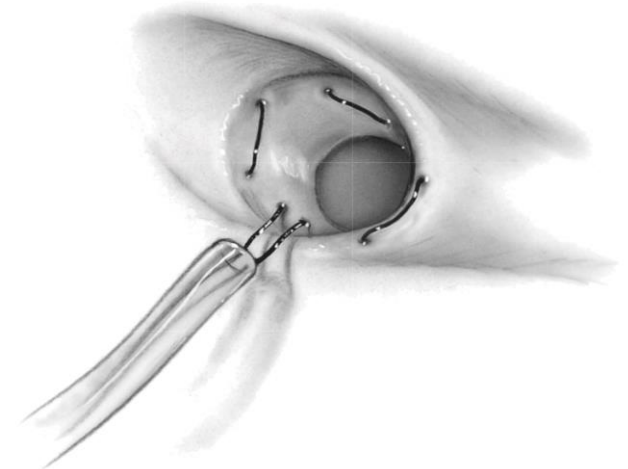
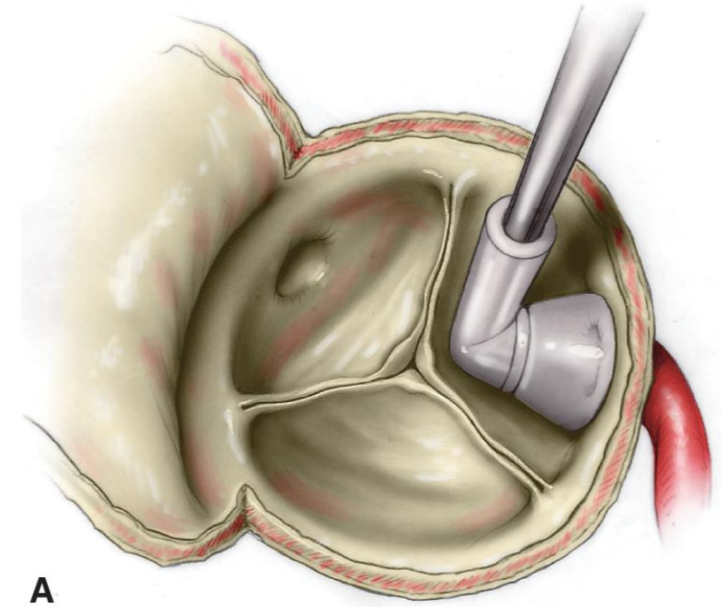
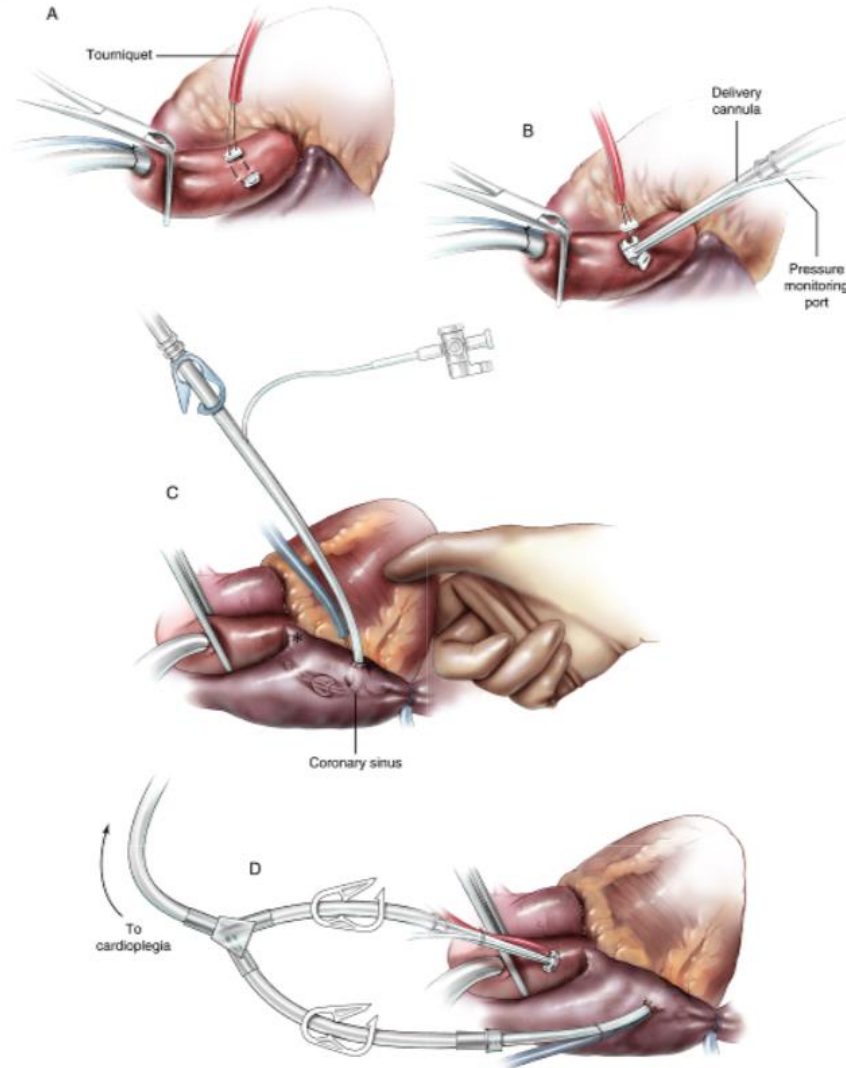


