

Lunge Transplantation Rigshospitalet

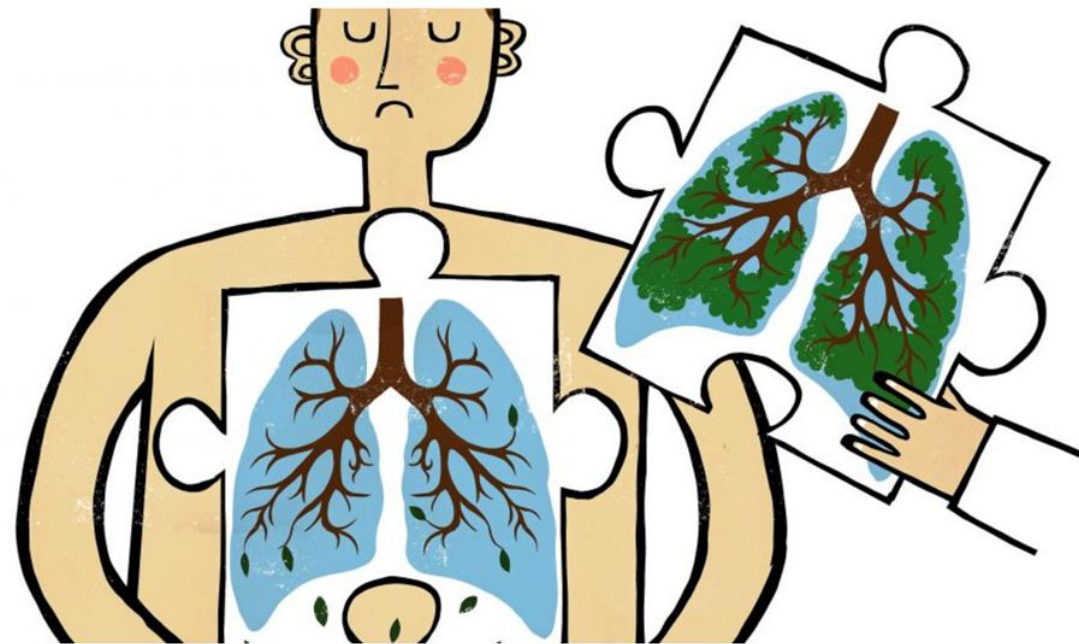


Foto: Tegning: Rasmus Juul

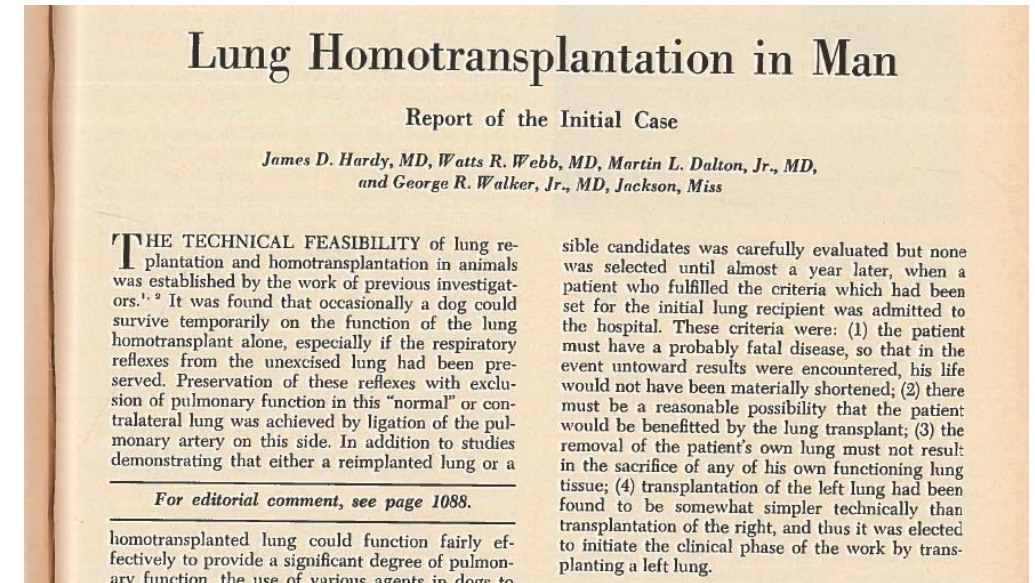
**Livet er den største
gave, man kan give**

The first lung transplant was performed by Hardy, in 1964.

Patient died of renal failure after 3 weeks

The lung was donated after circulatory death (DCD)

Early function of the lung was excellent.





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The New England Journal of Medicine

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Volume 306

MARCH 11, 1982

Number 10

HEART-LUNG TRANSPLANTATION

Successful Therapy for Patients with Pulmonary Vascular Disease

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MARGARET E. BILLINGHAM, M.B., PHILIP E. OVER, M.D., PH.D., EDWARD B. STINSON, M.D.,
AND NORMAN E. SHUMWAY, M.D., PH.D.

Abstract We report our initial experience with three patients who received heart-lung transplants. The primary immunosuppressive agent used was cyclosporin A, although conventional drugs were also administered.

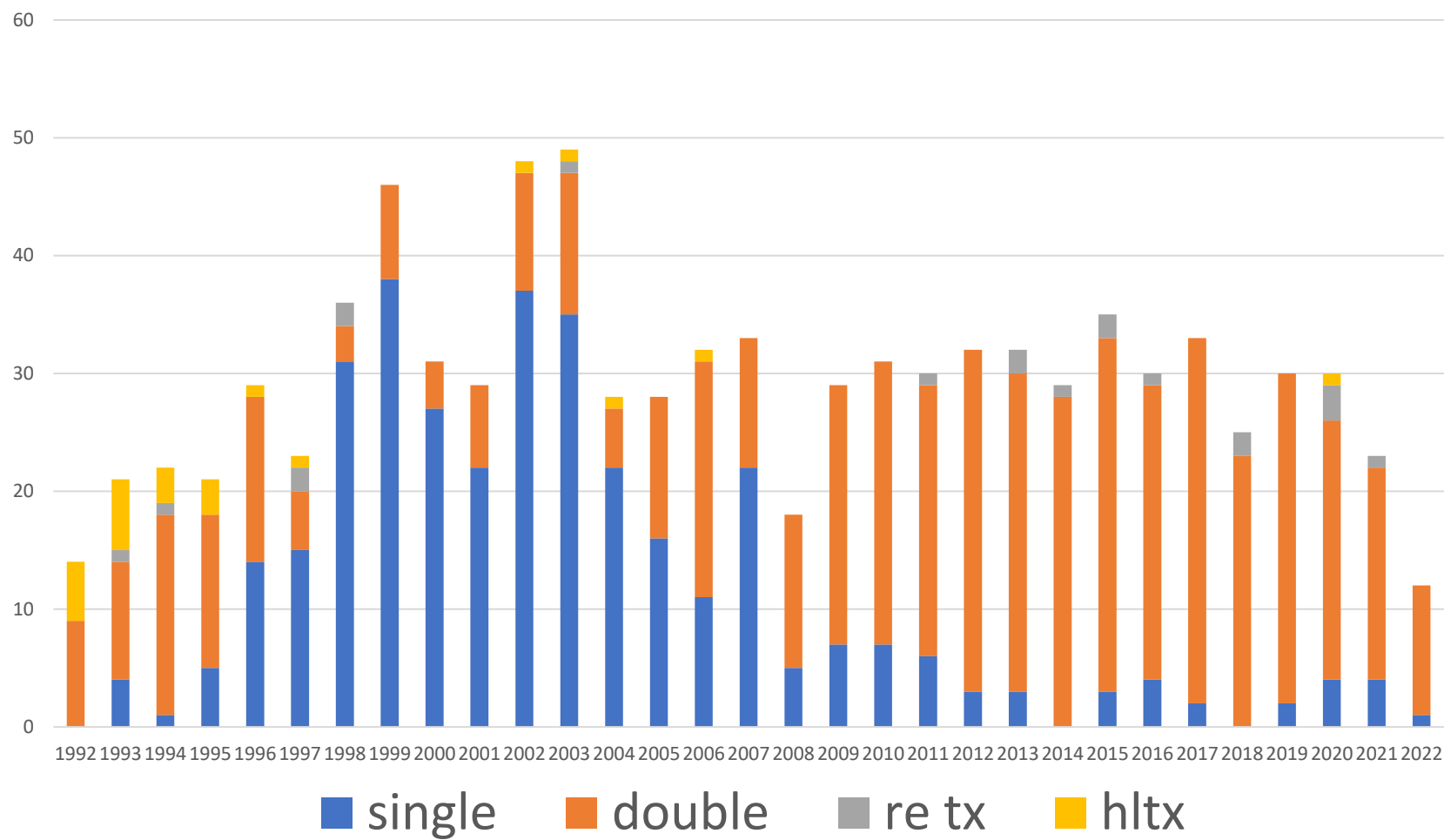
In the first patient, a 45-year-old woman with primary pulmonary hypertension, acute rejection of the transplant was diagnosed 10 and 25 days after surgery but was treated successfully; this patient still had normal exercise tolerance 10 months later. The second patient, a 30-year-old man, underwent transplantation for Eisenmenger's syndrome due to atrial and ventricular septal defects. His graft was not rejected, and his condition was markedly improved

eight months after surgery. The third patient, a 29-year-old woman with transposition of the great vessels and associated defects, died four days postoperatively of renal, hepatic, and pulmonary complications.

We attribute our success to experience with heart-lung transplantation in primates, to the use of cyclosporin A, and to the anatomic and physiologic advantages of combined heart-lung replacement. We hope that such transplants may ultimately provide an improved outlook for selected terminally ill patients with pulmonary vascular disease and certain other intractable cardiopulmonary disorders. (N Eng J Med. 1982; 306:557-64.)

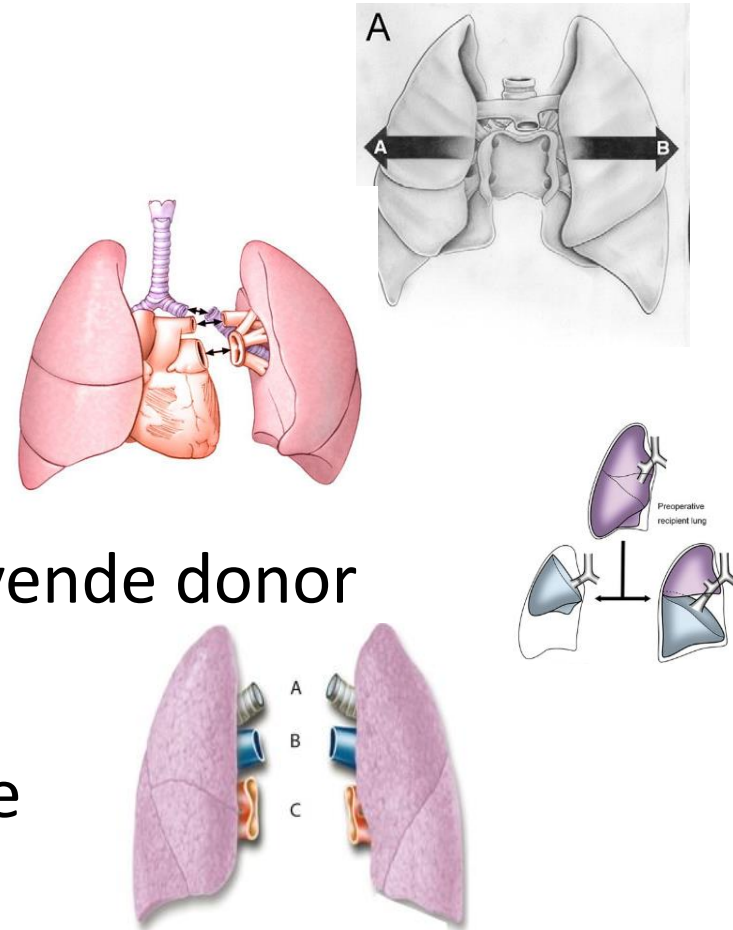


Lungetransplantation
Danmark



Typer af Lunge transplantationer udført i DK

- Maj 28, 1992: Én bloc dobbelt lunge
- Juni 19, 1993: Enkelt lunge
- September 21, 1993: Lunge lap fra levende donor
- Juli 23, 1998: Sekventiel dobbelt lunge



Bronkial Arterie Revaskularisering (BAR)

CURRENT REVIEWS

Revascularization of the Bronchial Arteries in Lung Transplantation: An Overview

Martin A. Nørgaard, MD, Peter S. Olsen, MD, PhD, Ulrik G. Svendsen, MD, PhD, and Gösta Pettersson, MD, PhD

Departments of Cardiovascular Surgery and Cardiology, Copenhagen University Hospital, Copenhagen, Denmark

