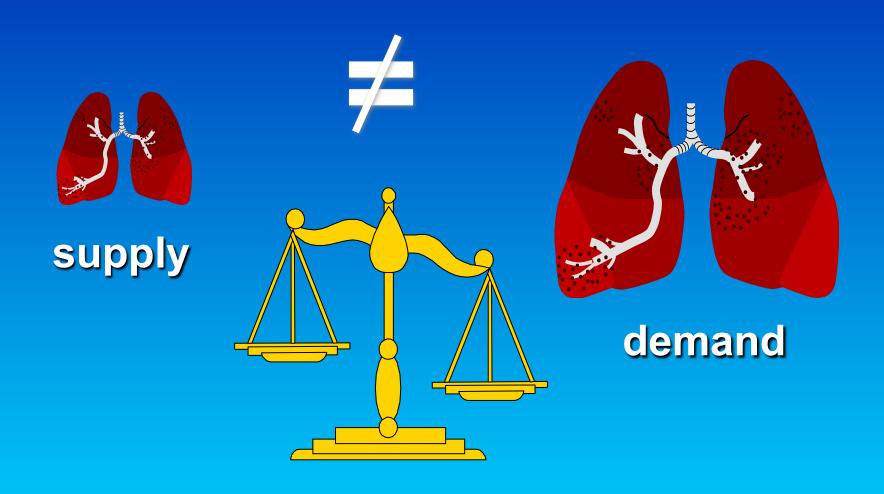


# Newsweek





# Who is an Organ Donor?

- brain-dead individuals on ventilator
  - intubated before brain death
- living donors
  - usually related, not always
  - kidney, liver, bilateral lung lobes
- donation after cardiac death (DCD)
  - kidney, liver, a few lungs

# How many U.S. transplants?

Organ #/year

Kidney 16,000 (6,000 living)

Liver 6,000 (300 living)

**Heart** 2,200

Lung 1,400

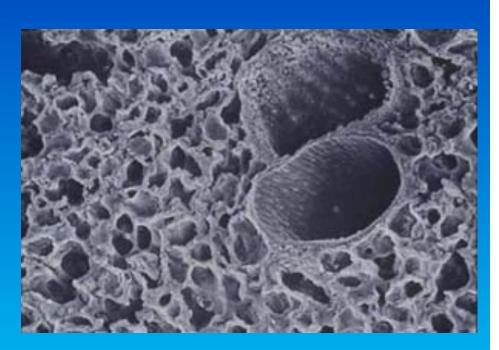
**Corneas** 50,000

# What is a Non-Heart-Beating Donor (NHBD)?

- someone who dies because circulation has stopped
- sometimes organs can be retrieved from NHBDs
- limitation for most organs is ischemic time
- the lung is different

### Rationale

- the lung parenchyma does not rely on blood flow for cellular respiration
- few metabolic functions means low energy requirements
- huge blood reserve after arrest



# **Hypothesis**

If lung cells remain viable after circulatory arrest, then the lung may be suitable for transplant, even if retrieved at substantial intervals after circulatory arrest and death

# **Proof of Concept**

#### A Strategy to Increase the Donor Pool: Use of Cadaver Lungs for Transplantation

Thomas M. Egan, MD, C. Jake Lamber Karl S. Ulicny, Jr, MD, Blair A. Keagy,

Division of Cardiothoraux Surgery, University of North C

A shortage of suitable donors is a scrious obstacle is widespread application of isolated lung transplant for end-stage lung disease. We hypothesized that tissue likely remains viable for a sufficient period of to allow for safe postmortem retrieval of lung transplantation. Studies were conducted in a nonsuimodel of canine lung allotransplantation. Donor an were sacrificed, and subsequent lung harvest wa layed for 1 hour, 2 hours, or 4 hours. Fulmonary retiwas then performed in a standard fashion, flushin lung block with modified Euro-Collins solution. I were then stored for 4 hours before single allotrans tation. Recipient animals were maintained anesthe and followed up for 8 hours. By occlusion of the put

#### Cadaver Lung Donors: Effect of Preharvest Ventilation on Graft Function

Karl S. Ulicny, Jr., MD, Thomas M. Egan, MD, C. Jake Lambert, Jr., MD, Robert L. Reddick, MD, and Benson R. Wilcox, MD

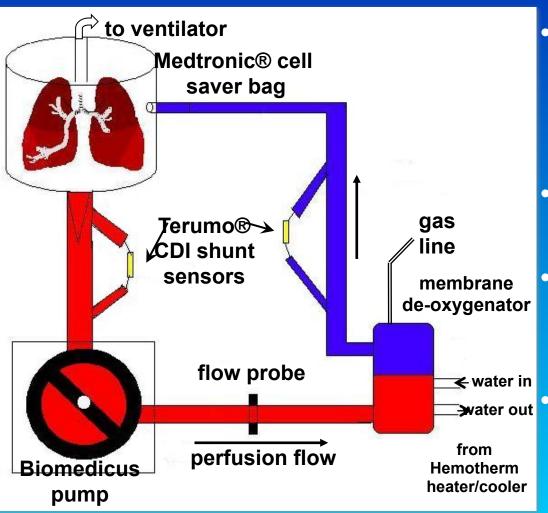
Departments of Surgery and Pathology, The University of North Carolina School of Medicine, Chapel Hill, North Carolina.

