Barriers to living donor kidney transplantation in the United Kingdom: a national observational study

Diana A. Wu¹, Matthew L. Robb², Christopher J.E. Watson³, John L.R. Forsythe^{1,2}, Charles R.V. Tomson⁴, John Cairns⁵, Paul Roderick⁶, Rachel J. Johnson², Rommel Ravanan⁷, Damian Fogarty⁸, Clare Bradley⁹, Andrea Gibbons⁹, Wendy Metcalfe¹, Heather Draper¹⁰, Andrew J. Bradley³ and Gabriel C. Oniscu¹

¹Transplant Unit, Royal Infirmary of Edinburgh, Edinburgh, UK, ²NHS Blood and Transplant, Bristol, UK, ³Department of Surgery, University of Cambridge and the NIHR Cambridge Biomedical Research Centre, Cambridge, UK, ⁴Department of Renal Medicine, Freeman Hospital, Newcastle upon Tyne, UK, ⁵Department of Health Services Research and Policy, London School of Hygiene and Tropical Medicine, London, UK, ⁶Primary Care and Population Sciences, Faculty of Medicine, University of Southampton, Southampton, UK, ⁷Department of Renal Medicine, Southmead Hospital, Bristol, UK, ⁸Regional Nephrology and Transplant Centre, Belfast Health and Social Care Trust, Belfast, UK, ⁹Health Psychology Research Unit, Royal Holloway, University of London, Egham, UK and and ¹⁰Health Sciences, University of Warwick, Conventry, UK (author has moved institutions since acceptance of the article)

Correspondence and offprint requests to: Gabriel C. Oniscu; E-mail: gabriel.oniscu@ed.ac.uk

ABSTRACT

Background. Living donor kidney transplantation (LDKT) provides more timely access to transplantation and better clinical outcomes than deceased donor kidney transplantation (DDKT). This study investigated disparities in the utilization of LDKT in the UK.

Methods. A total of 2055 adults undergoing kidney transplantation between November 2011 and March 2013 were prospectively recruited from all 23 UK transplant centres as part of the Access to Transplantation and Transplant Outcome Measures (ATTOM) study. Recipient variables independently associated with receipt of LDKT versus DDKT were identified.

Results. Of the 2055 patients, 807 (39.3%) received LDKT and 1248 (60.7%) received DDKT. Multivariable modelling demonstrated a significant reduction in the likelihood of LDKT for older age {odds ratio [OR] 0.11 [95% confidence interval (CI) 0.08–0.17], P < 0.0001 for 65–75 years versus 18–34 years}; Asian ethnicity [OR 0.55 (95% CI 0.39–0.77), P = 0.0006 versus White]; Black ethnicity [OR 0.64 (95% CI 0.42–0.99), P = 0.047 versus White]; divorced, separated or widowed [OR 0.63 (95% CI 0.46–0.88), P = 0.030 versus married]; no qualifications [OR 0.55 (95% CI 0.42–0.74), P < 0.0001 versus higher education qualifications]; no car ownership [OR 0.51 (95% CI 0.37–0.72), P = 0.0001] and no home ownership [OR 0.65 (95% CI 0.85–

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0.79), P = 0.002]. The odds of LDKT varied significantly between countries in the UK.

Conclusions. Among patients undergoing kidney transplantation in the UK, there are significant age, ethnic, socio-economic and geographic disparities in the utilization of LDKT. Further work is needed to explore the potential for targeted interventions to improve equity in living donor transplantation.

Keywords: inequity, kidney transplantation, living donor, preemptive transplantation, sociodemographic disparities

INTRODUCTION

For patients with end-stage renal disease (ESRD), living donor kidney transplantation (LDKT) provides better clinical outcomes and more timely access to transplantation than deceased donor kidney transplantation (DDKT) [1–3]. Current UK Renal Association guidelines recommend that LDKT be considered the treatment of choice for all patients suitable for kidney transplantation, whenever an appropriate living donor is available [4]. In contrast to the lengthy waiting time for DDKT, the LDKT procedure can be scheduled without delay, thereby minimizing the time that patients are exposed to pre-transplant dialysis and its associated morbidity, or enabling avoidance of dialysis entirely (pre-emptive transplantation). Pre-emptive LDKT is considered by many to be an optimal treatment, providing superior graft and patient survival compared with kidney transplantation following a period of dialysis [2, 4–6].