Right lateral branch

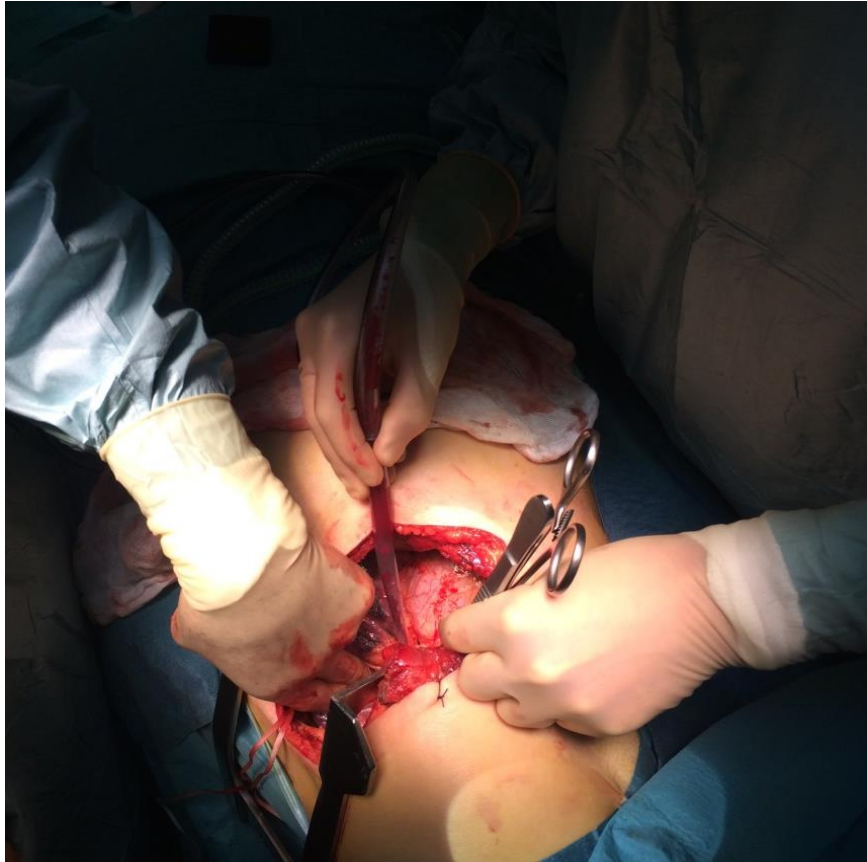
Right medial branch

Broncho-esophageal artery

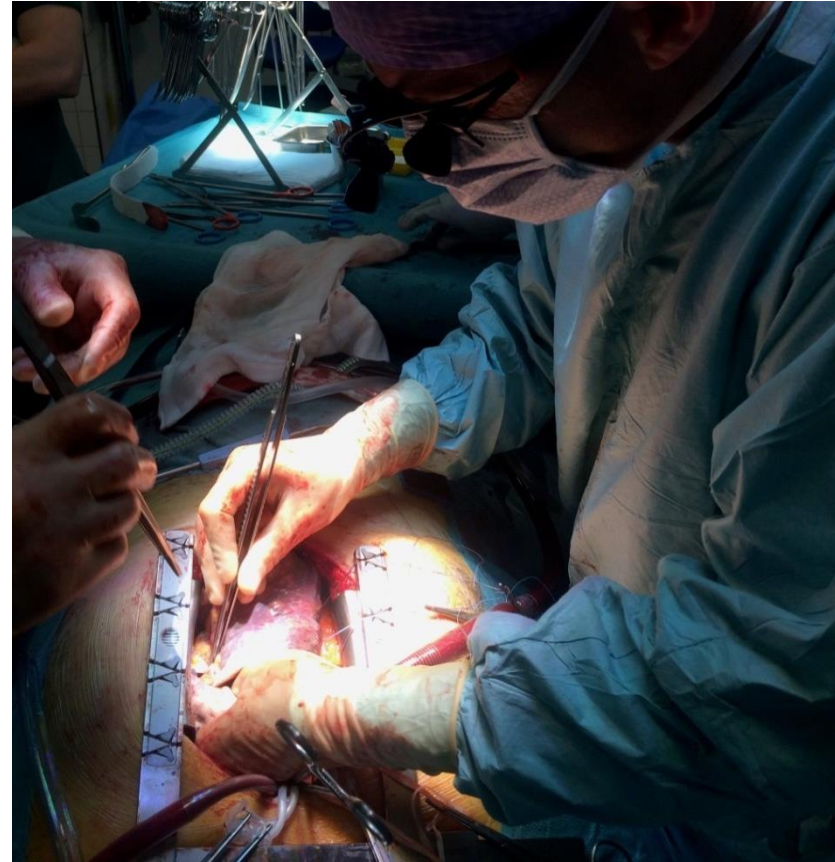
Left lateral branch

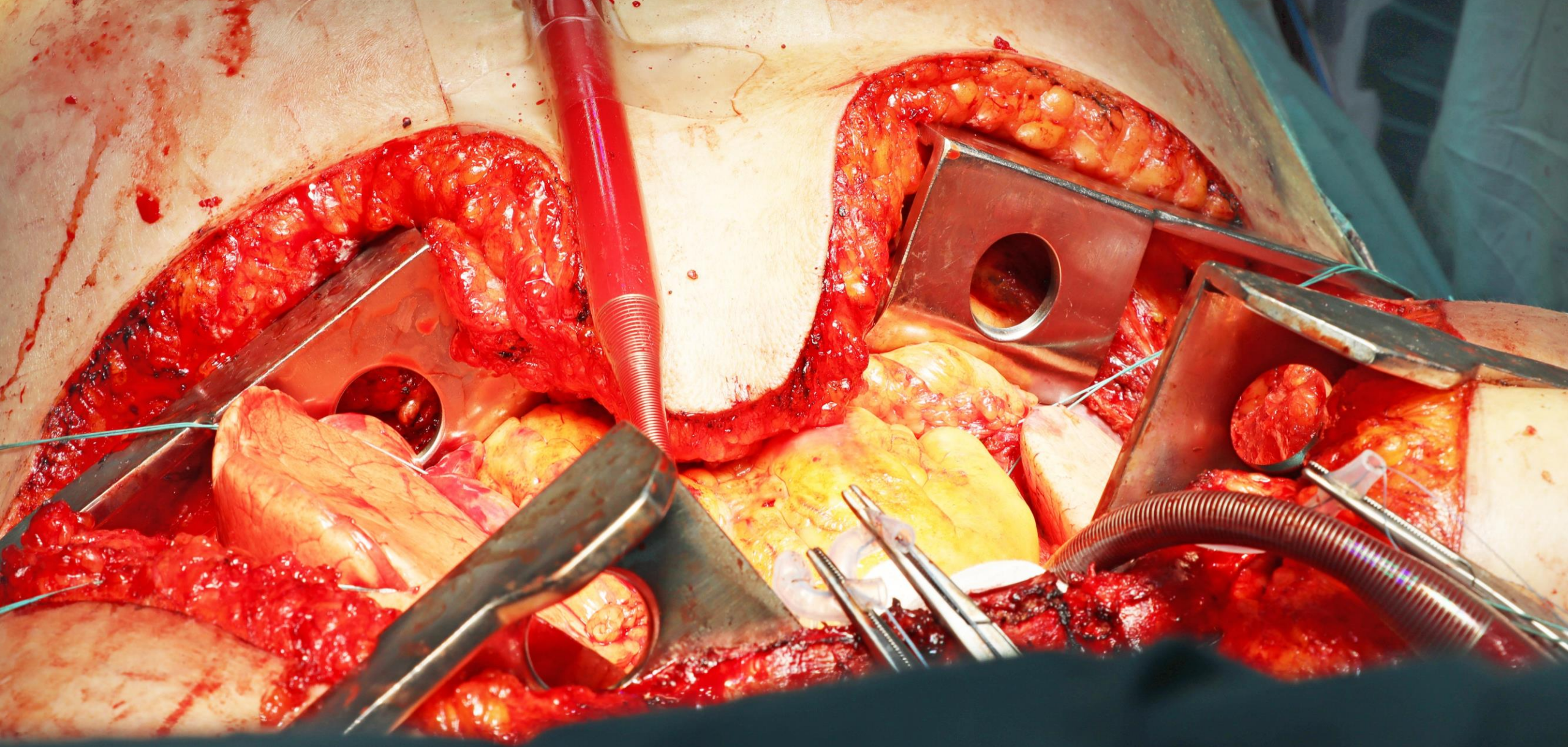
Left medial branch

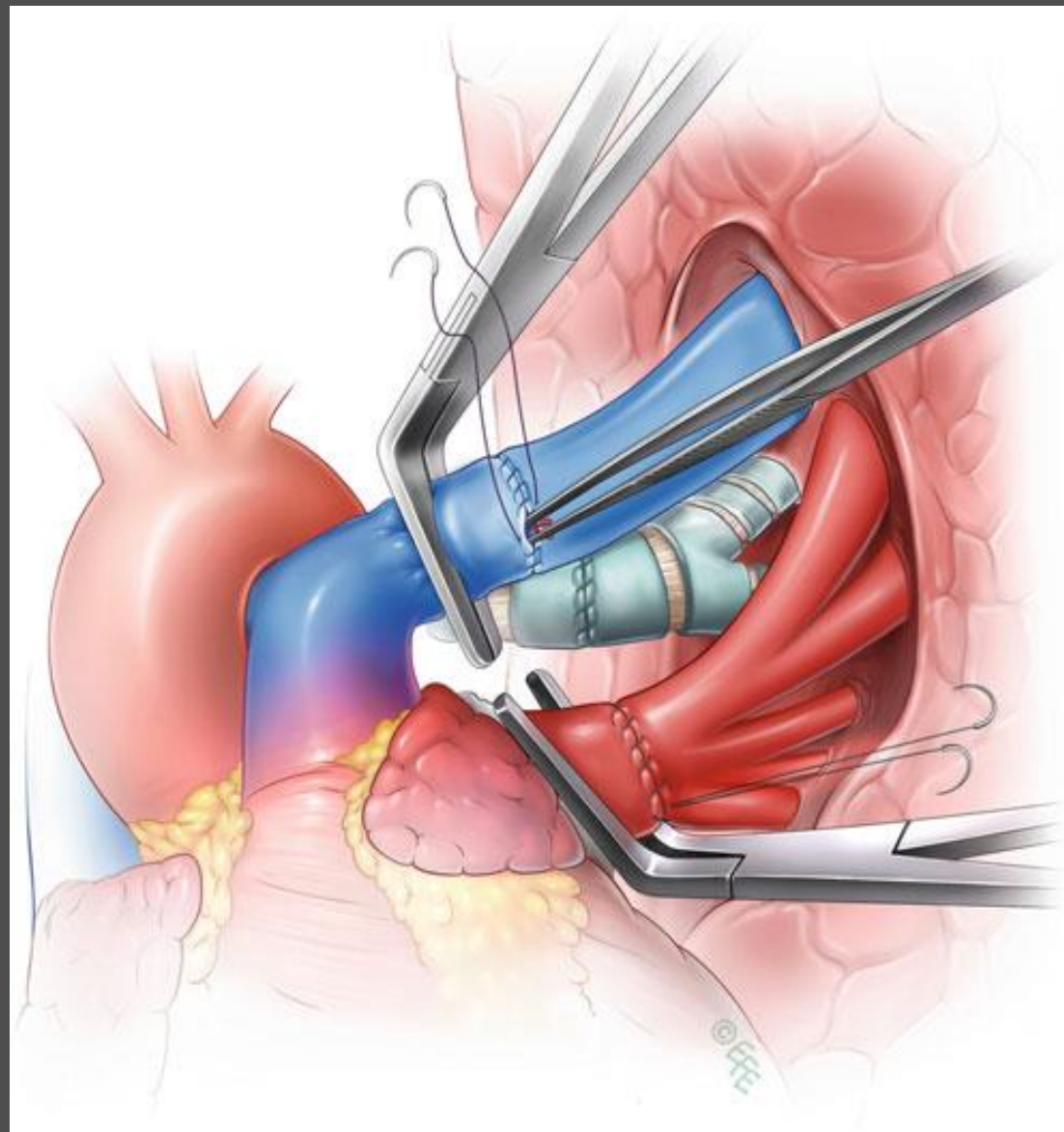
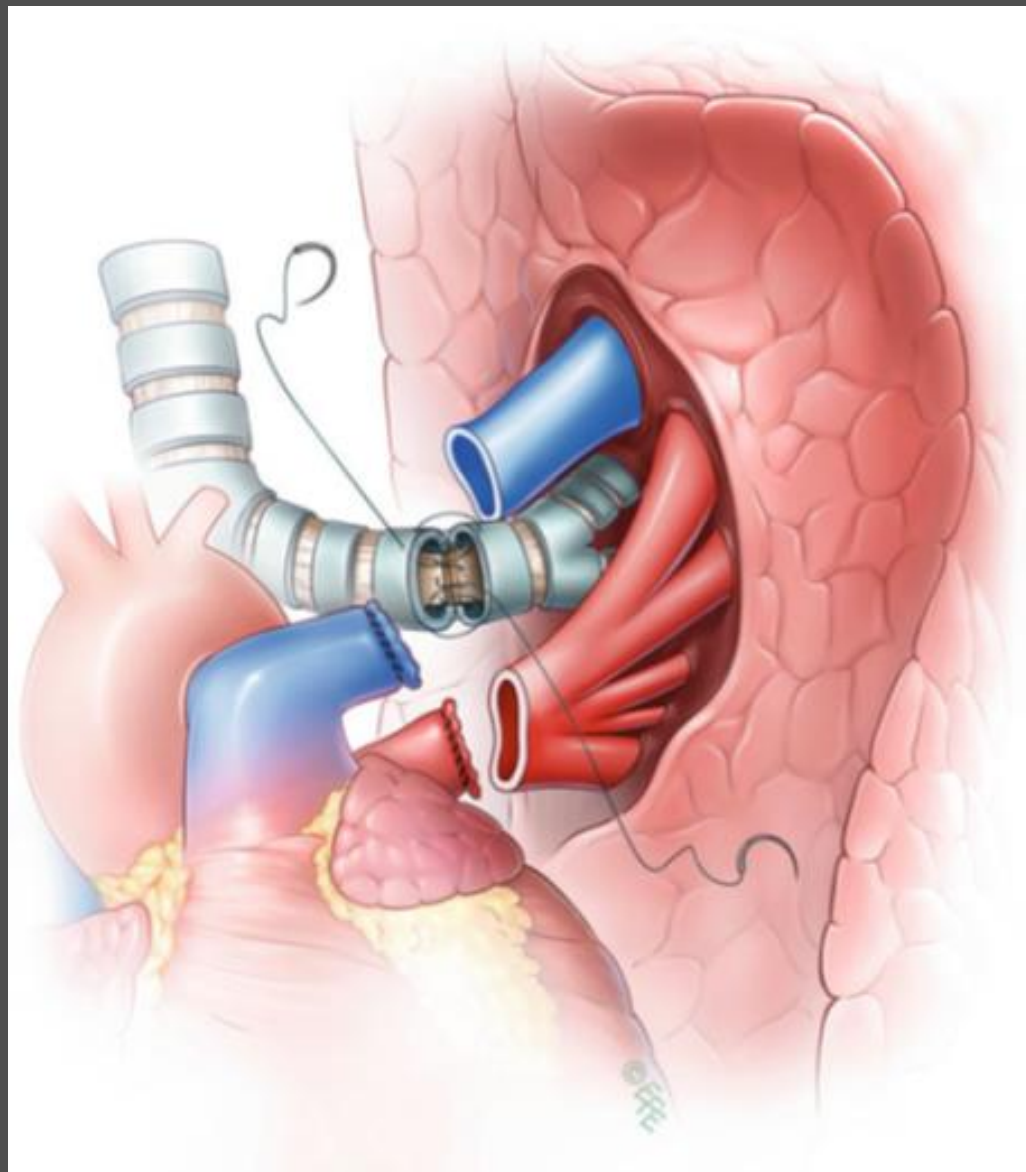
Lateral thorakotomi



