Cambridge Transplant Centre

Cambridge University Hospitals



Living Donor Kidney Transplantation is the Best Treatment for All Patients with ESRD?

FICTION

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- Outcomes following living kidney donor transplantation are NOT invariably better than those following deceased donor transplantation
- Risks to the donor
- But there are important benefits to living donation
- Reality as opposed to fantasy

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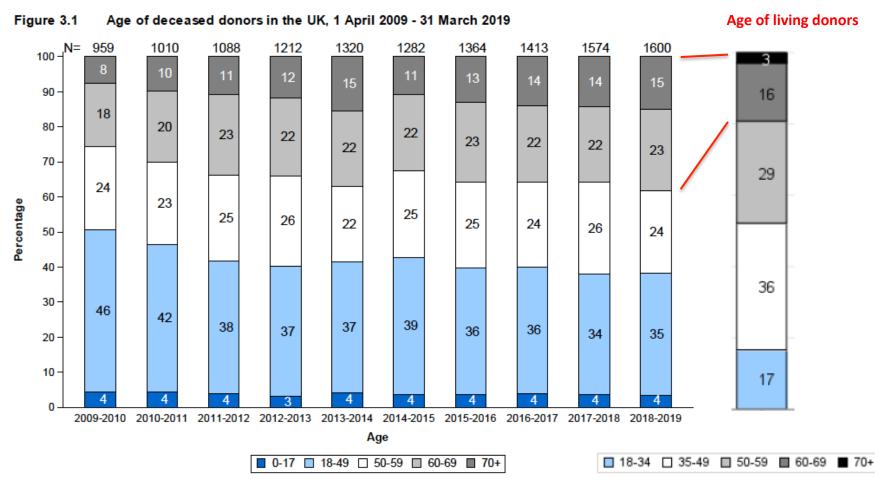
Living donor kidney transplantation is the optimal treatment for end-stage kidney disease (ESKD) but confers a risk upon the donor, both in the short term and many years after donation.

Living Donor Kidney Transplants are Best? Headline Figures May Be Misleading

Table 11.1 Graft survival after first adult kidney only transplant from a DBD						Table 11.5	ble 11.5 Graft survival after first adult living donor kidney transplant										
Year of transplant	No. at risk on day 0	% (One year	Fraft survival Two yea	l (95% confide ar Five		•	year	Year of transplant	No. at risk on day 0	On	% Gra e year		vival (95% o year		dence inte e year		n year
2005-2007 2008-2010 2011-2013 2014-2017	2208 2184 2318 3772	93 (92-94 94 (93-95 94 (93-95 95 (94-96	92 (91- 92 (91-	I-93) 87	(84-87) (85-88) (85-88)	76	(74-78)	2005-2007 2008-2010 2011-2013 2014-2017	1579 2230 2229 2609	96 97 97 98	(95-97) (96-97) (96-98) (98-99)	95 96 96	(94-96) (95-96) (95-97)	91 92 91	(90-93) (91-93) (90-92)	82	(80-84)

Table 11.2 Patient survival after first adult kidney only transplant from a DBD									Table 11.6	Patient surv	ival af	ter first ad	ult livi	ng donor	kidney	r transplan	t		
Year of transplant	No. at risk on day 0					Year of transplant	No. at risk on day 0	On	% Patient survival (95% confidence interval) One year Two year Five year Ten y			n year							
2005-2007 2008-2010 2011-2013 2014-2017	2210 2185 2319 3773	97 96 96 97	(96-98) (95-97) (95-97) (96-98)	95 95 94	(94-96) (94-95) (93-95)	89 90 88	(88-91) (88-91) (87-89)	77	(75-79)	2005-2007 2008-2010 2011-2013 2014-2017	1579 2230 2228 2609	99 99 99 99	(98-99) (98-99) (99-99) (99-99)	98 98 98	(97-99) (97-98) (97-99)	96 94 95	(95-97) (93-95) (94-96)	90	(88-91)

Living Donors are Younger (and with Fewer Comorbidities) than Deceased Donors



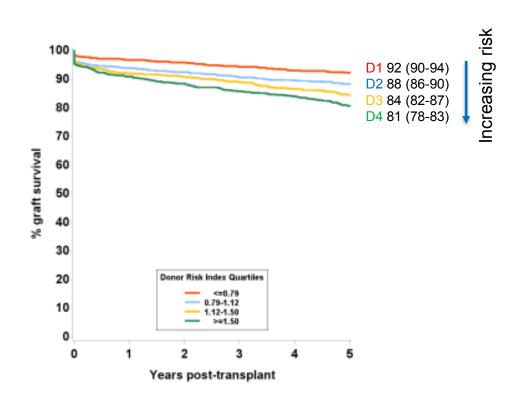
Donor Risk Index (DRI)



