

# The Future of Living Donor Transplantation

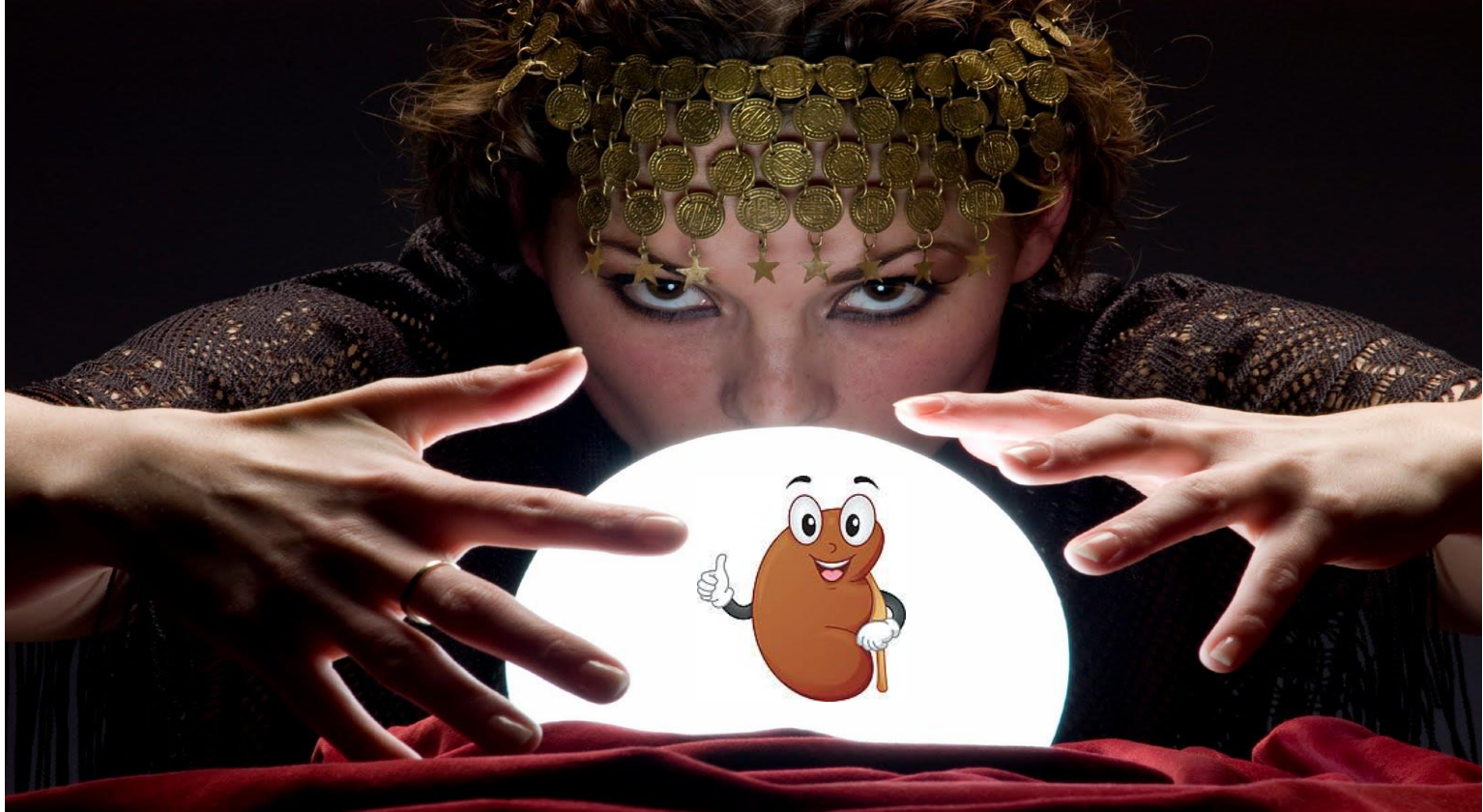


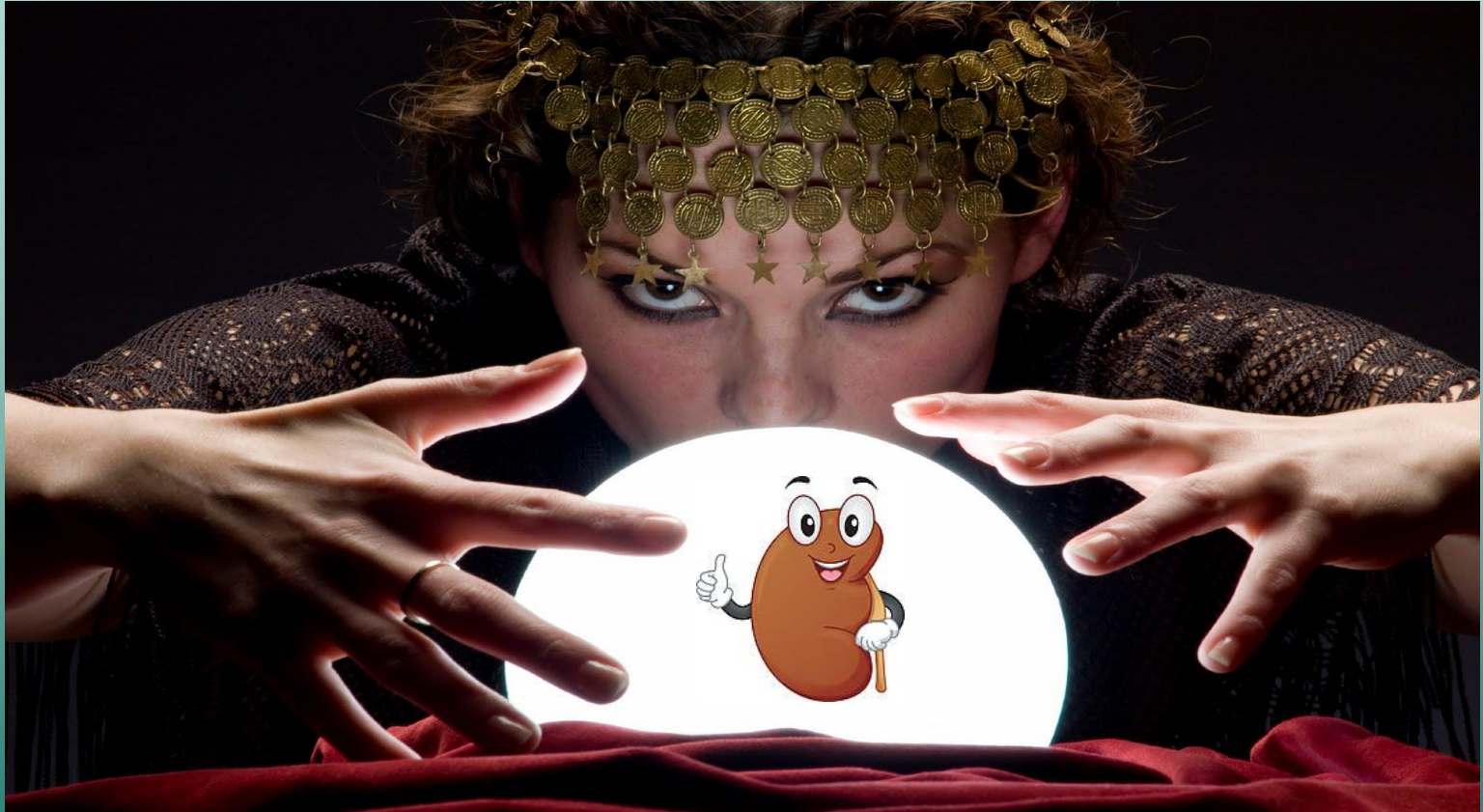
Matthew Cooper, MD  
Chief of Transplantation  
Director, Solid Organ Transplant Line  
Professor of Surgery  
Medical College of Wisconsin  
Past President, OPTN/UNOS  
President, AFDT



# Disclosures

- Surgical Director and Medical Advisor – National Kidney Registry





**Disclaimer!!**



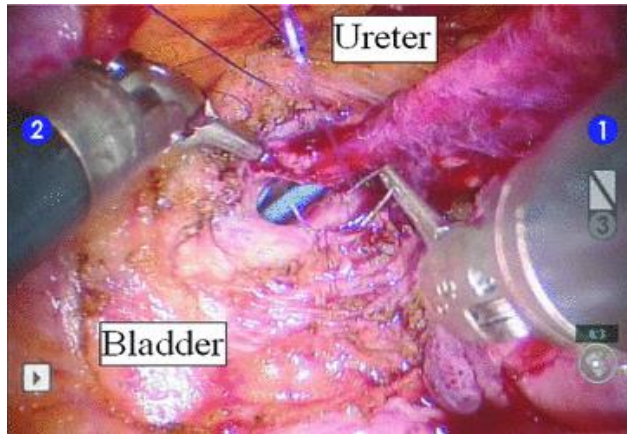
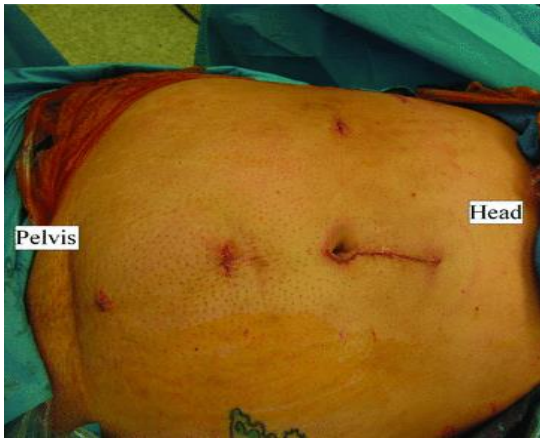
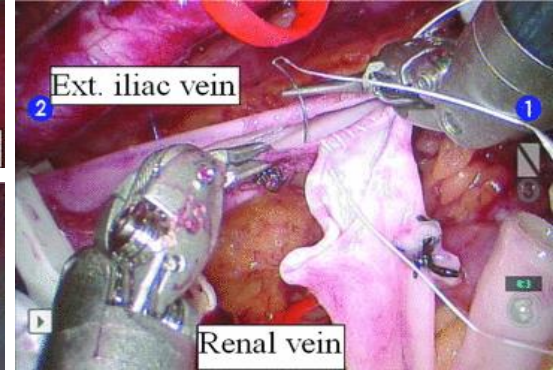
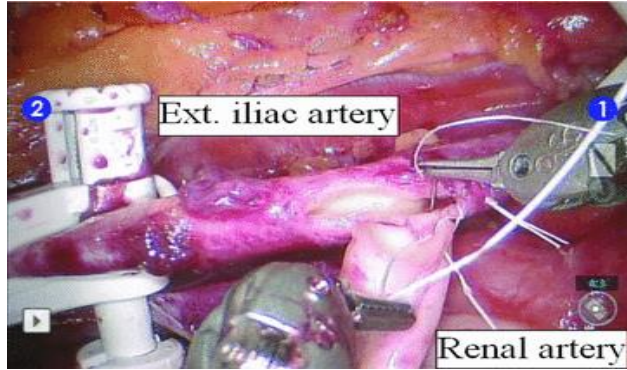
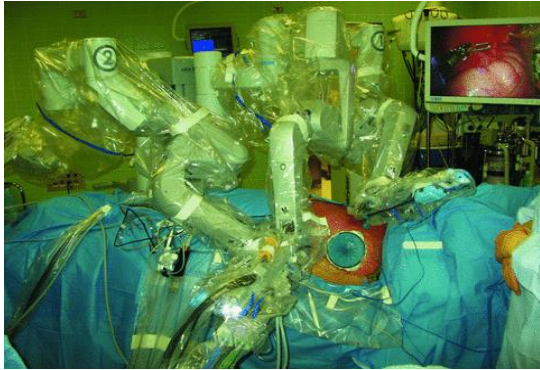
# Robotics – Donors and Recipients



# Robotic Trans-abdominal Kidney Transplantation



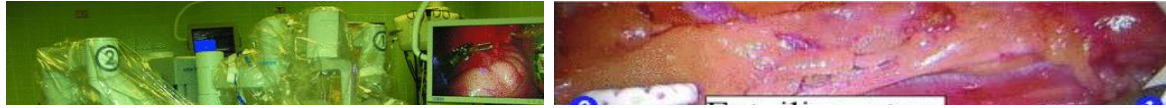
# Robotic Trans-abdominal Kidney Transplantation



Giulianotti et al: AJT 2010, 10(6):1478-1482



# Robotic Trans-abdominal Kidney Transplantation



## Robotic Assisted Living Donor Nephrectomies A Safe Alternative to Laparoscopic Technique for Kidney Transplant Donation

Spaggiari, Mario MD\*; Garcia-Roca, Raquel MD†; Tulla, Kiara A. MD\*; Okoye, Obi T. MD\*; Di Bella, Caterina MD\*;  
Oberholzer, José MD‡; Jeon, Hoonbae MD§; Tzvetanov, Ivo G MD\*; Benedetti, Enrico MD\*

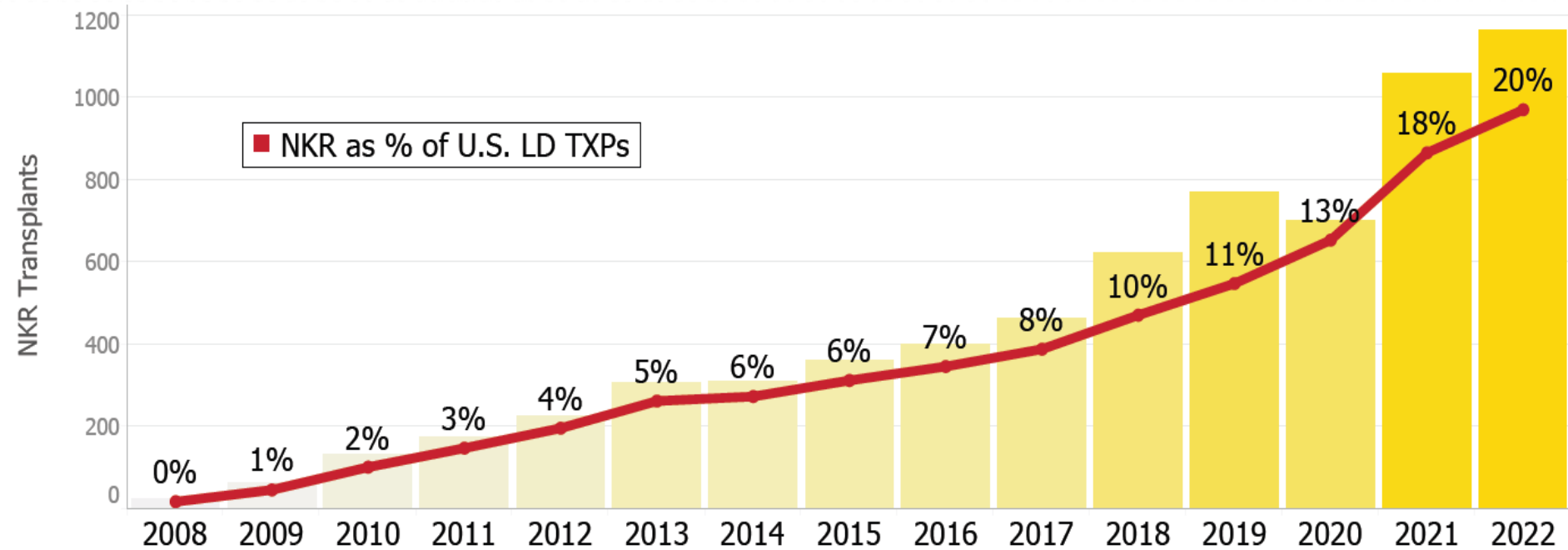


Giulianotti et al: AJT 2010, 10(6):1478-1482





# Mandatory Paired Kidney Exchange Options



NKR Facilitated TXPs*	22	61	131	175	226	308	310	360	399	462	621	770	700	1,058	1,164
U.S. LD TXPs**	5,968	6,387	6,278	5,773	5,619	5,733	5,538	5,628	5,629	5,811	6,442	6,867	5,234	5,974	5,863

# The NKR Voucher Program

Potential donors could be incompatible with their intended recipient based on:

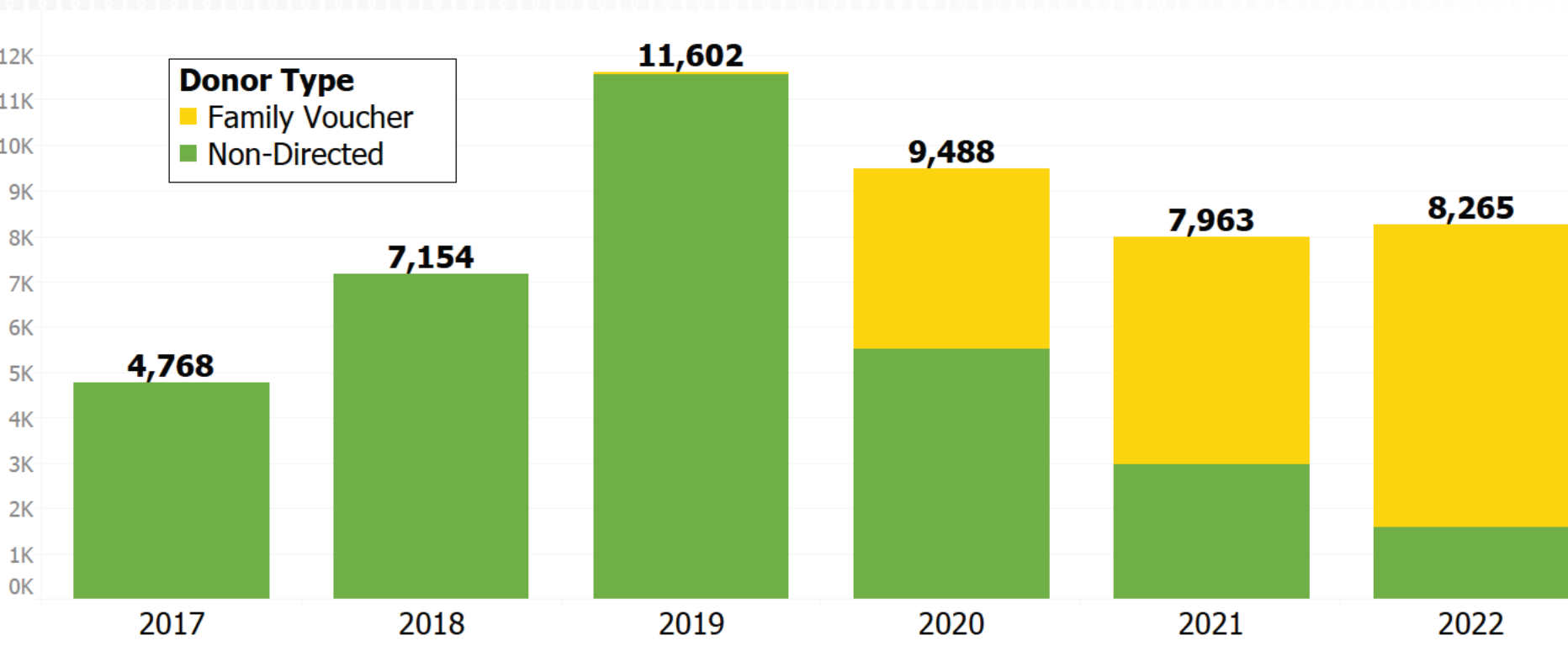
- Blood Type
- DSA
- Anatomy
- Time



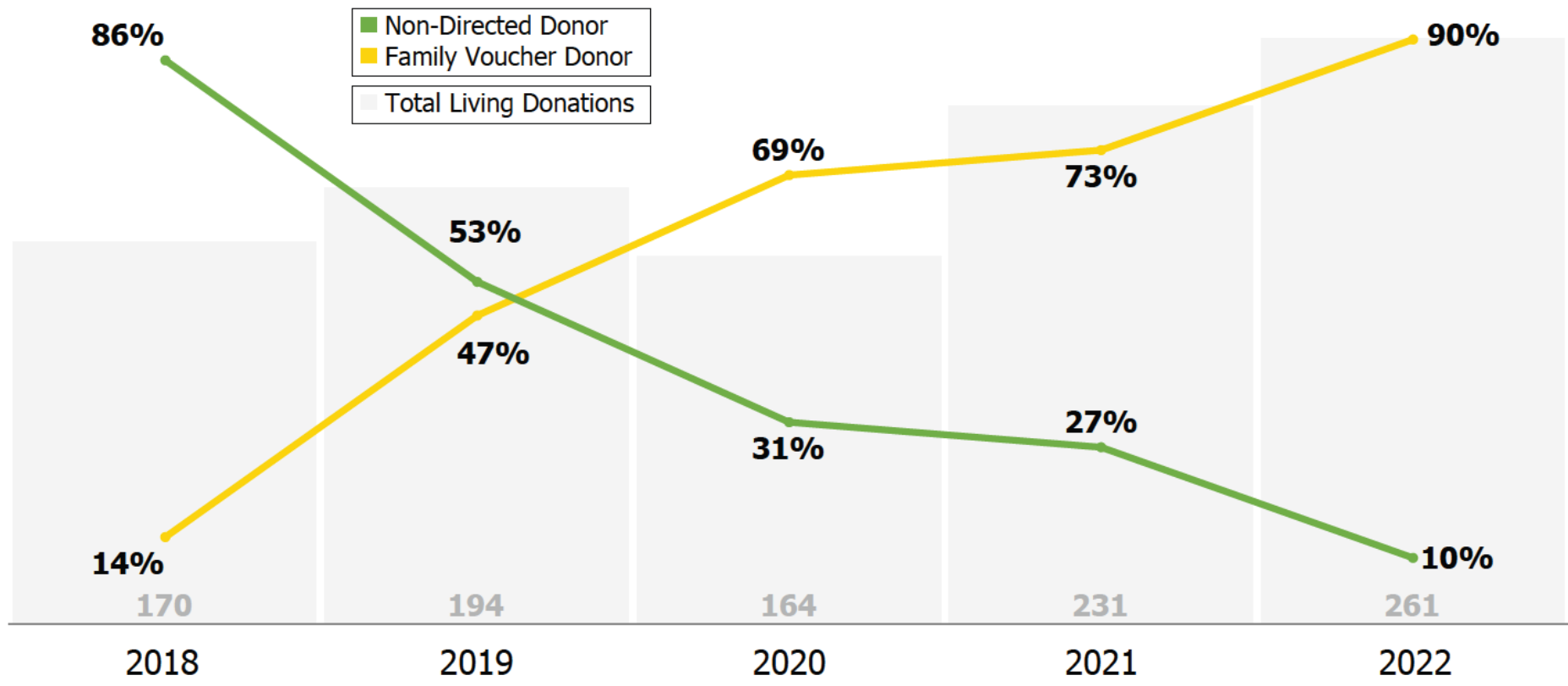
First 'voucher donor' Judge Broadman and first 'voucher holder' his grandson Quinn (UCLA 2014)

**Family Voucher** allows identification of up to 5 immediate family members to hold vouchers if EVER needing kidney!

# Non-Directed Donors as a National Resource

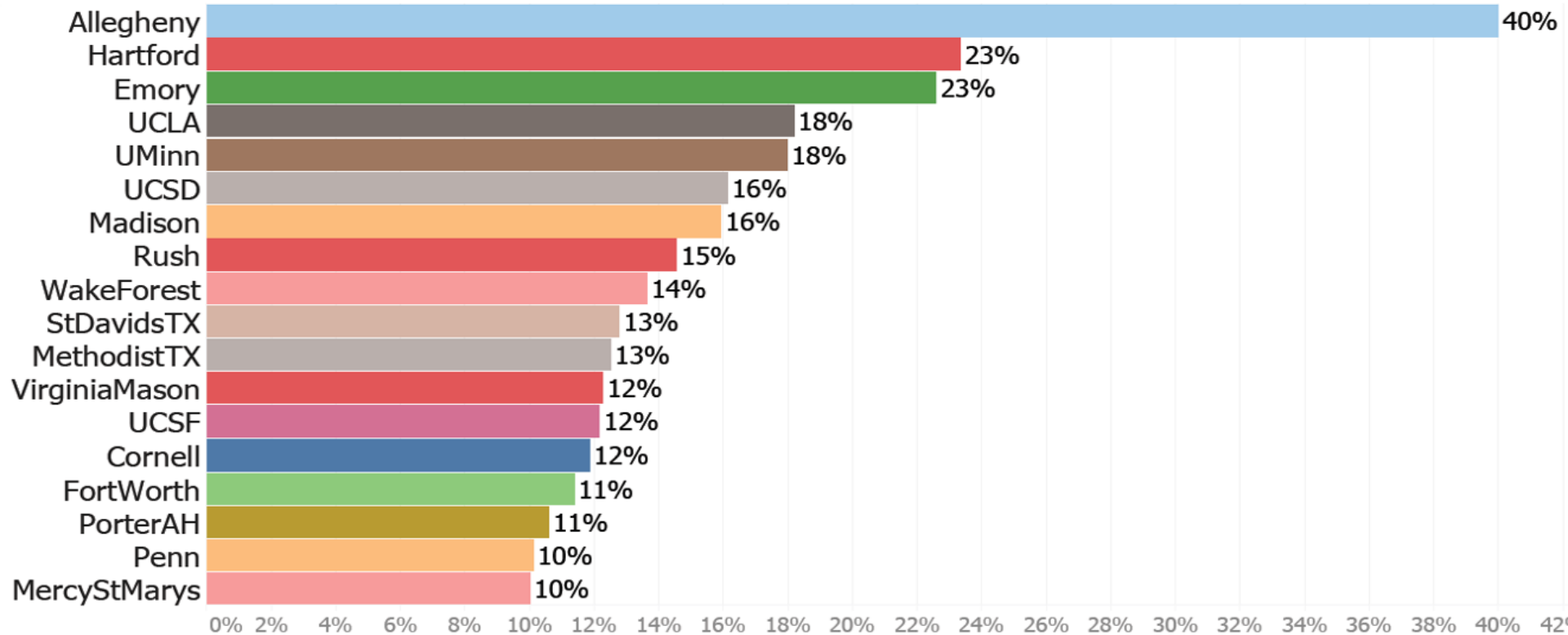


# Transition to Family Voucher Donation



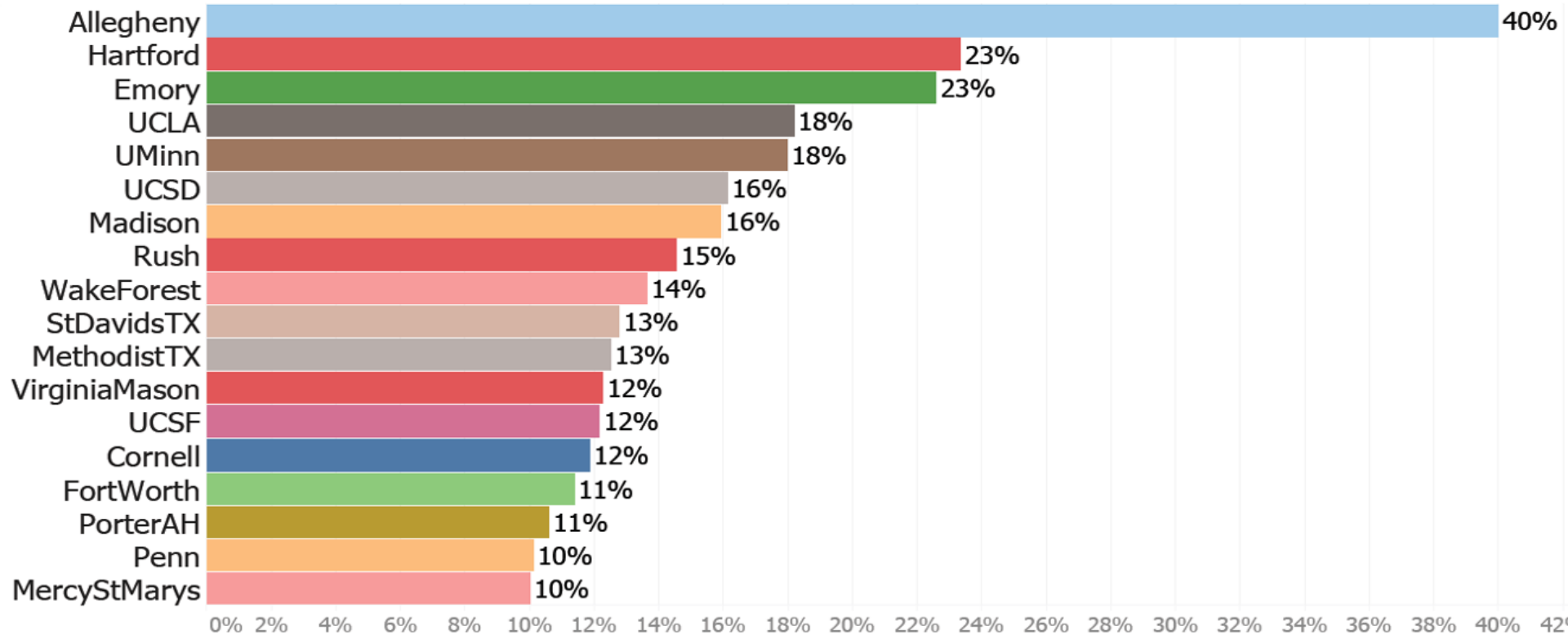


# LD Conversion Rates Becomes a Metric

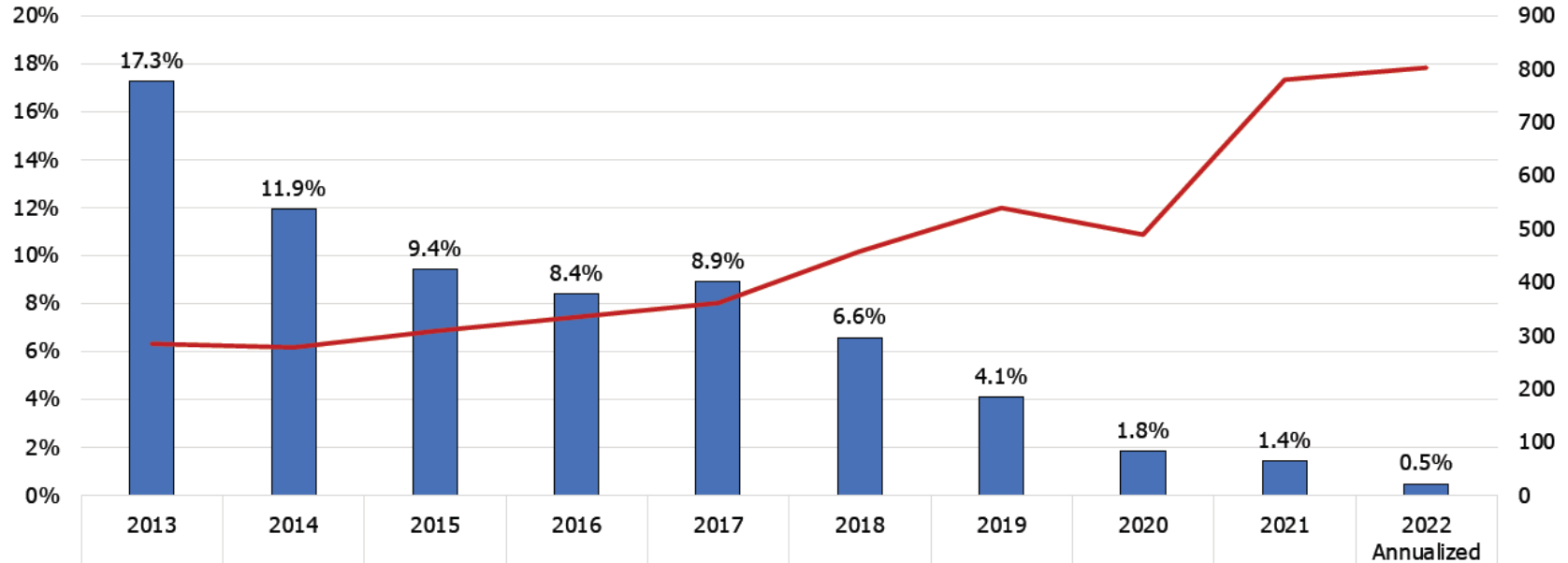




# LD Conversion Rates Becomes a Metric



# NKR Desensitization Cases Decline as Volume Grows



Desensitization Cases	49	33	29	28	32	30	22	9	11	4
NKR Transplants	284	277	308	334	360	457	539	489	779	802

