

IMMAGINARE IL CUORE

Rigenerare, ricostruire, sostituire
Il sogno delle staminale, la realtà del
trapianto e dell'assistenza cardiaca

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Advanced Heart Failure

➤ Europe: 10.000.000 with AHF

➤ USA: 400.000 new cases/year

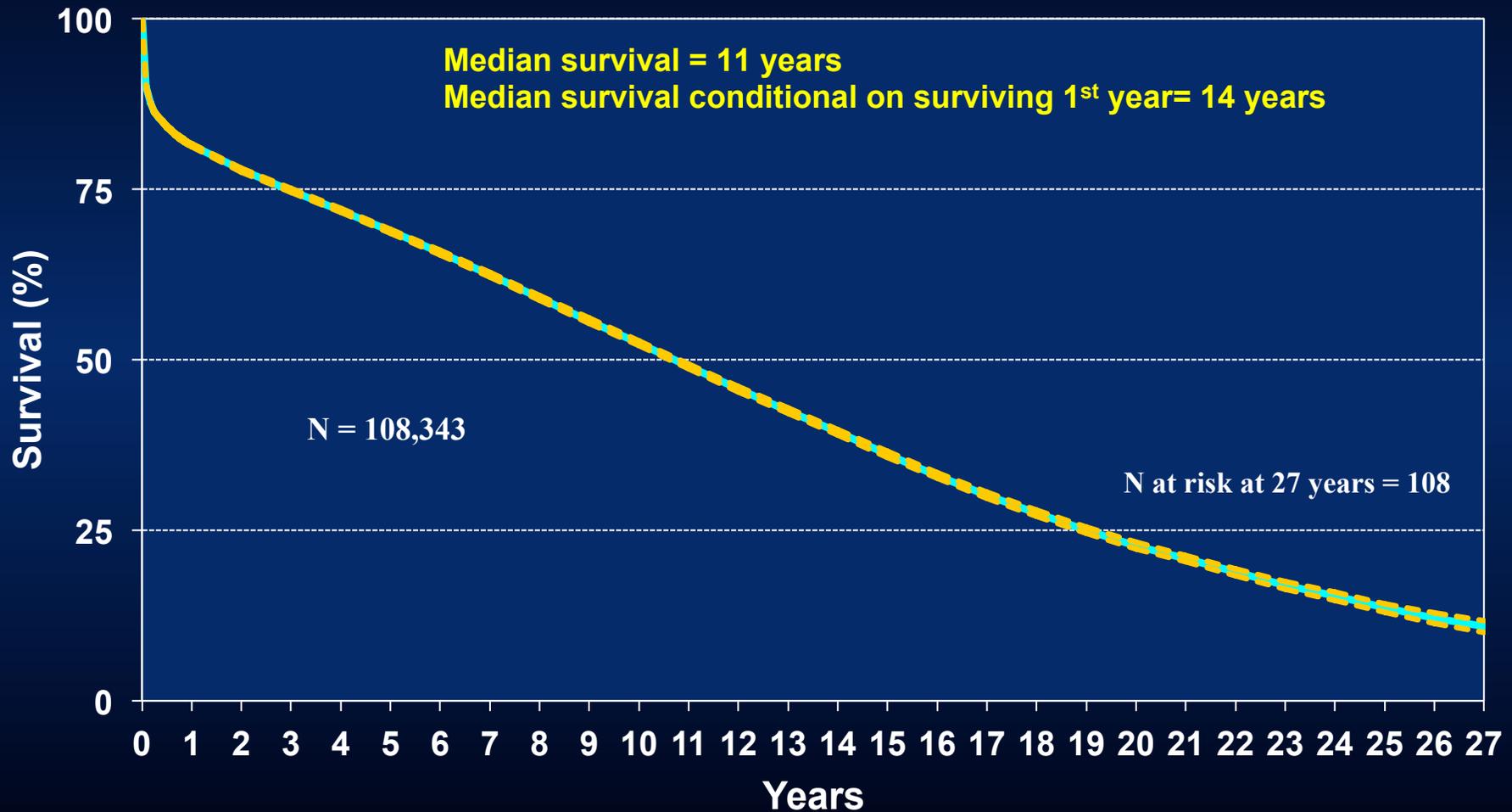
➤ 34 billions \$/year

Heart transplantation



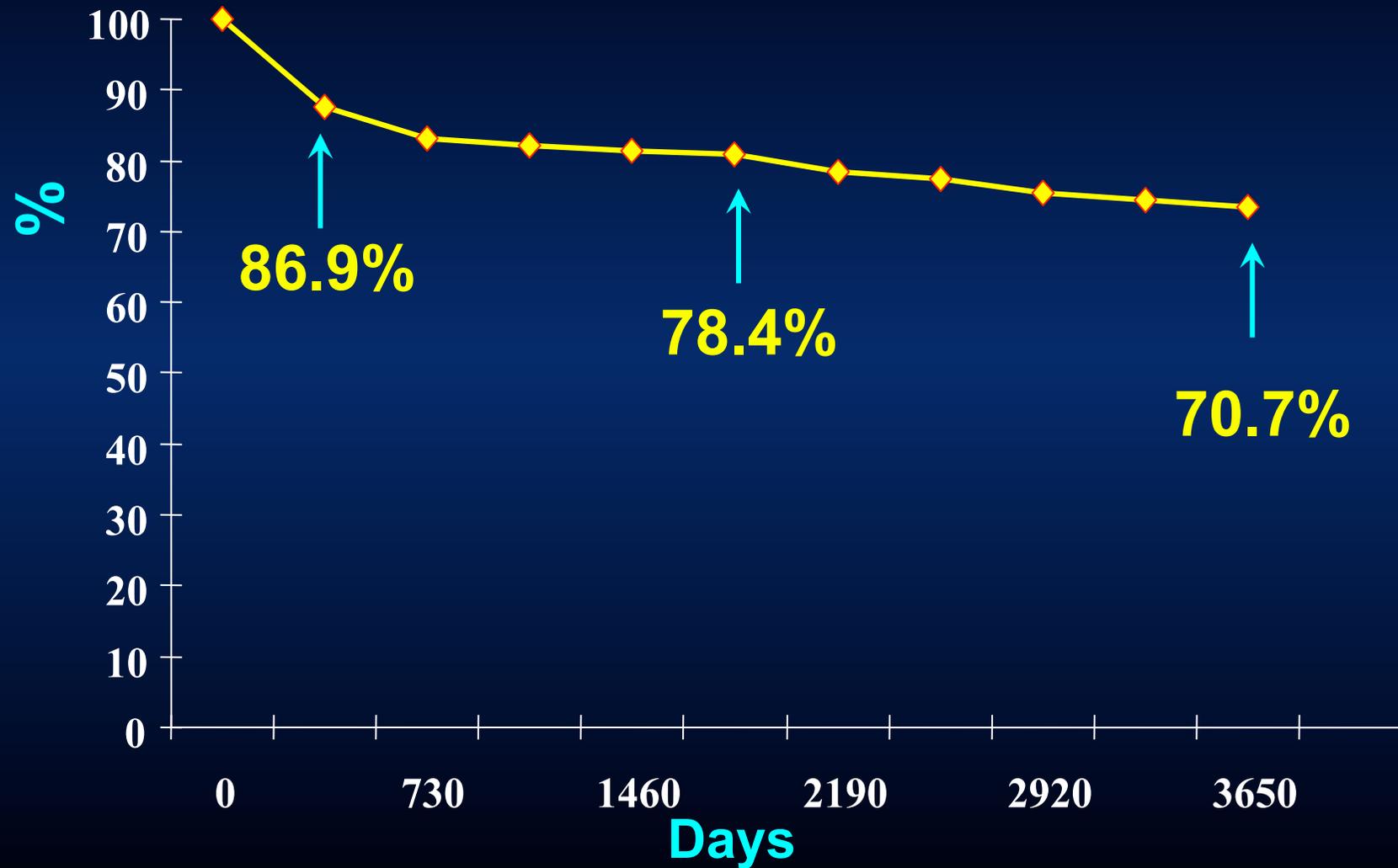


Adult and Pediatric Heart Transplants Kaplan-Meier Survival (Transplants: January 1982 – June 2012)



