



**Azienda  
Ospedaliero  
Universitaria  
Careggi**

# Percorso donatore a cuore battente e a cuore fermo

**Dtt.ssa Guetti Cristiana**  
SODc Cure intensive del trauma e  
delle gravi insufficienze d'organo

*Dipartimento Neuromuscoloscheletrico e degli Organi di senso*



## Liste di Attesa al 31 Dicembre 2017\*

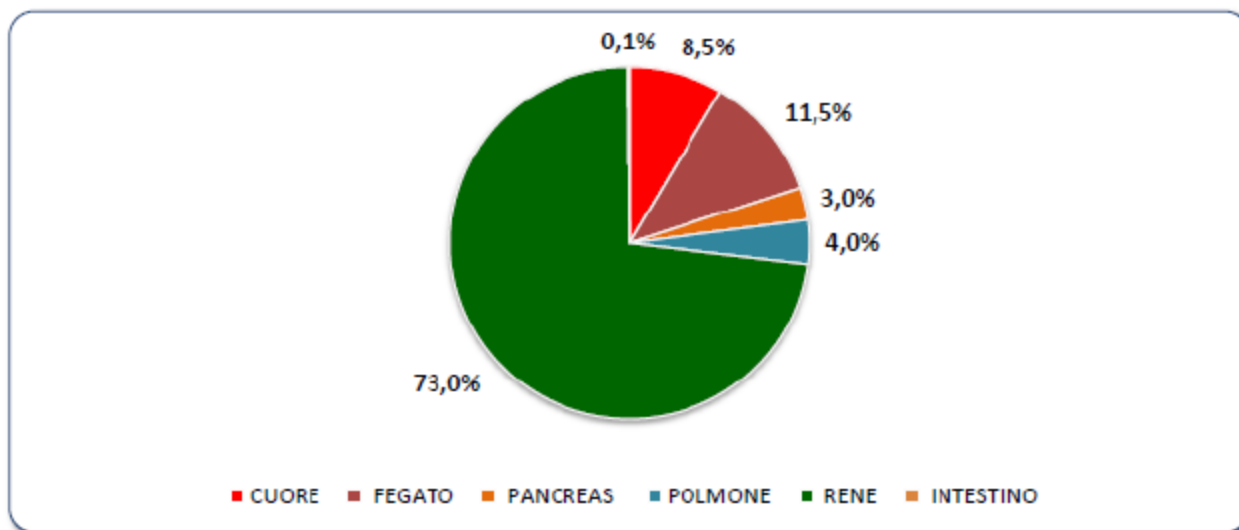
PAZIENTI in lista d'attesa in ITALIA al 31/12/2017 :

**8743**

Rene	6492**
Fegato	1019
Cuore	757
Polmone	354
Pancreas	264
Intestino	12

Iscrizioni rene  
**8449\*\***

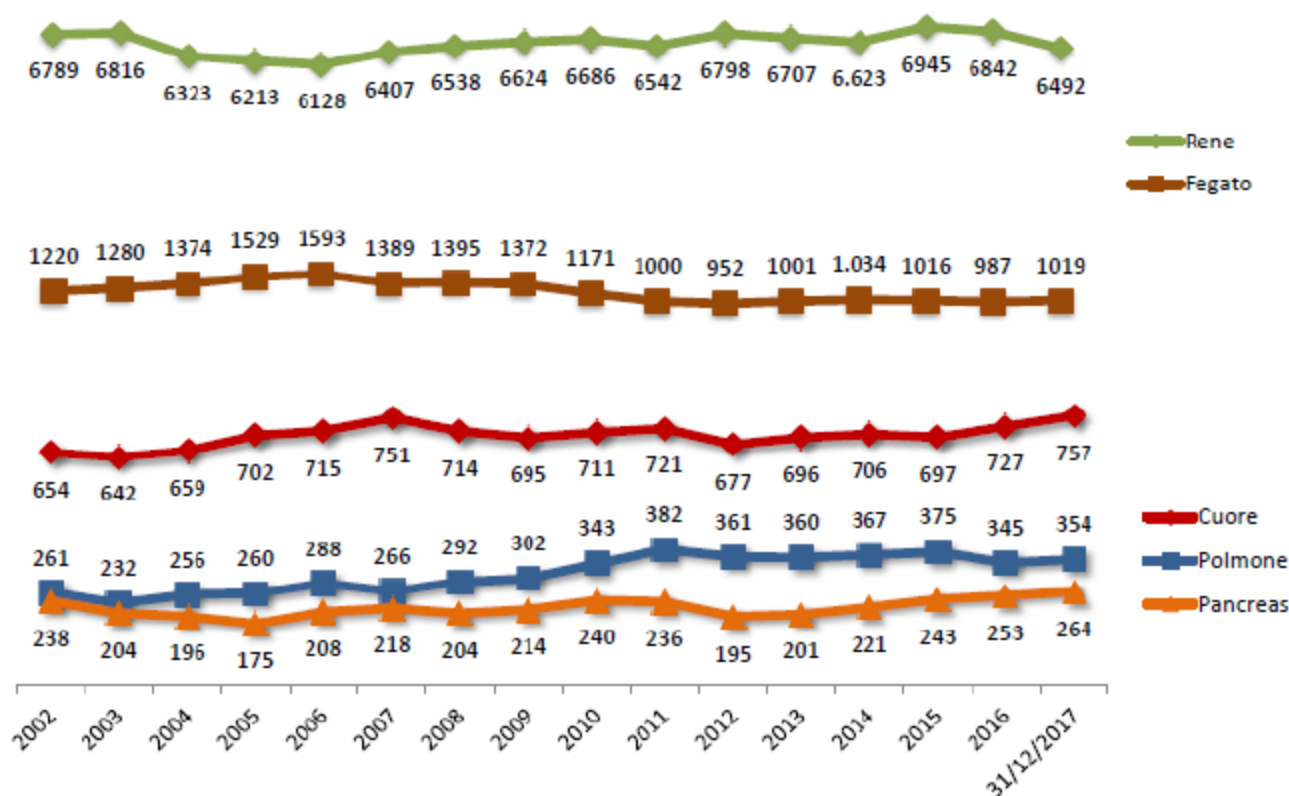
\*\* Per il rene ogni paziente può avere più di una iscrizione