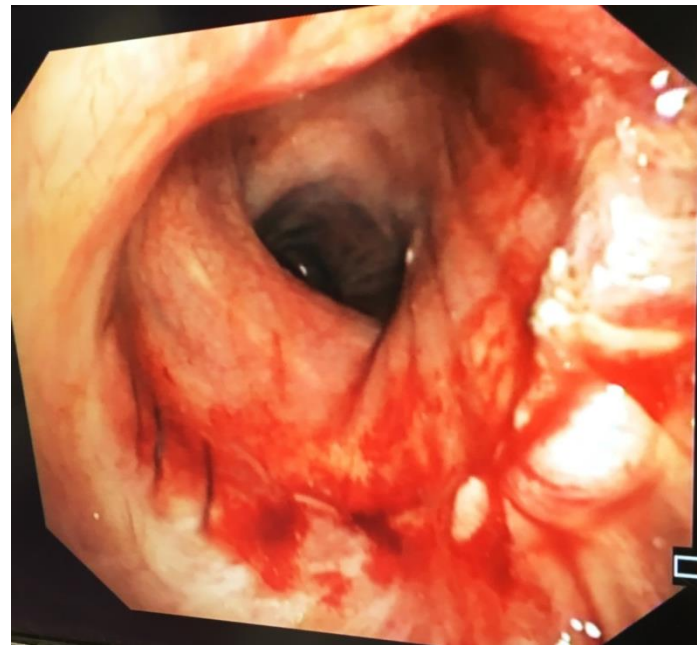
Median sternotomi



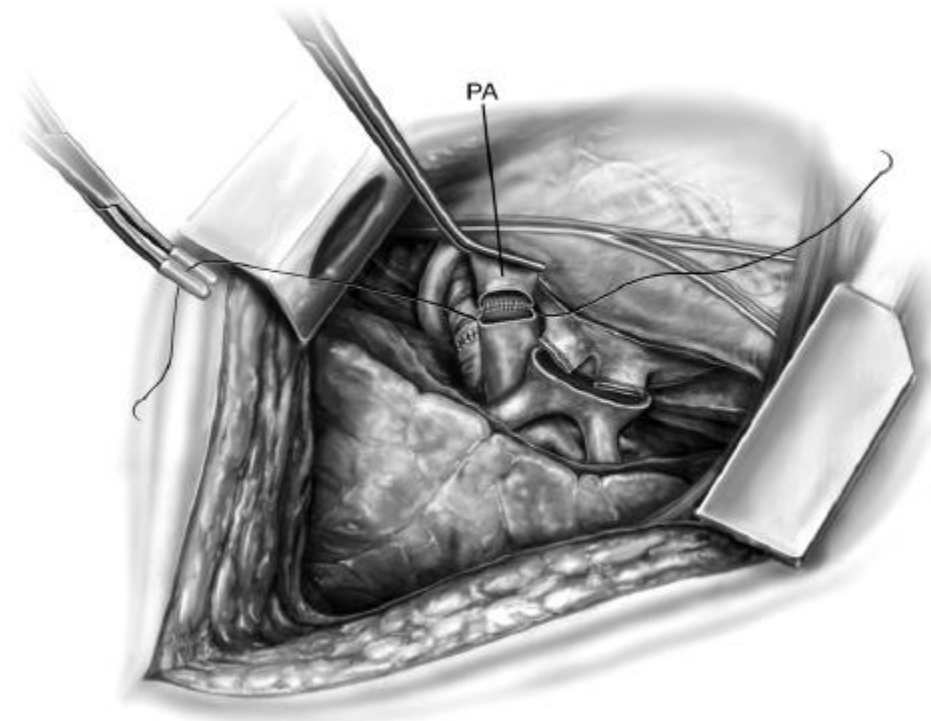


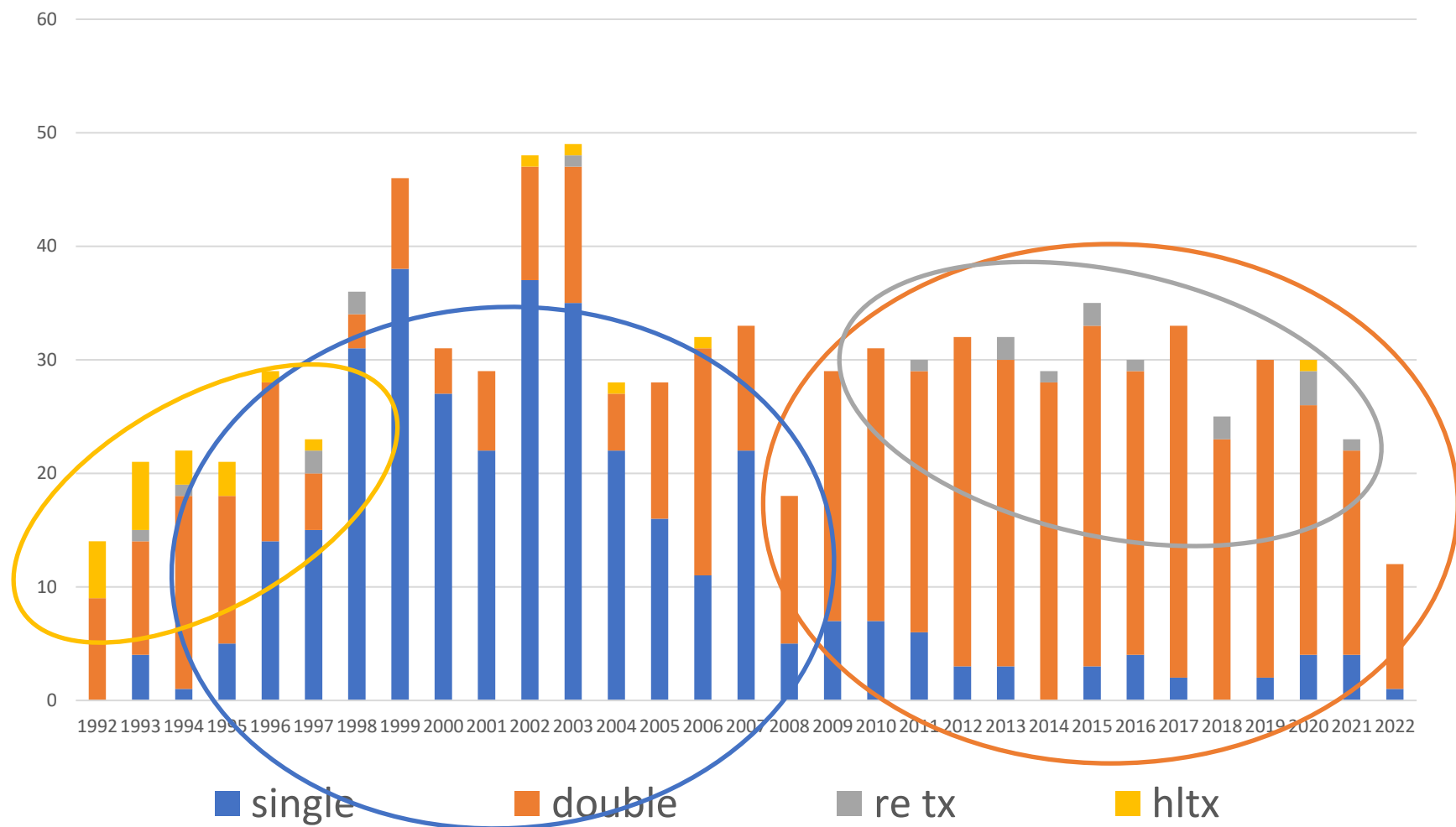


Anastomose af bronchus



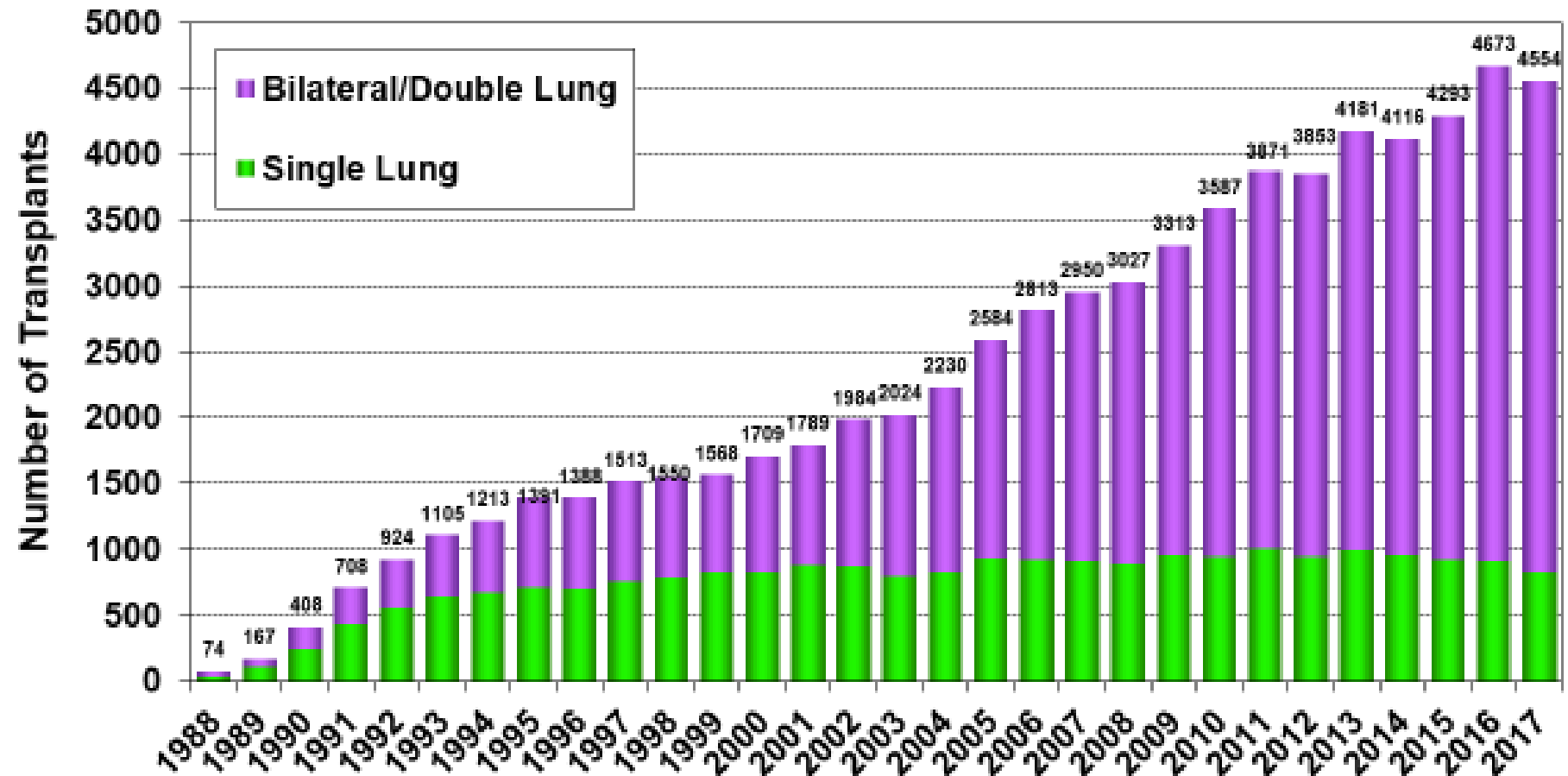
Anastomose af pulmonal arterien





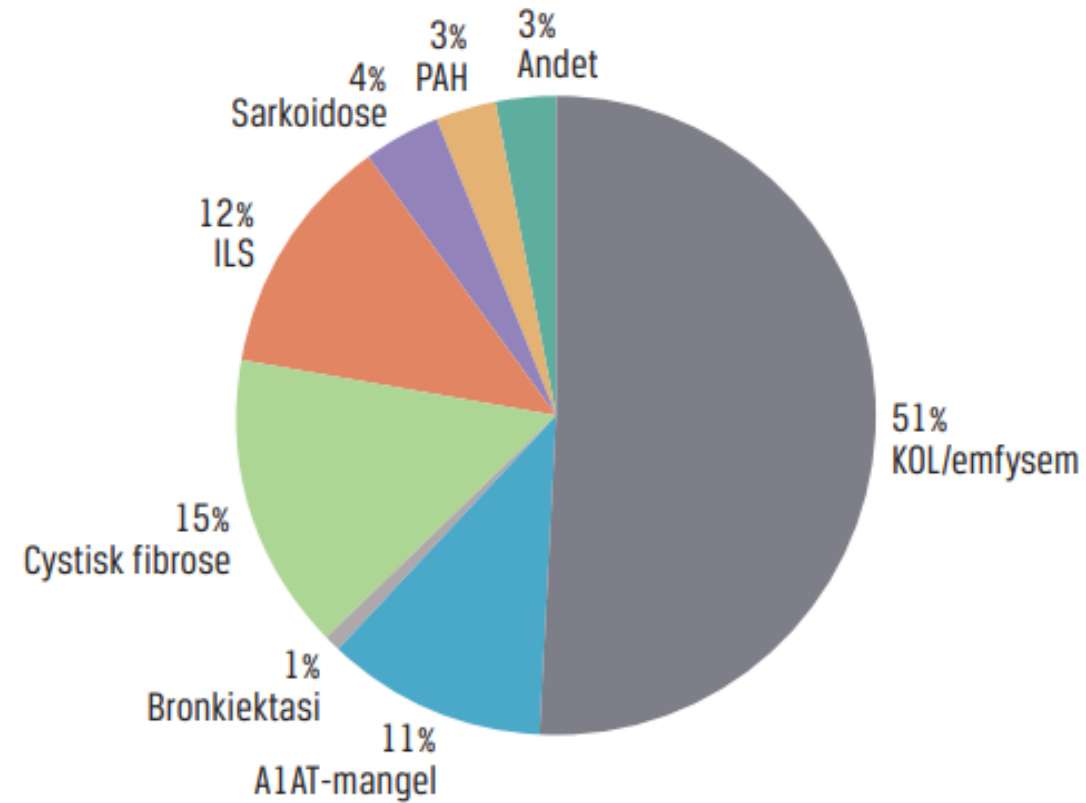
Adult and Pediatric Lung Transplants

Number of Transplants by Year and Procedure Type



NOTE: This figure includes only the lung transplants that are reported to the ISHLT Transplant Registry. As such, this should not be construed as representing changes in the number of lung transplants performed worldwide.