# Trend of Cardioplegia Temperature



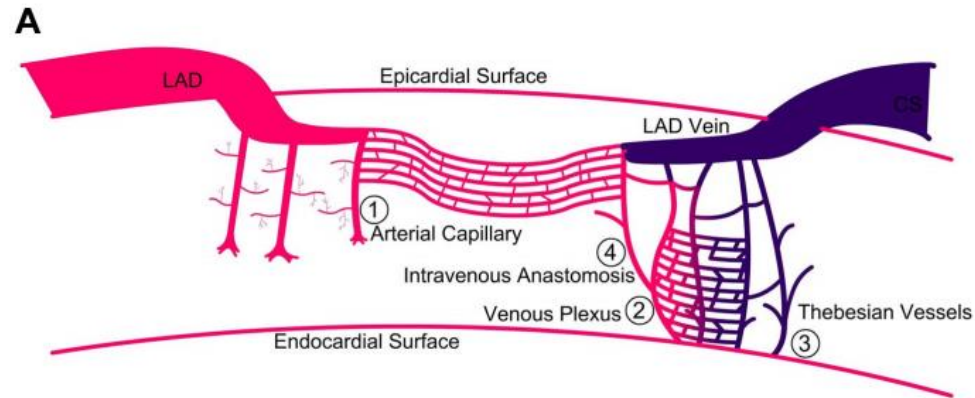
# Delivery Systems

- Antegrade
- Retrograde
- Combined

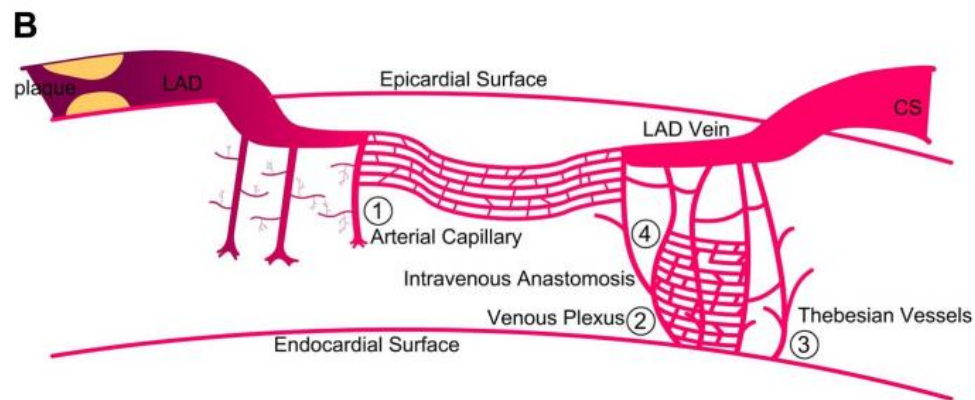


# Coronary Circulation

Antegrade

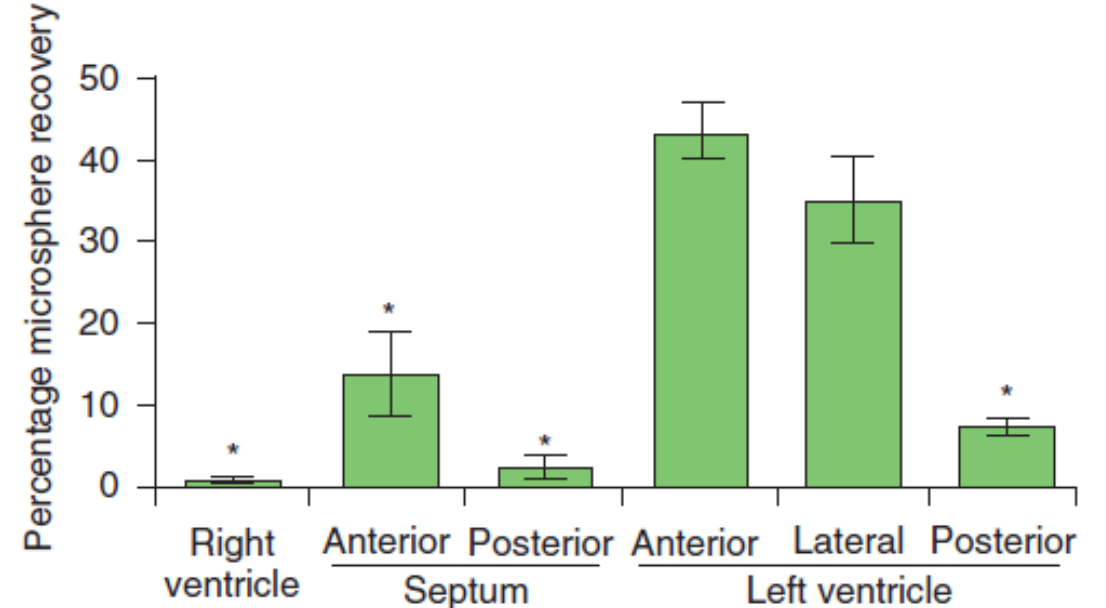


Retrograde



# Antegrade vs Retrograde

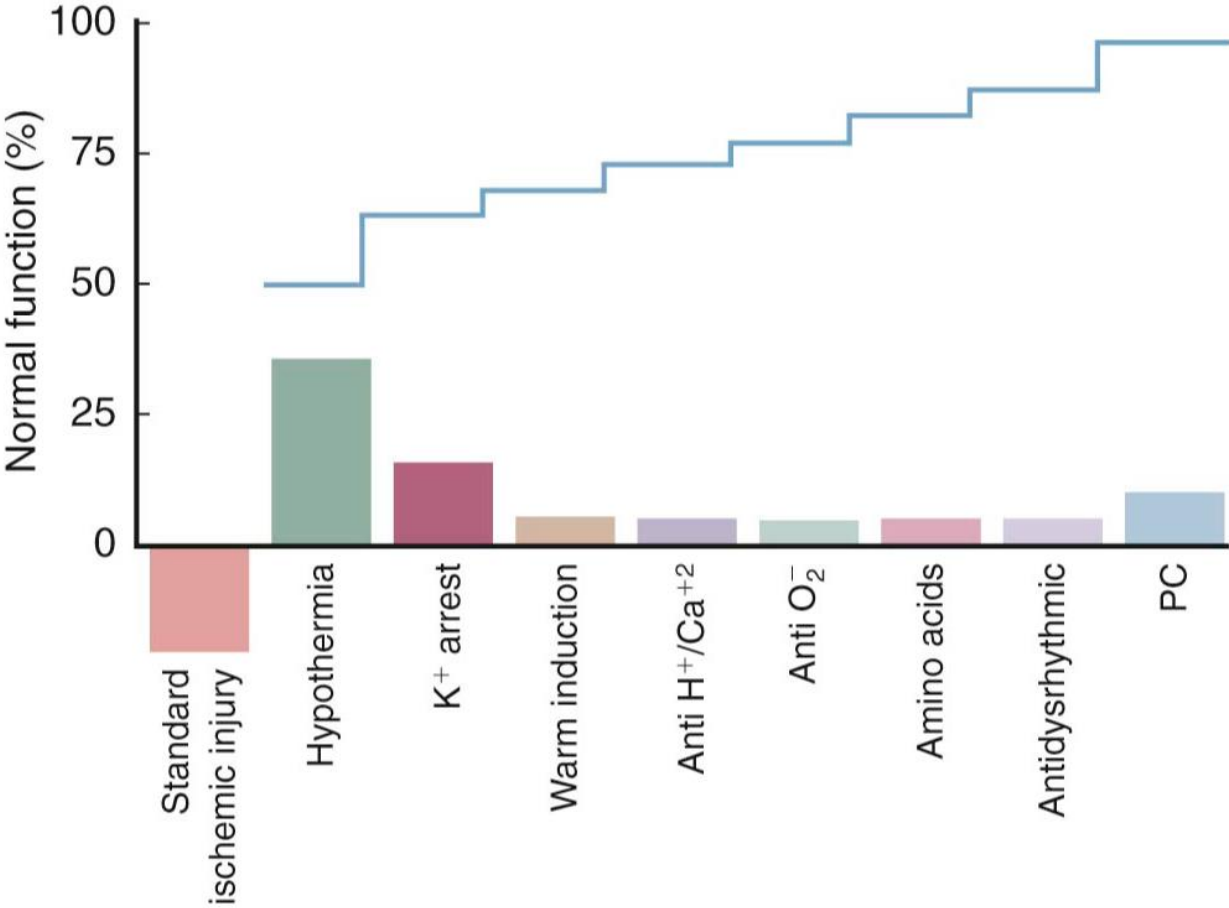
- Antegrade
  - Possible aortic injury
  - Limits coronary artery occlusion or aortic insufficiency
- Retrograde
  - Convenient for mitral valve surgery
  - Provide continuous cardioplegia
  - Prevent coronary thromboembolism
  - Easy dislocation
  - Possible injury of coronary sinus
    - Perfusion pressure < 30~40mmHg
  - Inadequate perfusion of RV



# Continuous vs Intermittent

- No differences in patient outcomes
- Most use intermittent cardioplegia in order to have a bloodless field