Despite these advantages, only one-third of kidney transplants undertaken in the UK are from living donors [7]. Internationally, the UK falls behind many other countries in terms of LDKT activity [8]. A recent strategy set out by National Health Service Blood and Transplant (NHSBT) aims to increase LDKT activity in the UK from the current rate of 17 transplants per million population (pmp) to 26 transplants pmp by 2020 [9].

There are limited data on the factors that may prevent or enable patients to receive LDKT in the UK. A better understanding of these factors will facilitate the identification of target patient groups and aid the development of appropriate interventions to improve LDKT rates. The principal aim of this study was to identify the recipient characteristics associated with achieving LDKT compared with DDKT in a national sample of UK kidney transplant recipients. The study was conducted as part of the Access to Transplantation and Transplant Outcome Measures (ATTOM) research programme.

MATERIALS AND METHODS

Study population

ATTOM is a national prospective cohort study investigating the factors that influence access, clinical and patientreported outcomes and cost-effectiveness of renal transplantation in the UK. A full description of the ATTOM study methods and protocol has been reported previously [10]. As part of the ATTOM study, incident kidney transplant recipients were recruited at the time of transplantation from all 23 UK renal transplant centres. In each centre, recruitment took place over a 12-month period, between 1 November 2011 and 31 March 2013. Patients 18-75 years of age were eligible for inclusion. A total of 3002 patients received kidney-only transplants in the UK within the recruitment period; 134 were outside the study age criteria and 775 declined to participate or were not able to be approached for recruitment. In all, 38 of 2093 recruited patients were excluded from the analysis due to missing data for the main outcome variable (living or deceased donor). Thus the final analysis cohort of 2055 patients represented 72% of eligible study participants (Figure 1). There were no significant differences in the age, gender or ethnicity distributions between study participants and the national registry adult kidney transplant recipient population (data not shown) [11].

Data collection

Extensive demographic, socio-economic, clinical and comorbidity data were collected for each patient at the time of transplantation. Trained research nurses collected uniformly defined data items from patient interviews, case notes and local electronic patient information systems.

Ethnicity was coded as White, Black, Asian or other (including patients of Chinese and mixed origin). The level of highest educational attainment was coded as no qualifications, qualifications at the secondary education level or equivalent [e.g. General Certificate of Secondary Education (GCSE), General Certificate of Education Advanced level (A-level), "National Vocational Qualification (NVQ) level 1-3]" or qualifications at



FIGURE 1: Study population (asterisk refers to recruitment that took place over a 12-month period in each centre between 1 November 2011 and 31 March 2013).

the higher education level or equivalent (e.g. bachelor's degree, higher degree, "NVQ level 4–5)". Employment status was coded as employed (including full time, part time or self-employed), unemployed, long-term sick/disabled, retired or other (including those looking after the family home, those not in work for some other reason and students). The primary renal diagnosis was classified by ERA-EDTA codes [12]. Donor details and recipient calculated reaction frequency (cRF) were obtained from linkage to UK Transplant Registry data. The cRF is a measure of recipient human leucocyte antigen (HLA) sensitization, calculated as the percentage of 10 000 recent donors to which the recipient has pre-formed HLA antibodies. A comorbidity score was calculated for each patient using a modified Charlson comorbidity index for patients with ESRD [13]. The index consists of weighted scores assigned to 14 comorbid conditions (myocardial infarction, congestive heart failure, peripheral vascular disease, cerebrovascular disease, dementia, chronic pulmonary disease, rheumatological disease, peptic ulcer disease, diabetes without complications, diabetes with complications, leukaemia, lymphoma, moderate-severe liver disease and metastatic disease). Our data set did not include two of the conditions (rheumatological disease and peptic ulcer disease). Scores were therefore calculated from the remaining 12 variables.

Statistical methods

Baseline characteristics of LDKT and DDKT recipients and donors were compared by chi-squared tests for categorical data and Wilcoxon tests for non-parametric continuous data.