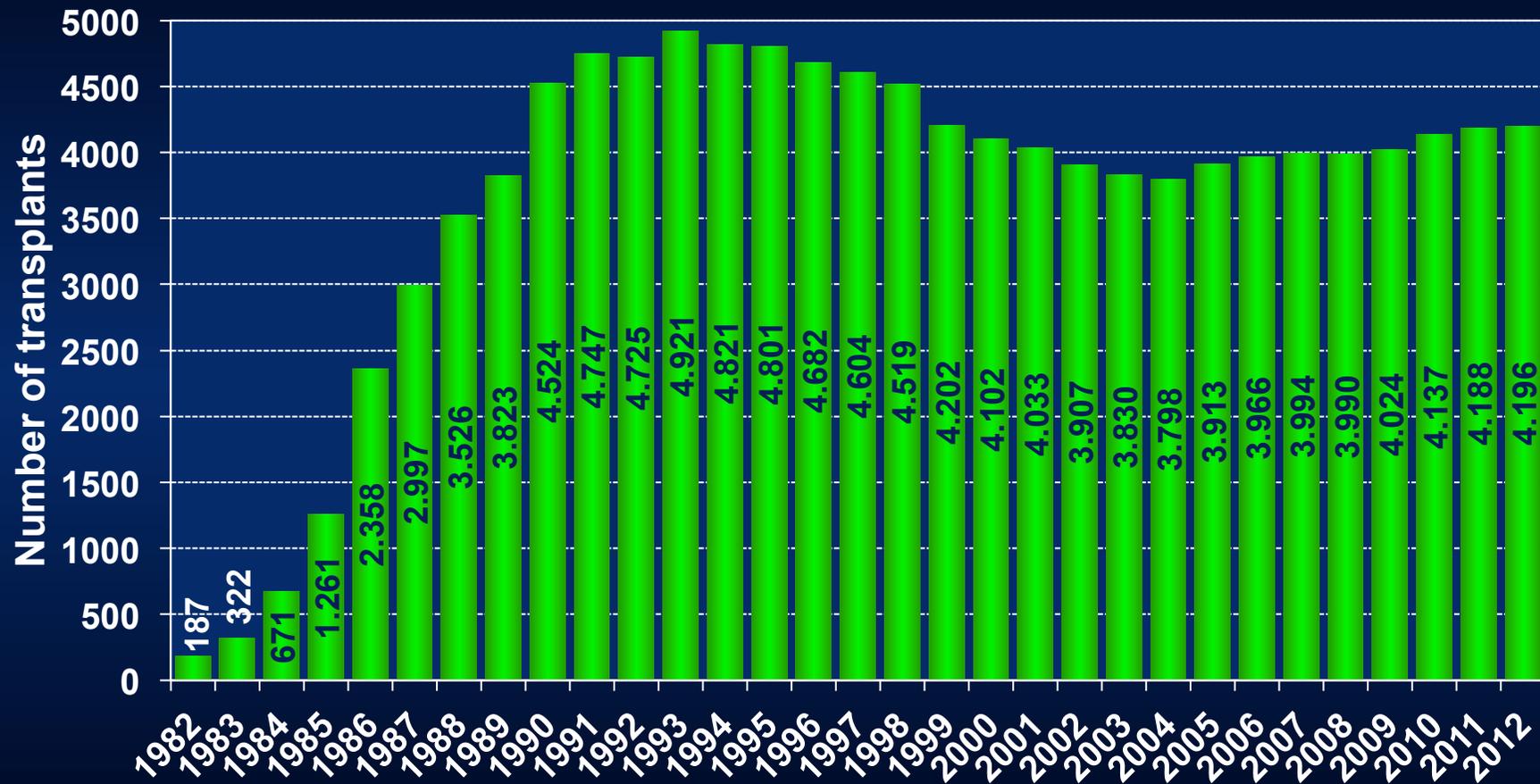
Survival after HTx

Milan Experience - 805 pts



Adult and Pediatric Heart Transplants

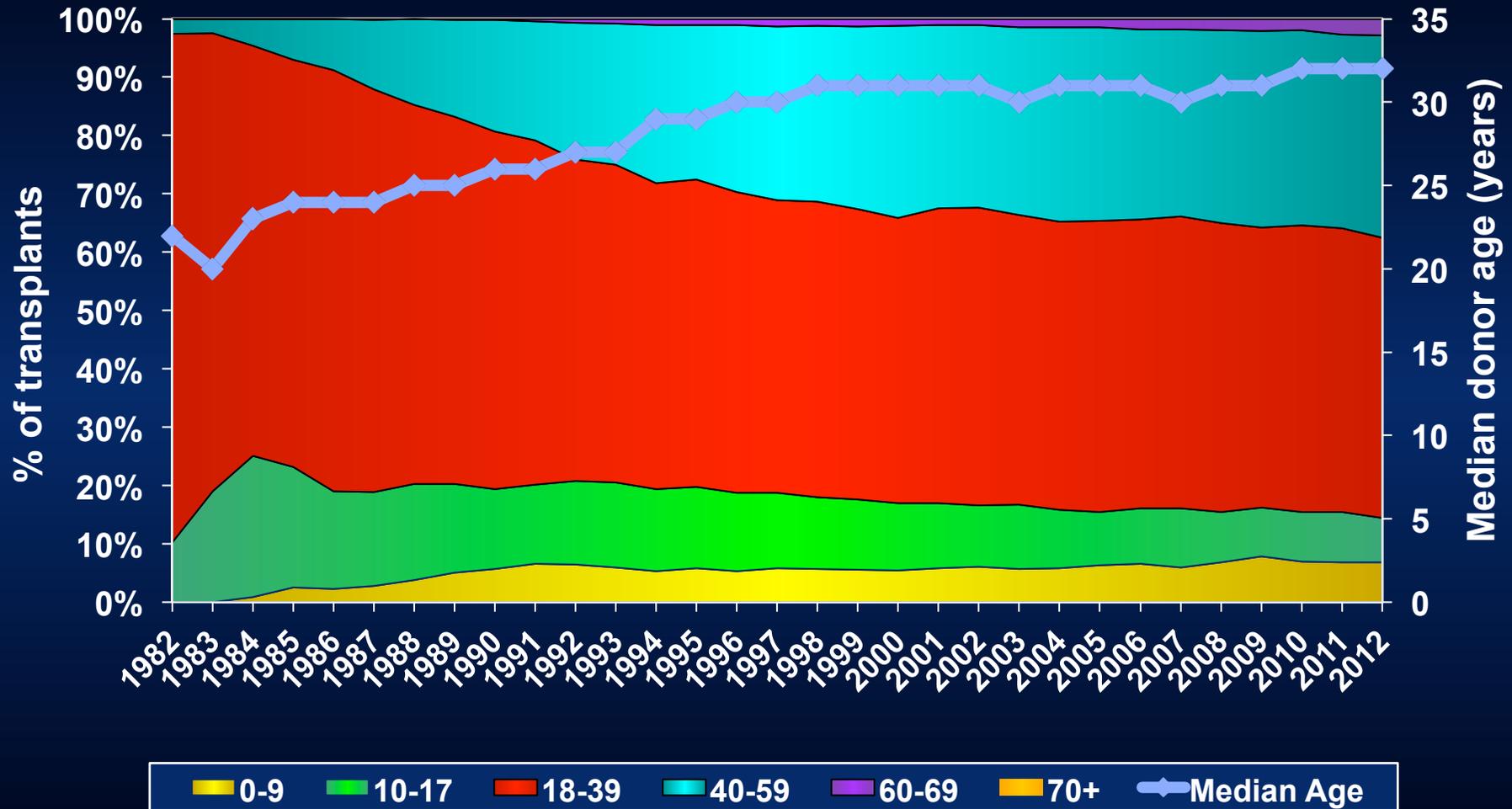
Number of Transplants by Year



NOTE: This figure includes only the heart transplants that are reported to the ISHLT Transplant Registry. As such, the presented data may not mirror the changes in the number of heart transplants performed worldwide.