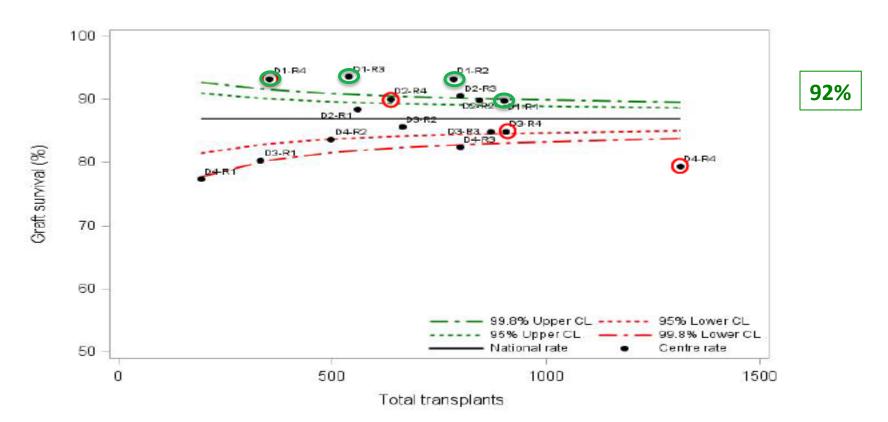
Validation dataset

Donor Factor	Hazard Ratio	p-value		
Age	1.02	<0.0001		
Height	0.86	0.0005		
Hypertension	1.15	0.1		
CMV	1.20	0.02		
Hospital stay	1.02	0.006		
eGFR	0.98	0.02		
Female	0.83	0.04		

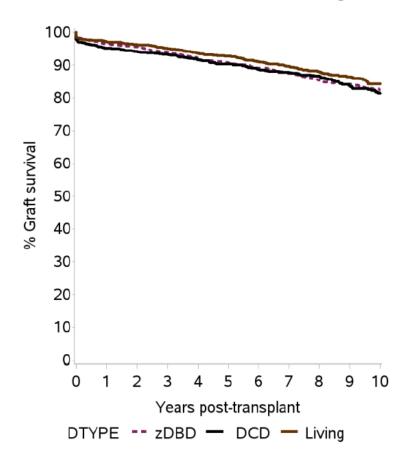
C-statistic = 0.64



5 Year Graft Survival after Deceased Donor Kidney Transplantation – Kidneys from Deceased Donors Similar to Living Donors have Equivalent Outcomes



10 Year Graft Survival after Living and Deceased Donor Kidney Transplantation From Donors Aged < 40 – NO DIFFERENCE



Summary of the Number of Censored and Uncensored Values									
					Percent				
Stratum	DTYPE	Total	Failed	Censored	Censored				
1	DCD	1449	172	1277	88.13				
2	Living	1543	158	1385	89.76				
3	zDBD	2493	290	2203	88.37				
Total		5485	620	4865	88.70				

Fiction (1)

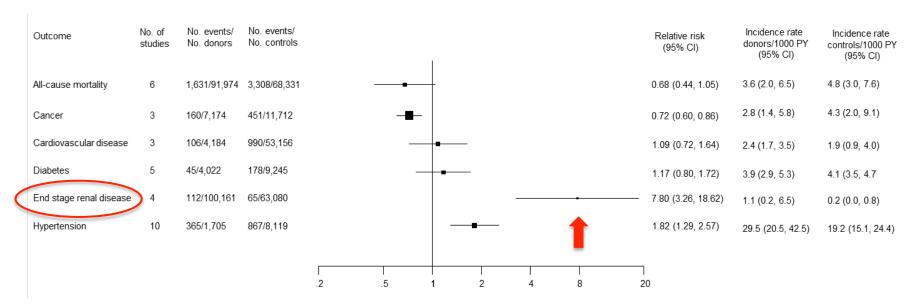
- Living donor kidney transplant outcomes are NOT invariably better than those from deceased donors
- Graft survival is equivalent when comparing living and deceased donor transplants from 'matched' donors
- There are as many, if not MORE, deceased donor kidney transplants (D1 and D2)
 with outcomes comparable to living donor transplants
- It is misleading to imply that, by not donating, a potential donor is condemning their recipient to an inferior transplant outcome