■ Desensitization Rate    — NKR Transplants

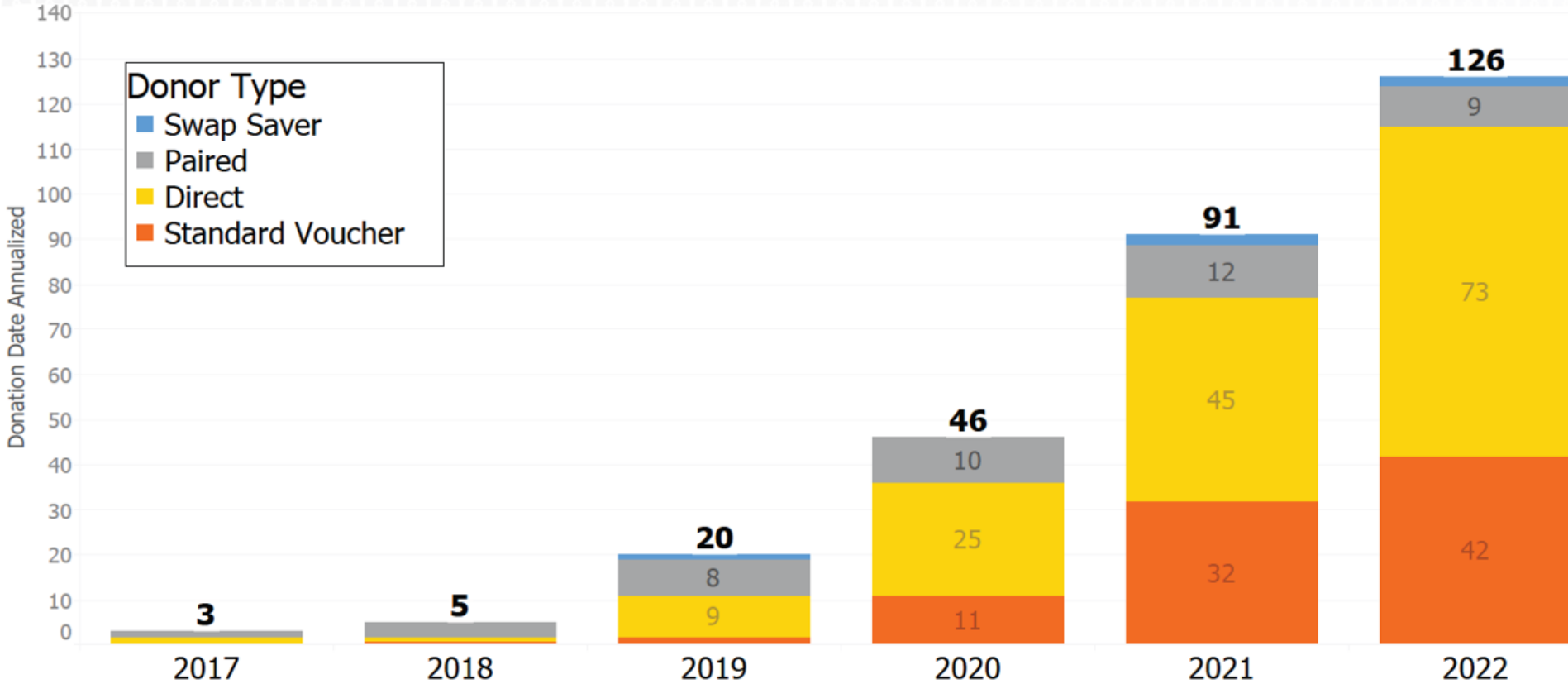


# Using KPD to Transplant the 'Untransplantable'

## Desensitization

		EASY	HARD
KPD	EASY	Try KPD for a few months If match -> KPD If no match -> <u>Desens.</u>	Wait in KPD
	HARD	Look in KPD pool <i>Prob. Not Worth Waiting</i> If match -> KPD If no match -> Desens.	<b>COMBINE</b> <b>KPD and Desensitization</b>

# Remote Donations → Donor Convenience

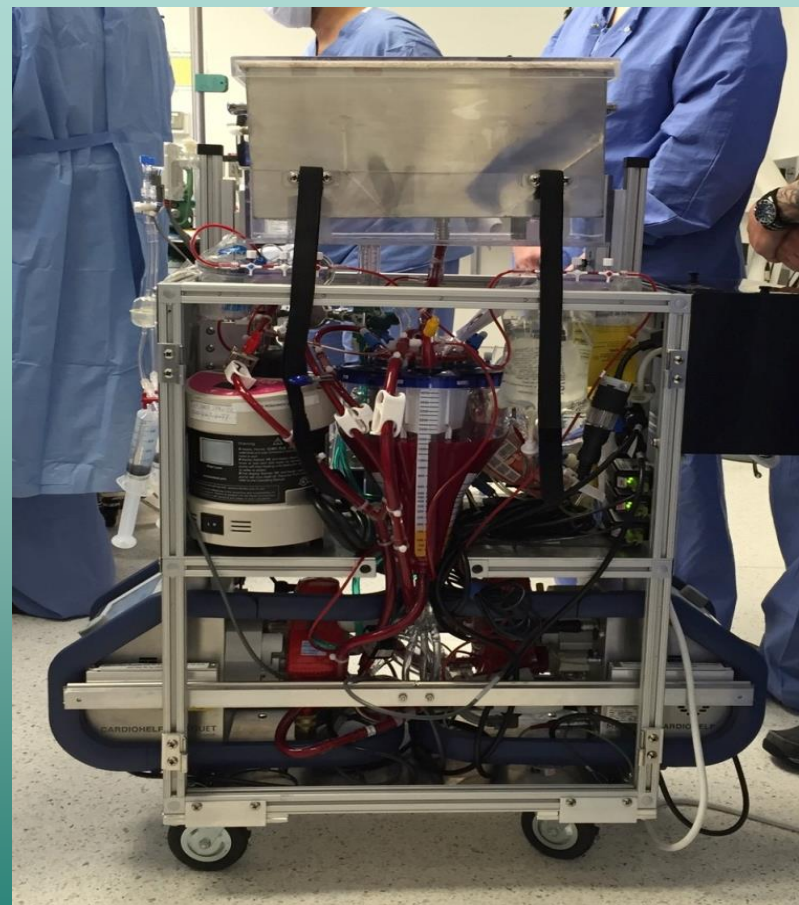
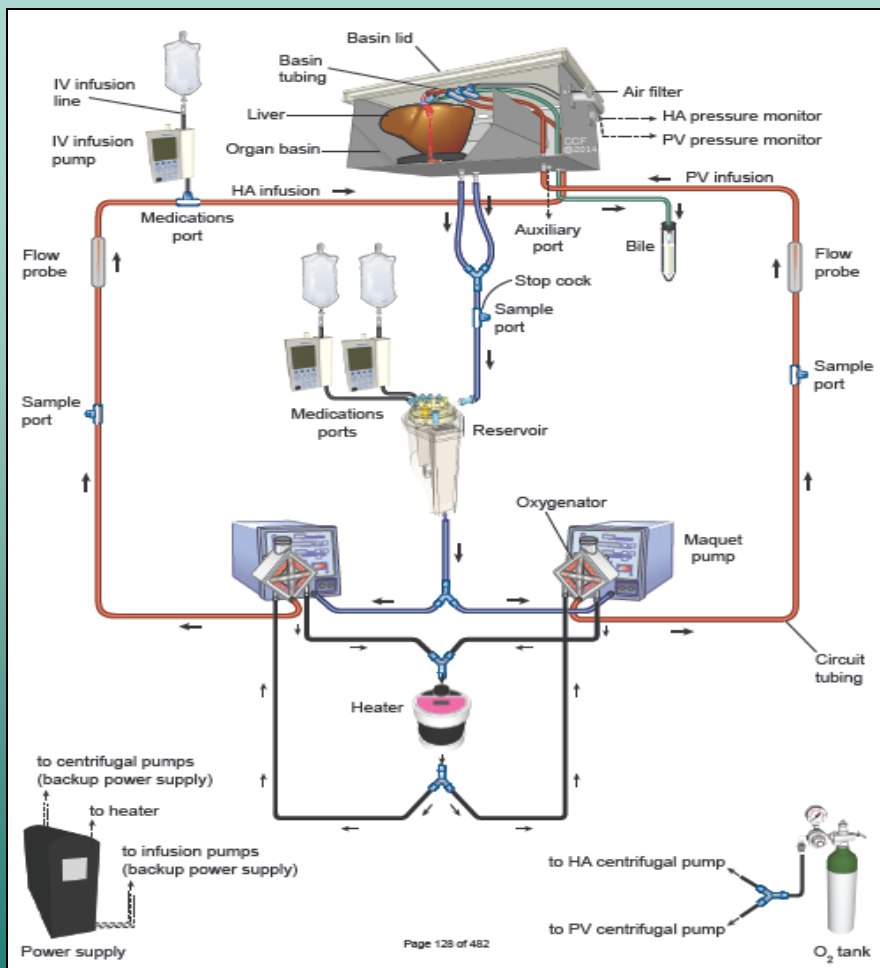


# Normothermic Preservation



Vs







# Compatible Pairs → Optimize Outcomes

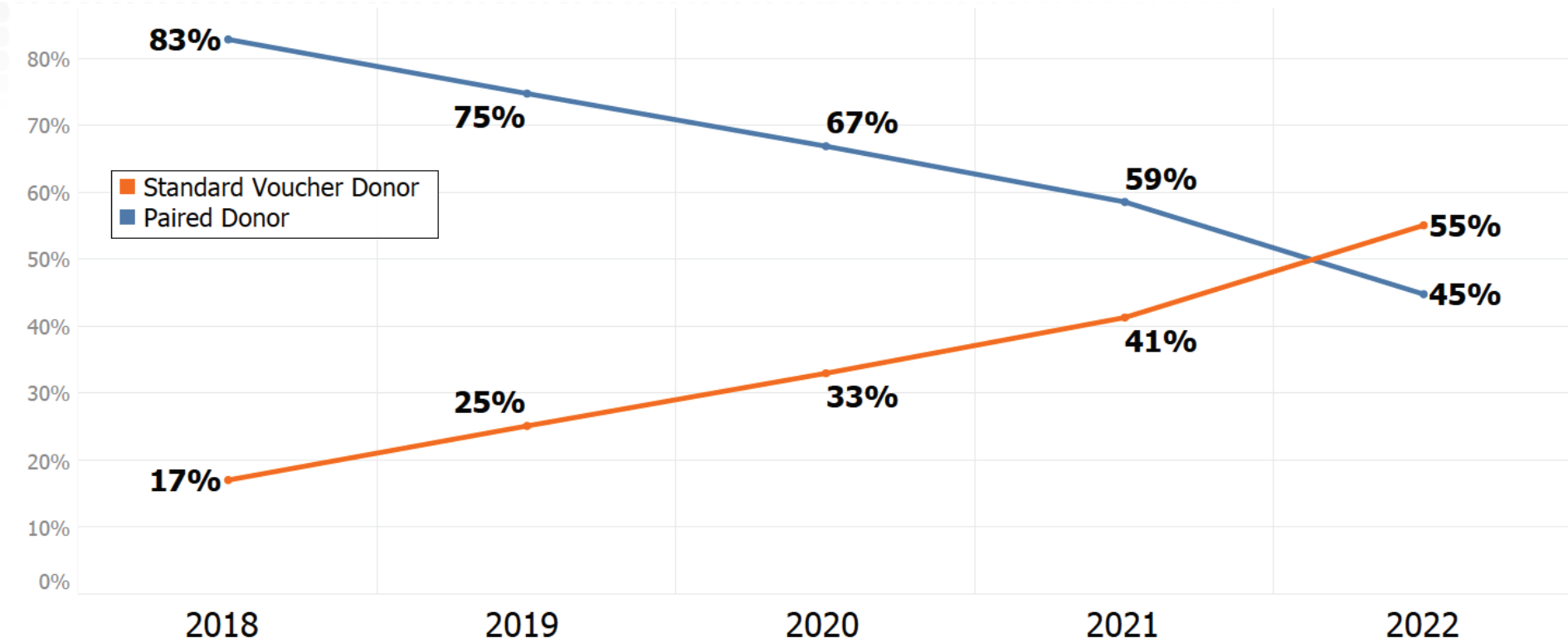
- What a compatible pairs gain:
  - Younger donor
  - Better size match (nephron mass)
  - O-mismatch
  - Ability to be one another's support
  
- What the incompatible pool gains if compatible pairs participate:
  - Higher match rates
  - More O-recipients match



get it How to Find a Kidney Donor

# Decoupling Trend

Transition from Paired to Standard Voucher Donation



# www.transplantmodels.com

The Epidemiology Research Group for Organ Transplantation is a research group focused on organ transplantation at the Johns Hopkins School of Medicine. Below are some of the decision models we have developed.

For more information, please visit our website, [www.transplantepi.org](http://www.transplantepi.org)

## Living Kidney Donor Risk Index (LKDPI)

This model predicts recipient risk of graft loss after living donor kidney transplantation based on donor characteristics, on the same scale as the KDPI ...

Massie AB, Leanza J, Fahmy LM, Chow EK et al. A Risk Index for Living Donor Kidney Transplantation. AJT 2016 (epub ahead of print)

[Continue to model »](#)

## Transplant Candidacy for Patients 65+

This prediction model is intended for adults with ESRD on dialysis aged 65 and above; it provides the predicted probability of 3-year survival after kidney transplantation (KT). Patients with predicted 3-year post-KT survival in the top quintile are deemed "excellent" candidates ...

Grams M E., Kucirka L M., Hanrahan C F., Montgomery R A., Massie A B., & Segev D L. (2012). Candidacy for kidney transplantation of older adults. *Journal of the American Geriatrics Society*, 60(1), 1-7.

[Calculate your score »](#)

## ESRD Risk Tool for Kidney Donor Candidates

This model is intended for low-risk adults considering living kidney donation in the United States. It provides an estimate of 15-year and lifetime incidence of end-stage renal disease...

Grams ME, Sang Y, Levey AS, Matsushita K, Ballew S, Chang AR et al. Kidney-Failure Risk Projection for the Living Kidney-Donor Candidate. *NEJM* 2015 (epub ahead of print)

[Continue to model »](#)

## Pediatric Transplant: Living or deceased donor first?

Most pediatric kidney transplant recipients live long enough to require retransplantation. The most beneficial timing for living donor transplantation in candidates with one living donor is not clear...

Van Arendonk, K J., Chow, E. K., James, N. T., Orandi, B. J., Ellison, T. A., Smith, J. M., Colombani, P. M., & Segev, D. L. (2012). Choosing the Order of Deceased Donor and Living Donor Kidney Transplantation in Pediatric Recipients: A Markov Decision Process Model. *Am J Transplant*, 99(2):360-6.

[Continue to model »](#)

## Infectious Risk Donors

When a patient with end stage renal disease (ESRD) on the waitlist for a kidney is offered an Infectious Risk Donor (IRD) kidney, they need to decide whether they will accept the IRD kidney and the associated infectious risk, or if they will decline it and continue to wait for the next available infectious-risk free kidney ...

Chow, E. K. H., Massie, A. B., Muzaale, A. D., Singer, A. L., Kucirka, L. M., Montgomery, R. A., ... & Segev, D. L. (2013). Identifying appropriate recipients for CDC infectious risk donor kidneys. *American Journal of Transplantation*, 13(5), 1227-1234.

[Continue to model »](#)

## Postdonation Risk of ESRD in Living Kidney Donors

Risk estimation is critical for appropriate informed consent and varies substantially across living kidney donors.

Massie, Allan B., et al. "Quantifying Postdonation Risk of ESRD in Living Kidney Donors." *Journal of the American Society of Nephrology* (2017): ASN-2016101084.

[Continue to model »](#)

# www.transplantmodels.com

The Epidemiology Research Group for Organ Transplant at Johns Hopkins School of Medicine. Below are

For more information, please visit

Characteristic	aHR <sup>a</sup>	P Value
Men (at age 40)	1.88 (95% CI, 1.50 to 2.35)	<0.001
black race (at age 40)	2.96 (95% CI, 2.25 to 3.89)	<0.001
Age per 10 yr: nonblack	1.40 (95% CI, 1.23 to 1.59)	<0.001
Age per 10 yr: black	0.88 (95% CI, 0.72 to 1.09)	0.3
BMI per 5 kg/m <sup>2</sup>	1.61 (95% CI, 1.29 to 2.00)	<0.001
First-degree biologically related to recipient	1.70 (95% CI, 1.24 to 2.34)	<0.01

... the top quintile are deemed "excellent" candidates ...