# Andamento Liste di Attesa 2002 – 31/12/2017\*

## Pazienti iscritti in lista



## DEFINITION

### Donazione / donatore a cuore battente:

Heart-beating donation / donor

**HBD**

Donation / donor after brain death

**DBD**

### Donazione / donatore a cuore non battente in asistolia

Non heart-beating donation / donor

**NHBD**

Donation / donor after circulatory death

**DCD**



# LA MORTE SI IDENTIFICA CON LA CESSAZIONE IRREVERSIBILE DI TUTTE LE FUNZIONI DELL'ENCEFALO

Art. 1 legge 23/12/1993 n°578



**Arresto  
respiratorio e  
circolatorio**



**Lesione  
cerebrale  
primitiva**



Danno cerebrale  
**secondario** irreversibile



Danno cerebrale  
**primario** irreversibile



# UNA MORTE SOLA

la morte è sempre la morte DI TUTTO l'ENCEFALO

DUE MODI PER ACCERTARLA

- ACCERTAMENTO CON CRITERI CARDIOLOGICI
- ACCERTAMENTO CON CRITERI NEUROLOGICI





# Il percorso del donatore DBD



# Il percorso del donatore uDCD







**1. CERTEZZA** della **DIAGNOSI EZIOPATOGENETICA**  
**DELLA LESIONE CEREBRALE**

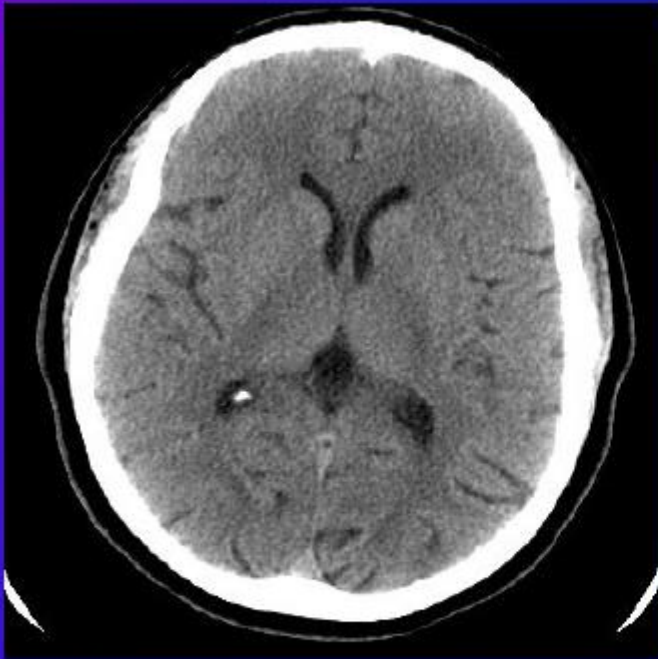
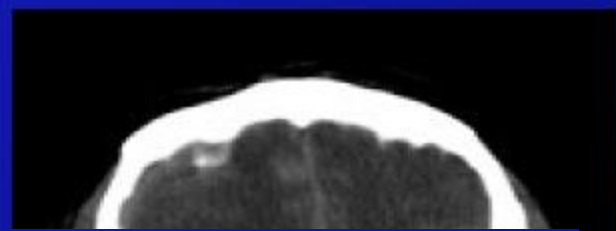
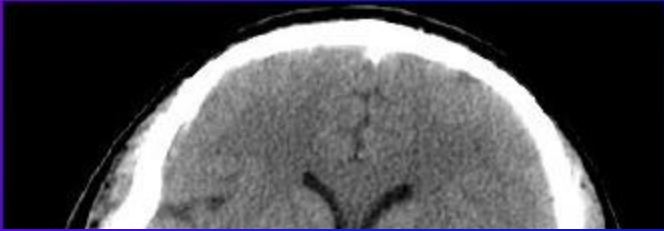
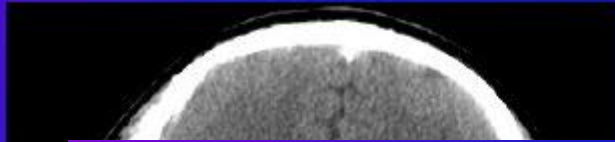
**2. NEUROIMAGING COERENTE** per gravità ed evoluzione