- Outcomes following living kidney donor transplantation are NOT invariably better than those following deceased donor transplantation
- Risks to the donor
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- Reality as opposed to fantasy

Clearly Demonstrated Risks to Living Donors

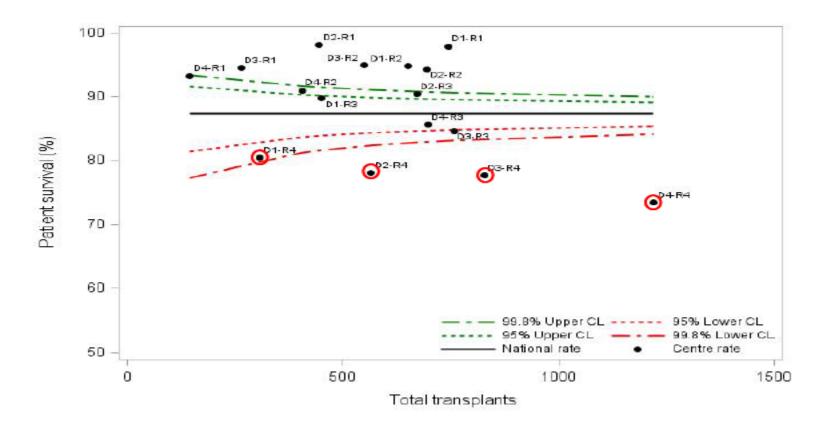
- Small (but not zero) perioperative mortality 1:3000
- Increased risk of ESRD 3-5 fold above pre-donation risk
 - 1:000 healthy non-donors vs 1:200-300 donors vs 1:100 general population, with greater relative risk in:
 - Young donors
 - Male donors
 - Black African ancestry
 - First degree relatives of patients with ESRD
- Long term medical complications (gout, hypertension)
- Pregnancy-related complications (hypertension, pre-eclampsia)

Figure 3. Meta-analysis of relative risks for selected clinical endpoints in living kidney donors compared to non-donor controls



Asking Donors to Accept Risks Cannot Be Justified by Claiming that Transplant Recipients will Invariably Receive a 'Better Kidney' or Live Longer Following Living (as opposed to Deceased) Donor Kidney Transplantation

5 Year Patient Survival after Deceased Donor Kidney Transplantation – Healthy Recipients Live Longer



Fiction (2)

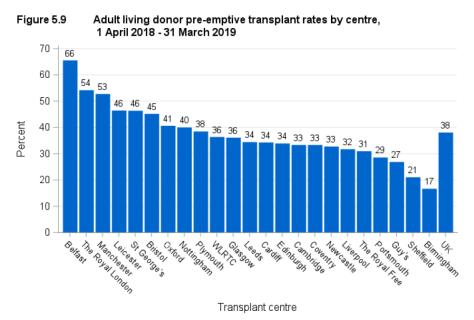
- As with graft survival, using 'headline' figures to suggest transplant recipients live longer following a living donor transplant are misleading
- Recipient survival is determined by the health and co-morbidities of the recipient
- A living donor kidney transplant does not convey mystical properties to extend life
- It is misleading to imply that, by not donating, a potential donor is condemning their recipient to an inferior transplant outcome

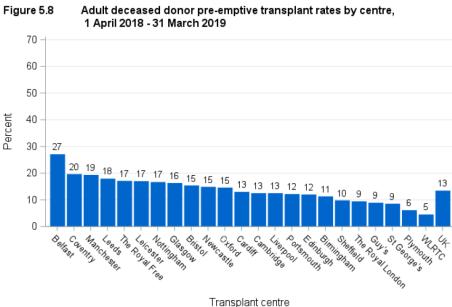
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Benefits of Living Donation Kidney Transplantation

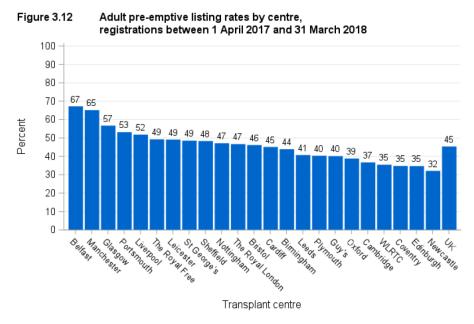
- Careful donor kidney evaluation
- Planned procedure:
 - Consultant surgeon and anaesthetist on elective operating lists
 - Pre-emptive transplant
- UK Living Kidney Sharing Scheme