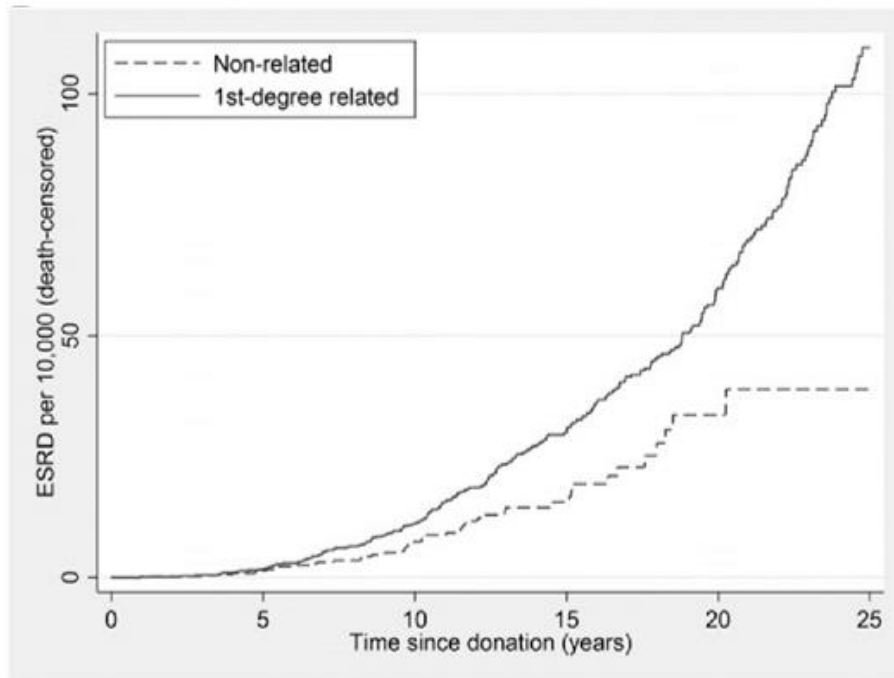
Grams, M. E., Kucirka, L. M., Hanrahan, C. F., Montgomery, R. A., Massie, A. B., & Segev, D. L. (2012). Candidacy for kidney transplantation of older adults. *Journal of the American Geriatrics Society*, 60(1), 1-7.

Calculate your score »

... one living donor is not clear

Van Arendonk, K. J., Chow, E. K., Smith, J. M., Colombani, P. M., & ... of Deceased Donor and Living Donor Recipients: A Markov Decision Process

Continue to model »



The Epidemiol  
Jc

### Characteristic

Men (at age 40)

black race (at age 40)

Age per 10 yr: nonblack

Age per 10 yr: black

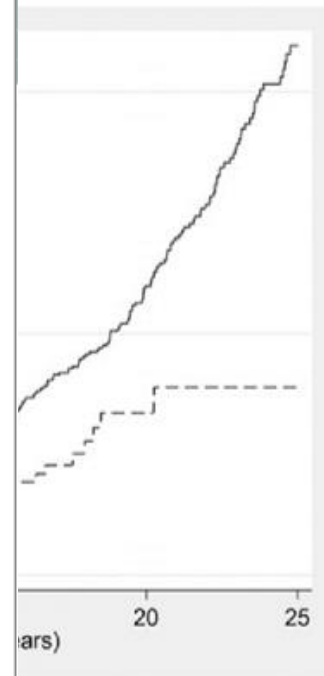
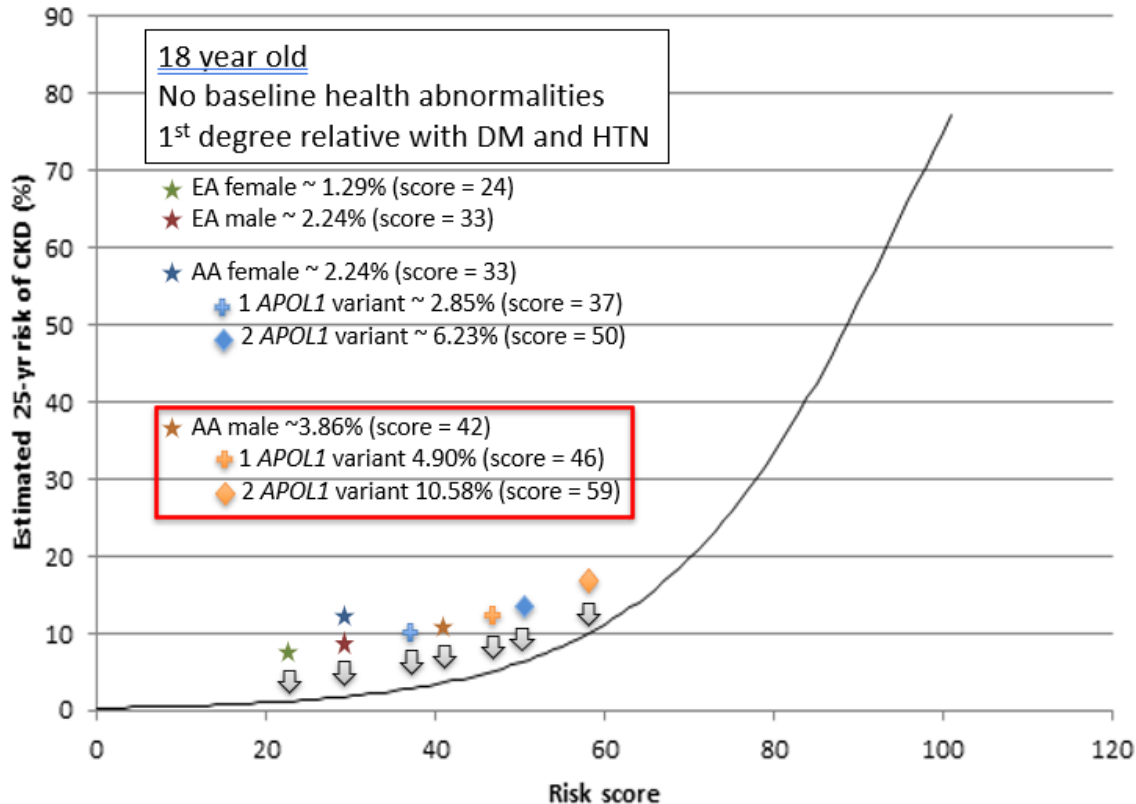
BMI per 5 kg/m<sup>2</sup>

First-degree biological  
related to recipient

the top quintile are de

Grams, M. E., Kucirka, L. M.,  
A. B., & Segev, D. L. (2012).  
adults. *Journal of the America*

Calculate your score



# Cumulative graft loss by LKDPI

## The Epidemiology of Living Kidney Donation

### Living Kidney Donor

This model predicts recipient donor kidney transplant characteristics, on the basis of recipient characteristics.

Massie AB, Leanza J, Fahmy LM. Donor Kidney Transplantation. *Am J Transplant*. 2017;17(12):1411-1420.

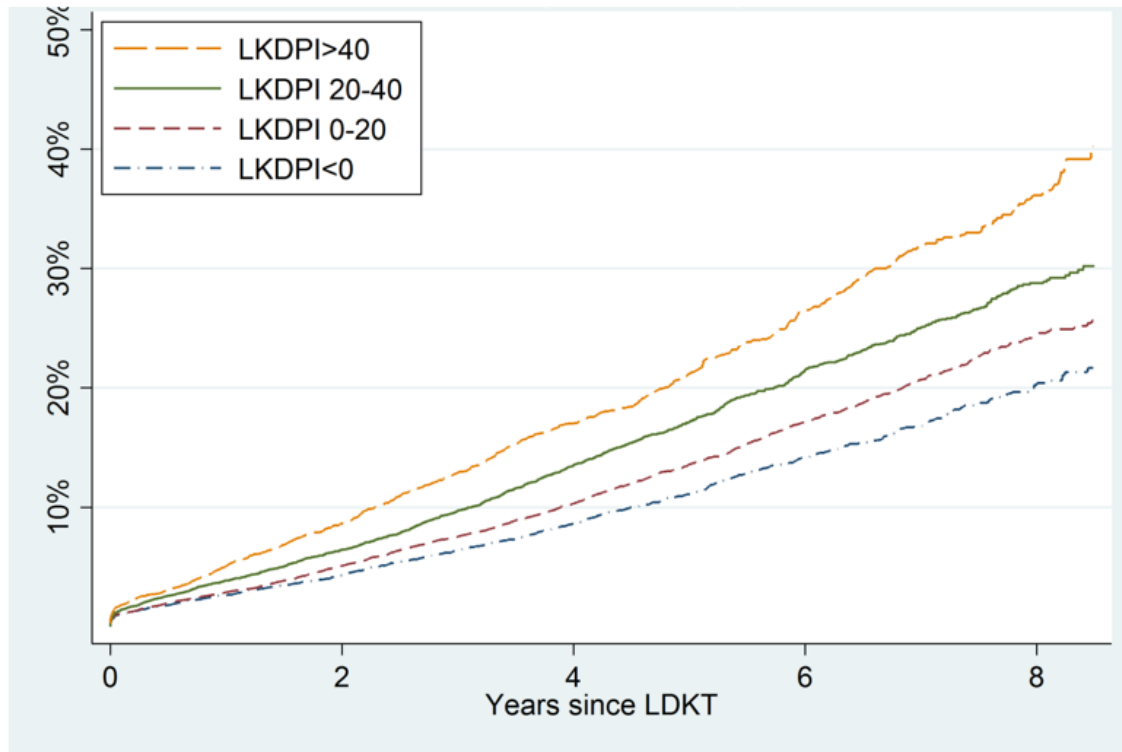
[Continue to model »](#)

### Transplant Candidate

This prediction model is for dialysis aged 65 and above with a probability of 3-year survival (KT). Patients with predicted top quintile are deemed high risk.

Grams, M. E., Kucirka, L. M., Ha, A. B., & Segev, D. L. (2012). Care of high-risk transplant candidates. *Journal of the American Society of Nephrology*, 23(11), 2111-2118.

[Calculate your score »](#)



transplantation at the pediatric level.

renal disease (ESRD) on the basis of recipient characteristics. Whether they will accept the risk, or if they will wait for the next donor.

Singer, A. L., Kucirka, L. M., et al. (2013). Identifying appropriate living kidney donors. *American Journal of Transplantation*, 13(12), 2811-2818.

### ESRD in Living Kidney Donors

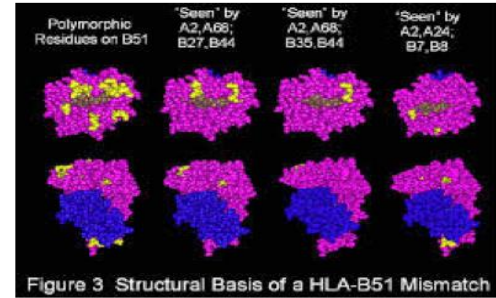
appropriate informed consent for living kidney donors.

Donor Risk of ESRD in Living Kidney Donors. *Society of Nephrology* (2017): 1-10.



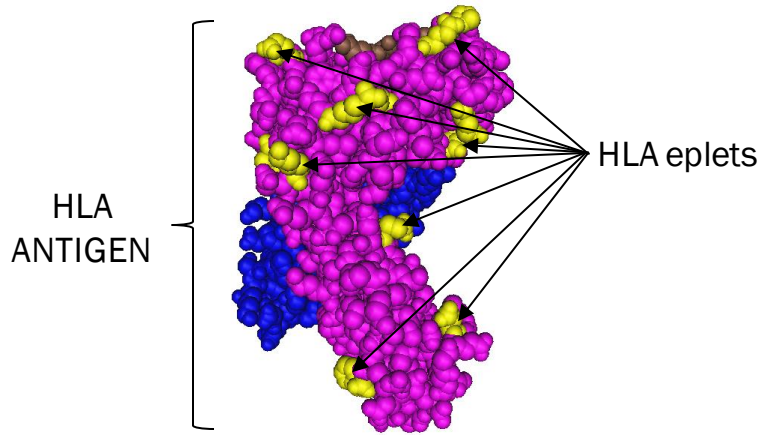
# Eplet Matching in the NKR

- Compatible pairs
- May benefit from closer immunologic matching
- Longer life for kidney
- Possibly less immunosuppression needed

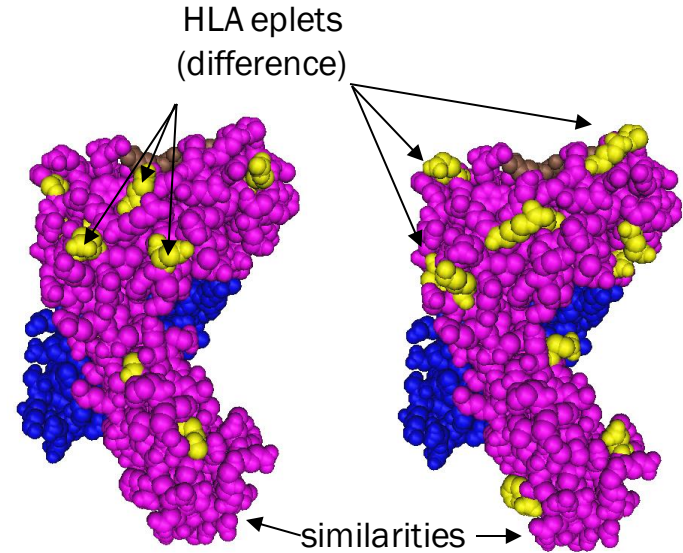


**NKR** KIDNEY FOR LIFE  
PROGRAM

# HLA EPLET: the “immunogenic” unit of HLA antigens



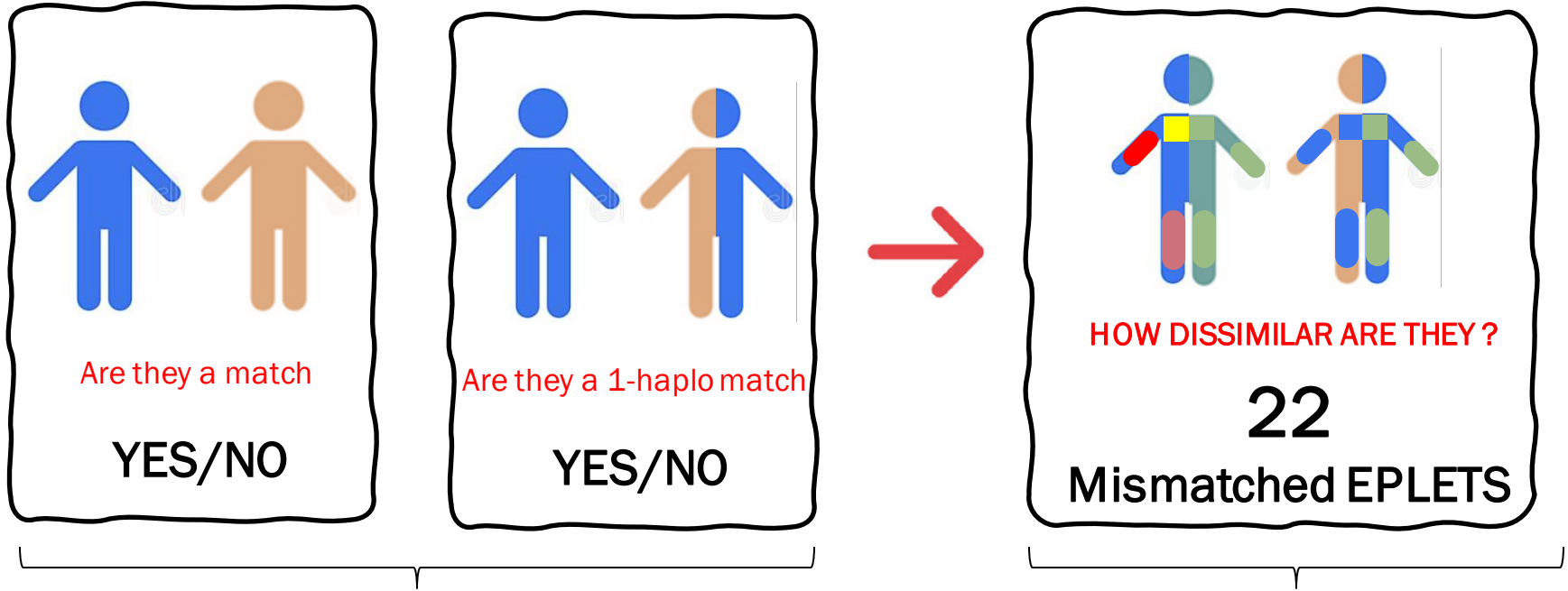
The HLA antigens are complex amino-acid structures expressing several immunogenic hot-spots, known as “HLA eplets”