# Incremental Beneficial Effects

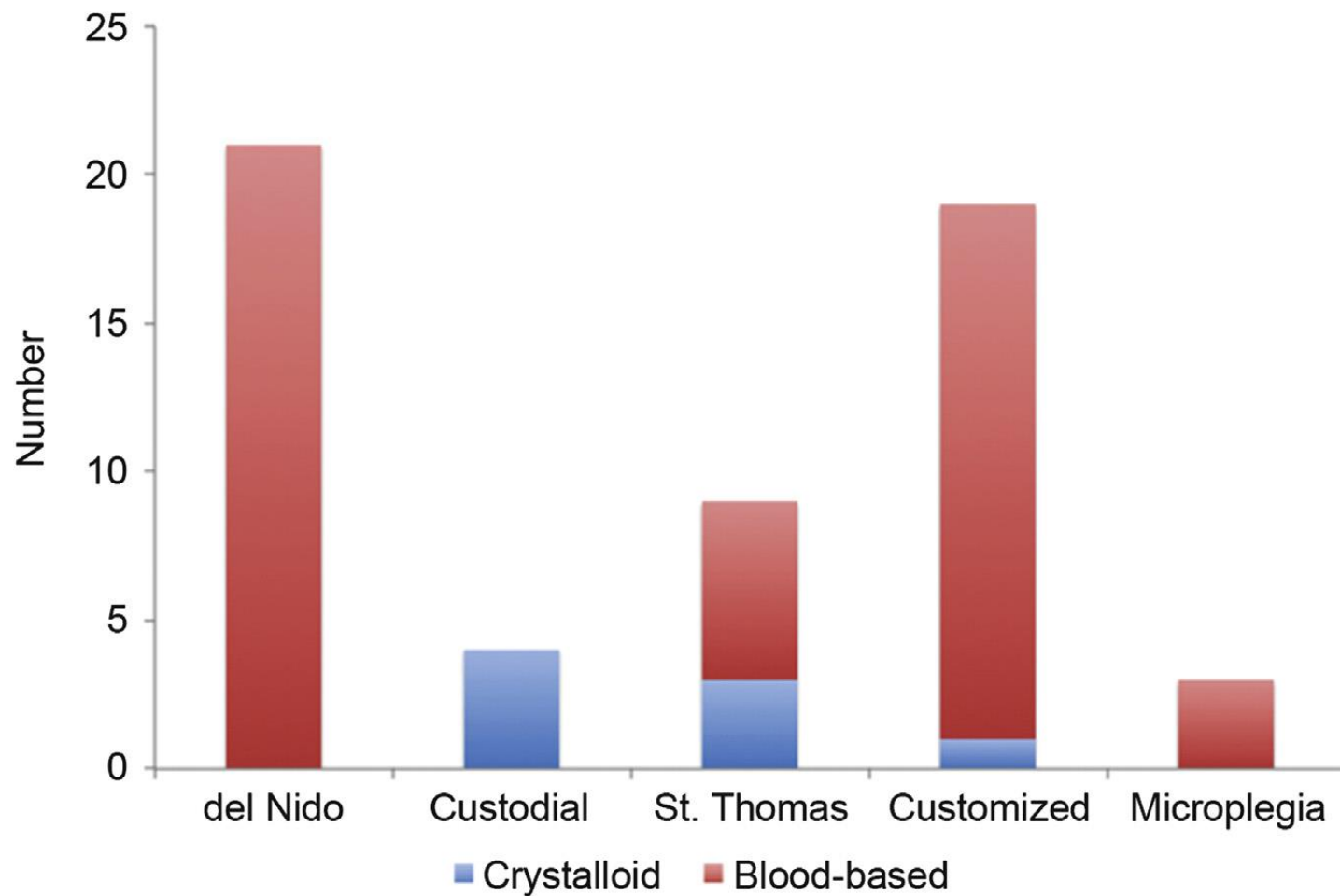


# Current Status of Cardioplegic Technique

- Controversy persists
  - Blood cardioplegia vs Crystalloid cardioplegia
  - Ideal temperature ?
  - Best method of delivery ?
- No international consensus regarding the ideal cardioplegic solution or its use