Recipient variables associated with receiving LDKT versus DDKT were analysed using logistic regression. Variables leading to a change in log likelihood at P < 0.15 on univariable analysis were entered into the multivariable model. The importance of each variable in the multivariable model was tested by examining the difference in log likelihood between the model with and without the variable. If the difference was not significant (P > 0.05) the variable was removed. Each time a variable was removed, the effect of removing each of the remaining variables was retested until the most parsimonious model was achieved. Potential interactions between variables were tested, none were significant. Less than 7% of values were missing for any variable. For modelling purposes, missing values were imputed using the fully conditional specification logistic regression method. In all, 10 imputed data sets were modelled separately then combined to produce final parameter estimates. Sensitivity analysis using casewise deletion of missing values did not change conclusions.

Complex links between socio-economic deprivation and ethnicity with respect to access to and outcomes from renal replacement therapy (RRT) have previously been reported [14, 15]. To avoid any confounding and/or interaction from ethnicity, a subgroup analysis was undertaken in White patients only, using the same multivariable modelling methods as described above.

A second subgroup analysis examined the recipient variables associated with receiving a transplant pre-emptively versus post-initiation of dialysis in the LDKT cohort. Multivariable modelling methods were the same as described above. All data were analysed using SAS 9.4 (SAS Institute, Cary, NC, USA).

RESULTS

Type of transplant received

Of 2055 kidney transplant recipients, 1248 (60.7%) received DDKT (583 donors after brain death and 665 donors after circulatory death) and 807 (39.3%) received LDKT. A significantly higher proportion of LDKT recipients received pre-emptive transplants compared with DDKT recipients (35.5% versus 12.0%; P < 0.0001).

Recipient characteristics

There were considerable differences in the characteristics of LDKT versus DDKT recipients (Table 1). LDKT recipients were significantly younger than DDKT recipients (median age 46 versus 53 years) and a higher proportion were of White ethnicity (87.1 versus 79.5%) and married or living with a partner (65.1 versus 60.5%). LDKT recipients were more likely to have obtained qualifications at the secondary education level (53.0 versus 47.9%) and at the higher education level (27.3 versus 18.3%). Compared with DDKT recipients, LDKT recipients had higher rates of employment (43.7 versus 31.3%), car ownership (91.0 versus 80.2%) and home ownership (66.1 versus 62.0%), suggesting they were a less socio-economically deprived population. The cause of renal failure was less likely to be diabetes, hypertension or renal vascular disease in the LDKT group. LDKT recipients had a significantly lower prevalence of comorbidity compared with DDKT recipients. The proportion of kidney transplants that were LDKTs was significantly higher in Northern Ireland (NI) at 68.5%, compared with 39.0% in England, 36.6% in Wales and 31.2% in Scotland.

Donor characteristics

Characteristics of the donors are shown in Tables 2 and 3. Living donors were significantly younger and more likely to be female than deceased donors. A higher proportion of deceased donors were of White ethnicity compared with living donors. A total of 354 (43.9%) living donors were not genetically related to the recipient. Parent, child, other blood relative and spouse living donors were more likely to be female. Pooled/altruistic living donors had the highest proportion of White donors.

Factors associated with the probability of LDKT among transplant recipients

Associations between recipient variables and the likelihood of LDKT versus DDKT were characterized using univariable and multivariable logistic regression (Table 4, Figure 2). The multivariable model demonstrated that with each sequential increase in age group, there was a marked reduction in the probability of LDKT versus DDKT, such that patients 65–75 years of age were ~90% less likely to undergo LDKT compared with patients 18–34 years of age {odds ratio [OR] 0.11 [95% confidence interval (CI) 0.08–0.17], P < 0.0001}. Compared with White patients, Asian patients [OR 0.55 (95% CI 0.39–