**3. STORIA CLINICA COERENTE** con l'andamento infausto

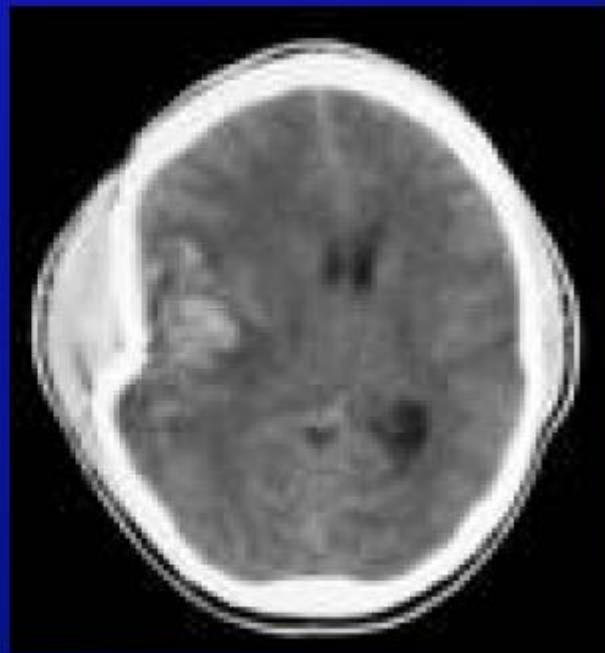
**4. ESCLUSIONE** di **TUTTE LE CAUSE DI COMA**  
**REVERSIBILE, RECENTI CRISI CONVULSIVE** ed  
**ATTIVITA' CRITICA SUBCLINICA**



itiva



normale



trauma cranico

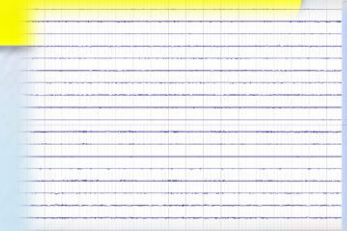


cerebrale

**ESAME CLINICO  
NEUROLOGICO**



**EEG**



**Le indagini strumentali  
non sostituiscono la  
diagnosi clinica ma completano la  
valutazione neurologica**

# ESAME CLINICO NEUROLOGICO

## RIFLESSI DEL TRONCO ENCEFALICO

VIA AFFERENTE: NERVO TRIGEMINO (5° PAIO N.C.)  
VIA EFFERENTE: NERVO FACCIALE (7° PAIO N.C.)

VIA AFFERENTE: NERVO VESTIBOLARE (8° PAIO N.C.)  
VIA EFFERENTE: NERVO OCULO MOTORE, ABDUCENTE (3°, 6° PAIO N.C.)



CORNEALE



OCULO  
VESTIBOLARE

FOTOMOTRE



VIA AFFERENTE: N. OTTICO (II)  
VIA EFFERENTE: N. OCULOMOTORE (III)



FARINGEO  
CARENALE

Via Afferente: N. Glossofaringeo (IX PAIO N.C.)  
Via Effrente: N. Vago (X PAIO N.C.)

RESPIRO SPONTANEO



Centro Bulbare



## Morte cerebrale: collegio medico



**COLLEGIO  
MEDICO**



**MEDICO LEGALE  
(O MEDICO DELLA DIREZIONE SANITARIA)**

**MEDICO ANESTESISTA/RIANIMATORE**

**MEDICO NEUROLOGO ESPERTO IN  
ELETTROENCEFALOGRAFIA**



# PRINCIPALI ALTERAZIONI FUNZIONALI NELLA MORTE ENCEFALICA

- **ALTERAZIONI CARDIOCIRCOLATORIE**  
(ipovolemia relativa, vasoplegia, ipotensione arteriosa)
- **PERDITA DELLA RESPIRAZIONE SPONTANEA**
- **SQUILIBRI IDROELETTRICI**
- **ALTERAZIONI ORMONALI E METABOLICHE** (deficit ADH, diabete insipido, ipotiroidismo, ipocorticosurrenalismo, iperglicemia)
- **PERDITA DELLA TERMOREGOLAZIONE**
- **GRAVE COAGULOPATIA**