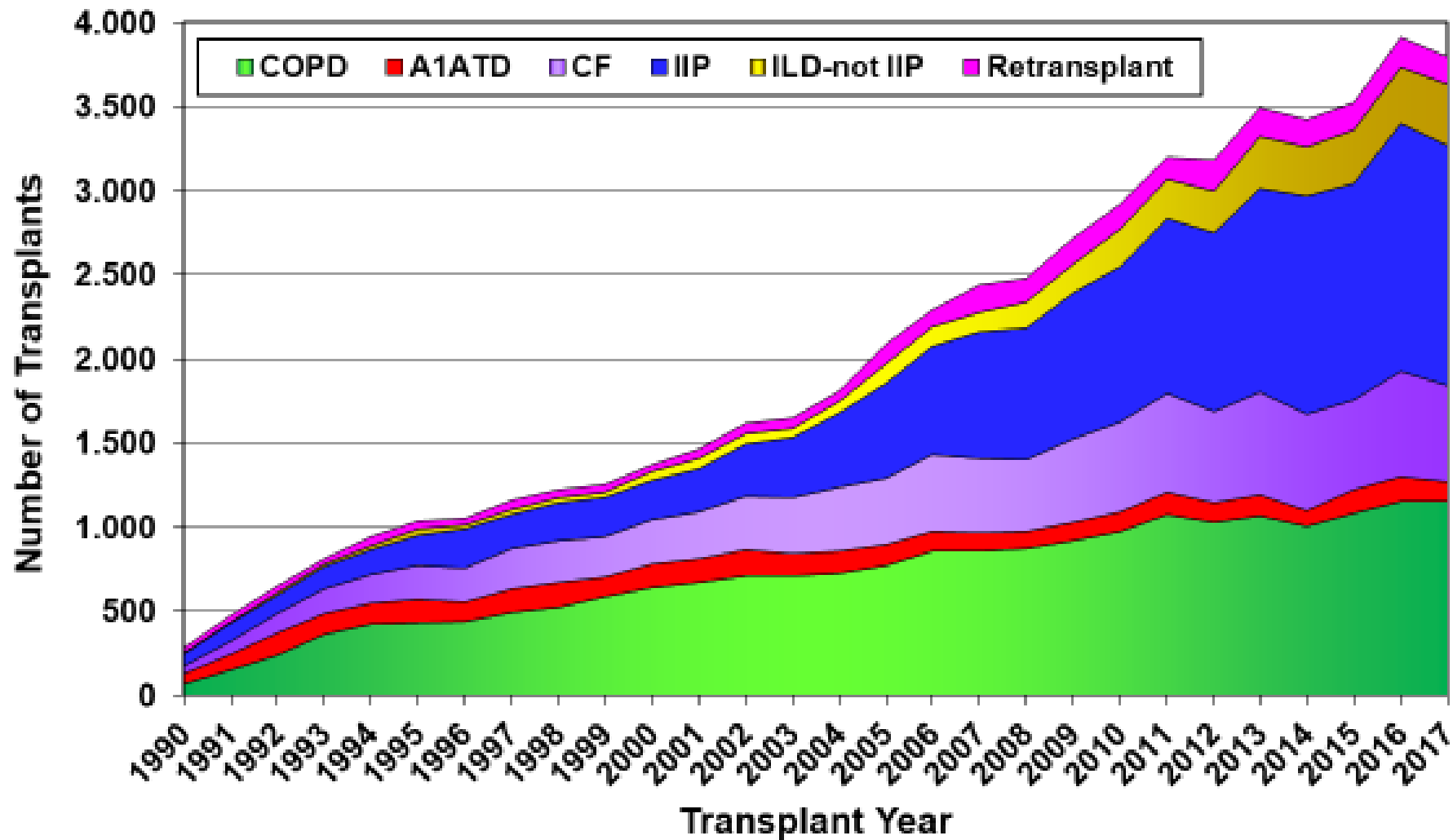
Hyppigheder af typer lungesygdom forud for lungetransplantation.



A1AT = alfa-1-antitrypsin; ILS = interstitiel lungesygdom;
KOL = kronisk obstruktiv lungesygdom; PAH = pulmonal arteriel
hypertension.

Adult Lung Transplants

Major Diagnoses by Year (Number)



LUNGER



Det er overraskende positive erfaringer, overlægerne Michael Perch og Tacjana Pressler har gjort sig i mødet med patienter med cystisk fibrose, siden en ny behandling kom til landet for et år siden. Udover langt bedre livskvalitet ser man hos nogle forbedret lun

Læger måber over ny behandling af cystisk fibrose

For omkring et år siden kom en ny behandling til personer med cystisk fibrose til Danmark. Ifølge læger på Rigshospitalet er de erfaringer, man siden har gjort sig med behandlingen, hidtil uset positive.

Kristian Sjøgren | 01/10/2021

En kombination af lægemidlerne Kafrio og Kalydeco har fuldstændigt ændret livet for rigtig mange danskere med cystisk fibrose.

NYHEDER



Det er overraskende positive erfaringer, overlægerne Michael Perch og Tacjana Pressler fra Rigshospitalet har gjort sig i mødet med patienter med cystisk fibrose, siden en ny behandling kom til landet for et år siden. Udover langt bedre livskvalitet ser man hos nogle forbedret lungfunktion på op til 40 pct. Nu håber de at behandlingen også bliver godkendt til børn i alderen 6-11 år. Foto: Rigshospitalet

Overlæge glæder sig over mulig behandling til børn med cystisk fibrose

Kafrio kan være på trapperne som behandling til børn med cystisk fibrose. Udsigten er meget positiv, siger dansk overlæge.

Kristian Sjøgren | 24/11/2021

Del: [Facebook](#) [Twitter](#) [LinkedIn](#) [Email](#)

- 07/10 Fund af 69 genetiske varianter for stroke åbner op for nye behandlingsmuligheder
- 07/10 Overlægeforeningen er bekymret for presset på radiologien
- 07/10 Chef læges ønske til politikerne: Behandlingsgaranti skal være fleksibel
- 07/10 Yngre læge i psykiatrien: Jeg gider ikke arbejde på den måde
- 07/10 Praktiserende læge: Politikerne skal finde løsninger til ældre

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BRIDGE - Translational Excellence Programme | Ansøgningsfrist: 28/02/2022

Konferencer

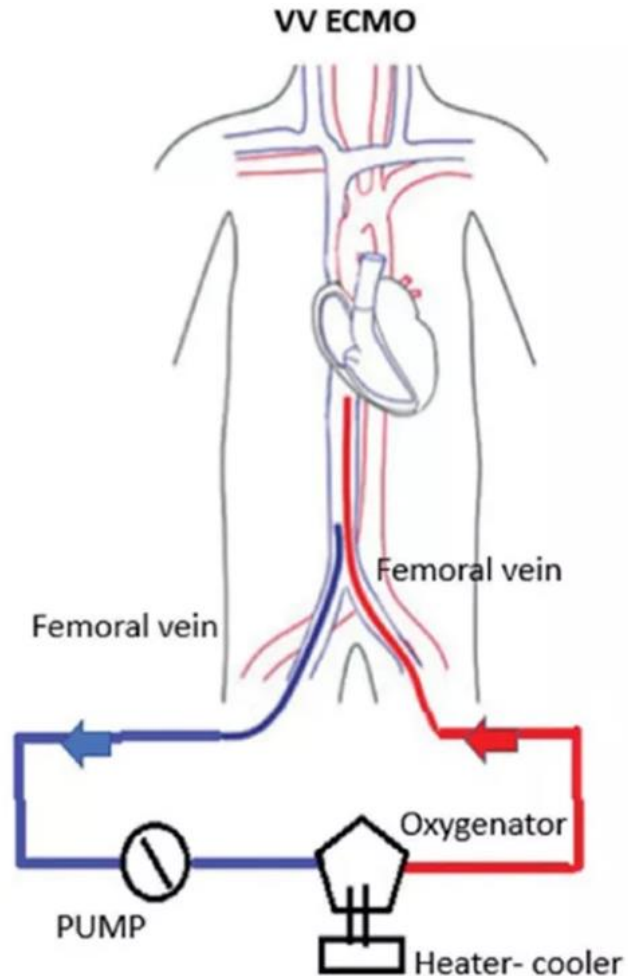
Der er opstået en fejl, feedet er sandsynligvis nede. Prøv igen senere.



For anmodning om en tilgængelig PDF send venligst en e-mail til production@Med.com

Medicinrådets protokol for vurdering af human alfa-1-antitrypsin til behandling af alvorlig alfa-1-antitrypsinmangel

Fra lungesvigt til transplantation



Toronto protokol

- Transplantation af recipienter med pre-formerede Lymfocytotoksiske antistoffer
- Transplantation mod positiv crossmatch
- Ved anvendelse af PLEX

American Journal of Transplantation 2015; 15: 417–426
Wiley Periodicals Inc.

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doi: 10.1111/ajt.13076

Survival in Sensitized Lung Transplant Recipients With Perioperative Desensitization

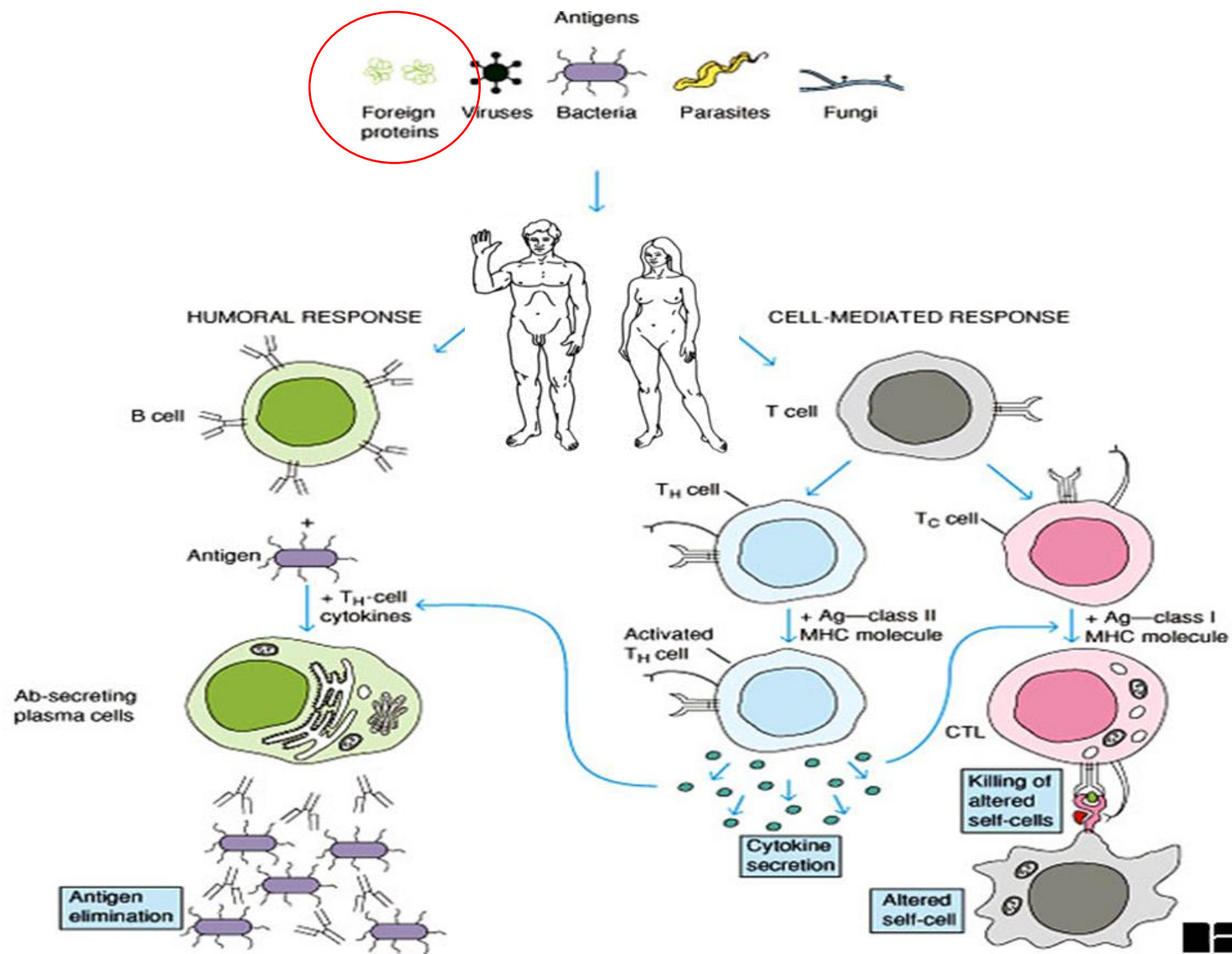
K. J. Tinckam², S. Keshavjee¹, C. Chaparro¹,
D. Barth^{2,3}, S. Azad¹, M. Binnie¹, C. W. Chow¹,
M. de Perrot¹, A. F. Pierre¹, T. K. Waddell¹,
K. Yasufuku¹, M. Cypel¹ and L. G. Singer^{1,*}