Table 1. Kidney transplant recipient characteristics by type of donor

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Long-term side/disability 182 (24.0) 343 (29.7) Retired 112 (14.7) 287 (24.9) Other 75 (9.9) 71 (6.2) Car ownership* 691 (91.0) 928 (80.2) <0.0001	Unemployed	59 (7.8)	92 (8.0)	
Retired 112 (14.7) 28 (24.9) Other 75 (99) 71 (6.2) Car ownership* 691 (91.0) 928 (80.2) <0.0001	Long-term sick/disability	182 (24.0)	343 (29.7)	
Other 75 (9.9) 71 (6.2) Car ownership* 691 (91.0) 288 (80.2) <0.0001 Home ownership* 501 (66.1) 716 (62.0) 0.068 Clinical variables <0.0001 Primary rend diagnosis* <0.0001 Glomerulonephritis 229 (28.5) 311 (24.9) <0.0001 Polycysit kidney disease 113 (14.1) 020 (16.8) Pyelonephritis 127 (15.8) 133 (10.7) Hypertensive nephropathy 37 (4.6) 86 (6.9) Renal vascular disease 10 (1.2) 27 (2.2) Other 156 (19.4) 193 (15.5) Uncertain 84 (10.5) 156 (12.5) 2 59 (7.3) 136 (10.9) 23 23 29 (3.6) 90 (7.2) Pre-transplant 117 (14.5) 157 (12.6) 0.212 114 (17) 60.001 Pre-transplant treatment modality* 351 (43.7) <td>Retired</td> <td>112 (14.7)</td> <td>287 (24.9)</td> <td></td>	Retired	112 (14.7)	287 (24.9)	
Car ownership* 691 (91.0) 928 (80.2) <0.0001	Other	75 (9.9)	71 (6.2)	
Hone ownersnip* 501 (66.1) 716 (62.0) 0.0068 Clinical variables	Car ownership"	691 (91.0)	928 (80.2)	< 0.0001
Primary renal diagnosis ⁴ <0.0001	Home ownership	501 (66.1)	716 (62.0)	0.068
Primary renar useries	Clinical variables			<0.0001
Datch:HermiopanityHermiopanityHermiopanityGlomerulonephritis22 (28.5)311 (24.9)Polycystic kidney disease113 (14.1)209 (16.8)Pyelonephritis127 (15.8)133 (10.7)Hypertensive nephropathy37 (4.6)86 (6.9)Renal vascular disease10 (1.2)27 (2.2)Other156 (19.4)193 (15.5)Uncertain84 (10.5)166 (12.5)Charlson comorbidity score ^a 0625 (77.7)851 (68.4)191 (11.3)168 (13.5)259 (7.3)136 (10.9) ≥ 3 29 (3.6)90 (7.2)Previous transplant117 (14.5)157 (12.6)0.0086Pre-transplant treatment modality ^a <0.001	Diabetic nenhronathy	48 (6 0)	132(10.6)	< 0.0001
bill of the term of term	Glomerulonenhritis	40 (0.0) 229 (28 5)	311 (24.9)	
InterfaceInterfaceInterfacePyelonephritis127 (15.8)133 (10.7)Hypertensive nephropathy37 (4.6)86 (6.9)Renal vascular disease10 (1.2)27 (2.2)Other156 (19.4)193 (15.5)Uncertain84 (10.5)156 (12.5)Charlson comorbidity score ^a <0.0001	Polycystic kidney disease	113 (14 1)	209 (16.8)	
Hypertensive nephropathy37 (4.6)86 (6.9)Renal vascular disease10 (1.2)27 (2.2)Other156 (19.4)193 (15.5)Uncertain84 (10.5)156 (12.5)Charlson comorbidity score ^a <0.0001	Pvelonephritis	127 (15.8)	133 (10.7)	
Renal vascular disease10 (1.2)72 (2.2)Other156 (19.4)193 (15.5)Uncertain84 (10.5)156 (12.5)Charlson comorbidity score ^a <0.0001	Hypertensive nephropathy	37 (4.6)	86 (6.9)	
$\begin{array}{cccccccccccccccccccccccccccccccccccc$	Renal vascular disease	10 (1.2)	27 (2.2)	
$ \begin{array}{cccccccccccccccccccccccccccccccccccc$	Other	156 (19.4)	193 (15.5)	
$\begin{array}{c c c c c } Charlson comorbidity score^a & <0.0001 \\ \hline 0 & 625 (77.7) & 851 (68.4) \\ \hline 1 & 91 (11.3) & 168 (13.5) \\ \hline 2 & 59 (7.3) & 136 (10.9) \\ \hline 2 & 29 (3.6) & 90 (7.2) & 0.212 \\ \hline 1 & 90 (7.2) & 0.086 & 0.212 \\ \hline 1 & 117 (14.5) & 157 (12.6) & 0.212 \\ \hline 1 & 119 (9.5) & 0.086 & 0$	Uncertain	84 (10.5)	156 (12.5)	
	Charlson comorbidity score ^a			< 0.0001
	0	625 (77.7)	851 (68.4)	
$\begin{array}{cccccccccccccccccccccccccccccccccccc$	1	91 (11.3)	168 (13.5)	