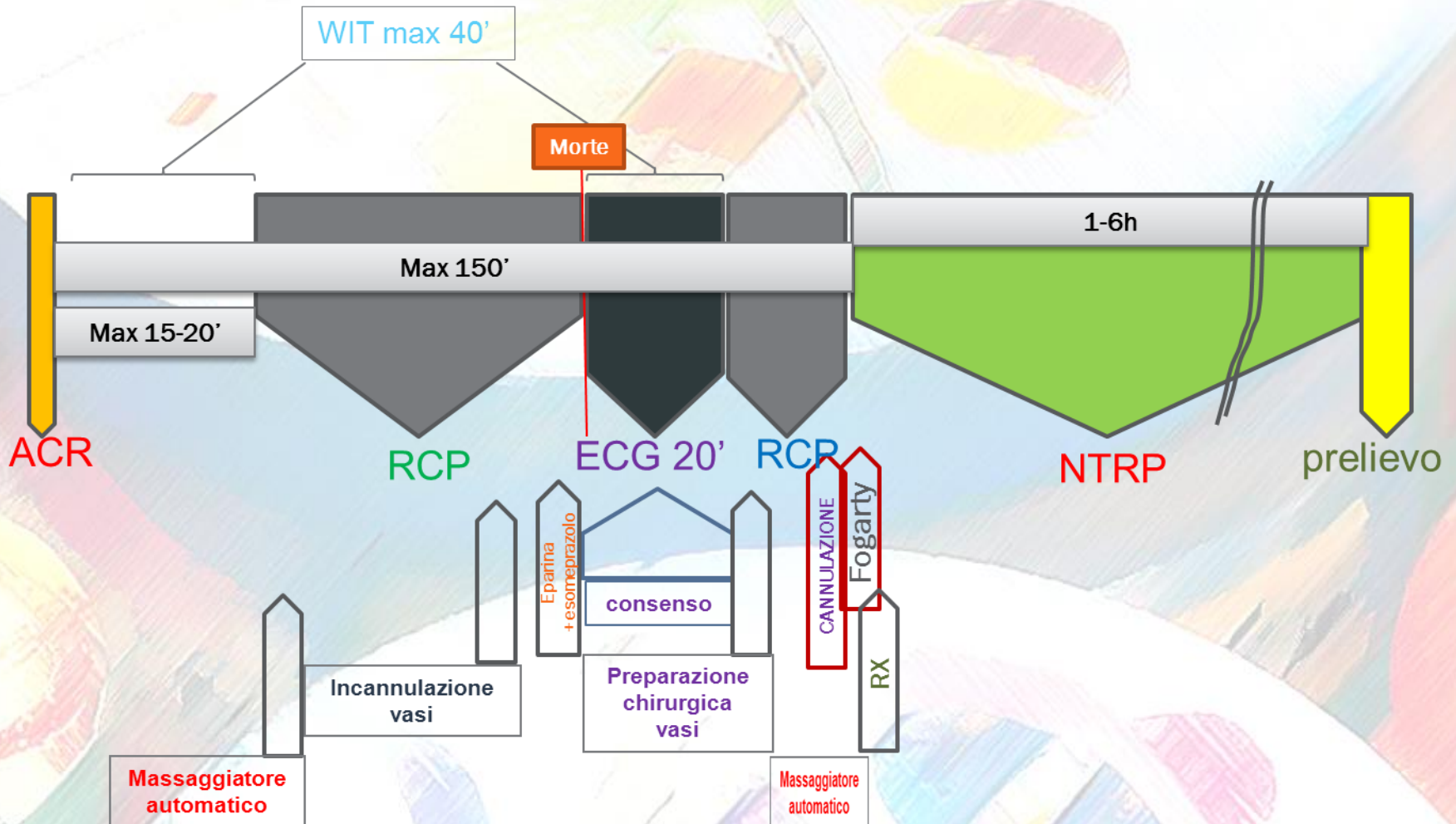




## Modified Maastricht Classification on DCD donors (Paris 2013)

<b>I</b> Uncontrolled	Found dead IA. Out-of-hospital IB. In-hospital
<b>II</b> Uncontrolled	Witnessed cardiac arrest IIA. Out-of- hospital IIB. In-hospital
<b>III</b> Controlled	Withdrawal of lifesustaining therapy
<b>IV</b> Uncontrolled Controlled	Cardiac arrest while brain death  <i>Thuong M, et al. Transplant Int 2016; 29:749-59</i>

# PROCEDURA NHBD





# Protocollo AOUC : criteri di arruolamento

15-65 aa

Donazione

ACR-RCP <20'

ACR to Hospital <90'

RCP complessiva <120'

ACR-no touch-NTRP <150'

**ACR testimoniato**  
**Paziente identificabile**  
**Parenti rintracciabili**

15-70aa

ECLS terapeutico

ACR-RCP <5'

FV come primo ritmo (qualunque tempo stimato ACR-RCP)

ACR to Hospital <40'

ACR-ECLS <60'