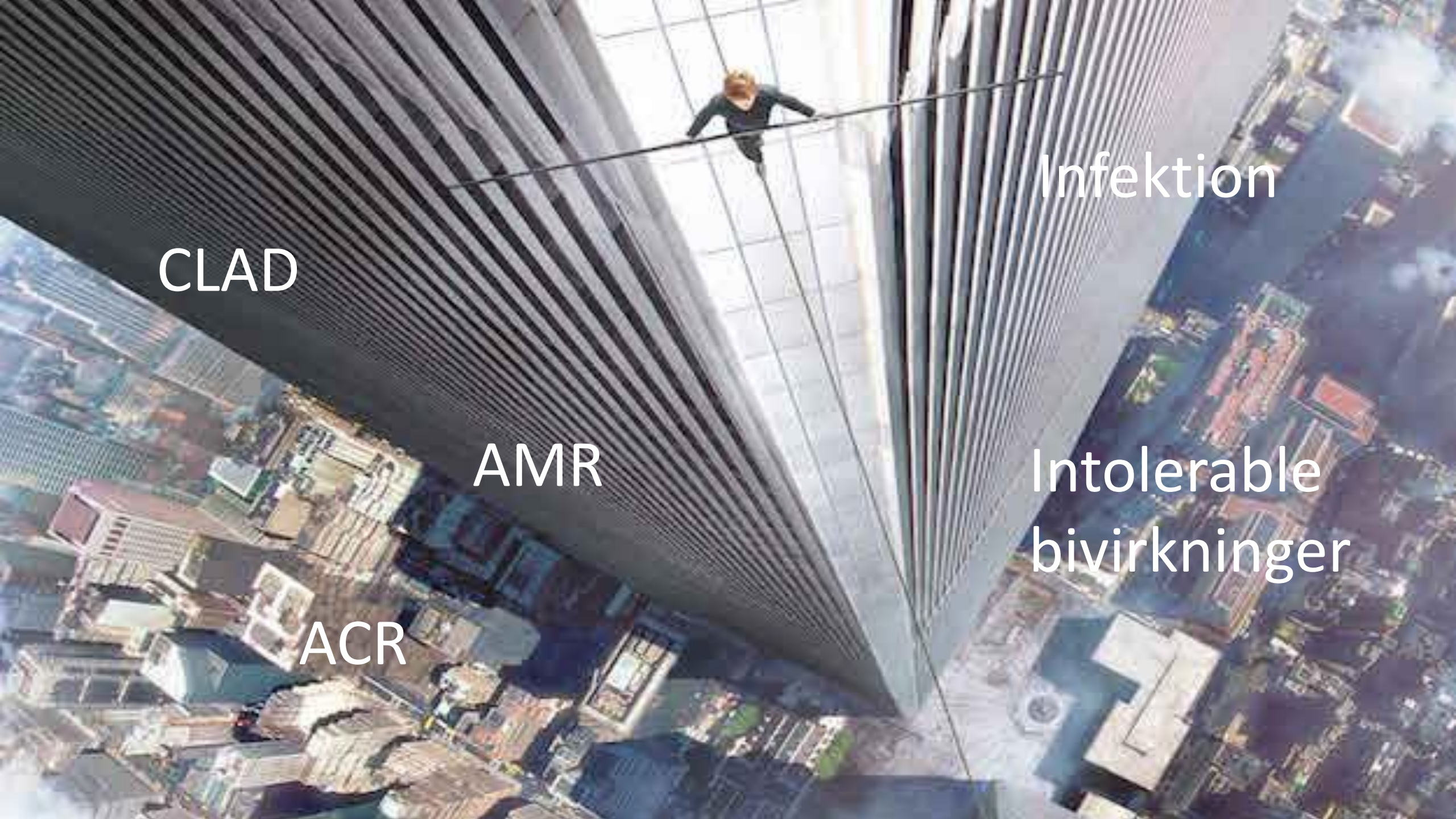
¹Toronto Lung Transplant Program, University Health Network, University of Toronto, Toronto, ON, Canada
²Laboratory Medicine Program and Department of Medicine, University Health Network, University of Toronto, Toronto, ON, Canada
³Therapeutic Apheresis Program, University Health Network, University of Toronto, Toronto, ON, Canada

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specific antibody; FCXM, flow crossmatch; FEV₁, forced expiratory volume (1 s); HLA, human leukocyte antigen; IgG, immunoglobulin G; IVIG, intravenous immune globulin; MPA, mycophenolic acid; MFI, mean fluorescence intensity; PCR, polymerase chain reaction; PGD, primary graft dysfunction; PLEX, plasma exchange; PRA, panel reactive antibody; SSO, sequence-specific oligonucleotide; SSP, sequence specific primer; UHN, University Health Network; XM, crossmatch

Received 24 January 2014, revised 19 August 2014 and
accepted for publication 06 September 2014





CLAD

Infektion

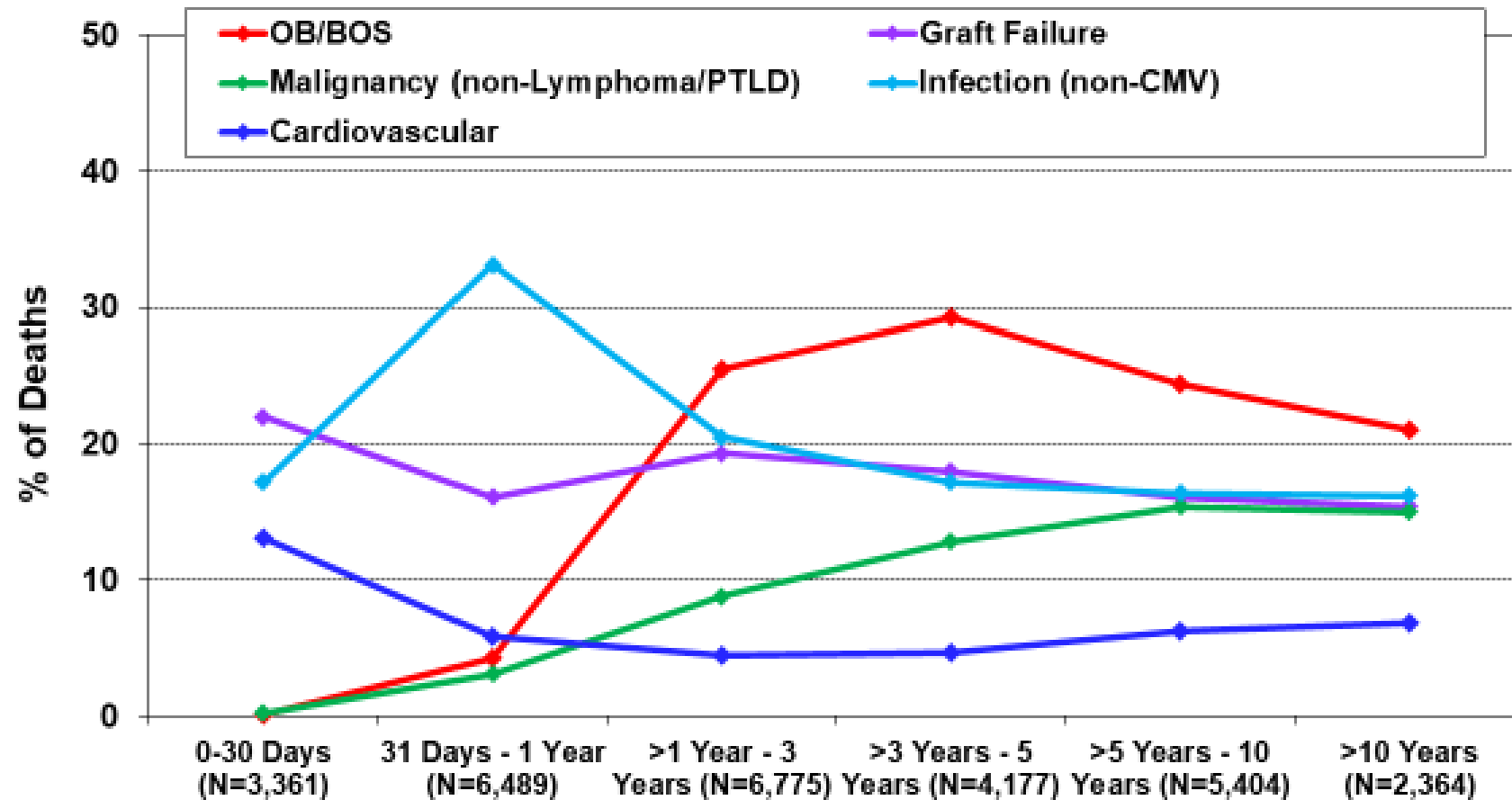
AMR

Intolerable
bivirkninger

ACR

Adult Lung Transplants

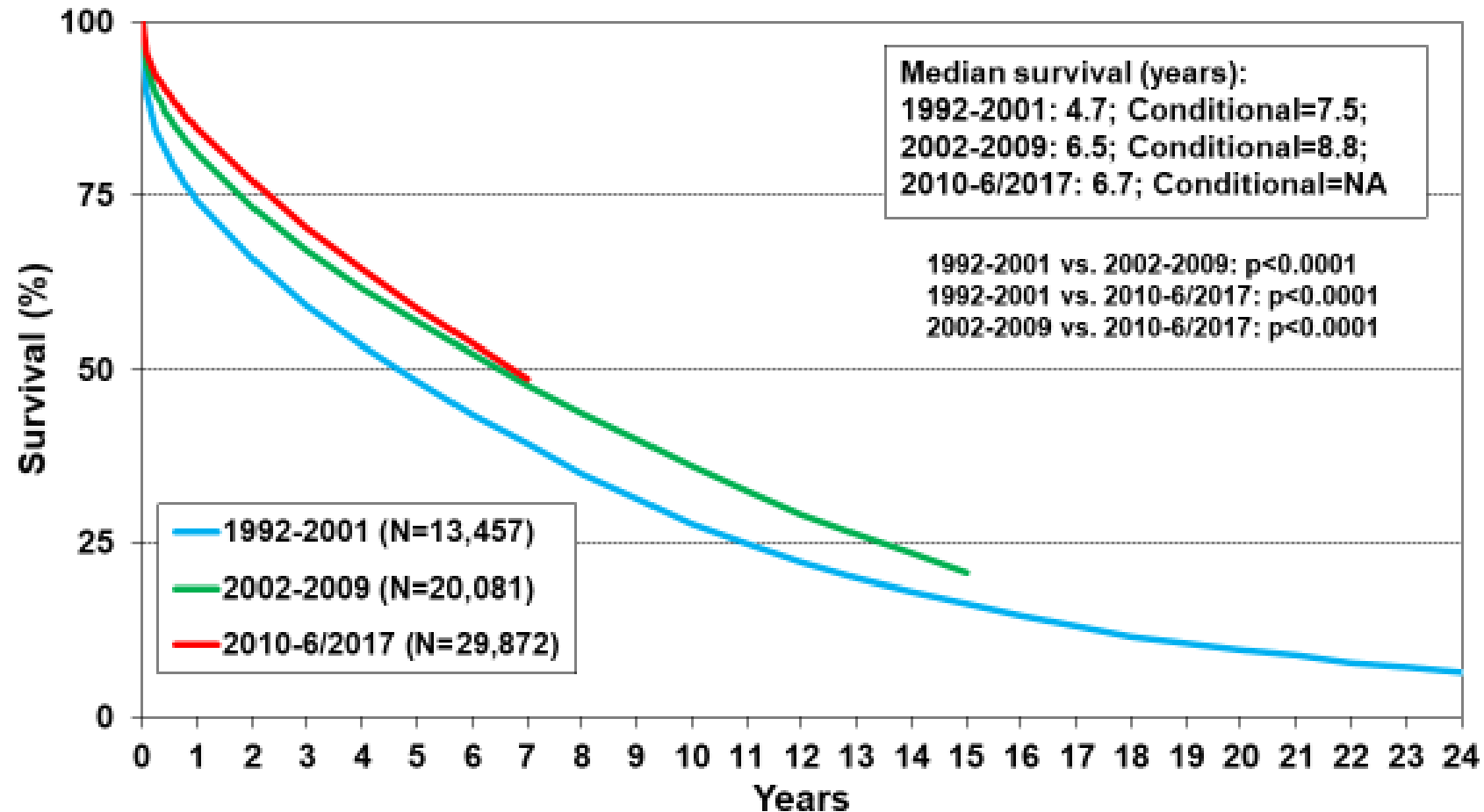
Relative Incidence of Leading Causes of Death (Deaths: January 1995 – June 2018)



Adult Lung Transplants

Kaplan-Meier Survival by Era

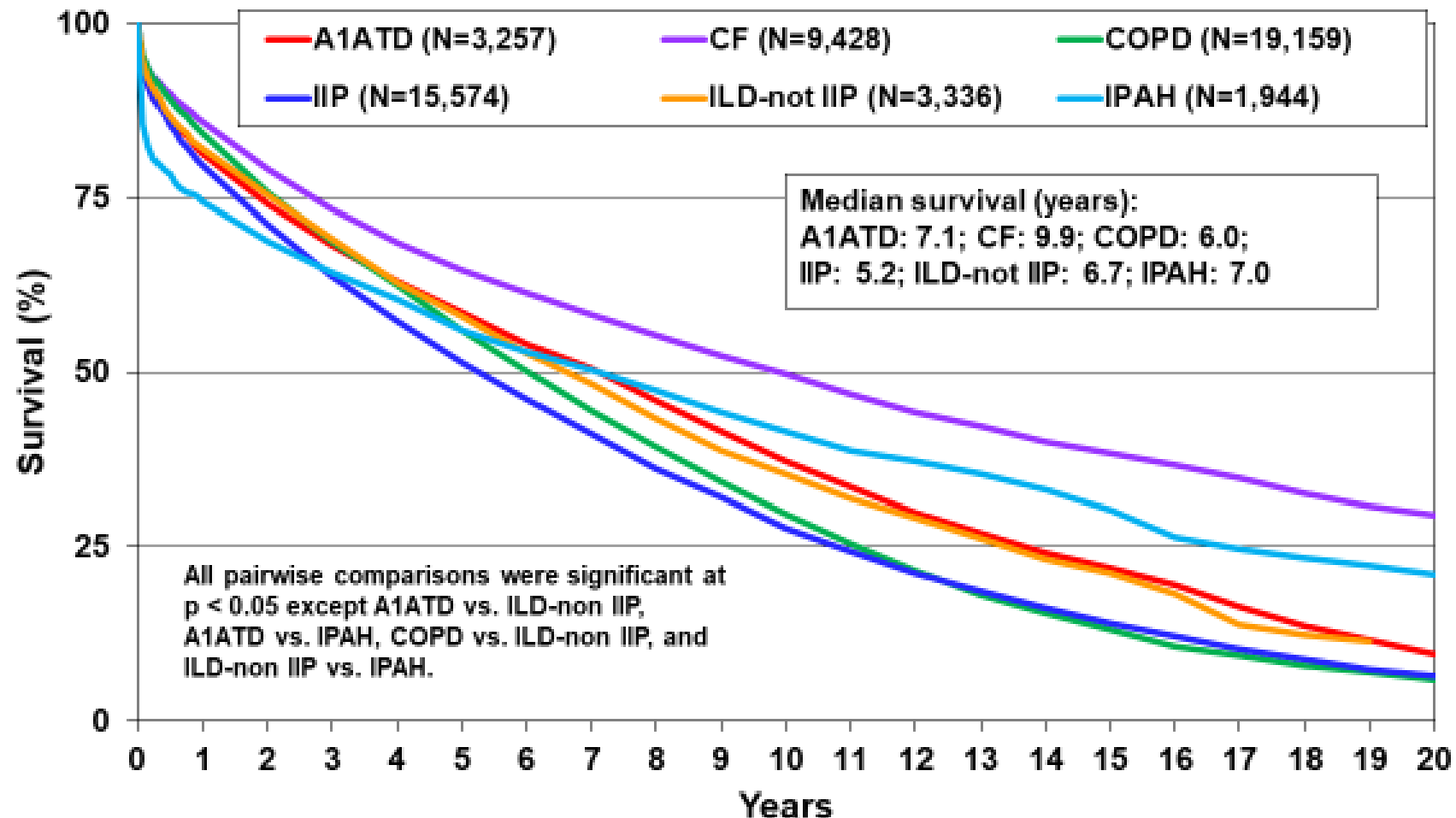
(Transplants: January 1992 – June 2017)