≥3 29 (3.6) 90 (7.2) Previous transplant 117 (14.5) 157 (12.6) 0.212 Highly sensitized (cRF > 85%)a 96 (11.9) 119 (9.5) 0.086 Pre-transplant treatment modalitya Pre-transplant treatment modalitya Haemodialysis 351 (43.7) 718 (57.6) (57.6) (718 (57.6) (71	2	59 (7.3)	136 (10.9)	
Previous transplant117 (14.5)157 (12.6)0.212Highly sensitized $(cRF > 85\%)^a$ 96 (11.9)119 (9.5)0.086Pre-transplant treatment modalityaHaemodialysis351 (43.7)718 (57.6)Haemodiafiltration14 (1.7)39 (3.1)Continuous ambulatory peritoneal dialysis67 (8.3)130 (10.4)Failing transplant14 (1.7)6 (0.5)Pre-emptive285 (35.5)150 (12.0)Geographic variablesCountryEngland670 (83.0)1049 (84.1)Wales34 (4.2)59 (4.7)Northern Ireland50 (6.2)23 (1.8)Scotland53 (6.6)117 (9.4)	≥ 3	29 (3.6)	90 (7.2)	
Highly sensitized (cRF > 85%) ^a 96 (11.9)119 (9.5)0.086Pre-transplant treatment modality ^a <	Previous transplant	117 (14.5)	157 (12.6)	0.212
Pre-transplant treatment modality ⁴ <0.0001	Highly sensitized (cRF $> 85\%$) ^a	96 (11.9)	119 (9.5)	0.086
Haemodialysis 351 (43.7) 718 (57.6) Haemodiafiltration 14 (1.7) 39 (3.1) Continuous ambulatory peritoneal dialysis 73 (9.1) 204 (16.4) Automated peritoneal dialysis 67 (8.3) 130 (10.4) Failing transplant 14 (1.7) 6 (0.5) Pre-emptive 285 (35.5) 150 (12.0) Geographic variables Country England 670 (83.0) 1049 (84.1) Wales 34 (4.2) 59 (4.7) Northern Ireland 50 (6.2) 23 (1.8) Scotland 53 (6.6) 117 (9.4)	Pre-transplant treatment modality ^a			< 0.0001
Haemodialitration 14 (1.7) 39 (3.1) Continuous ambulatory peritoneal dialysis 73 (9.1) 204 (16.4) Automated peritoneal dialysis 67 (8.3) 130 (10.4) Failing transplant 14 (1.7) 6 (0.5) Pre-emptive 285 (35.5) 150 (12.0) Geographic variables 20001 England 670 (83.0) 1049 (84.1) Wales 34 (4.2) 59 (4.7) Northern Ireland 50 (6.2) 23 (1.8) Scotland 53 (6.6) 117 (9.4)	Haemodialysis	351 (43.7)	718 (57.6)	
Continuous ambulatory peritoneal dialysis 73 (9.1) 204 (16.4) Automated peritoneal dialysis 67 (8.3) 130 (10.4) Failing transplant 14 (1.7) 6 (0.5) Pre-emptive 285 (35.5) 150 (12.0) Geographic variables 204 (16.4) 204 (16.4) Country 285 (35.5) 150 (12.0) Geographic variables 20001 20001 England 670 (83.0) 1049 (84.1) Wales 34 (4.2) 59 (4.7) Northern Ireland 50 (6.2) 23 (1.8) Scotland 53 (6.6) 117 (9.4)	Haemodiafiltration	14(1./)	39(3.1)	
Failing transplant 14 (1.7) 6 (0.5) Pre-emptive 285 (35.5) 150 (10.4) Geographic variables Country England 670 (83.0) 1049 (84.1) Wales 34 (4.2) 59 (4.7) Northern Ireland 50 (6.2) 23 (1.8) Scotland 53 (6.6) 117 (9.4)	Automated peritoneal dialysis	67 (8 3)	204(10.4) 130(104)	
Pre-emptive 285 (35.5) 150 (12.0) Geographic variables	Failing transplant	14 (1 7)	6 (0.5)	
Geographic variables 150 (12.0) Country <0.0001	Pre-emptive	285(355)	150 (12.0)	
Country <0.0001 England 670 (83.0) 1049 (84.1) Wales 34 (4.2) 59 (4.7) Northern Ireland 50 (6.2) 23 (1.8) Scotland 53 (6.6) 117 (9.4)	Geographic variables	200 (33.3)	130 (12.0)	
England 670 (83.0) 1049 (84.1) Wales 34 (4.2) 59 (4.7) Northern Ireland 50 (6.2) 23 (1.8) Scotland 53 (6.6) 117 (9.4)	Country			< 0.0001
Wales 34 (4.2) 59 (4.7) Northern Ireland 50 (6.2) 23 (1.8) Scotland 53 (6.6) 117 (9.4)	England	670 (83.0)	1049 (84.1)	20.0001
Northern Ireland 50 (6.2) 23 (1.8) Scotland 53 (6.6) 117 (9.4)	Wales	34 (4.2)	59 (4.7)	
Scotland 53 (6.6) 117 (9.4)	Northern Ireland	50 (6.2)	23 (1.8)	
	Scotland	53 (6.6)	117 (9.4)	