Consentiti tempi più lunghi se ipotermia o intossicazione da betabloccanti

ACR testimoniato

# CRITERI ASSOLUTI DI ESCLUSIONE

Evidenza o alto sospetto di patologie neoplastiche\*: es cachessia, presenza di colostomia, cicatrici visibili recenti  
Sepsi, evidenza di malattie infettive e trasmissibili acute o croniche\*: es AIDS, cirrosi epatica, epatite

Malattie autoimmuni con diagnosi certa\*

Emodialisi cronica

Decadimenti psico-cognitivi se con sospetto di una encefalopatia spongiforme

Donazione

Peso > 1 kg per cm  
Dissezione aortica/insufficienza grave  
Aterosclerosi arti inf. nota sintomatica  
«Comorbidità»

ECLS terapeutico

*Criteria in H*  
pH < 6,9  
Lattati > 20 mmol  
Adrenalina > 3 mg  
EtCO<sub>2</sub> < 10

\*informazioni disponibili e attendibili

arresto cardiaco

cessazione della circolazione

24 h

TESSUTI

WIT:  
WARM ISCHEMIA  
TIME

ORGANI

30'

120'

Morte della persona

> 5'

ENCEFALO

danno irreversibile

assenza di perfusione

ischemia

anossia

sofferenza

danno



REVIEW

Maximum tolerable warm ischaemia time in transplantation from non-heart-beating-donors

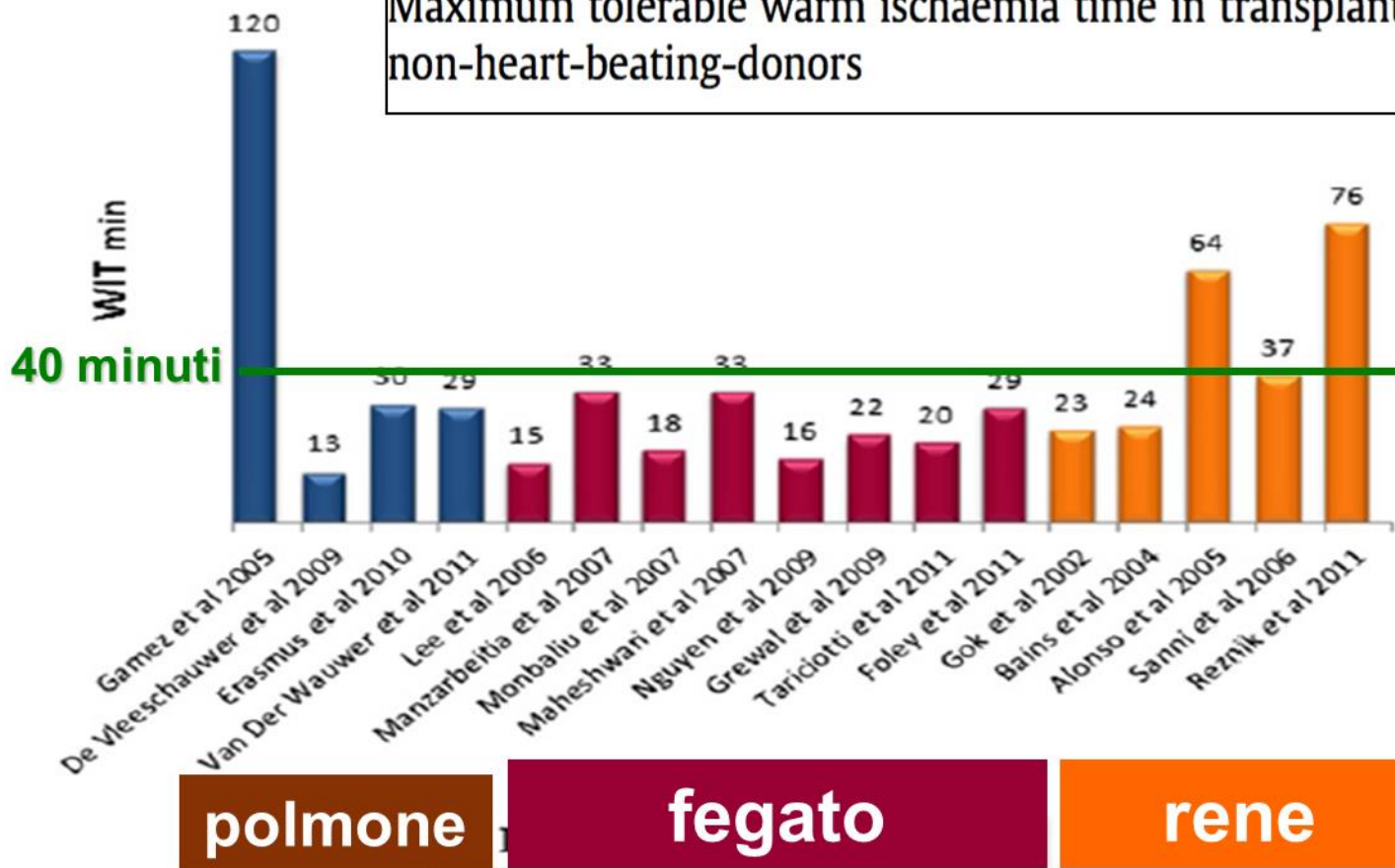
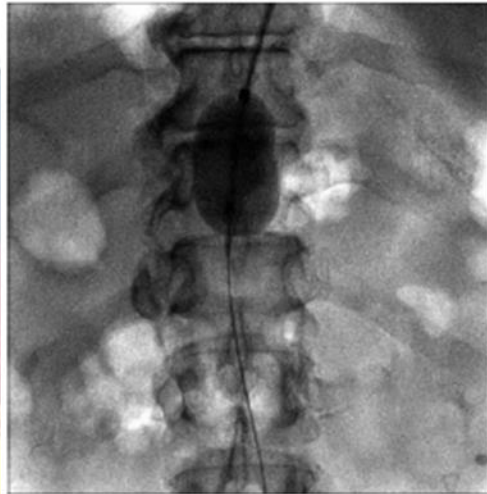