Adult Lung Transplants

Kaplan-Meier Survival by Major Diagnosis

(Transplants: January 1992 – June 2017)



Kristeligt Dagblad

INTERVIEW | 20.10.17 KL. 00:00

Overlæge: Vi har pligt til at bruge donor-organer med stor omhu

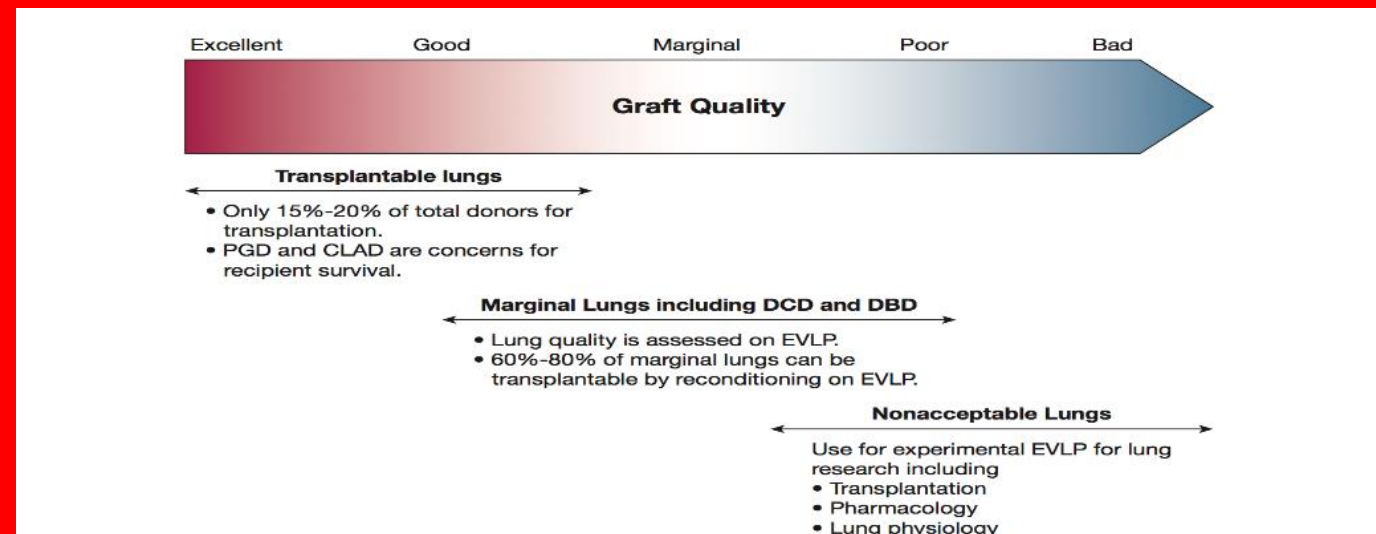


Generel knaphed på donor organer

Kun 15-25% of donor lunger accepters ved donations forløb af SOT

Laveste accept rate ved SOT

Undersøgelser har vist, at 40% af afviste donor lunger måske var egnede til transplantation





Opdateret 09.11.2012 Nyheder: Mediko-teknik

Maskine reparerer kassable donorlunger



Thoraxkirurg Bodil Brandt kigger til lungerne for at se, om processen forløber, som den skal. Foto: Casper Seldelin

Ny maskine på Rigshospitalet reparerer lungerne så effektivt, at de bliver bedre end andre donorlunger. Foreløbig går tre danskere rundt med renoverede lunger.

Ledige job

Fleres annoncer

Bornholms Hospital søger overlege/afdelingslæge i ortopedisk kirurgi - Bornholm

Bornholms Hospital søger overlege i kirurgi - Bornholm

Psykiater søges til privatklinik i Dubai - Dubai, De Forenede Arabiske Emirater

Sundhedsfaglig medarbejder - København

Vi søger en speciallæge i oto-rhino-laryngologi til vikariat - Nuuk

Klinikkleder for medicinsk gastroenterologi - Aabenraa

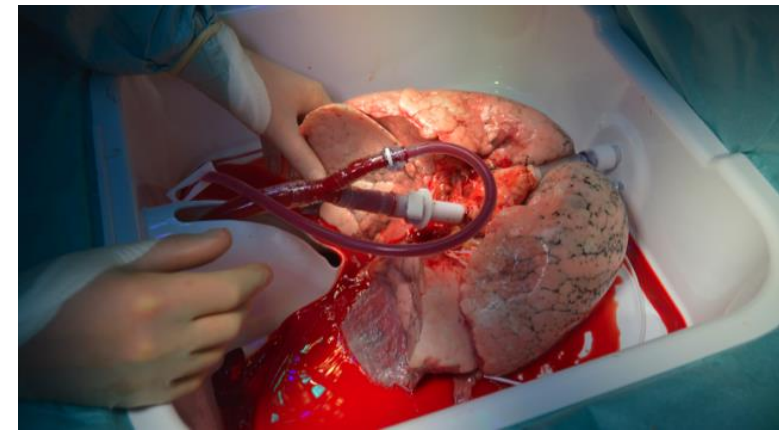
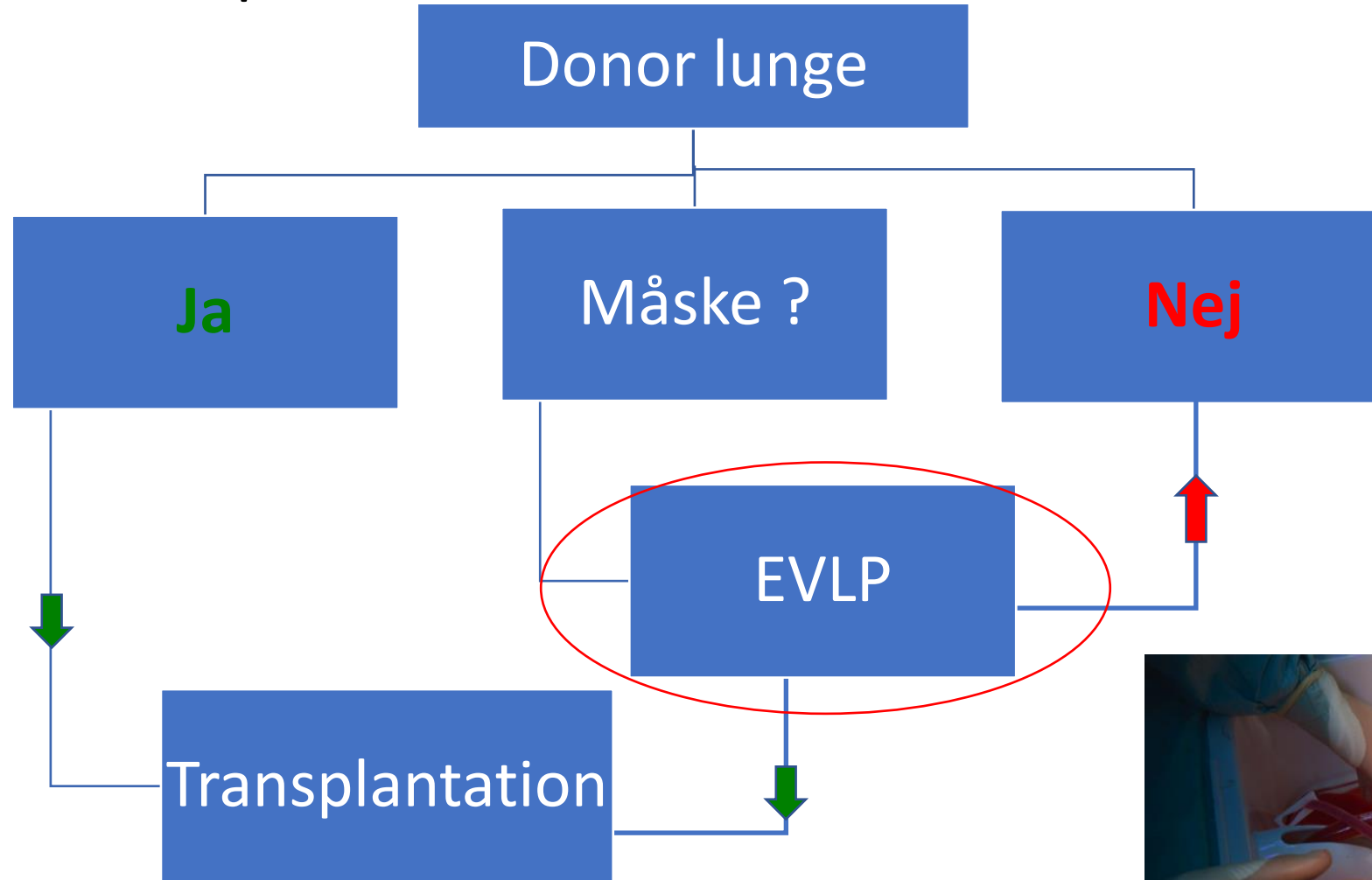
Overlege, Diagnostisk Radiologi, Billeddiagnostisk Afdeling - Viborg

Er du vår nye psykolog/psykologspesialist? - Harstad

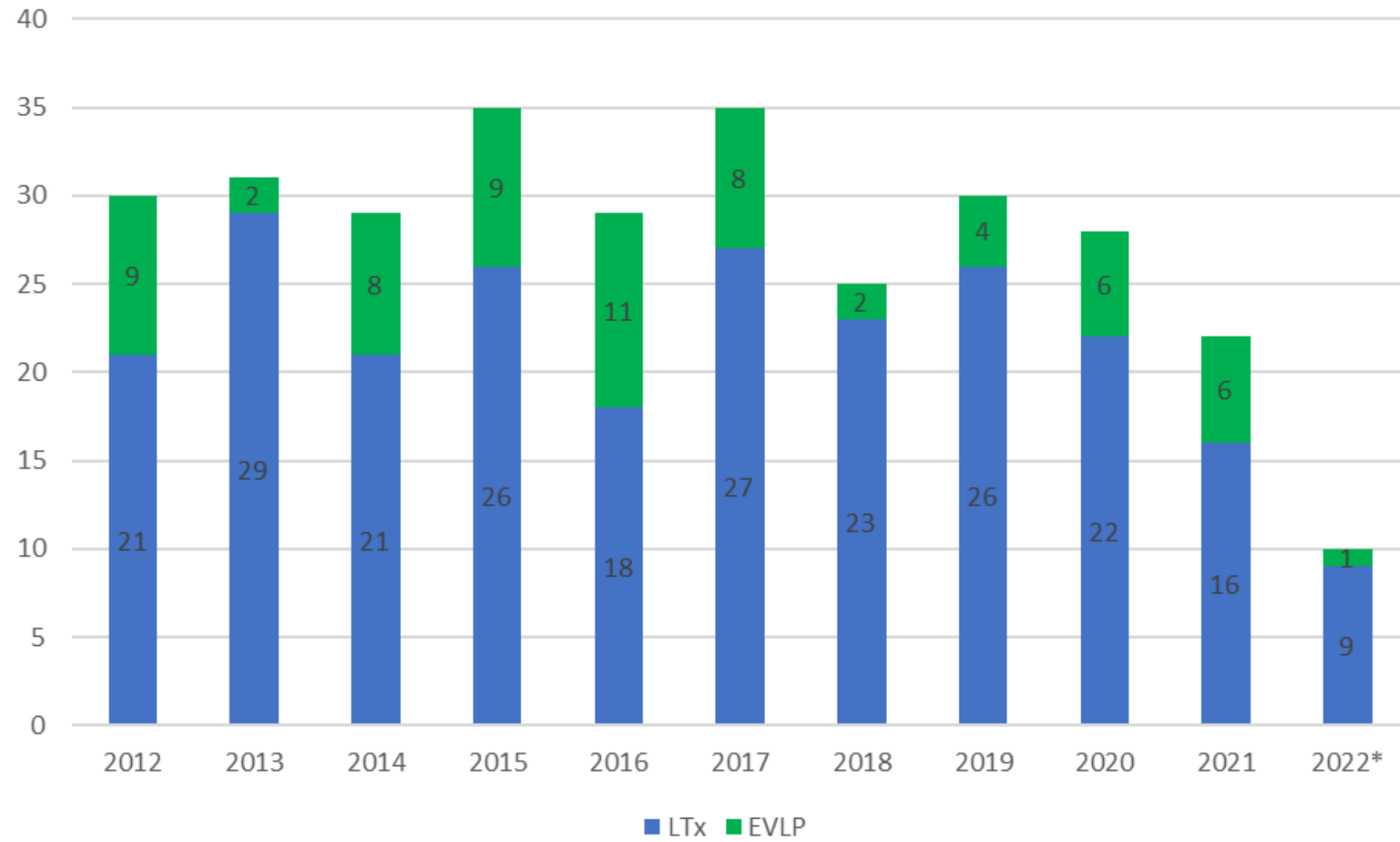
Danmarks største anæstesi- og intensivafdeling søger ledende overlege - Odense