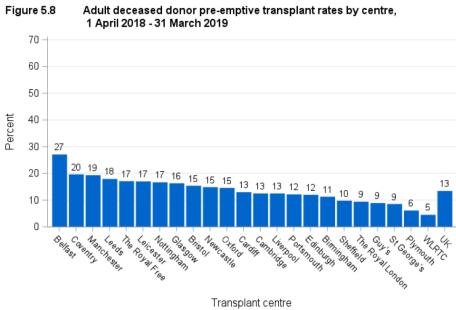
Greater Chance of Pre-emptive Transplant with Living Donation





Pre-emptive Listing vs Transplant with Deceased Donation



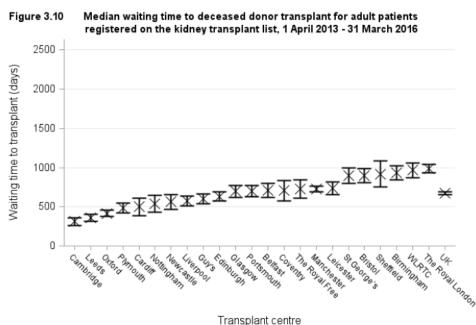


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Figure 5.1 Adult kidney only transplants, 1 April 2009 - 31 March 2019 Number 1000 -2010/11 2011/12 2012/13 2013/14 2014/15 2015/16 2016/17 2017/18 2009/10 2018/19 Year DBD ■ DCD □ Living

Falling Median Waiting Time for Deceased Donor Kidney Transplant

Figure 3.10 Median waiting time to deceased donor transplant for adult patients registered on the kidney transplant list, 1 April 2008 - 31 March 2011 2500 Waiting time to transplant (days) 2000 $\mathbf{x}^{\mathbf{x}}$ 1500 1000 Transplant centre



2013-14 Kidney Report

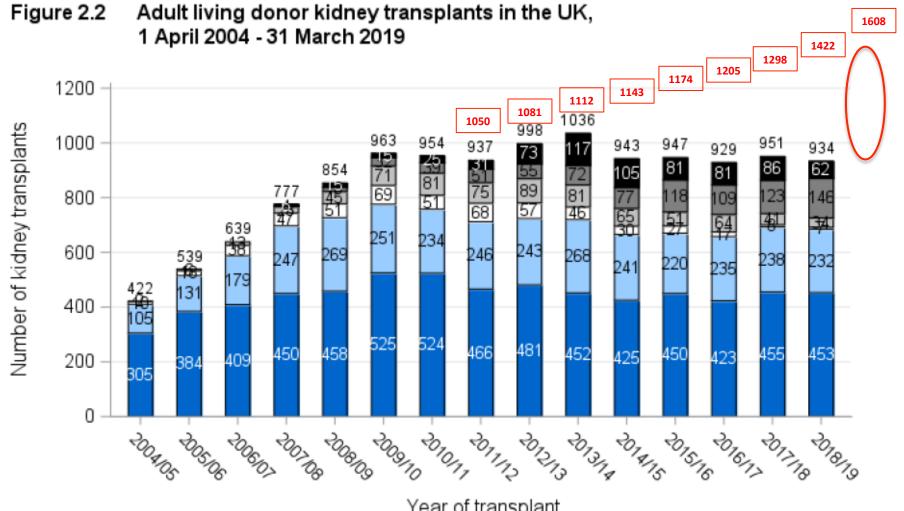
2018-19 Kidney Report

Facts (1)

- The number of kidney transplants from deceased donors is increasing year on year
- And will likely continue to do so with changes to organ donation legislation
- The waiting list and waiting times are falling year on year
- There remain important opportunities:
 - Organ reconditioning and assessment
 - Allocation matching donor and recipient
 - Allocation promoting pre-emptive deceased donor transplantation

Living Donor Transplants are Best – A FICTION, But

- An important component in the care of patients with progressive
 CKD:
 - Management of CKD to slow disease progression
 - Preparation for dialysis
 - Preparation for pre-emptive transplant listing
 - Identification of potential living donors
- For some patients a pre-emptive living donor transplant will be the optimum treatment
- But only with absolute regard to donor safety



Facts (2)

- The mismatch between the number of living donor transplants and the aspirations in the NHSBT Strategy are NOT a failing
- Instead this mismatch represents the transplant community placing donor safety at the centre of living donation
- Deceased donation presents tremendous opportunities
- The success of deceased donation means that taking additional risks to increase living donation (younger donors, donors with comorbidities) is NOT justified
- Doing deceased donor transplantation badly is not an excuse to justify living donation