- How to provide optimal myocardial protection is understanding of the various protection techniques

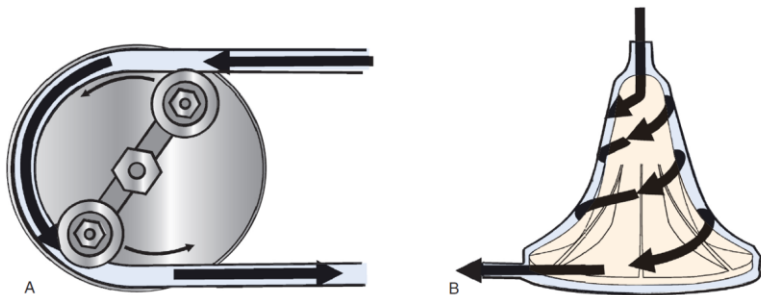


# North American Survey



# Summary

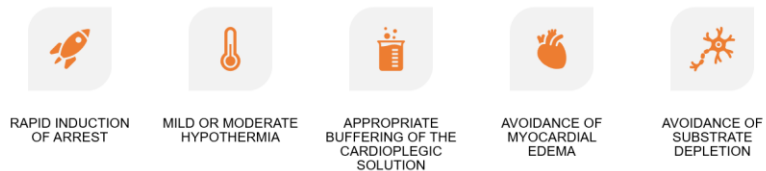
## Types of Pump



## Determinants of Safe Perfusion

- Blood flow rate
- Arterial pressure
- Hematocrit
- Temperature
- Acid/Base management