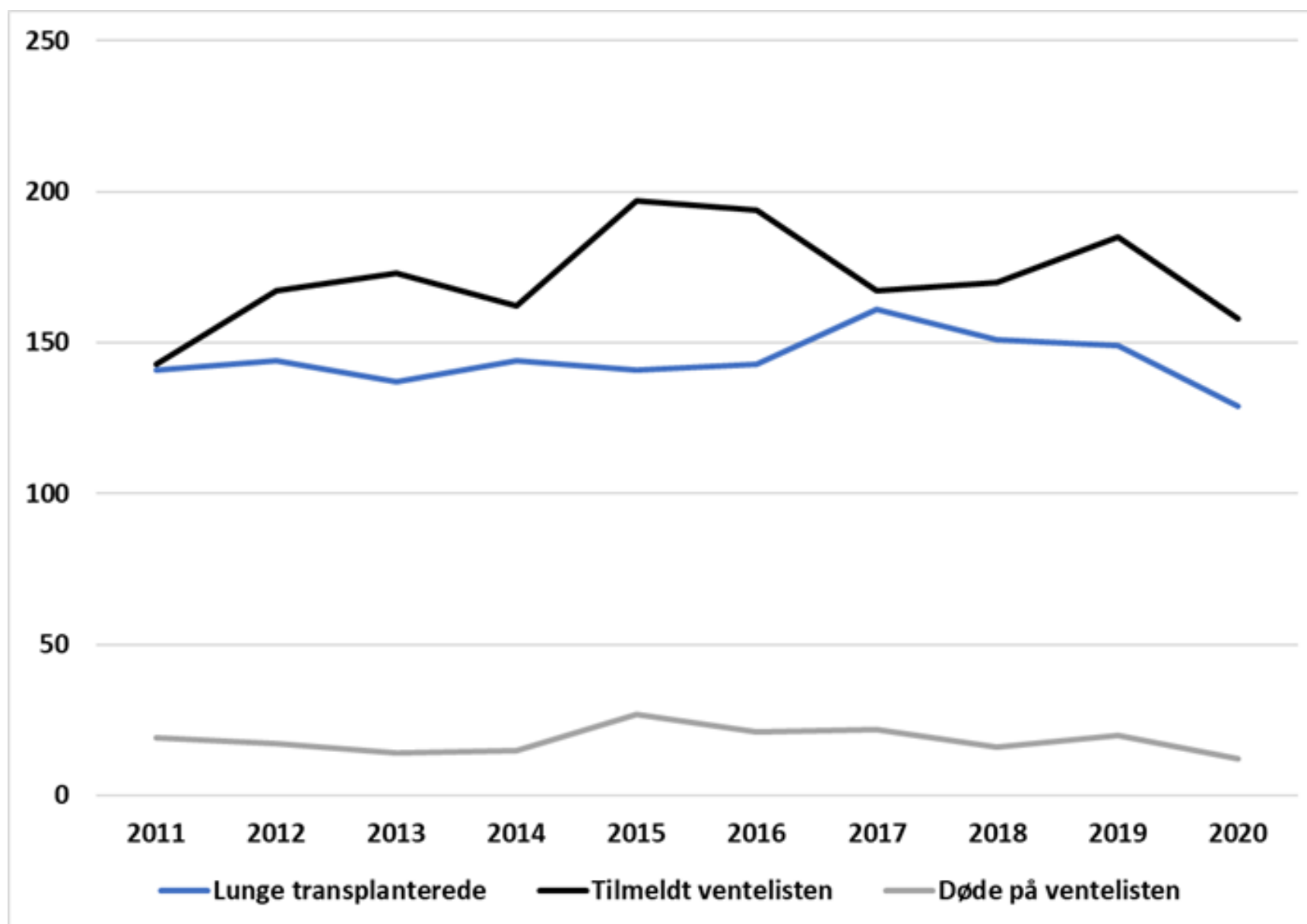
Afdelingslæge til Frømandskorpset/ Specialoperationsstyrkeelementet - Holbæk

EVLP konceptet

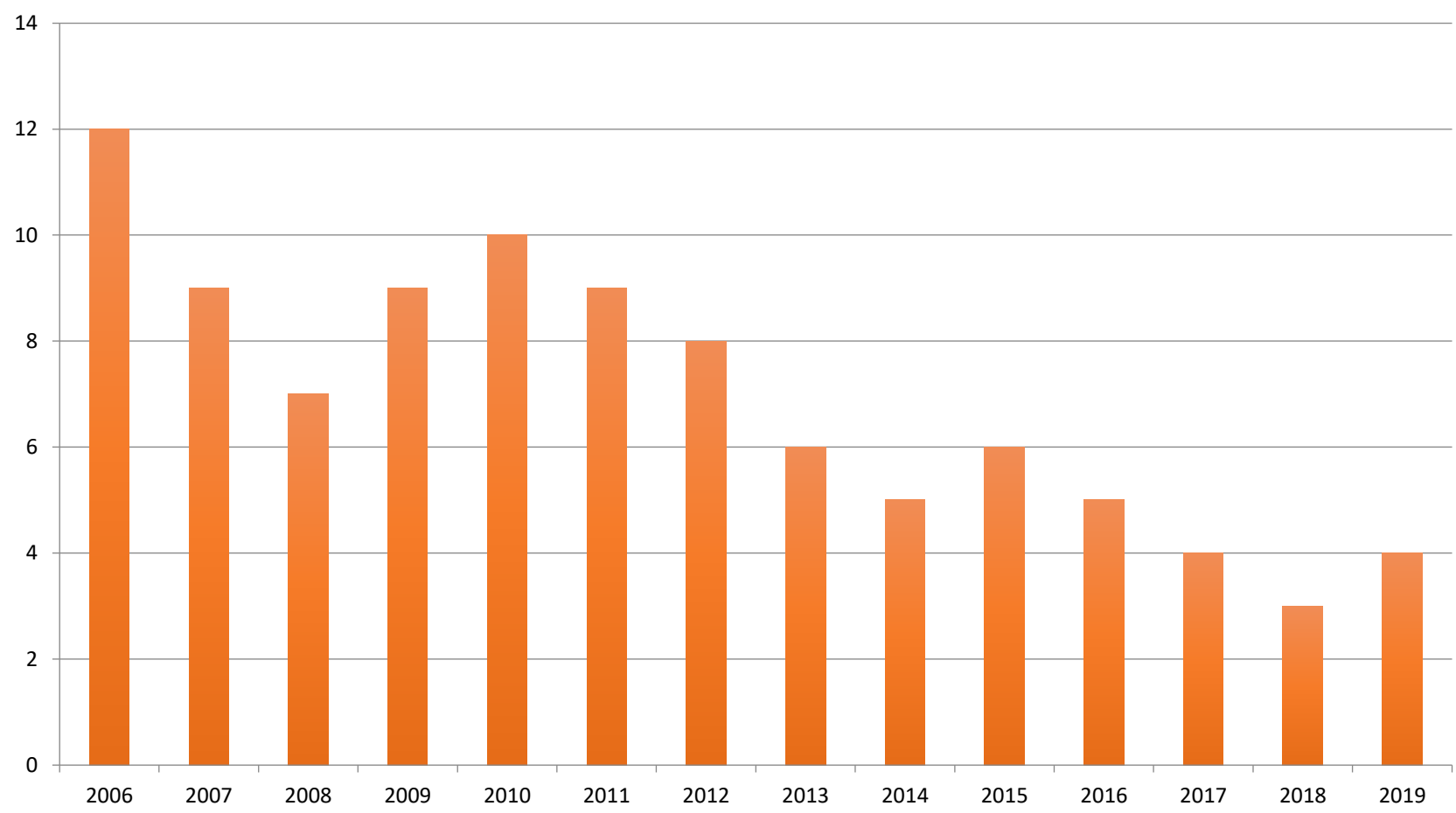


LTx Rigshospitalet





Death on the waiting list Copenhagen / year



Perspektiver for EVLP

- Øge antallet af donor organer
 - Non-Heart beating donor (DCD)
- Forskning:
 - Stam Celler
 - EVLP som behandling: cytokineabsorption; IL-10 ; anti-biotics/viral (CMV, Hepatitis)/fungal
 - Fjernelse af blood gruppe karakteristika
 - Evolution og forståelse af transplantations immunologi
- Logistic og transplantation



The Future of Transplantation is here...
The "Organ Repair Center"



Lung

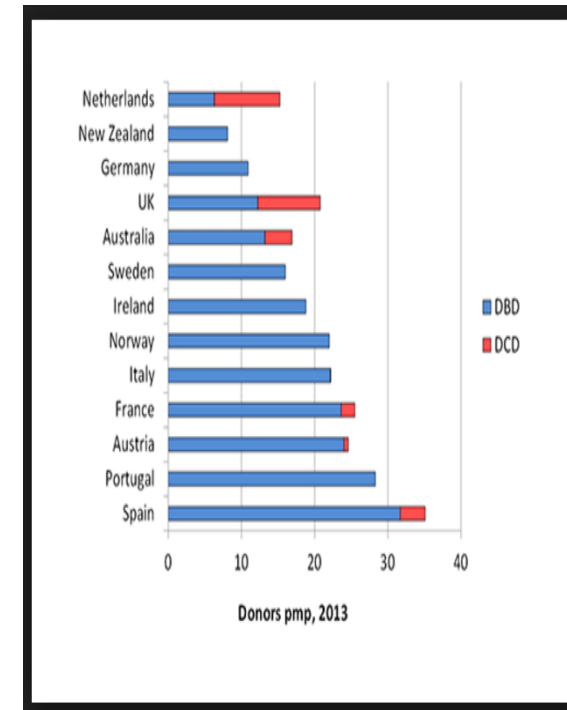
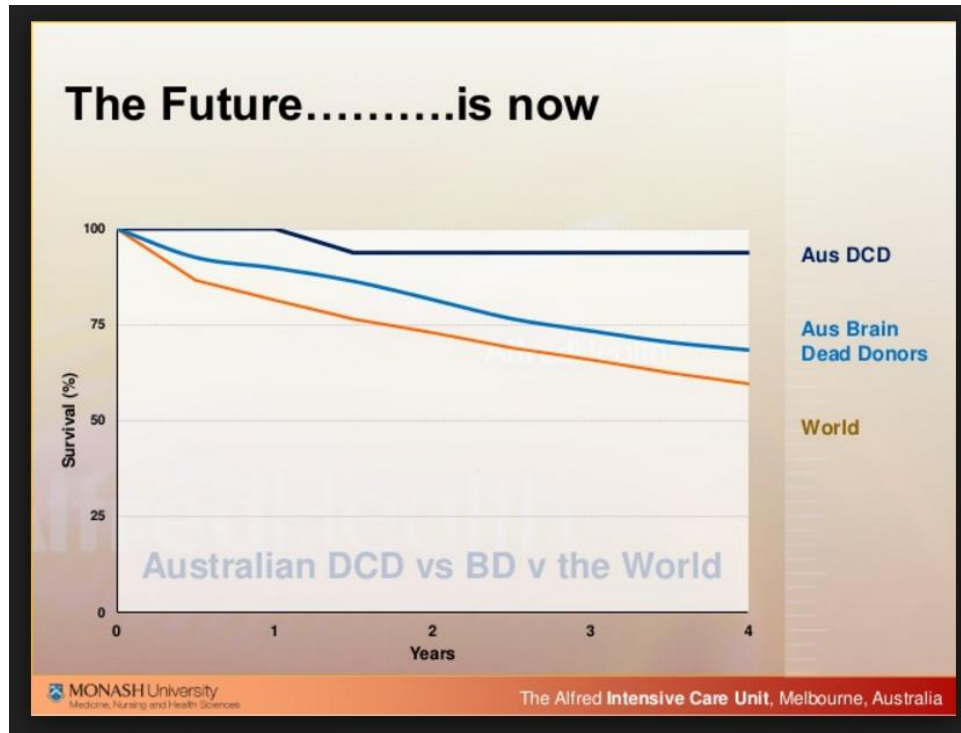
Heart

Liver

Kidney



DCD –organ donation-



Kunstige Lungen

Artificial lung basics

Fundamental challenges, alternative designs and future innovations

Heather Nolan,¹ Dongfang Wang² and Joseph B. Zwischenberger^{1*}

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Key words: artificial lung, paracorporeal lung, lung transplant, mechanical ventilation, pulmonary function, organ replacement

Abbreviations: ECMO, extracorporeal membrane oxygenator; IVOX, intravascular oxygenator; W-Z DLC, Wang-Zwischen double lumen cannula

There exists a growing demand for new technology that can take over the function of the human lung, from assisting an injured or recently transplanted lung to completely replacing the native lung. Many investigators are endeavoring to achieve the lofty goals and expectations of such a device. An artificial lung need be able to sustain the gas exchange requirements of a normal functioning lung. Pursuant to this purpose, the device must maintain appropriate blood pressure, decrease injury to blood cells and minimize clotting and immunologic response. Attachment methods vary and, ideally, researchers want the device to fit the way the native body tissue fits the native lung and utilizes the inherent properties of the native lung. The currently proposed methods include the parallel, in-series and venous double-lumen cannula configurations. For the time being, current research focuses on the extracorporeal (i.e., outside the body) placement, but ultimate long-term goals look toward total implantation.

Meeting the Need

The classic medical drama would be incomplete without the aspen-filled meadow-to-mountain rescue scene, one person breathing for another in peak-of-life. Hardly a ventilator device in itself, the concept still plays an essential role in current thoracic studies: develop a means of respiration, outside of an individual's own power, that can assist or completely take over breathing for an impaired individual. Currently in use is the mechanical ventilator with a wide variety of techniques, calculations and adjustments aimed at improving its success. On the horizon is the mechanical or artificial lung, a device that is implantable in a human for respiratory support.