Adult and Pediatric Heart Transplants Donor Age by Year of Transplant

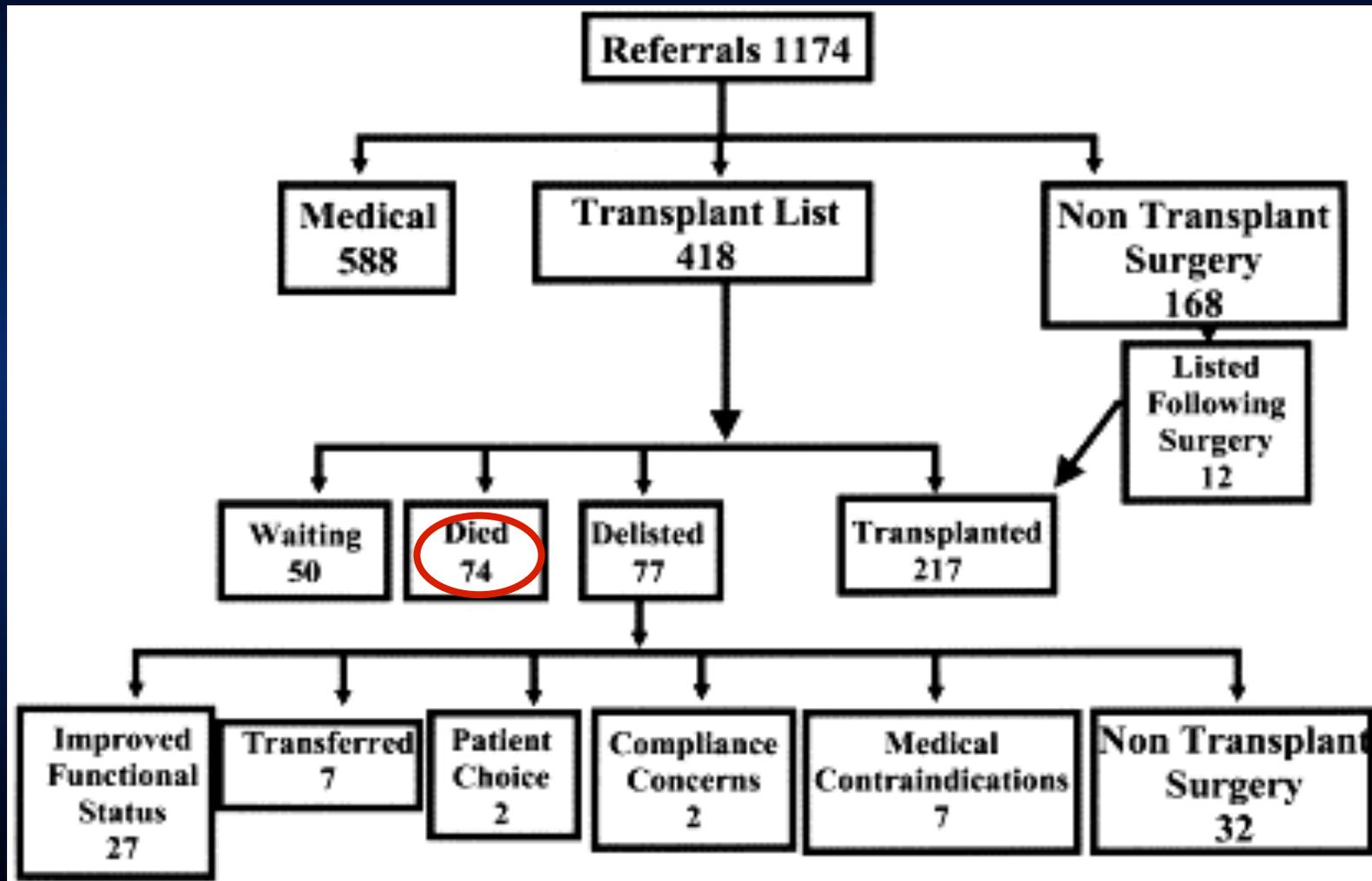


Comment

- HTx excellent options in end stage cardiac failure
- The better the pre-HTx clinical status the better results
- Early and late results related to comorbidities
- Donor shortage requires proper pts selection
- **Alternative** surgical procedures necessary :
 - to reduce the need of HTx
 - to delay it
 - to stabilize the clinical status
 - **instead of HTx**



”Listing a patient for transplantation is not an immediate therapy and waiting list mortality is 18%”

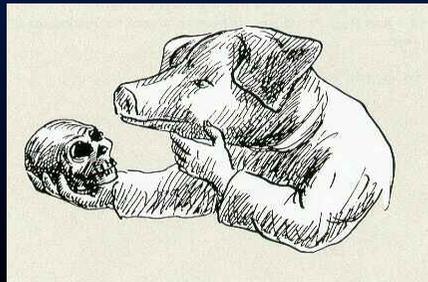
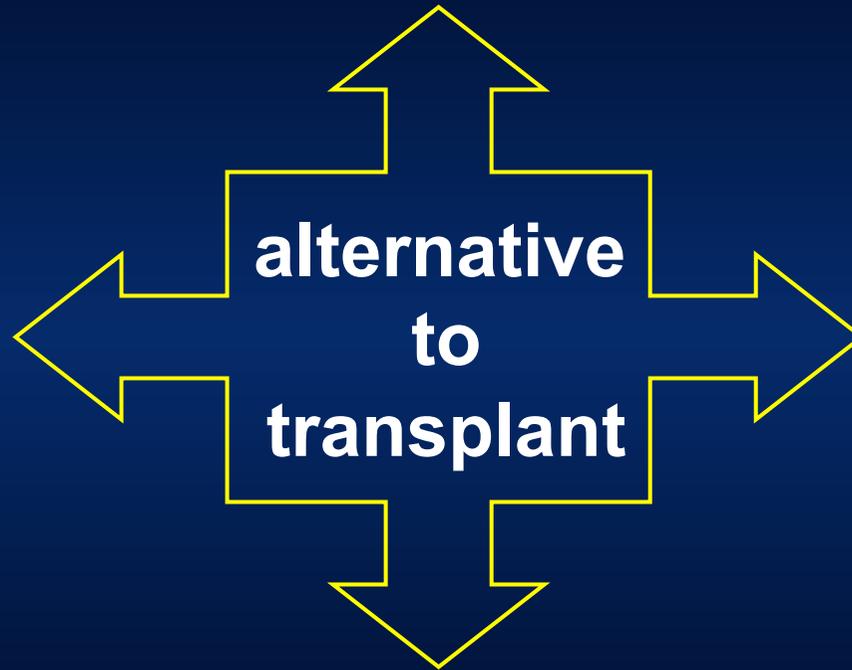
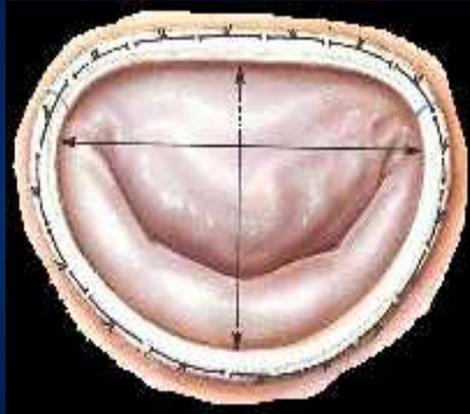
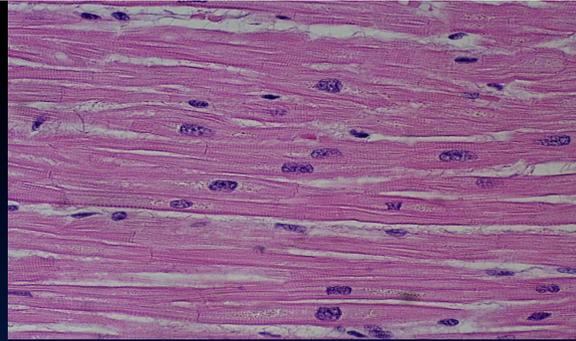