The HLA antigens of two individual share a large degree of homology (magenta areas) but differ at some or all the HLA eplets (yellow patches)

# HLA EPLET: the new HLA compatibility “currency”

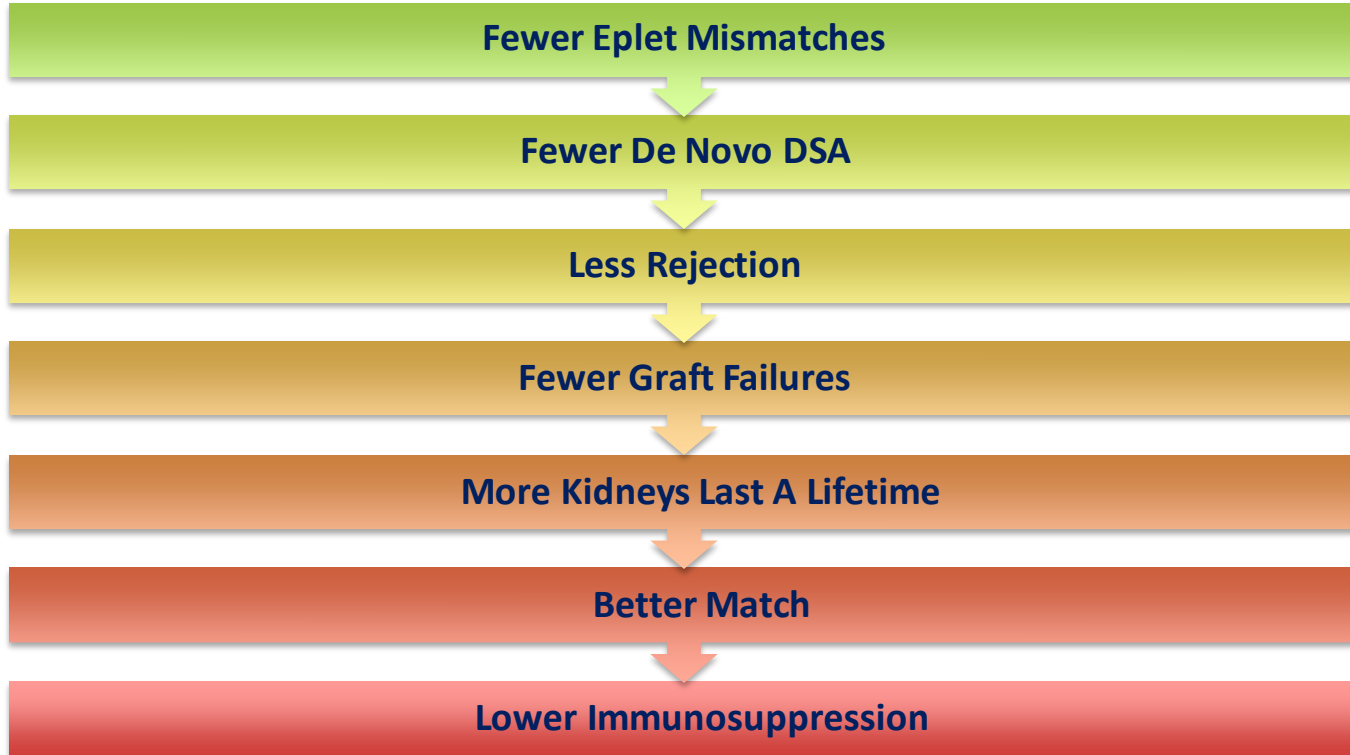
We need to change the way we define HLA compatibility ....



CONVENTIONAL HLA MATCH DOES NOT QUANTIFY THE IMMUNOLOGICAL RISK

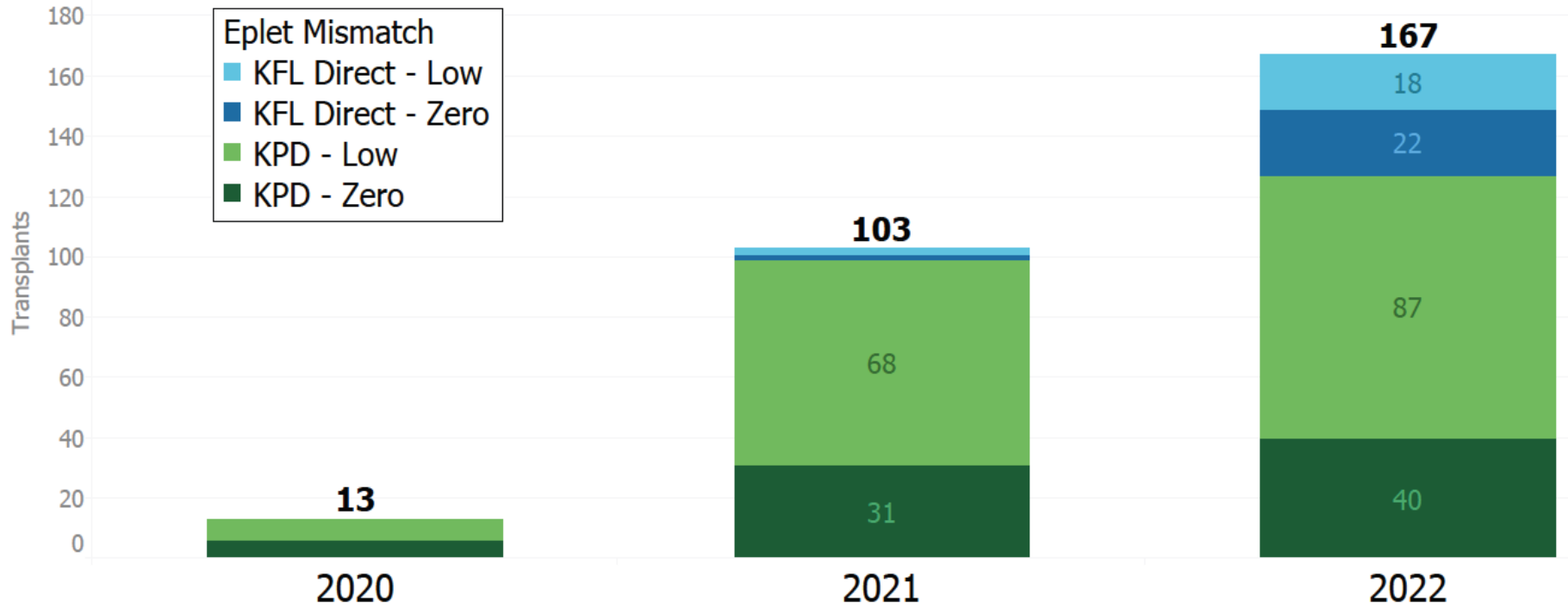
BETTER REPRESENTATION OF THE IMMUNOLOGICAL RISK

# Kidney For Life Basics



# Kidney for Life Low Eplet Mismatch Transplants

## Compatible and Incompatible Pairs

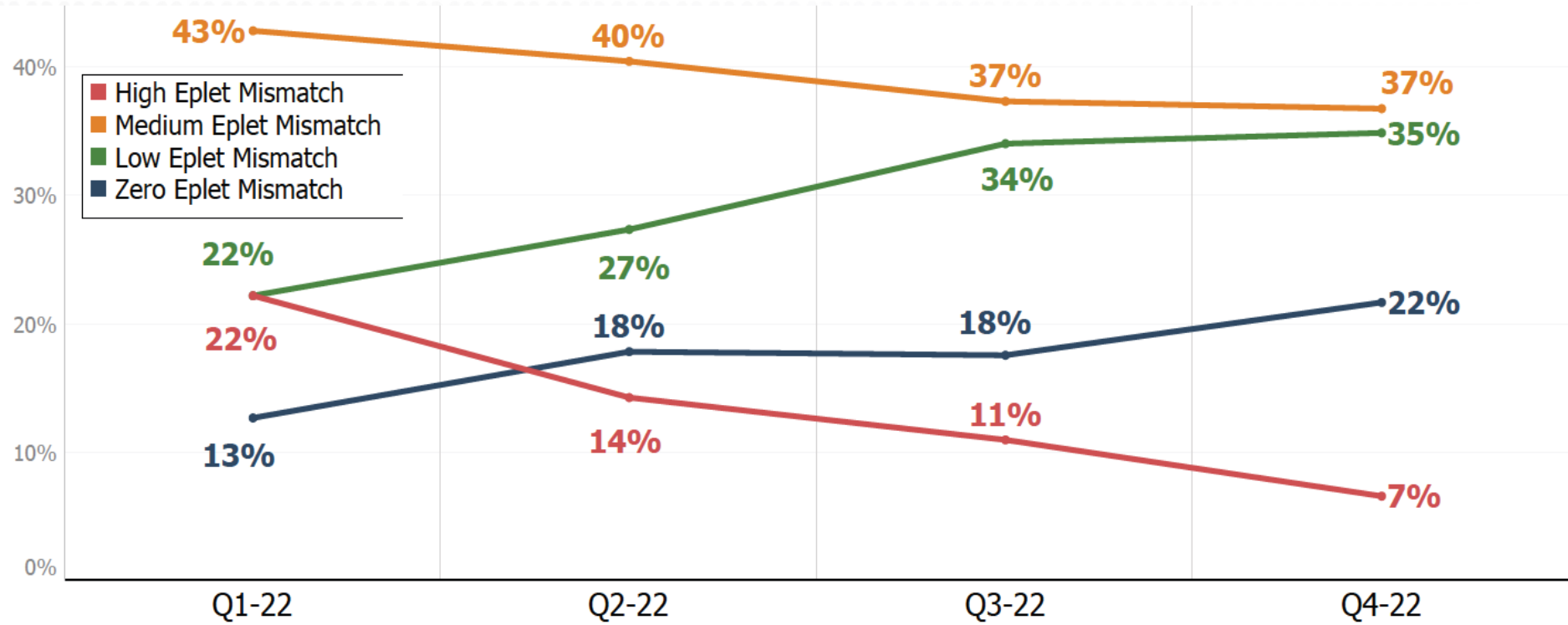


Eplet Mismatch

- KFL Direct - Low
- KFL Direct - Zero
- KPD - Low
- KPD - Zero

# Percentage of Transplants with Eplet Mismatch Data

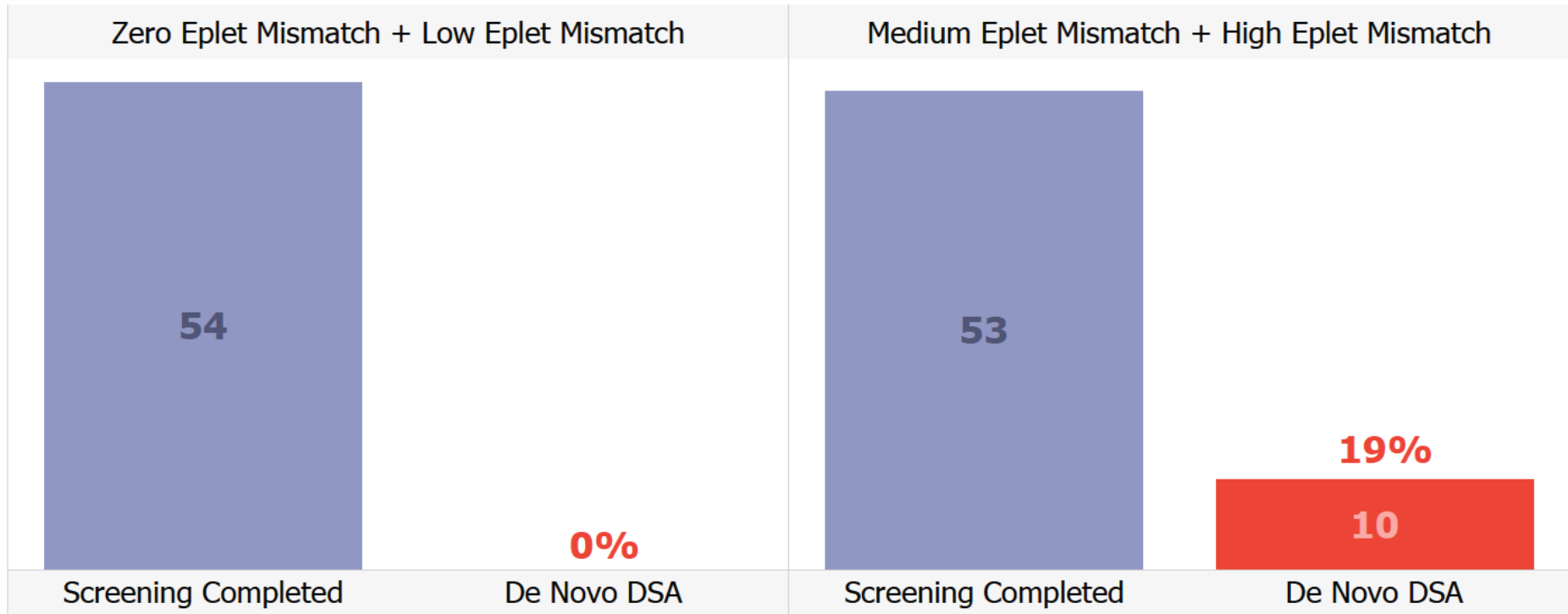
Continually Improving Low & Zero Eplet Mismatch Percentage



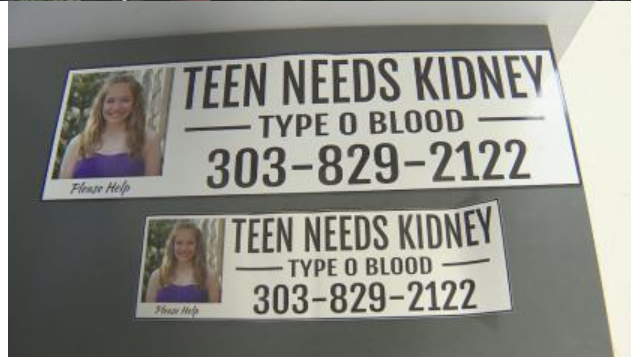


# Kidney for Life Results

## One Year Antibody Screening



# Patients/Families Tell Their Story



# Champion Microsites

- ✓ Center invites patient to setup site
- ✓ Patient creates site profile
- ✓ Center & NKR approve site
- ✓ Microsite link posted to the web
- ✓ NKR prints/ships business cards to patient

National **Kidney** Registry

Khadijah Sabir

**I NEED A KIDNEY DONOR**

If you are interested in learning more about my story, kidney donation or in being tested to see if you are eligible to donate, please visit the URL below.

[WWW.NKR.ORG/CEN498](http://WWW.NKR.ORG/CEN498)

National **Kidney** Registry

Nathaniel Aiken

**I NEED A KIDNEY DONOR**

If you are interested in learning more about my story, kidney donation or in being tested to see if you are eligible to donate, please visit the URL below.