Data are median (IQR) or number (%).

^aData are missing for some participants and excluded from percentage calculations. Numbers of missing data are shown in Supplementary data, Table S1.

*Wilcoxon test for age. All others chi-squared test.

0.77), P = 0.0006] and Black patients [OR 0.64 (95% CI 0.42– 0.99), P = 0.047] were less likely to undergo LDKT than DDKT. Patients who were divorced, separated or widowed had a lower probability of LDKT compared with patients who were married or living with a partner [OR 0.63 (95% CI 0.46–0.88), P = 0.03]. Having no formal qualifications [OR 0.55 (95% CI 0.42–0.74), P < 0.0001] and having only secondary education qualifications [OR 0.76 (95% CI 0.59–0.97), P = 0.01] reduced the odds of LDKT compared with patients with higher education qualifications. Not owning a car [OR 0.51 (95% CI 0.37–0.72), P < 0.0001] and not owning a home [OR 0.65 (95% CI 0.49–0.85), P = 0.002] decreased the odds of LDKT versus DDKT. With adjustment for recipient variables, the odds of LDKT versus DDKT were >3-fold higher for patients in NI [OR 3.25 (95% CI 1.89–5.57), P < 0.0001] compared with patients in

Table 2. Donor characteristics

	Living donor $(n = 807)$	Deceased donor $(n = 1248)$	P-value*
Median age, years	48 (39–57)	54 (42-64)	< 0.0001
Age group ^a (years)			< 0.0001
<18	0 (0.0)	28 (2.2)	
18-34	141 (17.5)	156 (12.5)	
35-49	295 (36.6)	296 (23.7)	
50-64	307 (38.1)	497 (39.8)	
65-75	61 (7.6)	236 (18.9)	
>75	2 (0.3)	35 (2.8)	
Gender ^a			0.002
Male	376 (46.7)	671 (53.8)	
Female	429 (53.3)	577 (46.2)	
Ethnicity ^a			< 0.0001
White	716 (88.8)	1169 (95.0)	
Asian	50 (6.2)	22 (1.8)	
Black	28 (3.5)	22 (1.8)	
Other	12 (1.5)	17 (1.4)	

Data are median (IQR) or number (%).