The advantages of the “alternative procedures” vs. HTx

- Availability*
- Pre - and perioperative “tune-up” feasible
- Standard post-op management
- No immunosuppression-related risks
- “Bridge” to HTx

* *HTx candidacy is not a therapy*





“Conventional” Alternatives to Transplant

“Riparare il cuore”

risk

- operation
- reoperation



risk

- Waiting list
- transplant



“Conventional” Alternatives to Transplant

- Better patients selection
- Pre-postoperative treatment improvements
- Safer surgical technique
- “*postcardiotomy*” mechanical support

LOWER OPERATIVE RISK



Severe Ischemic CMP (FE < 20-30%)

“Estreme” CABG Surgery

- technical improvements
- better myocardial protection
- appropriate patient selection

- | | |
|----------------------|---------|
| • Hospital mortality | 1,7-11% |
| • 5 years survival | 57-75% |



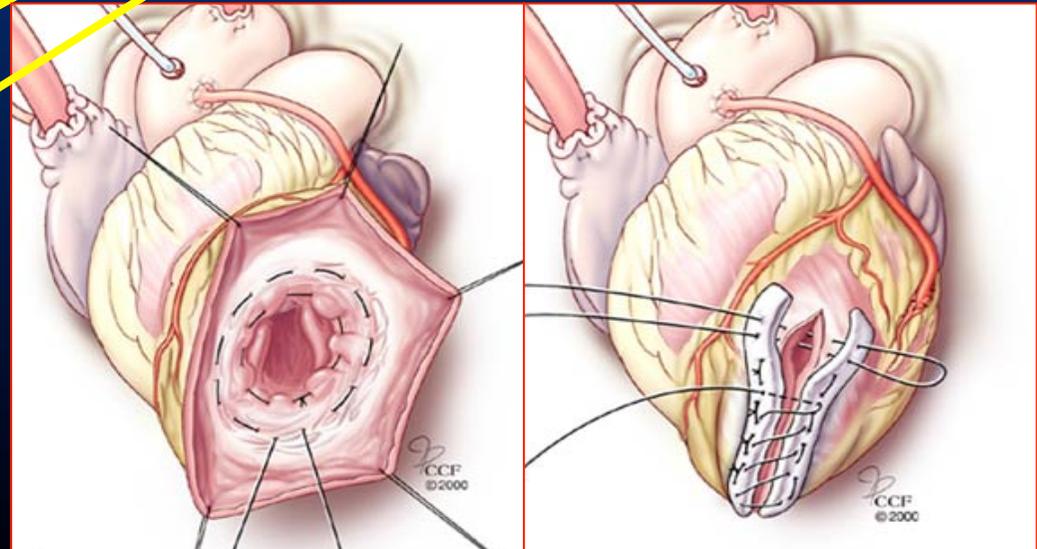
Severe Ischemic MCP

“Left Ventricular Restoration” (Dor)

Goals

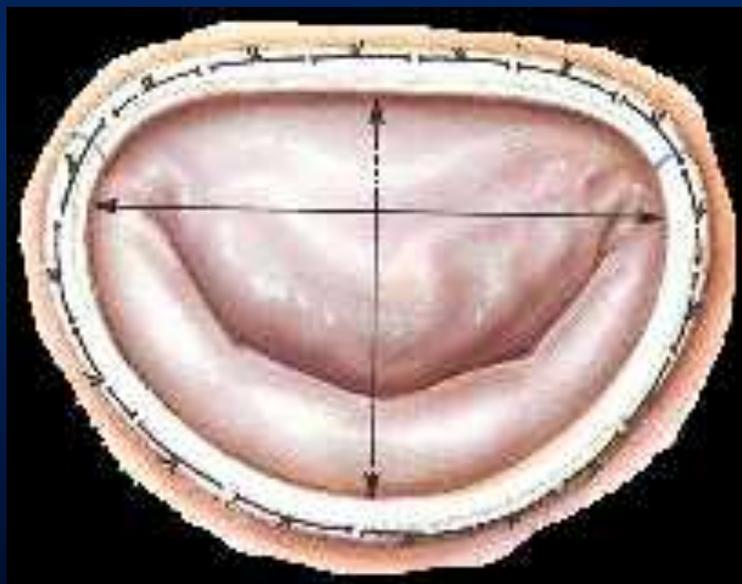
- ✓ functional infarcted area exclusion
- ✓ volume reduction
- ✓ LV reshaping

remodelling



Functional MR in ischemic or primitive dilatative cardiomyopathy

Undersizing Annuloplasty



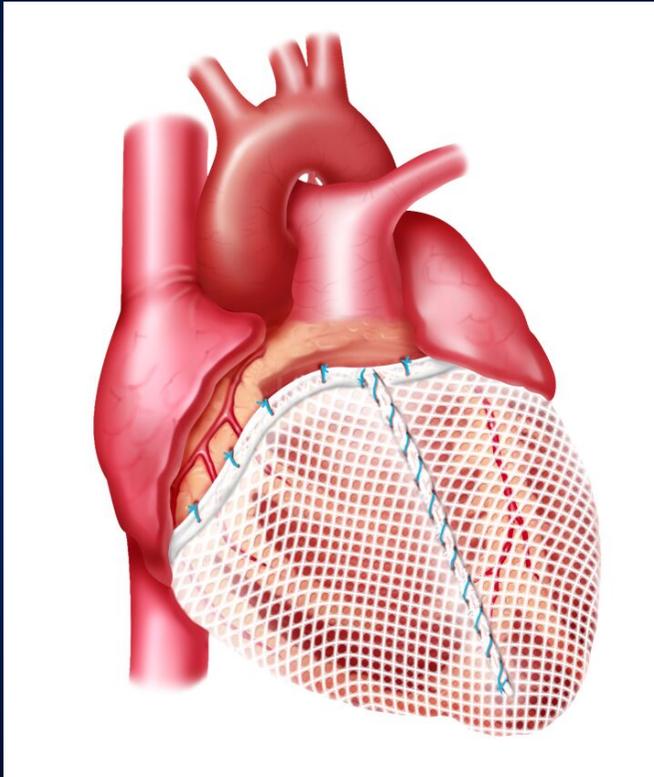
Mitral valve reconstruction for the correction of secondary MR in patients with end-stage dilated cardiomyopathy and refractory CHF has an acceptable 1-year mortality and results in improved symptomatic status accompanied by evidence of improvement in left ventricular shape and performance.

Bolling SF Am J Cardiol 1996



Cardiac Support Device (ACORN)

Goal



- LV passive end diastolic support reducing wall stress and myocardial *stretch*
- *reverse remodelling*
- recovery of cardiac function and functional capacity



ACORN CorCap

Randomized Multicentric Study

300 pts - LVEF 27%, LVEDV 72

Status	treated	not treated
better	38%	27%
worse	37%	45%

Improvements of quality of life, cardiac volumes and sphericity index.