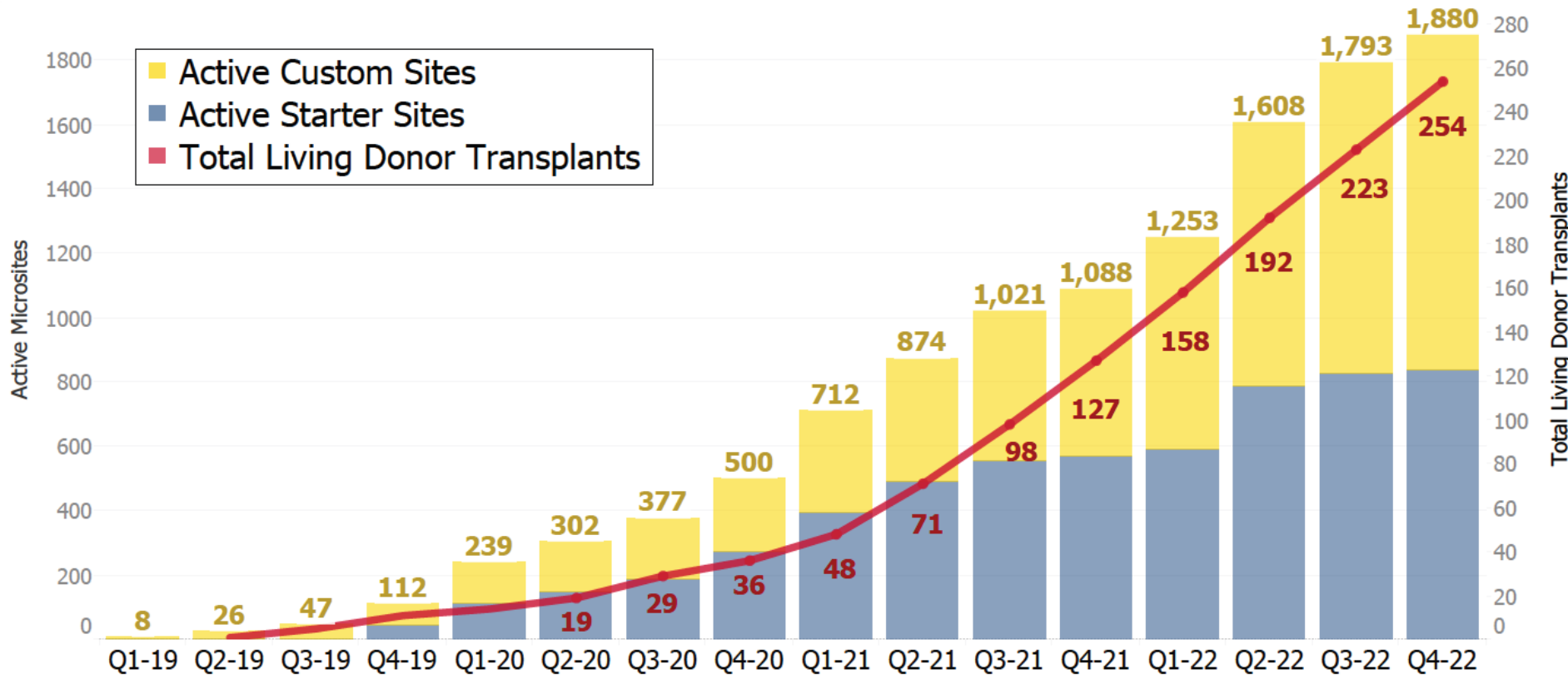
[WWW.NKR.ORG/ZHS796](http://WWW.NKR.ORG/ZHS796) 498

## Written policy for what is NOT allowable on microsite:

- Disclosure of financial information of any kind
- Material items in photos / videos
- Anything that can be construed as an attempt to compensate potential donors

The screenshot shows a microsite for Nathaniel Aiken on the National Kidney Registry. At the top, there is a navigation bar with the NKR logo, the name 'Nathaniel Aiken', a link 'Learn about living donation', and a red button 'Register to donate for Nathaniel'. Below the navigation bar is a circular profile picture of Nathaniel, a young boy, smiling. To the right of the photo, his name 'Nathaniel Aiken' and age 'Age 8' are listed, along with his 'Need level: Pre-Emptive'. Below this is a 'My Story' section with a paragraph of text. Underneath is a 'Why I Need a Kidney' section with another paragraph. A 'Nathaniel's Photos' section contains four small images of him in various settings. At the bottom, there is a red banner with the NKR logo, the text 'Can you help Nathaniel by donating a kidney?', a short paragraph, and two buttons: 'Learn about living donation' and 'Register to donate for Nathaniel'.

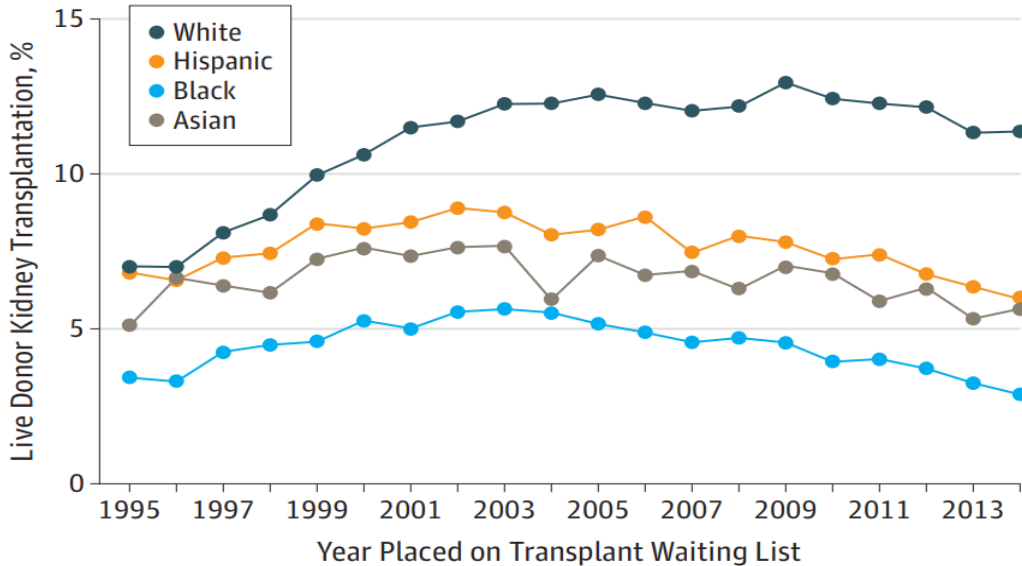
# Smarter Use of Social Media



# Association of Race and Ethnicity With Live Donor Kidney Transplantation in the United States From 1995 to 2014

Tanjala S. Purnell, PhD, MPH; Xun Luo, MD, MPH; Lisa A. Cooper, MD, MPH; Allan B. Massie, PhD; Lauren M. Kucirka, MD, PhD, ScM; Macey L. Henderson, JD, PhD; Elisa J. Gordon, PhD, MPH; Deidra C. Crews, MD, ScM; L. Ebony Boulware, MD, MPH; Dorry L. Segev, MD, PhD

**A** Live donor kidney transplantation



1995-1999  
 0.83 Hispanic  
 0.45 Black  
 0.56 Asian

2014-2019  
 0.52 Hispanic  
 0.27 Black  
 0.42 Asian

JAMA 2018

# Current State

## Go out and find a living donor



### Live Donor Champion: Finding Live Kidney Donors by Separating the Advocate From the Patient

*Jacqueline M. Garonzik-Wang,<sup>1</sup> Jonathan C. Berger,<sup>1</sup> Reside Lorie Ros,<sup>1</sup> Lauren M. Kucirka,<sup>1</sup>  
Neha A. Deshpande,<sup>1</sup> Brian J. Boyarsky,<sup>1</sup> Robert A. Montgomery,<sup>1</sup> Erin C. Hall,<sup>1</sup>  
Nathan T. James,<sup>1</sup> and Dorry L. Segev<sup>1,2,3</sup>*

**Future State**  
**Let us help you find a living donor!**

Not 'Do you have' but 'Who is your living donor?'



# Making House Calls Increases Living Donor Inquiries and Evaluations for Blacks on the Kidney Transplant Waiting List

James R. Rodrigue,<sup>1,5</sup> Matthew J. Paek,<sup>1</sup> Ogo Egbuna,<sup>2</sup> Amy D. Williams,<sup>3</sup> Lisa D. Gill,<sup>4</sup> Martha Pavlakis,<sup>1</sup> and Didier A. Mande

# Transplant Center Provision of Education and Culturally and Linguistically Competent Care: A National Study

E. J. Gordon<sup>a,b,\*</sup>, J. C. Caicedo<sup>b</sup>, D. P. Ladner<sup>a,b</sup>,

Received 14 July 2010, revised 13 August 2010, accepted for publication 04 September 2010

ORIGINAL CLINICAL SCIENCE—GENERAL

# Financial Impact of a Culturally Sensitive Hispanic Kidney Transplant Program on Increasing Living Donation

Wang, Andrew PhD, MPH<sup>1,2</sup>; Caicedo, Juan Carlos MD<sup>3</sup>; Mathur, Amit K. MD, MS<sup>4</sup>; Ruiz, Richard M. MD<sup>5</sup>; Gordon, Elisa J. PhD, MPH<sup>1,3</sup>

# Financial Neutrality??



National Living Donor Assistance Center

Program ▾

Potential Donors ▾

Transplant Centers ▾

News & Updates

Resources

Login

*“This is a terrific program.  
Thank you so much.”*

– Lenora - Living Kidney Donor



## Living Donor Assistance Program

Providing financial assistance to those who want to donate an organ, priority is given to individuals not otherwise able to afford the travel and subsistence expenses associated with living organ donation.

[Learn more ▶](#)

## For Donors

Learn if you are eligible for reimbursement of travel and subsistence expenses and how to file an application.

[More information ▶](#)

OUR MISSION IS TO **REDUCE**  
FINANCIAL DISINCENTIVES  
to **LIVING ORGAN DONATION**



**350**  
participating  
transplant  
programs

**3,340**  
organ  
donations  
made possible

**90%**  
of applications  
have received funding

# Financial Neutrality → A Must!!



National Living Donor Assistance Center

Program ▾

Potential Donors ▾

Transplant Centers ▾

News & Updates

Resources

Login

*“This is a terrific program.  
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**3,340**  
organ  
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**90%**  
of applications  
have received funding

# Donor Shield Program

- ❖ Lost wage reimbursement up to \$1500/week for up to 6 weeks
- ❖ Travel and lodging reimbursement up to \$3000 for donor and a caregiver
- ❖ Kidney Prioritization should the donor ever need a transplant
- ❖ Donation life insurance with \$1,000,000 principal sum
  - ❖ Donation disability insurance
    - ❖ Free legal support
  - ❖ Complication Protection
  - ❖ Home blood draws



Personal Viewpoint

doi: 10.1111/ajt.13232

## Living and Deceased Organ Donation Should Be Financially Neutral Acts

F. L. Delmonico<sup>1,\*</sup>, D. Martin<sup>2</sup>, B. Domínguez-Gil<sup>3</sup>,  
E. Muller<sup>4</sup>, V. Jha<sup>5</sup>, A. Levin<sup>6</sup>, G. M. Danovitch<sup>7</sup>  
and A. M. Capron<sup>8</sup>

care units; LDCOP, Live Donor Community of Practice;  
LKD, living kidney donor; NLDAC, National Living  
Donor Assistance Center; NOTA, National Organ  
Transplant Act

Meeting Report

doi: 10.1111/ajt.13233

## AST/ASTS Workshop on Increasing Organ Donation in the United States: Creating an “Arc of Change” From Removing Disincentives to Testing Incentives

D. R. Salomon<sup>1,\*</sup>, A. N. Langnas<sup>2</sup>, A. I. Reed<sup>3</sup>,  
R. D. Bloom<sup>4</sup>, J. C. Magee<sup>5</sup> and R. S. Gaston<sup>6</sup>  
for the AST/ASTS Incentives Workshop Group  
(IWG)<sup>a</sup>

Transplant Act; OPTN, Organ Procurement and Trans-  
plantation Network

Received 14 October 2014, revised and accepted for  
publication 16 January 2015

Personal Viewpoint

doi: 10.1111/ajt.13234

## Between Scylla and Charybdis: Charting an Ethical Course for Research Into Financial Incentives for Living Kidney Donation

J. S. Fisher<sup>1</sup>, Z. Butt<sup>2</sup>, J. Friedewald<sup>3</sup>,  
S. Fry-Revere<sup>4,5</sup>, J. Hanneman<sup>6</sup>,  
M. L. Henderson<sup>7</sup>, K. Ladin<sup>8,9,10</sup>, H. Myser<sup>11</sup>,  
L. Preczewski<sup>12</sup>, L. A. Sherman<sup>13</sup>, C. Thiessen<sup>14</sup>  
and E. J. Gordon<sup>15,\*</sup>

financial compensation on living kidney donation rates, many fear that financial incentives will exploit vulnerable individuals and cast the field of transplantation in a negative public light, ultimately reducing donation rates. This paper provides an ethical justification for conducting a pilot study of a federally regulated approach to providing financial incentives to living



# Inducing Tolerance??

Human Immunology 79 (2018) 272–276



Contents lists available at ScienceDirect

Human Immunology

journal homepage: [www.elsevier.com/locate/humimm](http://www.elsevier.com/locate/humimm)



Tolerance induction in HLA disparate living donor kidney transplantation by facilitating cell-enriched donor stem cell infusion: The importance of durable chimerism



Joseph R. Leventhal<sup>a,\*</sup>, Suzanne T. Ildstad<sup>b</sup>

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<sup>b</sup> Institute for Cellular Therapeutics, University of Louisville, Louisville, KY, USA



# Tolerance as SOC!!

Human Immunology 79 (2018) 272–276



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Tolerance induction in HLA disparate living donor kidney transplantation by facilitating cell-enriched donor stem cell Infusion: The importance of durable chimerism



Joseph R. Leventhal<sup>a,\*</sup>, Suzanne T. Ildstad<sup>b</sup>

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# Value proposition for “One Transplant for Life”



**Human Costs:**  
Lost Productivity  
Impaired QoL



**Clinical & Economic Costs:**  
IS Complications  
IS Co-morbidities  
Graft Loss  
IS Costs

## Value From Eliminating Chronic IS

- **Improve outcomes**
  - Fewer rejections, graft losses
  - No IS co-morbidities or complications
  - Enhance patient’s QoL and freedom
- **Reduce systematic costs**
  - IS and meds to manage co-morbidities
  - Avoid return to dialysis or 2<sup>nd</sup> transplant
  - Bolster recipients’ productivity