^aData are missing for some participants and excluded from percentage calculations. Numbers of missing data are shown in Supplementary data, Table S1. ^{*}Wilcoxon test for age. All others chi-squared test.

Table 3. Living donor characteristics by donor-recipient relationship

England. Further analysis showed the odds of LDKT in NI were also higher compared with Wales [OR 3.77 (95% CI 1.88–7.56), P = 0.0002] and Scotland [OR 4.53 (95% CI 2.42–8.48), P < 0.0001], but there were no significant differences between patients in England, Wales and Scotland.

Factors associated with the probability of LDKT among White ethnicity transplant recipients

The same analysis was undertaken in a subgroup of White patients only (n = 1692) and confirmed that the effects of socio-economic factors on the likelihood of LDKT versus DDKT were independent of ethnicity (Table 5).

Factors associated with the probability of pre-emptive transplantation among living donor kidney transplant recipients

A further subgroup analysis in the LDKT group examined factors associated with achieving pre-emptive transplantation versus transplantation after the initiation of dialysis (Table 6). Patients with missing data for pre-transplant treatment modality (n = 3) and patients with a previous transplant (n = 117) were excluded, leaving a final cohort of 687 LDKT recipients. Multivariable analysis demonstrated a significantly decreased likelihood of pre-emptive LDKT for Asian patients [OR 0.45 (95% CI 0.23–0.86), P = 0.016], unemployed patients [OR 0.44 (95% CI 0.21–0.92), P = 0.029], patients unable to work due to long-term sickness/disability [OR 0.44 (95% CI 0.28–0.68), P = 0.0002], retired patients [OR 0.41 (95% CI 0.29–0.75), P = 0.002], not owning a car [OR 0.41 (95% CI 0.19–0.86), P = 0.018] and not owning a home [OR 0.65 (95% CI 0.44–0.96), P = 0.029].

DISCUSSION

Among patients undergoing kidney transplantation in the UK, there are significant age, ethnic, socio-economic and geographic

	Living donors ($n = 807$)						
	Parent [<i>n</i> = 147 (18.2%)]	Child [<i>n</i> = 75 (9.3%)]	Sibling [<i>n</i> = 196 (24.3%)]	Other blood relative $[n = 35 (4.3\%)]$	Spouse/partner [<i>n</i> = 188 (23.3%)]	Pooled/altruistic $[n = 93 (11.5\%)]$	Other non-related $[n = 73 (9.1\%)]$
Age group ^a (years)							
18-34	0 (0.0)	51 (68.0)	49 (25.0)	5 (14.7)	10 (5.3)	12 (12.9)	14 (19.2)
35-49	33 (22.5)	24 (32.0)	94 (48.0)	14 (41.2)	69 (36.7)	29 (31.2)	32 (43.8)
50-64	94 (64.0)	0 (0.0)	44 (22.5)	15 (44.1)	94 (50.0)	38 (40.9)	22 (30.1)
65-75	20 (13.6)	0 (0.0)	9 (4.6)	0 (0.0)	15 (8.0)	12 (12.9)	5 (6.9)
>75	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	0 (0.0)	2 (2.2)	0 (0.0)
Gender ^a							
Male	62 (42.2)	34 (45.3)	99 (50.5)	16 (47.1)	72 (38.3)	50 (53.8)	43 (59.7)
Female	85 (57.8)	41 (54.7)	97 (49.5)	18 (53.0)	116 (61.7)	43 (46.2)	29 (40.3)
Ethnicity ^a							
White	132 (89.8)	64 (85.3)	169 (86.2)	30 (88.2)	170 (90.4)	86 (92.5)	65 (89.0)
Asian	9 (6.1)	5 (6.7)	15 (7.7)	2 (5.9)	11 (5.9)	2 (2.2)	6 (8.2)
Black	2 (1.4)	5 (6.7)	10 (5.1)	2 (5.9)	4 (2.1)	4 (4.3)	1 (1.4)
Other	4 (2.7)	1 (1.3)	2 (1.0)	0 (0.0)	3 (1.6)	1 (1.1)	1 (1.4)

Data are number (%)

^aData are missing for some participants and excluded from percentage calculations. Numbers of missing data are shown in Supplementary data, Table S1.