No improvements of survival, NYHA class, LV EF



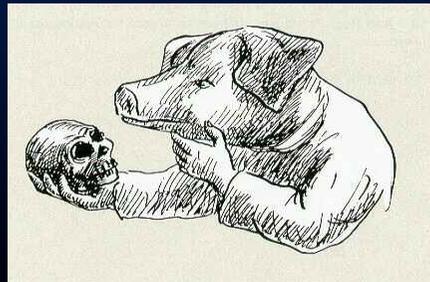
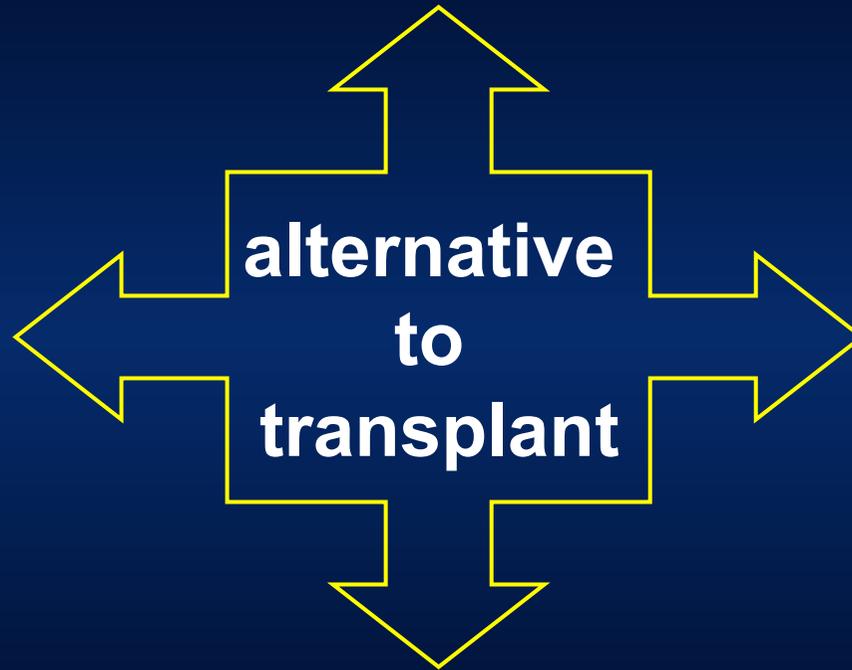
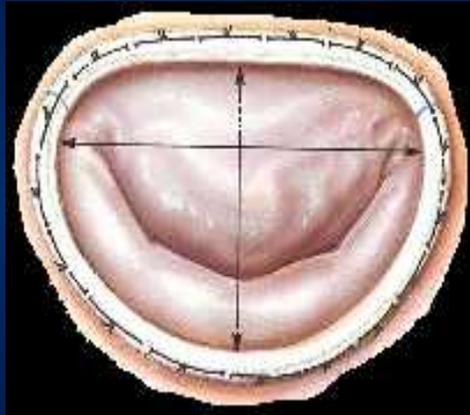
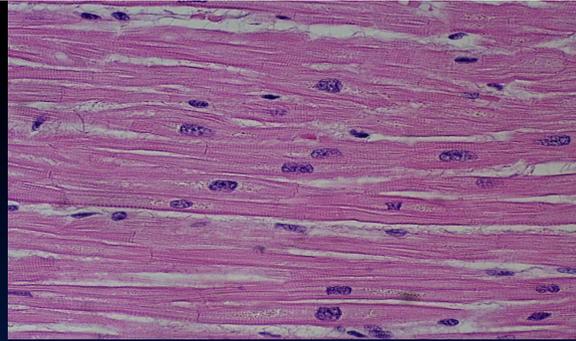
“Conservative” Surgery

In very sick pts not alternative to HTx

- mitral annuloplasty
- diastolic support passive device
- LV surgical “Reshaping”
- “extreme” CABG

results related to severity of disease





Stem cells therapy

“Rigenerare il cuore”

Goal

- Regeneration of cardiac myocytes
- neovascularization in infarcted area

Delivery

- intracoronary
- Transmiocardial
- bone marrow mobilization

Results

- Partial improvements
- Risk

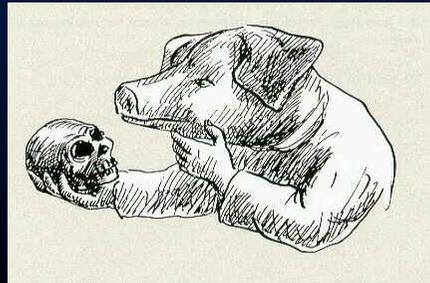
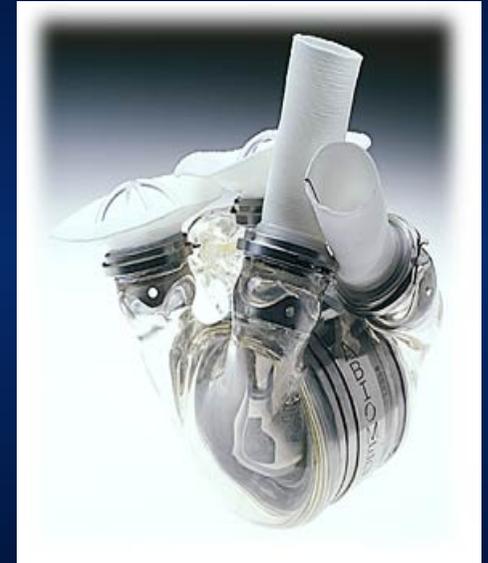
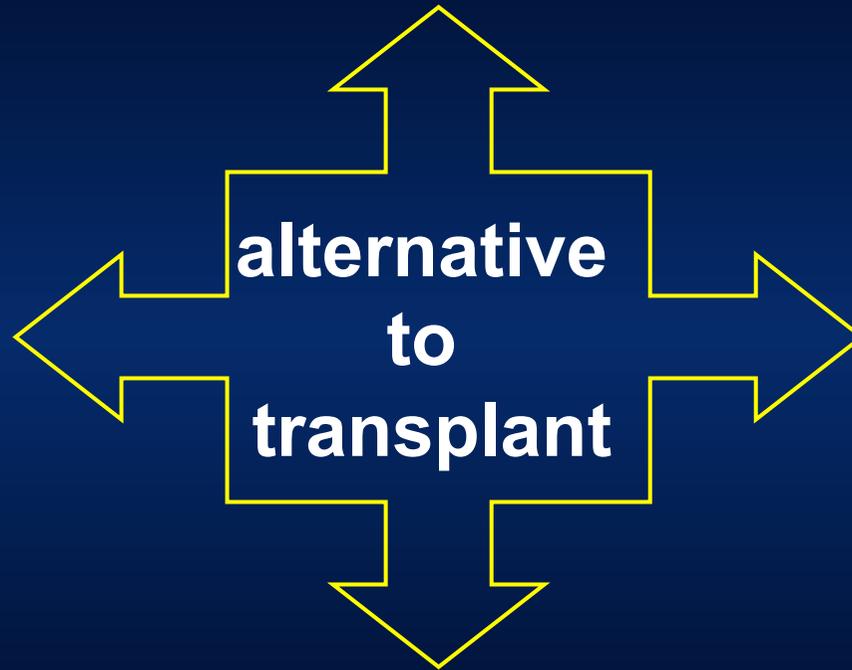
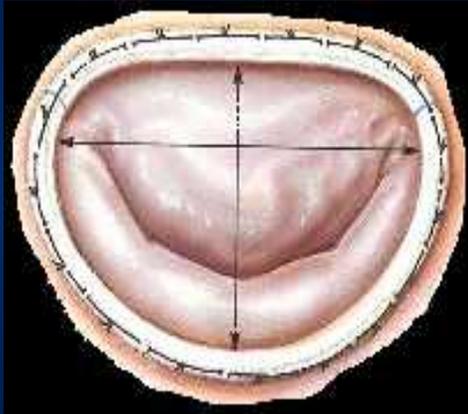
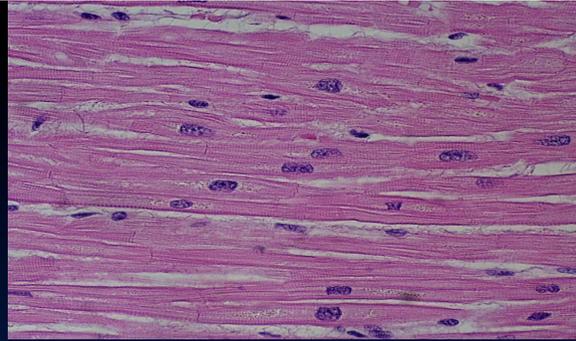
NOT AVAILABLE

Problems:

Type of cells/delivery

Patient selection





Xenotransplant

Pros

- Availability

Cons

- Anatomic problems
- Immunological aspects
- Infections (retrovirus)
- Ethics issues

Future options

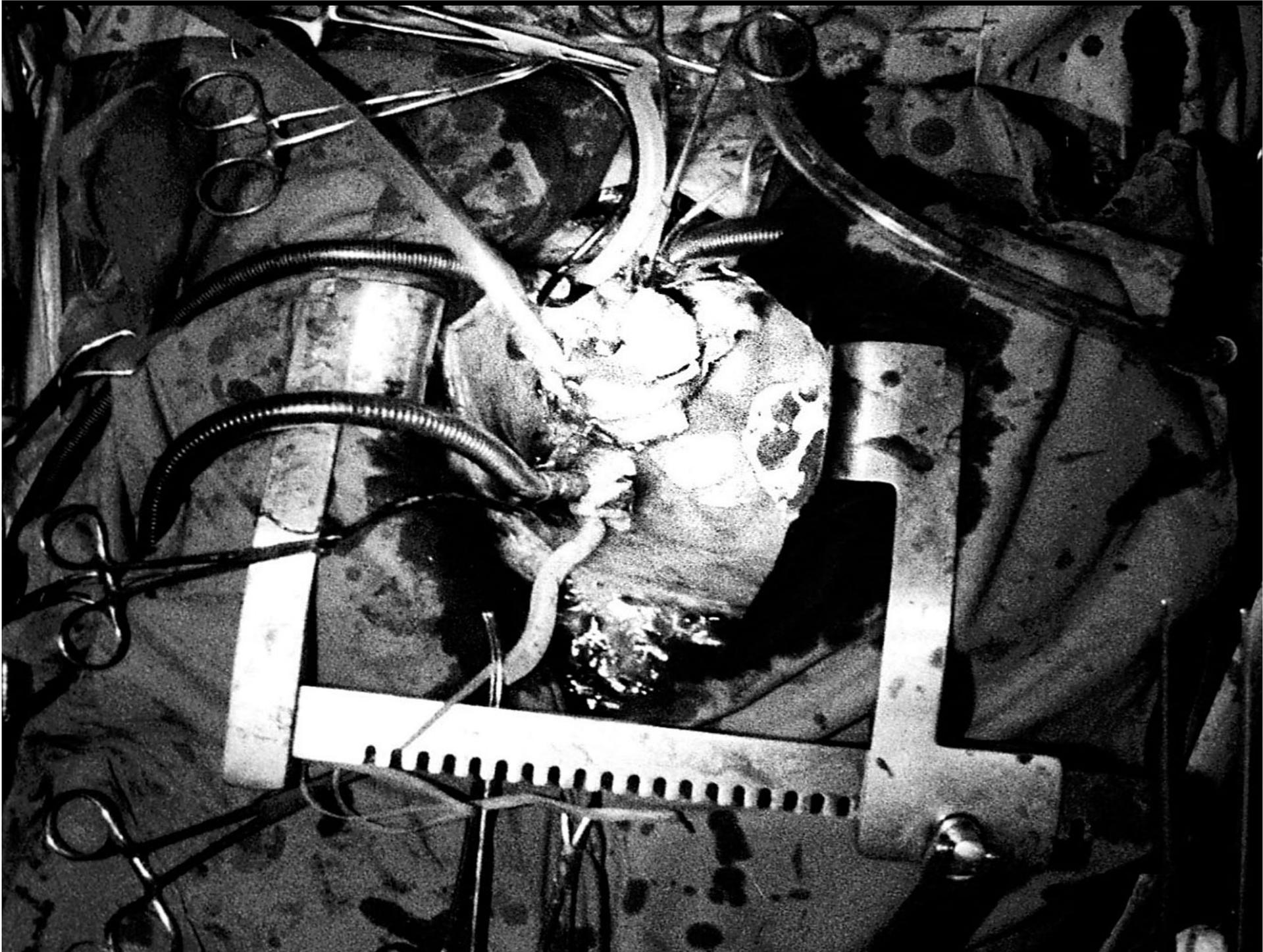
- Transgenic animals
- Immunosuppression

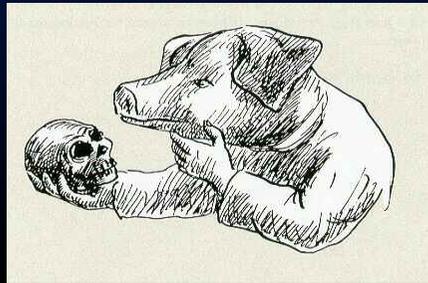
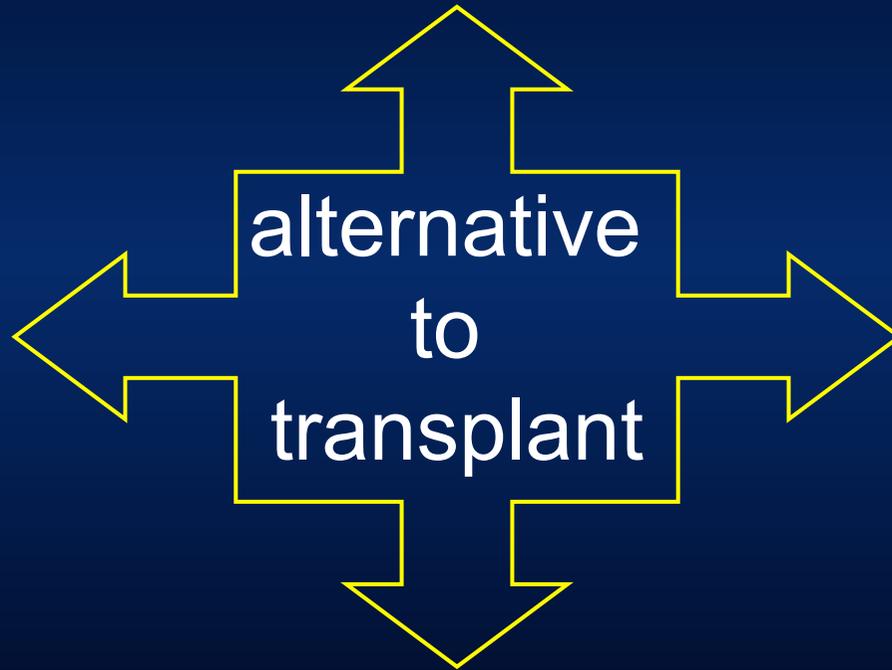
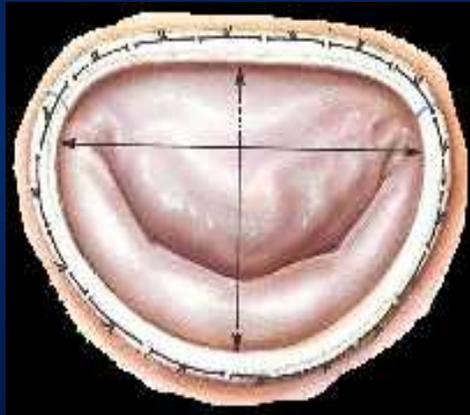
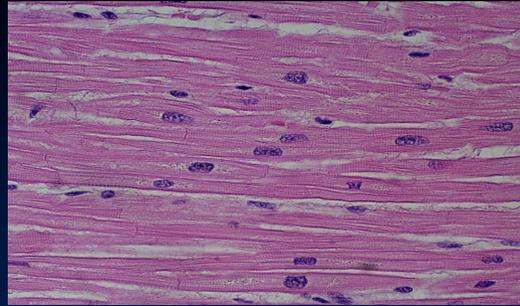