The pulmonary donor pool would increase substantially if lungs could be safely transplanted after cessation of circulation. To determine whether ventilation of cadaver lungs could improve graft function, canine donors were sacrificed and then ventilated with 100% oxygen (n = 6) or 100% nitrogen (n = 6); is served as nonventilated controls. Four hours after death, the lungs were flushed with modified Euro-Collins solution and harvested. Controls were ventilated with 100% oxygen only during flush and harvest. Recipients were rendered dependent on the transplanted lung by occlusion of the right pulmonary artery and bronchus 1 hour after transplantation. Ventilation was maintained at a constant inspired oxygen fraction of 0.4. Four controls died of pulmonary edema

shortly after occlusion of the native lung. The mean arterial oxygen tensions in the oxygen-ventilated, nitrogeo-ventilated, and control groups at the end of 8 hours were 81 mm Hg (n = 4), 88 mm Hg (n = 3), and 55 mm Hg in = 2), respectively. Postmortem oxygen ventilation improved early recipient survival and gas exchange. Postmortem nitrogen ventilation improved early gas exchange and delayed recipient death compared with non-ventilated controls. The mechanics of ventilation appears to confer a functional advantage independent of a continued supply of oxygen. Transplantation of lungs harvested from cadavers after cessation of circulation might be feasible.

(Ann Thorac Surg 1993,55:1185-91)

### Ex-vivo human lung perfusion circuit

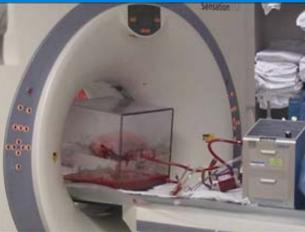


- perfusate is deoxygenated by ventilating membrane with CO2 / N2
- lungs slowly rewarmed to 37°C
- flow slowly increased to 40% donor cardiac output
- PA pressure < 20 mm</li>
  Hg ventilated with TV=
  7cc/kg donor weight

### Ex-vivo evaluation of human lungs

- gas exchange
- bronchoscopy
- CT scan









- HRSA awarded a grant to CDS and UNC to investigate the NHBD donor pool
- Ex-vivo Evaluation of Human Lungs from Non-Heart Beating Donors for Transplant











# Where are the challenges?

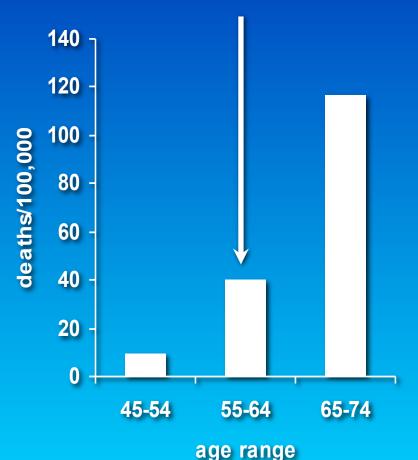




- Emergency Medical Services (EMS)
- County Medical Examiners (ME)
- Organ Procurement Organizations (OPO)

# How many recipients are there?

total COPD deaths = 13,000

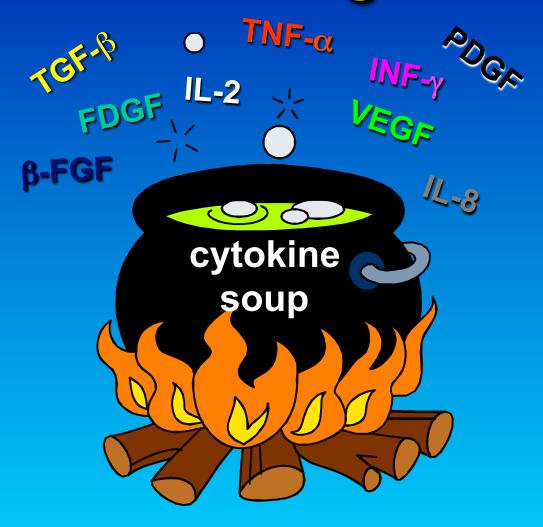


total CF deaths = 400/year

total IPF deaths = 40,000/year

total deaths related to pulmonary hypertension = 15,000/year

# Could NHBD lungs be better?



### Where are we now?

- UNC awarded ARRA grant to perfect ex-vivo lung perfusion and plan a multi-center study of lung tx from NHBDs
  - we will use unsuitable lungs from conventional donors and DCDs
- the grant funds a study to learn about EMS attitudes about organ donation
  - develop a web-based teaching tool for EMS personnel that can be used nationally

### What Was the Plan?

- All cardiac arrests that were terminated in the field and:
  - Not suspicious (by law enforcement)
  - Between 18 and 65 years of age
  - Drivers License with affirmed donation status

### What Was the Plan?

- Then, APP or DC:
  - Obtain family member phone numbers
  - Call Carolina Donor Services (CDS)
- · CDS:
  - Calls family to obtain consent and history
  - Communicates with surgeon to accept or deny

### What Was the Plan?

- If accepted, then:
  - Second EMS unit dispatched to move the patient
  - Crew will ventilate patient en route
  - Crew may remain for harvest
- Estimated 2 patients to be screened per week

# Trainwreckology...

The study of impending failure and the "I told you so"s that follow.

### Went for 3 Months

- 4 patients screened
- 2 accepted and had viable lungs
  - Size problems prohibited transplant (single hospital IRB)



# if you feel like a guinea pig...



you are... and we thank you!

# **Next Steps**

Memo to law enforcement completed

NIH funding application is scored

 We await funding to restart the project