Fig. 1. Clinical experiences of WIT in lung, liver and kidney transplantations from non-heart-beating-donors (WIT: warm ischaemia time).

# NTRP nella pratica clinica:

- Introduzione di cannule ECMO in vena e arteria femorale



- Catetere di fogarty nella arteria femorale controlaterale (cuffiato al di sopra del diaframma)



- Connessione cannule a circuito ECMO (BF 2-3 l/min)

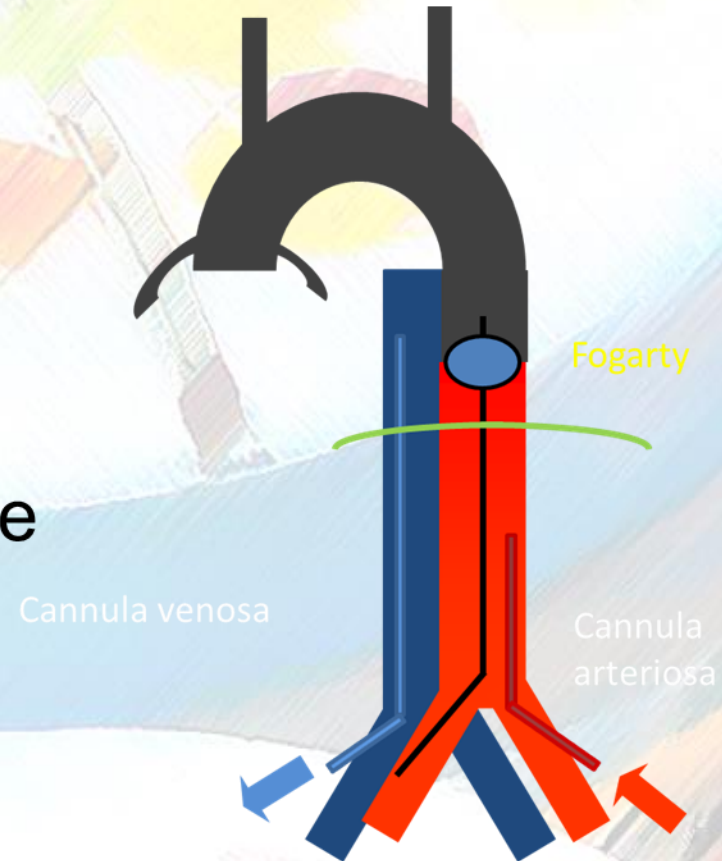


**Abdominal regional in-situ perfusion in donation  
after circulatory determination of death donors**

*Amelia J. Hessheimer, Juan C. García-Valdecasas,  
and Constantino Fondevila*

Curr Opin Organ Transplant 2016, 21:322–328

**ECLS**  
+  
**CATETERE FOGARTY**  
aorta toracica discendente



**NTRP:** Normo-thermic Regional Perfusion

**ANOR:** Abdominal Normothermic Oxygenated Recirculation

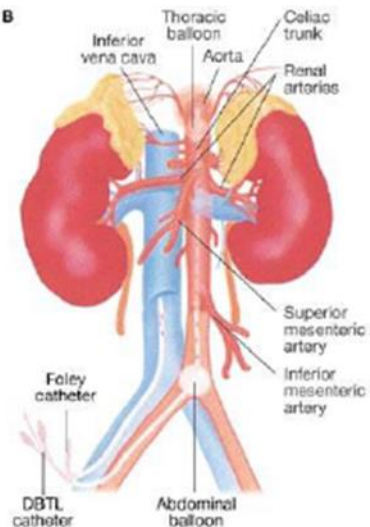


# DCD: Metodiche di mantenimento organi



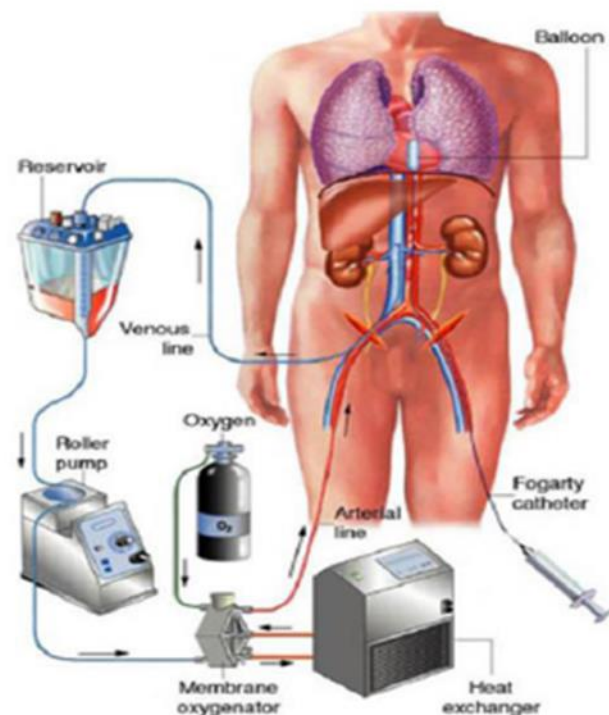
**SUPER  
RAPIDE  
TECHNIQUE**

## COLD IN SITU PERFUSION (ISP)



## ECMO:

- CORE COOLING (TOTAL BODY COOLING)
- N-ECMO: NORMOTHERMIC-ECMO



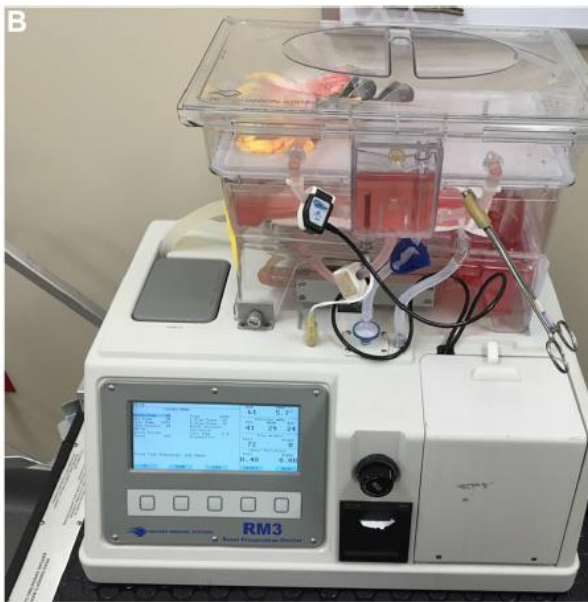
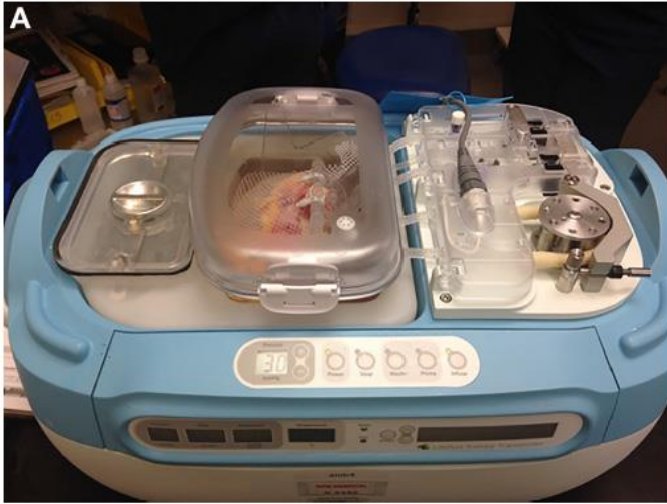