Hypothetical Indications:

bridge to recovery/Tx
contraindications to VAD







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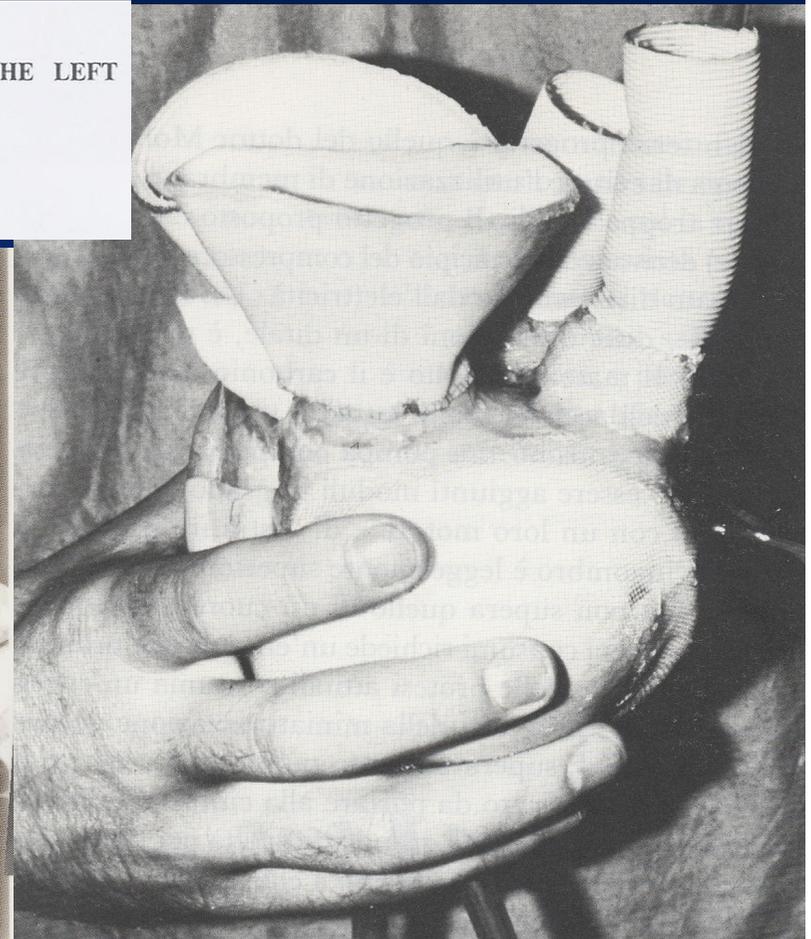
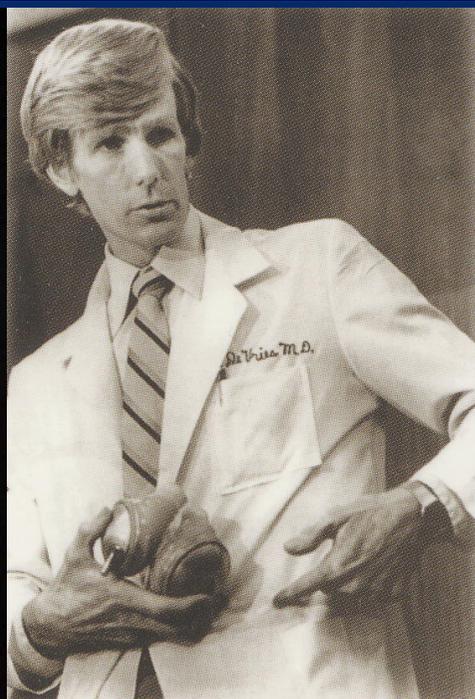
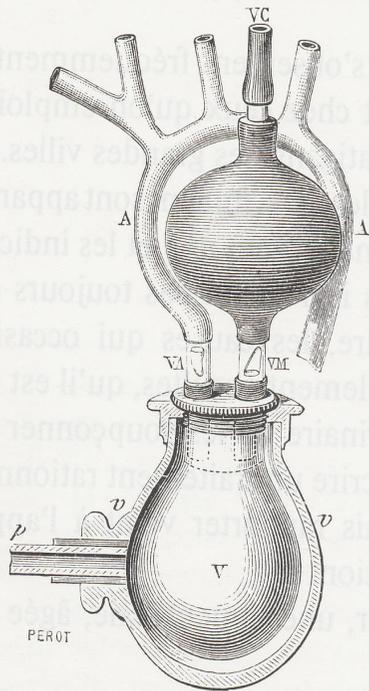
Number 5

CLINICAL USE OF THE TOTAL ARTIFICIAL HEART

WILLIAM C. DeVRIES, M.D., JEFFREY L. ANDERSON, M.D., LYLE D. JOYCE, M.D., FRED L. ANDERSON, M.D.,
ELIZABETH H. HAMMOND, M.D., ROBERT K. JARVIK, M.D., AND WILLEM J. KOLFF, M.D., PH.D.

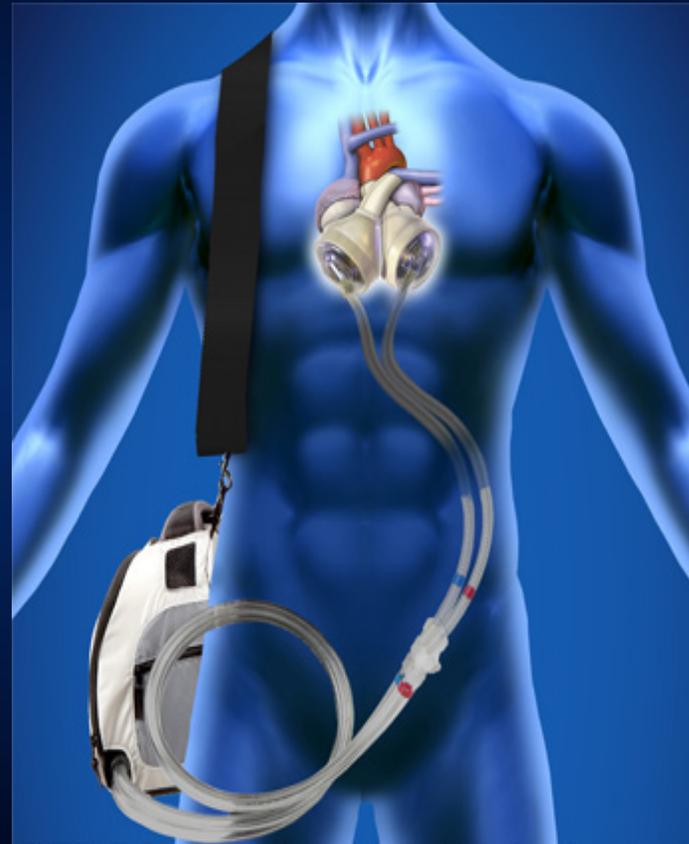
TEMPORARY MECHANICAL SUBSTITUTE FOR THE LEFT VENTRICLE IN MAN

Forest D. Dodrill, M.D., Edward Hill, M.D.
and
Robert A. Gerisch, M.D., Detroit



Total Artificial Heart (TAH)

- totally implantable
- replacing native heart
- biventricular support
- thromboembolism
- short-medium term supp.
- bulky system
- costs