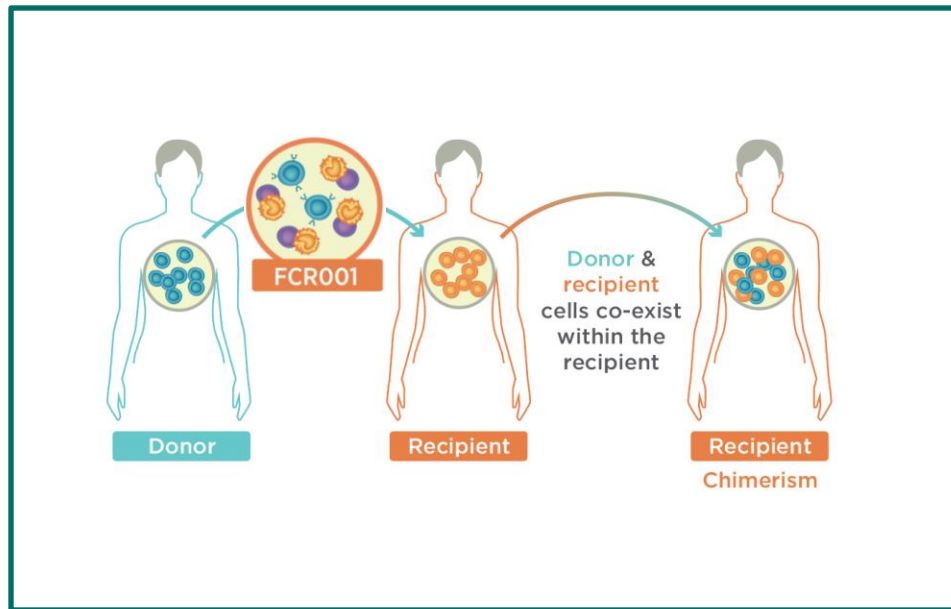
# Allogeneic Tolerance and Chimerism



Nobel Prize 1960\*

Goal: facilitate allogeneic tolerance by establishing durable chimerism

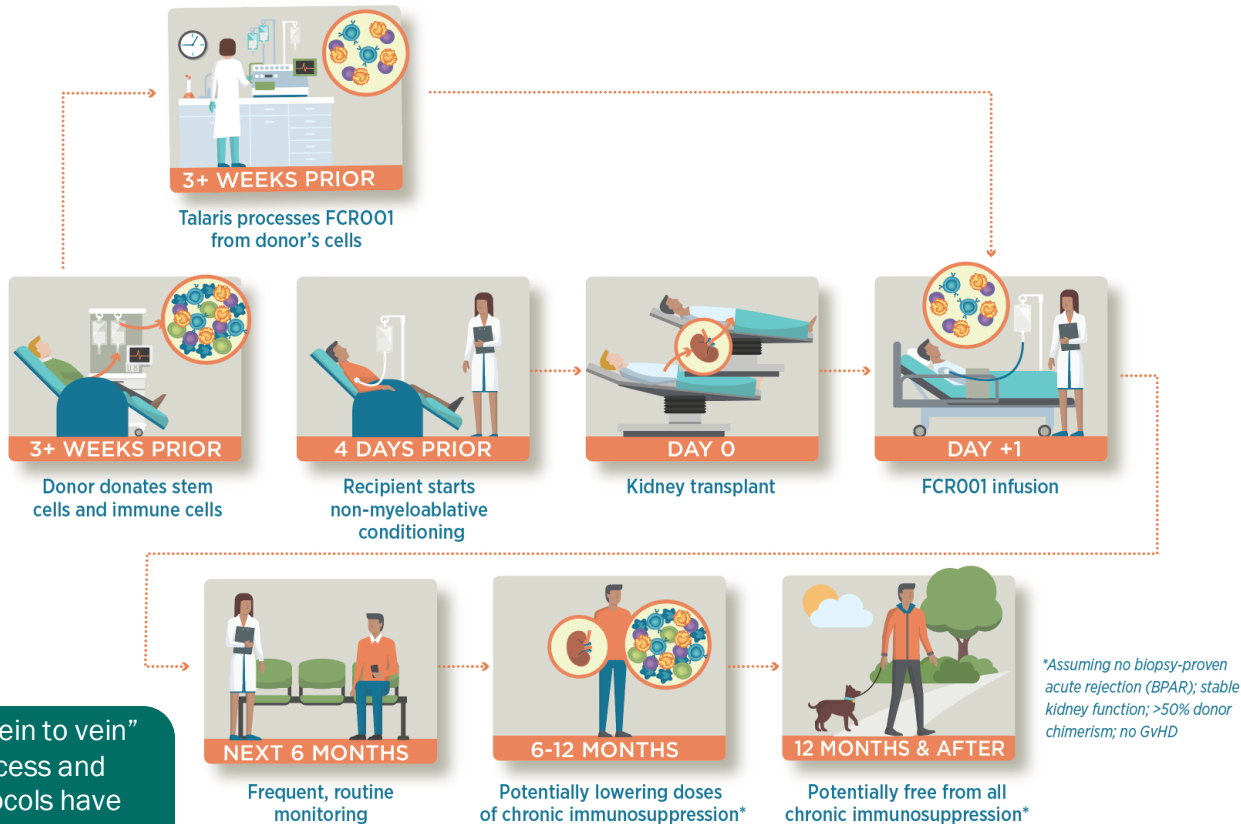
**Allogeneic tolerance:** An approach to enable donor HSCs to coexist with recipient HSCs in the recipient's bone marrow ("chimerism"), and mature into mutually-tolerated, functional immune cells and blood cells



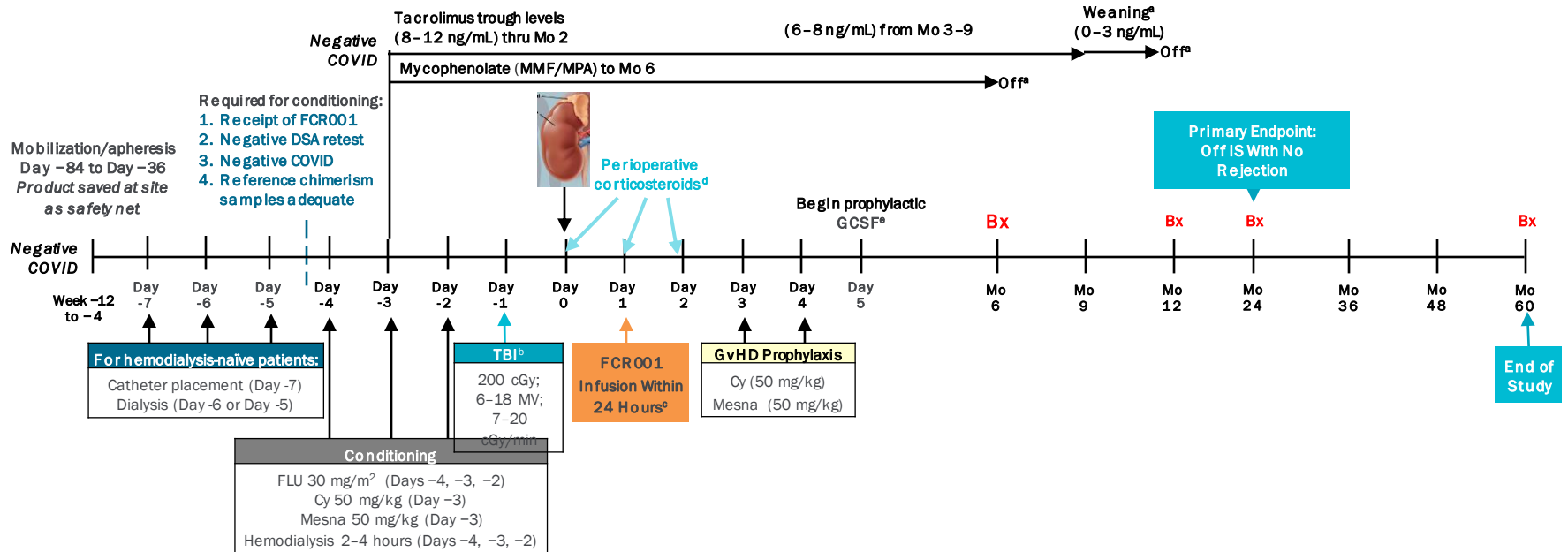
\*Nobel Prize in Physiology or Medicine 1960 was awarded jointly to Sir Frank Macfarlane Burnet and Peter Brian Medawar 'for discovery of acquired immunological tolerance.'

# FCR001: The Donor- Recipient Journey

Our “vein to vein”  
process and  
protocols have  
been fully  
proceduralized



# FREEDOM-1: FCR001 Protocol Overview



\*Patients demonstrating stable donor chimerism (>50%), no history of rejection, no DSA, no GvHD, not using corticosteroids, and adequate kidney function.

<sup>b</sup>TBI dose of 200 cGy delivered as a single fraction at 10-18 MV at a rate of 15-20 cGy/min are the preferred energy and rate parameters and should be followed when possible.

<sup>c</sup>Administered using a central or dedicated peripheral line; infusion by gravity.

<sup>d</sup>Methylprednisolone 500 mg IV on Day 0 in OR; 250 mg Day 1 and 125 mg Day 2.

<sup>e</sup>Until absolute neutrophil count is >1000/mm<sup>3</sup> for 3 consecutive days.

# Highlights from Phase 2 Study (+ long- term followup)

37 adult living donor kidney transplant (LDKT) patients were dosed with our therapy at two leading US transplant sites between 2009 - 2016



## 70%

(26 OF 37) OFF  
ALL IMMUNOSUPPRESSION  
THERAPIES \*\*

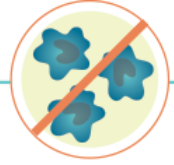
- Across all HLA-mismatches
- 82% success rate (14 of last 17) once key parameters were optimized



## 100%

TAKEN OFF  
IMMUNOSUPPRESSION  
REMAIN IS-FREE

- Median follow up: >7 yrs
- Six patients followed >10 years
- Longest follow up: >12 yrs

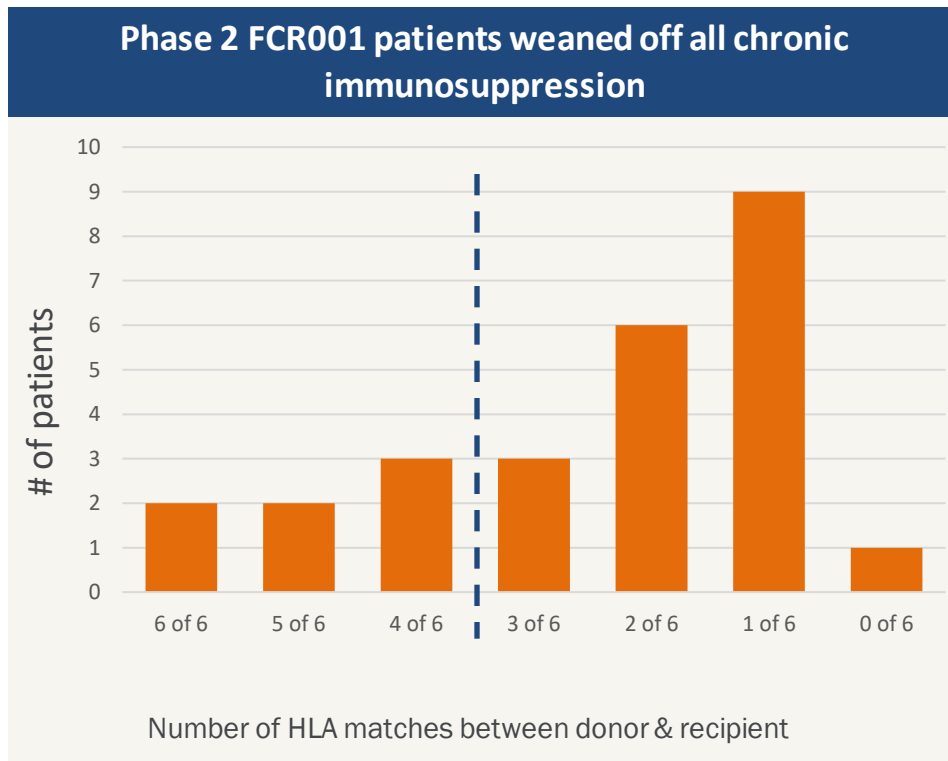


## 7/7

TOLERIZED PATIENTS WITH  
PRIOR KIDNEY AUTO-IMMUNE  
CONDITION HAD NO  
RECURRENCE

- Recurrence ordinarily seen in 20% - 60% of patients\*\*\*

# Phase 2 Results Robust Across All Degrees of HLA-Mismatch



19/26 (73%) durably off all chronic immunosuppression had HLA match of 3 or less between LDKT donor & recipient

Comparable kidney and patient survival for all FCR001 vs standard of care (SoC) LDKT patients

FCR001 safety & tolerability generally consistent with separate SoC kidney transplant + allogeneic HSCT with non-myeloablative conditioning

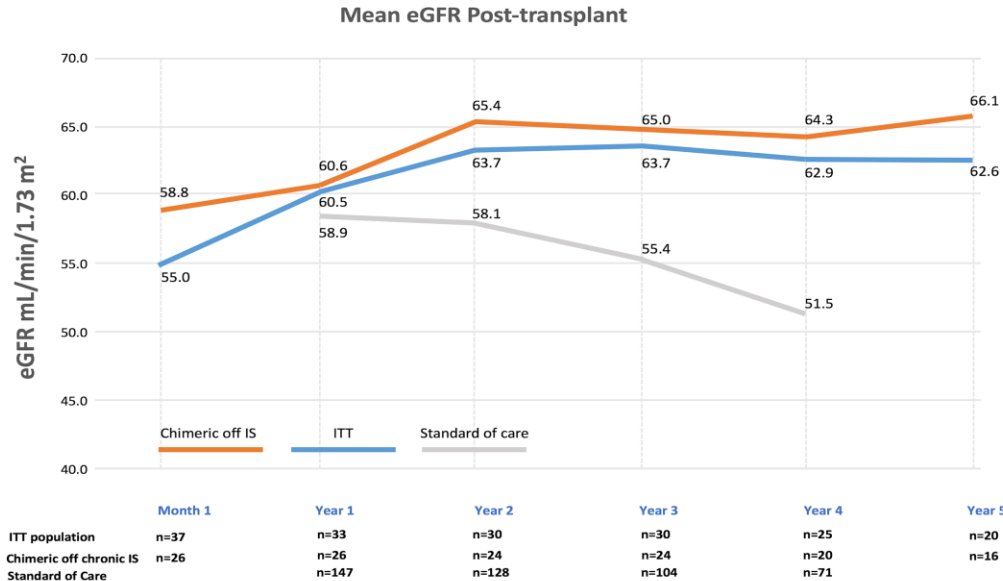
No acute rejection or donor-specific antibodies in FCR001 patients off immunosuppression



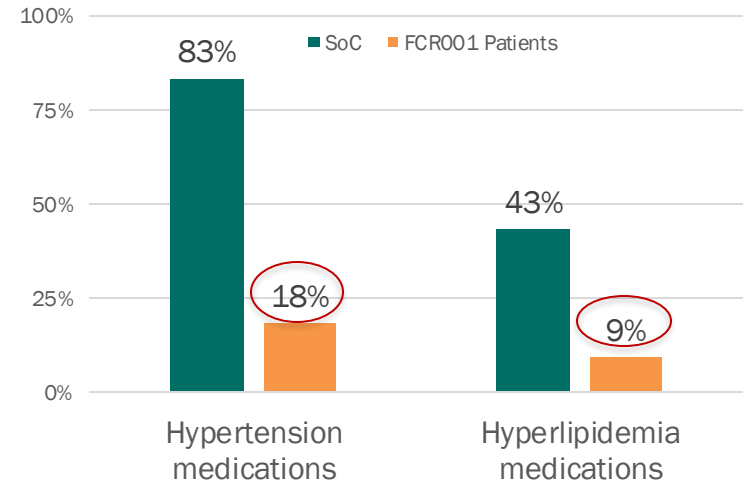
# Evidence of Potential Longer-Term Clinical Benefit

FCR001 improved Quality of Life, preserved kidney function and enabled lower reliance on cardiovascular medications

## Mean Estimated eGFR\* Over Time Post-Transplant



## Cardiovascular Medication Usage SoC vs Durably Chimeric FCR001 Patients



# Potential to Extend Across Solid Organ Transplant

## Living Donor Kidney Transplant Delayed Tolerance Induction

FREEDOM.2

- Phase 2 study - initiated October 2021
- **Goal:** Safely induce durable tolerance and eliminate immunosuppression in **prior recipients of LDKT** (those transplanted 3 – 12 months prior to FCRO01 administration)
- Potential to expand market to prevalent LDKT population

Potential US Market Opportunity:  
~6,000 – 10,000\*/yr

## Deceased Donor Kidney Transplant

FREEDOM.4

- Active research program to establish easibility of extracting same cells directly from deceased donor bone marrow
- Relationships established with KODA and other OPOs
- Product would be administered a few months after organ transplant