- Trasporto paziente in SO
- Esplorazione addominale



- Stop ECMO e utilizzo delle cannule per perfusione con liquido freddo e raffreddamento con ghiaccio
- Prelievo degli organi



# Perfusione ex-vivo del rene



**Machine Perfusion Versus Cold Storage for the Preservation of Kidneys Donated After Cardiac Death**  
*A Multicenter, Randomized, Controlled Trial*

Transplantation Reviews 32 (2018) 1–9  
 Contents lists available at ScienceDirect  
**Transplantation Reviews**  
 journal homepage: [www.elsevier.com/locate/trre](http://www.elsevier.com/locate/trre)

**Ex vivo machine perfusion for renal graft preservation**  
 J. Moritz Kathis<sup>a,b,c,\*</sup>, Andreas Paul<sup>d</sup>, Lisa A. Robinson<sup>e</sup>, Markus Selzner<sup>b</sup>

<sup>a</sup> Department of General, Visceral, and Transplantation Surgery, University Hospital Essen, University Duisburg-Essen, Essen, Germany  
<sup>b</sup> Adult Organ Transplant Program, Department of Surgery, Toronto General Hospital, University Health Network, Toronto, Ontario, Canada  
<sup>c</sup> Division of Nephrology, The Hospital for Sick Children, University of Toronto, Toronto, Ontario, Canada