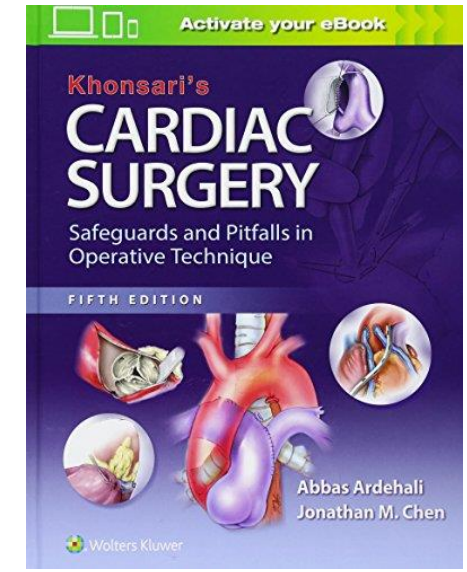
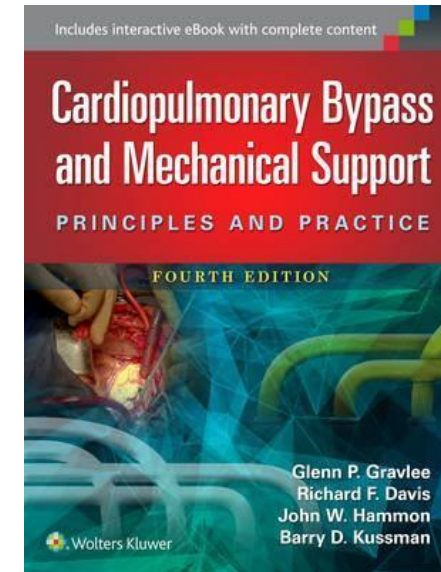
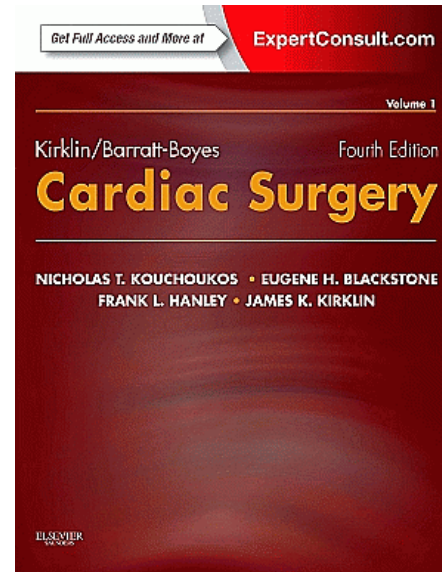
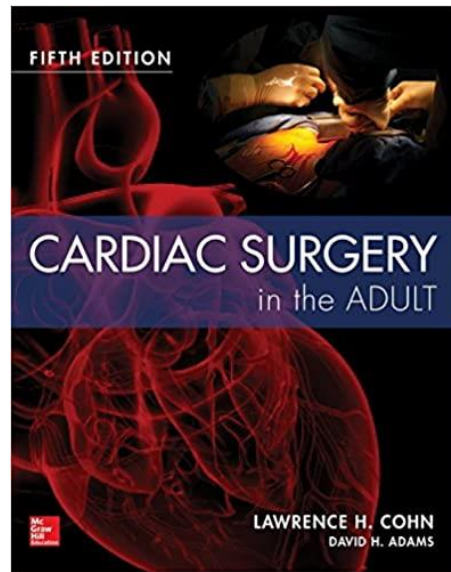
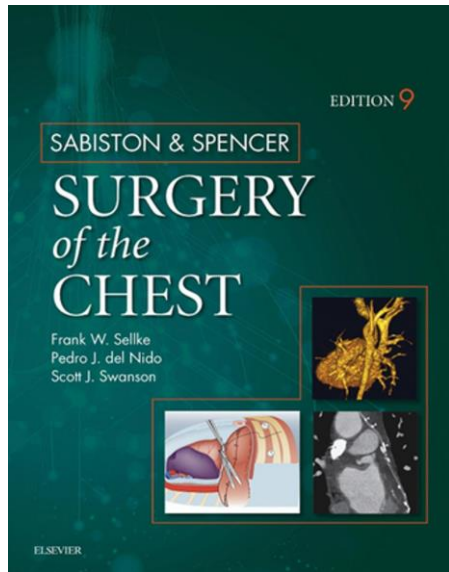
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***Thank you !***