Potential US Market Opportunity  
>~16,500 / yr

# Uterus Transplantation – Greater Interest

## Uterus Transplant - A Unique Type of Transplant

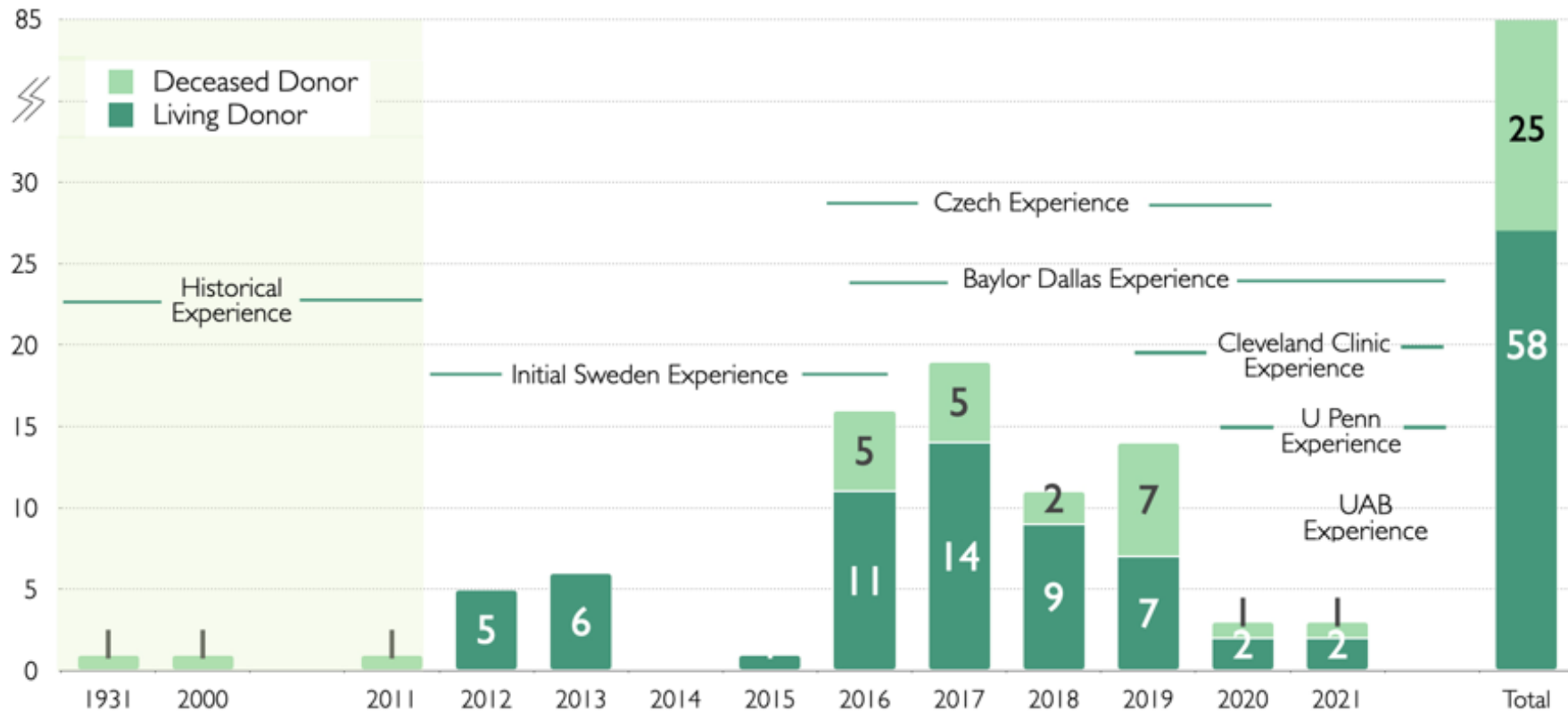
- 1) Non-lifesaving transplant (However can enhance QOL for parents)
- 2) Recipients are healthy w/out comorbidities (some MRKH have solitary kidney)
- 3) Temporary transplantation
- 4) Living or deceased donor option
- 5) Stepwise success determined years posttransplant
- 6) Living donors renounce an organ for transplantation after it has exhausted its function in the donor
- 7) One allograft shared by two recipients

The First Baby Born  
From A Transplanted  
Uterus in the US

November 2017



# World Uterus Transplant Experience



# Logistical Issues – Donor Organ Supply

Potential Annual  
Demand for Uterus

~ 3050  
Uterus Allograft

Mismatch  
≠

Potential Annual Supply  
Deceased Donor Uterus

<< 2000  
Deceased Donor  
Uterus Allograft

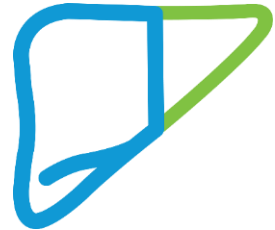
Deceased donor supply unlikely to demand → **Need living donors**



# Liver Paired Donation Program

Nation's first multi-hospital liver paired donation (LPD) program

- Increase access to living donor transplant
- Increase candidate access earlier, when in better health
- Determine how to create a nationally available program, how to sustain it, identify challenges
- Keep it algorithmically simple, 2-way only





# Insurance coverage

In partnership with the **American Foundation for Donation and Transplant**

- Accidental death and dismemberment
  - maximum, one year \$500,000
- Medical complications
  - \$5,000 deductible, maximum \$250,000
- Temporary disability
  - maximum \$100 per week
- Permanent disability
  - \$5000.00 per month up to maximum \$250,000
- Traveling companion benefit (one person)
  - One year \$500,000 accidental death and dismemberment benefit



# Center Requirements

All participating centers must:

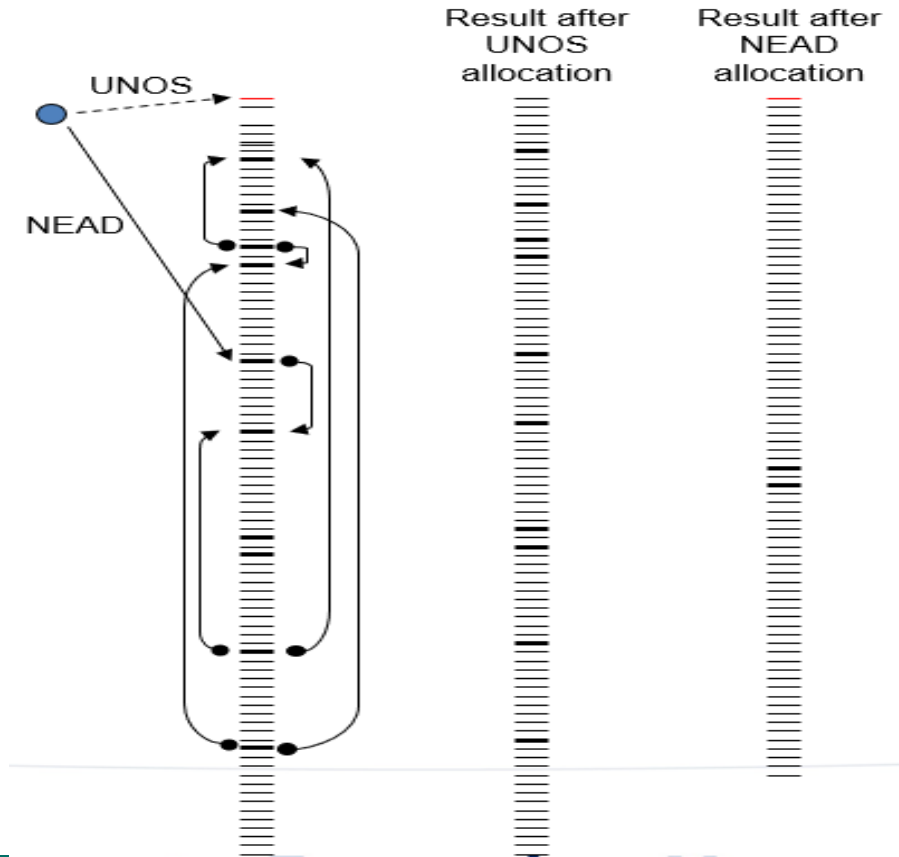
- ✓ Have performed ~20 adult LDLTs or more over the last 3 years or a children's hospital affiliated with a donor recovery hospital who meets criteria
- ✓ Have consistent liver transplant program directorship over the last 3 years;
- ✓ Not be under OPTN review for liver transplant or living liver donation-related outcomes

Centers must also:

- ✓ Agree to the Participation Agreement;
- ✓ Abide by the Liver Paired Donation Pilot Program Operational Guidelines;
- ✓ Be active OPTN and UNOS members and OPTN-approved to perform liver transplants and living liver recoveries;
- ✓ Abide by all relevant OPTN and UNOS Policies;
- ✓ Agree to share feedback with UNOS to facilitate improvements to the program;

# Expand Donor and Candidate Eligibility

# Deceased Donor-initiated Chains



— Patient on Deceased Donor Waiting List **without** an incompatible willing living donor

*Estimated Distribution by Ethnicity\**

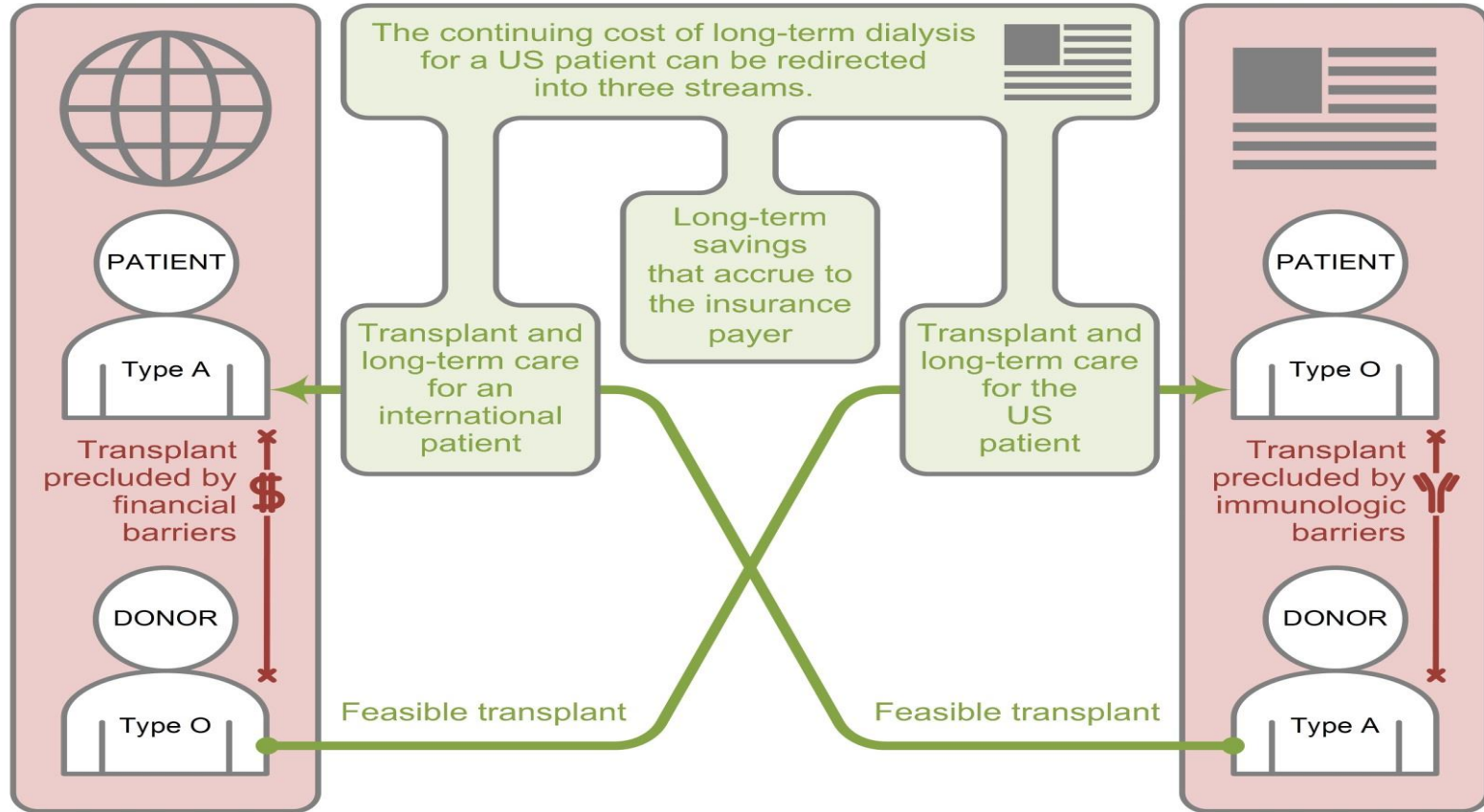
Caucasian	38.4%
African American	34.0%
Hispanic	18.1%
Asian	8.1%
Other	1.4%

— Patient on Deceased Donor Waiting List **with** an incompatible, but willing living donor

*Estimated Distribution by Ethnicity\**

Caucasian	70.3%
African American	12.5%
Hispanic	13.0%
Asian	2.9%
Other	1.2%

# Global Kidney Exchange



# Living Donor Registry

- More lives would be saved through living donation.
- The majority of the public expresses support of living donation; however, the number of living donor transplants has remained flat.
- Estimated that for every 35 living donors, there may be another 26 willing to donate.
- Transplant Centers may not be equipped with personnel and resources to efficiently conduct an abundance of living donor prospect testing.



Nearly **95,000** men, women and children await lifesaving kidney transplants



Every **10** minutes another person is added to the waiting list.



**17** people die every day waiting for a kidney



Only **6,442** living-kidney donor transplants in 2018



*America*

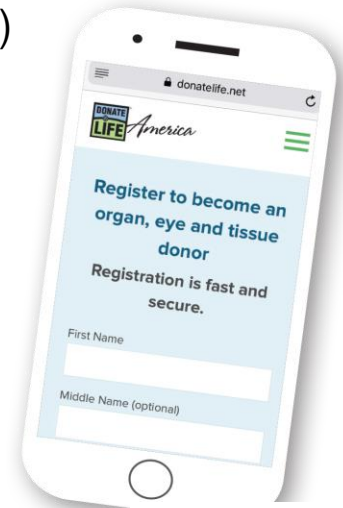
# National Donate Life Registry

All living donor prospects will enter the living donor registry pathway through the National Donate Life Registry (NDLR)

Here's why:

## More than 6 Million Registrations

- 4,000 NEW Registrations Every Day
- 8.5% Registration Conversion Rate
  - Exceeding national nonprofit benchmark of 1.5%
- 1,263 Campaign Pages with more than 700 organization pages
- National Partnerships with Apple, Walgreens and Android
- 1.9 million donor searches conducted in last 24 months
- All NDLR data is handled in accordance with HIPPA privacy standards and is only accessed by certified organ recovery organizations





# Living-Donor Prospect Experience



1. Express interest through the National Donate Life Registry



4. Return sample for lab analysis



2. Complete online health screen questionnaire



5. Potential donor shared with transplant program through UNOS connectivity

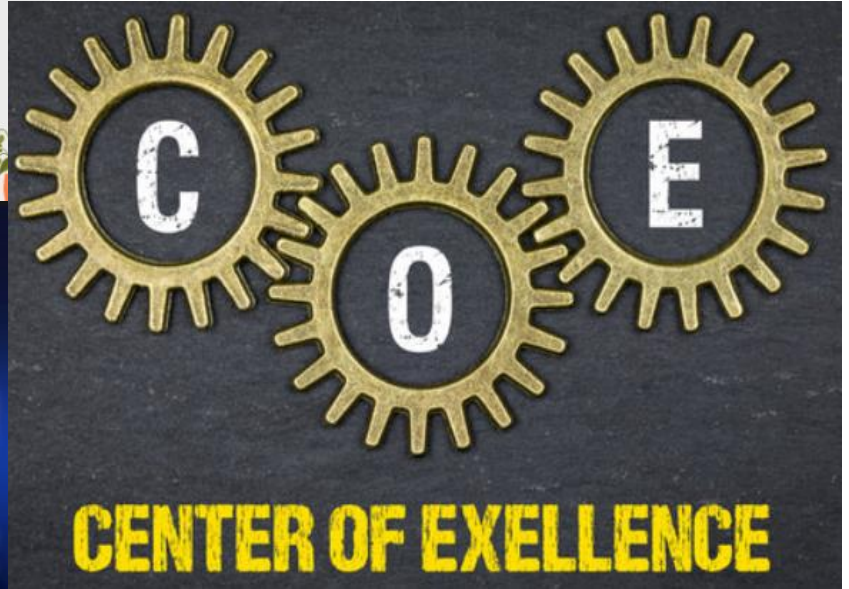


3. Collect saliva sample



6. Follow-up by transplant program

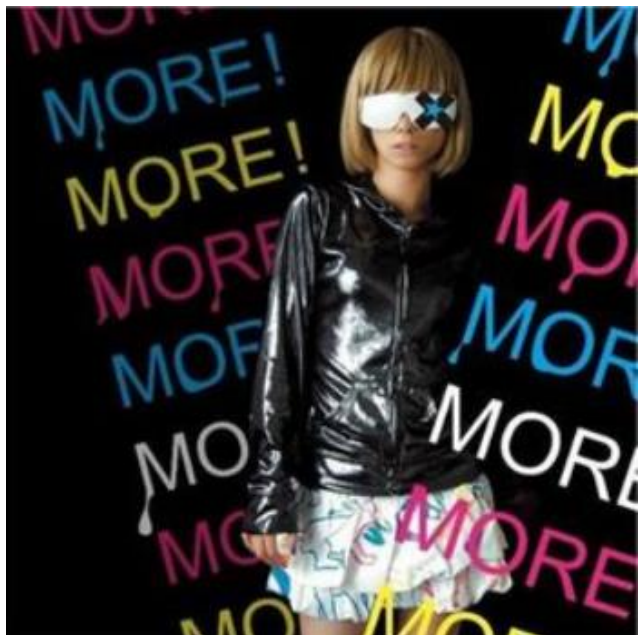




# Living Donor Transplant Centers of Excellence



# What is the Future of LD Transplant?



FOR