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Submitted: 08/07/10; Accepted: 10/25/11
DOI: 10.4236/ojs.2011.31405

the number of lungs available, the demand clearly outweighs the supply. Additionally, only 18% of the 13,154 lungs from organ donors were transplanted in 2006; 81% were not recovered.¹ The reason for this discrepancy was cited as "poor organ function," leading to an even greater disparity between needed and available lungs.¹ As such, research has focused not merely on an artificial lung as a replacement organ, but rather an artificial lung as a bridge to transplantation.^{1,2} Additionally, a successful artificial lung could be used as a support device following transplant or as supplemental support to mechanical ventilation.^{3,4}

Current Devices

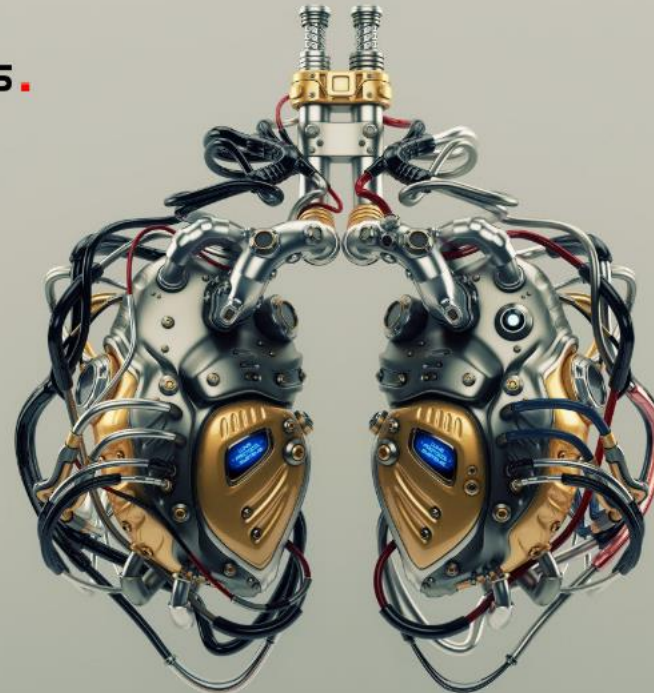
Historically, artificial lung technology has been limited to its use in cardiopulmonary bypass and extracorporeal membrane oxygenation (ECMO). ECMO, however, has inherent complications. It is very time-consuming, needing large amounts of monitoring and personnel, and the device itself consists of tubes and cannulae that can rupture and lead to fatal results.¹ ECMO therapy is noninflammatory and requires multiple transfusions; additional complications include infection, erythrocyte and platelet damage and embolization of tube fragments.^{2,3} For these reasons and others, the latest advances have deviated from ECMO therapy.

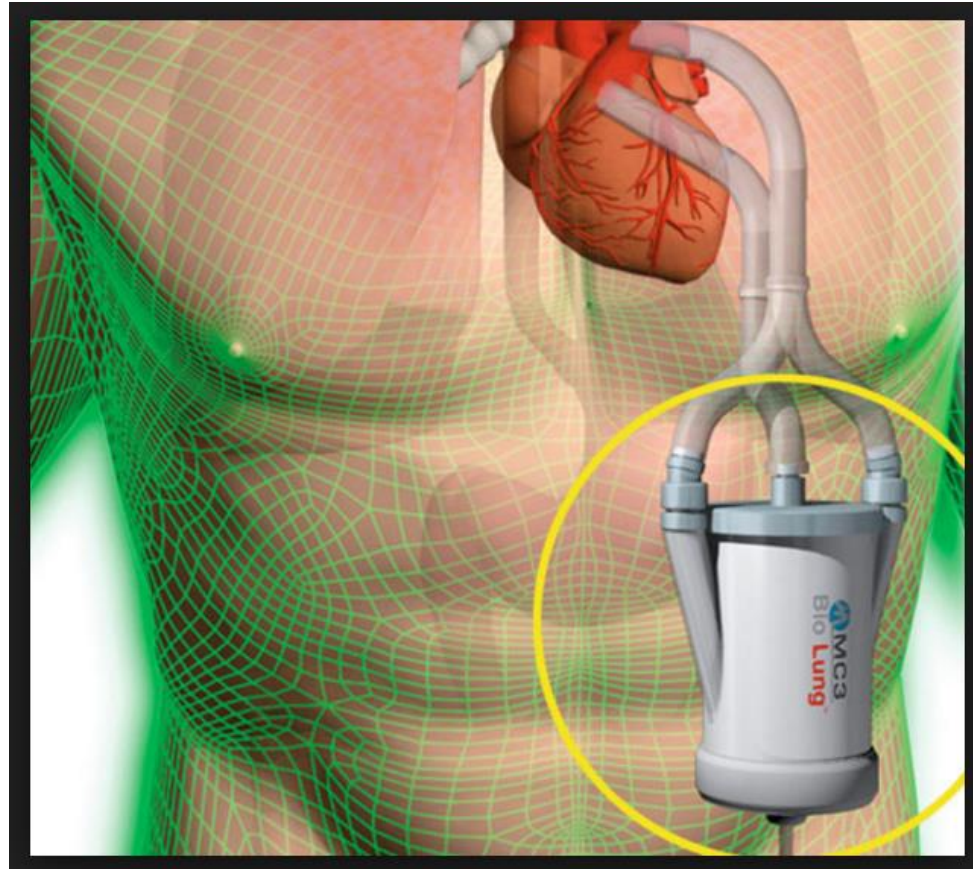
The intravascular oxygenator, IVOX, is one such alternative. Gas enters and leaves the system via conduits outside a small skin incision. The device, a membrane oxygenator, is placed in the vena cava and a vacuum pump pulls oxygen through the device fibers.¹ Another version has replaced the vacuum pump with a pulsating balloon that allows for optimal blood mixing across the gas exchange fibers.² Yet, IVOX does not have the surface area or the gas exchange capacity to meet the needs of a pre- or post-

Another alternative choice is a high-flow cannula placed in the femoral artery and vein that creates a pumpless shunt. This method, deemed interventional lung assist, allows for complete CO₂ removal and, due to the device's location in the peripheral vasculature, the natural lung can still provide oxygenation within its 'capabilities'. Conversely, the device will receive only partial cardiac output, which will limit its effectiveness, as both adequate

CYBER ORGANS.
CONCEPT DESIGN BY VLADISLAV OCIACIA

LUNG





Interview

Could 3D printing solve the organ transplant shortage?

By Tim Lewis

Scientists are racing to make replacement human organs with 3D printers. But while the technology's possibilities are exciting, already there are fears we could be 'playing God'

Tissue Engineering

REVIEW

Organogenesis 10(2), 196–207, April/May/June 2014; © 2014 Landes Bioscience

Future prospects for tissue engineered lung transplantation Decellularization and recellularization-based whole lung regeneration

Tomoshi Tsuchiya¹*, Anagh Sivarapatna², Kevin Rocco³, Atsushi Nanashima⁴, Takeshi Nagayasu⁵, and Laura E Niklason⁶¹Division of Surgery, Department of Surgery, Nagasaki University Graduate School of Biomedical Sciences, Nagasaki, Japan;
²Departments of Anesthesia and Biomedical Engineering, Yale University, New Haven, CT, USA**Keywords:** lung transplantation, decellularization, recellularization, extracellular matrix, stem/progenitor cells, induced pluripotent stem cells

The shortage of donor lungs for transplantation causes a significant number of patient deaths. The availability of laboratory engineered, functional organs would be a major advance in meeting the demand for organs for transplantation. The accumulation of information on biological scaffolds and an increased understanding of stem/progenitor cell behavior has led to the idea of generating transplantable organs by decellularizing an organ and recellularizing using appropriate cells. Recellularized solid organs can perform organ-specific functions for short periods of time, which indicates the potential for the clinical use of engineered solid organs in the future. The present review provides an overview of progress and recent knowledge about decellularization and recellularization-based approaches for generating tissue engineered lungs. Methods to improve decellularization, maturation of recellularized lung, candidate species for transplantation and future prospects of lung bioengineering are also discussed.

Introduction

Lung transplantation is the last option for the treatment of terminal lung disorders such as chronic obstructive pulmonary disorder (COPD), which is the third leading cause of death in the United States.^{1,2} More than 2000 lung transplants are performed annually in the United States.³ Whereas lung transplantation is recognized as an established therapy that improves survival and provides an improved quality of life for transplant recipients,⁴ there are several hurdles that make the transplantation of donor lungs a challenging endeavor.

One critical issue is the condition of the donor lungs. Donor lungs often suffer edema, atelectasis, or pneumonia originating from donor systemic problems or preoperative systemic control.⁵ The long ischemic time also causes donor lung damage, which

directly correlates with primary graft failure.⁶ The advantage of a lung transplant is the need for limitation of immunosuppression drugs in the recipient, which increases the risks of infections and cancer.⁷ Therefore, however, is the acute shortage of transplantable lungs. In 2011, there were 2280 patients waiting for lung in the US alone with the average waiting time of mortality for patients on the wait list is 15.7 years in the US.⁸ In Japan, the situation is worse; the average waiting time in Japan is about 2 y and 9 m and the mortality is 43 per 100 wait-list years.⁹ The shortage of lungs greatly exceeds the supply. Further, a long waiting time for a suitable organ results in deterioration of the patient's condition and increases operative risk.

Such unresolved problems, lung transplantation is necessary but currently imperfect treatment for lung disease. In order to address the problem of organ shortage, several approaches for making transplantable organs are attempted, including biomaterial improvement (3D) cell cultures, and ex vivo bioengineering.¹⁰ The progress of regenerative medicine and the development of stem cell advances, investigators have also begun to use decellularized native scaffolds for recellularization to generate transplantable bioorgans (Fig. 1).^{11,12}

This review describes the recent development of decellularization and recellularization strategies for generating lungs for transplantation. We will begin by discussing the concepts of decellularization and recellularization by the current status of the cell replacement approach for whole lung regeneration, including the use of cell lines and studies of extracellular matrix (ECM). Finally, we address other important aspects of lung tissue engineering, such as protocols for organ maturation, the choice of species, operation procedures and the perspective of recellularized lung transplantation.

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Organogenesis

HOME > SCIENCE > VOL. 329, NO. 5991 > TISSUE-ENGINEERED LUNGS FOR IN VIVO IMPLANTATION

RESEARCH ARTICLE

f t in g e m

Tissue-Engineered Lungs for in Vivo Implantation

THOMAS H. PETERSEN, ELIZABETH A. CALLE, LIPING ZHAO, EUN JUNG LEE, LIQIONG GUI, MICHASAM B. RAREDON, KSENIYA GAVRILOV, TAI YI, ZHEN W. ZHUANG, [...], AND

LAURA E. NIKLASON

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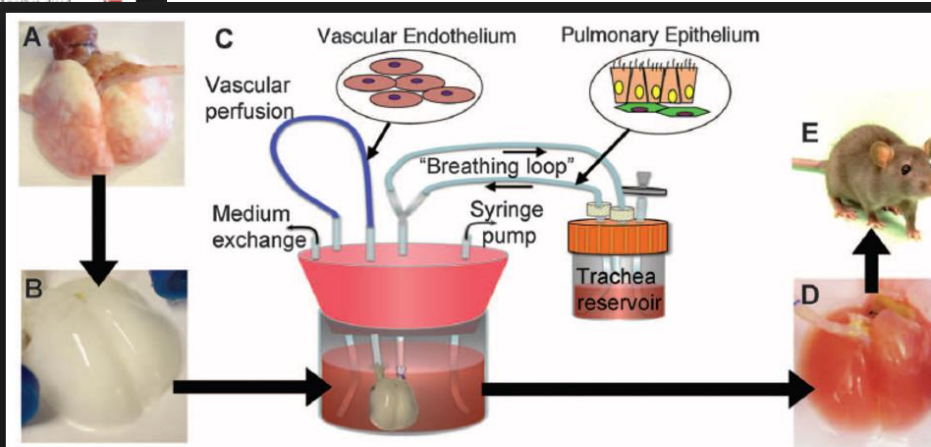


Fig. 1. Schema for lung tissue engineering. (A) Native adult rat lung is cannulated in the pulmonary artery and trachea for infusion of decellularization solutions. (B) Acellular lung matrix is devoid of cells after 2 to 3 hours of treatment. (C) Cellular matrix is mounted inside a biomimetic bioreactor that allows seeding of vascular endothelium into the pulmonary artery and pulmonary epithelium into the trachea. (D) After 4 to 8 days of culture, the engineered lung is removed from the bioreactor and is suitable for implantation into (E) the syngeneic rat recipient.





TISSUE ENGINEERING

Production and transplantation of bioengineered lung into a large-animal model

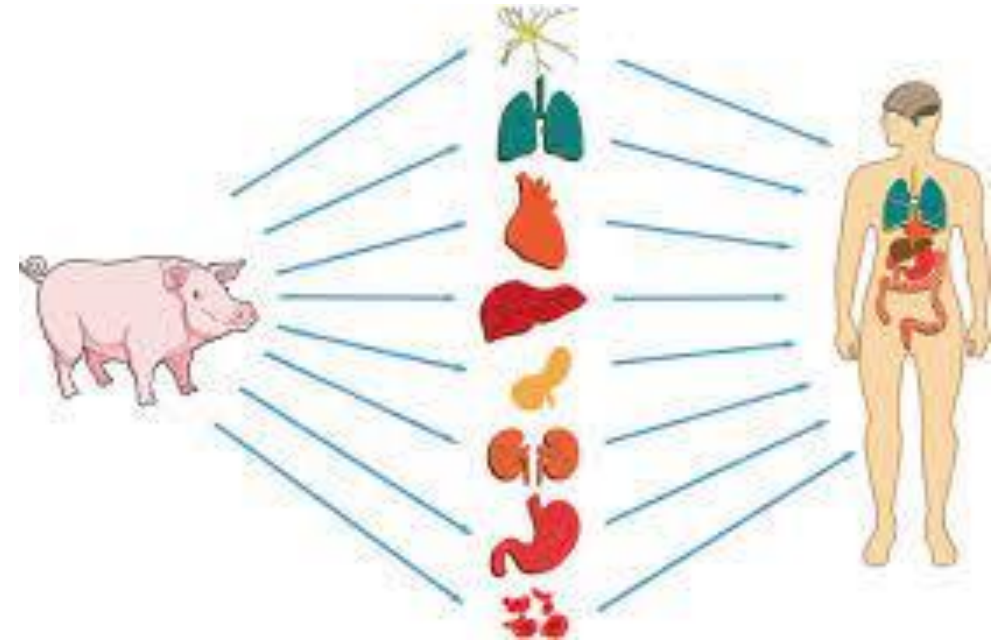
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Xenotransplantation: Are pigs the future of organ transplants?

13 March



Afsluttende bemærkninger

Transplantation kan være en fantastisk behandling

Transplantation kræver en hold indsats

Transplantation kræver gaven af organ donation
- så husk at tage stilling !

Tak for jeres opmærksomhed