**ABSTRACT**

Kidney transplantation is the treatment of choice for end-stage renal disease. Despite its superiority over dialysis, the persisting organ shortage remains a major drawback. Additional sources to increase the donor pool are grafts recovered from extended criteria donors (ECD) and donation after circulatory death (DCD). Although transplantation of marginal grafts demonstrates promising outcomes, increased rates of primary non-function, delayed graft function, and reduced graft survival have been reported. Cold ischemic injury, caused by static cold storage is a significant risk factor for poor outcome. Machine perfusion (MP) at various temperatures bears the potential to improve organ preservation, assessment, and repair. While hypothermic machine perfusion (HMP) is well established in clinical practice, modified HMP, subnormothermic machine perfusion (SMP), and normothermic machine perfusion (NMP) are novel emerging strategies with the potential to significantly improve the outcome of marginal kidney grafts. This review summarizes findings and recent advances from pre-clinical and clinical machine perfusion studies, organized by temperature, and discusses potential future developments for graft assessment and repair.

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- Machine Perfusion:**
- Reduced incidence and duration of DGF
  - Higher CrClearance 1 month after transplantation



PNF:  
PRIMARY  
NON  
FUNCTION



Non- and never functioning transplant, return to dialysis.

DGF:  
DELAYED  
GRAFT  
FUNCTION



Failure of renal graft function so there is a need for dialysis in the 1<sup>st</sup> week after kidney transplantation

# NTRP: ESPERIENZE CLINICHE

RENE

One of the main advantages of NRP-DCD kidney transplantation is marked reductions in DGF

Transpl Int 2000;13: 303-310. Clin  
Transplant 2011;25:511-516.  
J Am Coll Surg 2008;206:1028-1037

Reducing DGF diminishing greater risk of graft failure and rejection



Curr Anaesth Crit Care 2010;21:220-223.

The recirculation of blood at the homeostatic temperature (37°C) has been shown to reduce DGF and replenish antioxidant and ATP levels.

Am J Transplant 2010;10:1365-1374.  
Transplant Proc 2007;39:249-252.



## Postoperative Care in Kidney Transplantation: A Comparison Between Controlled and Uncontrolled Donation After Circulatory Death

Á.J. Roldán-Reina<sup>a,\*</sup>, J.J. Egea-Guerrero<sup>b</sup>, N. Palomo-López<sup>a</sup>, D.X. Cuenca-Apolo<sup>a</sup>,  
M. Adriaensens-Pérez<sup>a</sup>, M. Porrás-López<sup>a</sup>, Z. Ruiz de Azúa-López<sup>c</sup>, Y. Corcia-Palomo<sup>a</sup>,  
and L. Martín-Villén<sup>c</sup>

<sup>a</sup>Intensive Care Unit, Virgen del Rocío University Hospital, Seville, Spain; <sup>b</sup>Coordination of Transplants: Seville-Huelva Sector, IBiS/CSIC, Seville University, Seville, Spain; and <sup>c</sup>Coordination of Transplants: Seville-Huelva Sector, Seville, Spain

When comparing postoperative renal function between uDCD and cDCD transplants, some studies reported delayed graft function in uDCD [3,5], given that warm ischemic time is more prolonged in this patient subgroup.

However, different authors found that short- and long-term graft function normalizes in both donor groups [3,6], even when compared with grafts from donation after brain death. These studies demonstrate that DCD should remain a viable option in kidney transplantation, regardless of immediate postoperative renal function [4,5,7].

Hoogland et al [6] found that postoperative renal function could be similar in both patient groups, a finding that may be attributed to the use of expanded criteria in DCD.

Moreover, our results showed a relationship between higher lactate levels and the risk of poor outcome after kidney transplantation. Along this line, a recent study found that defective postreperfusion metabolic recovery, as determined by lactate levels, is a factor that could lead to delayed graft function [8].

hemodialysis [1]. In recent years, an increasing demand for organ transplantation, paired with a progressive decline in brain-dead donors, has made it necessary to seek new strategies for organ donation, including donation after circulatory death (DCD) [2]. However, recent research on



## CHAIN OF SURVIVAL



## CHAIN OF OPPORTUNITIES



*Courtesy: Dr. Juan J. Egea-Guerrero, Virgen del Rocío Hospital, Seville, Spain*



A vibrant, comic book-style graphic. The background is a bright yellow, filled with numerous thin, black, radiating lines that create a sense of energy and motion. Scattered throughout the scene are several stylized white clouds with black outlines, some appearing to be puffs of smoke or steam. The central focus is the text 'THANK YOU' written in a large, bold, red font. The letters are thick and have a slight 3D effect, with a white outline and a black drop shadow. The words are arranged in two lines: 'THANK' on top and 'YOU' below it. The overall aesthetic is classic and celebratory, reminiscent of a comic book's 'THANK YOU' panel.

**THANK  
YOU**