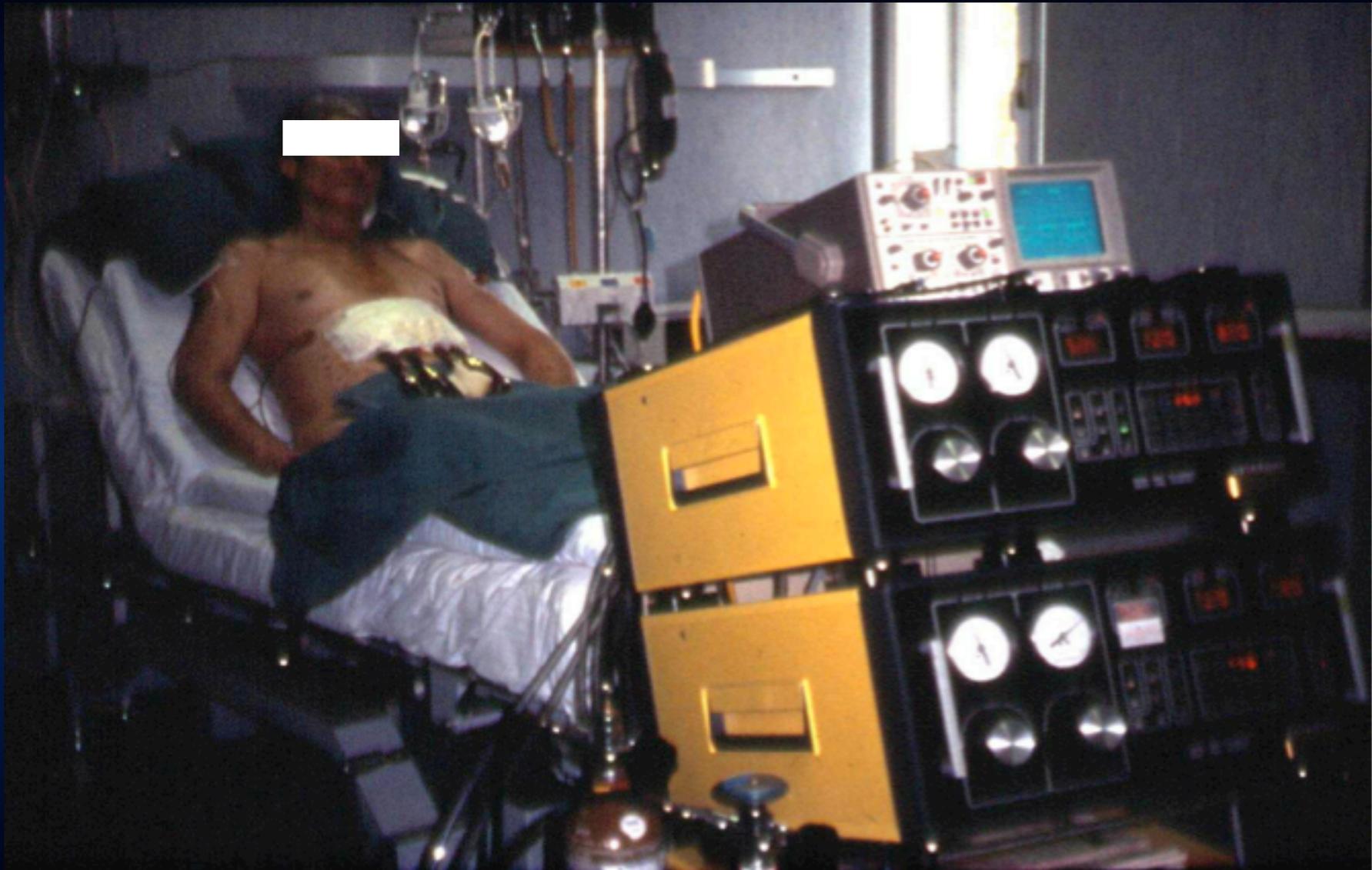
Ventricular Assist Device (VAD)

Mechanical system supporting a failing heart to optimize hemodynamic status and end-organs perfusion:

- pump
- inflow and outflow cannulas
- control panel
- power source

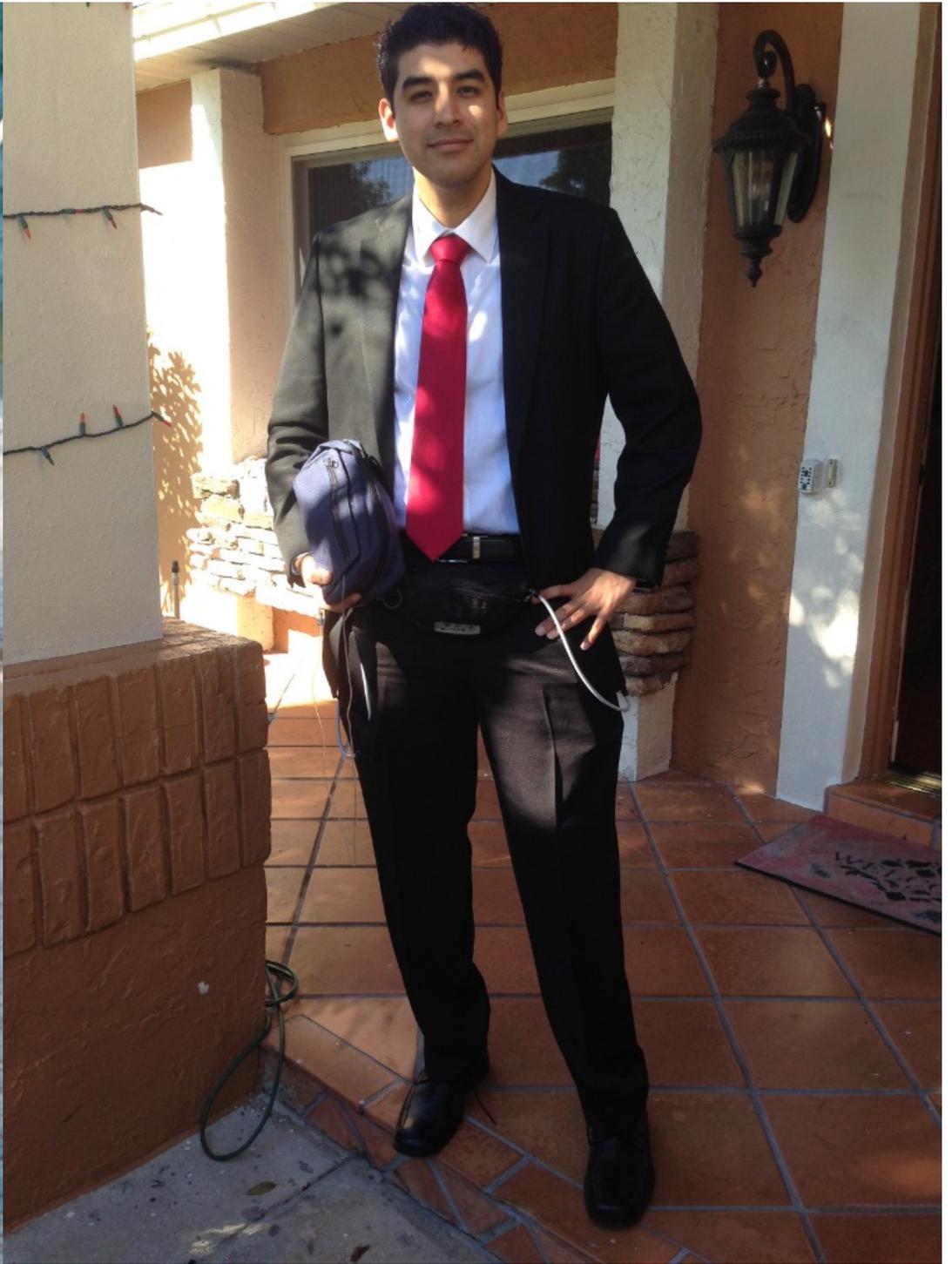












Indications to VAD

chronic heart failure

acute heart failure



Destination Therapy

- Not HTx eligible
- No contraindications to VAD
- Class IIIB or IV CHF (No cardiogenic shock)
- Inotropes dependent
- Maximal medical therapy with $\text{VO}_2 < 10$ ml/kg min
- No comorbidities with life expectancy < 2 yy
- No neurological impairment
- No psychosocial problems



.....beyond the imagination

VAD instead of Transplant

Small dimensions → *Easy to implant*

Length of support → *Years* → *destination*

Biocompatibility → *No anticoagulation*

Totally Implantable → *No transcut. cable*

Associated Therapy → ***Genetic or stem cells***





Conclusions

- HTx still the gold standard for severe CHF
- in selected patients, “conventional” surgery effective in replace or delay HTx
- stem cells are not a reality so far
- increasing role of mechanical support as “bridge” to Htx or as “destination therapy” or as tools for donor organ retrieval .





*The surgeon that
think to operate the
heart is not worthy
of respect by his
Colleagues*

T. Billroth - 1875