$Table \ 4. \ Univariable \ and \ multivariable \ logistic \ regression \ analysis \ of \ factors \ associated \ with \ LDKT \ versus \ DDKT$

	Univariable		Multivariable	
	OR (95% CI)	P-value	OR (95% CI)	P-value
Demographic variables				
Age group (years)				
18-34	1 (reference)		1 (reference)	
35-49	0.41 (0.31-0.53)	< 0.0001	0.34 (0.25-0.46)	< 0.0001
50-64	0.27 (0.20–0.34)	< 0.0001	0.19 (0.14–0.27)	< 0.0001
65-/5 Conden	0.16 (0.11-0.23)	<0.0001	0.11 (0.08–0.17)	< 0.0001
Gender	1 (reference)			
Female	1 13 (0.94 - 1.36)	0 192		
Ethnicity	1.15 (0.94 1.50)	0.172		
White	1 (reference)		1 (reference)	
Asian	0.62 (0.45-0.85)	0.003	0.55 (0.39-0.77)	0.0006
Black	0.52 (0.35-0.78)	0.001	0.64 (0.42-0.99)	0.047
Other	0.49 (0.22-1.10)	0.081	0.46 (0.19-1.11)	0.084
Socio-economic variables				
Civil status				
Married/living with partner	1 (reference)		1 (reference)	
Divorced/separated/widowed	0.46 (0.34–0.63)	< 0.0001	0.63 (0.46–0.88)	0.030
Single	1.10 (0.88–1.36)	0.406	0.77 (0.58–1.02)	0.067
Qualifications	1 (100601000 00)		1 (1060100000)	
Secondary education	1 (reference) 0.73 (0.58–0.92)	0.009	1 (reference) 0.76 (0.59–0.97)	0.010
No qualifications	0.75(0.38-0.92) 0.39(0.30-0.51)	< 0.009	0.55(0.42-0.74)	< 0.010
Employment status	0.59 (0.50-0.51)	<0.0001	0.55 (0.42-0.74)	<0.0001
Employed	1 (reference)			
Unemployed	0.71 (0.50–1.02)	0.064		
Long-term sick/disability	0.58 (0.46-0.73)	< 0.0001		
Retired	0.42 (0.33-0.55)	< 0.0001		
Other	1.12 (0.79–1.58)	0.542		
Car ownership				
Yes	1 (reference)		1 (reference)	
No	0.41 (0.31–0.55)	< 0.0001	0.51 (0.37–0.72)	0.0001
Home ownership				
Yes	1 (reference)	0.052	1 (reference) $(2, 5, 6, 40, 6, 95)$	0.002
NO Clinical variables	0.82 (0.68–1.00)	0.053	0.65 (0.49-0.85)	0.002
Primary renal diagnosis				
Diabetic nephropathy	1 (reference)			
Glomerulonephritis	2.03 (1.40-2.94)	0.0002		
Polycystic kidney disease	1.48 (0.99–2.22)	0.054		
Pyelonephritis	2.62 (1.74-3.95)	< 0.0001		
Hypertensive nephropathy	1.19 (0.72–1.98)	0.498		
Renal vascular disease	1.02 (0.46-2.26)	0.968		
Other	2.22 (1.50-3.29)	< 0.0001		
Uncertain	1.48 (0.97–2.27)	0.068		
Charlson comorbidity score	1 (
0	1 (reference)	0.021		
1	0.74(0.36-0.97) 0.59(0.43-0.82)	0.031		
>3	0.55(0.45-0.62) 0.45(0.30-0.70)	0.0002		
Previous transplant		010000		
No	1 (reference)			
Yes	1.18 (0.91–1.53)	0.212		
Highly sensitized (cRF $> 85\%$)				
No	1 (reference)			
Yes	1.28 (0.97–1.71)	0.087		
Geographic variables				
England	1 (reference)		1 (reference)	
Wales	0.90 (0.59–1.39)	0.642	0.86 (0.54–1.38)	0.539
Northern Ireland	3.40(2.06-5.63)	< 0.0001	3.25(1.89-5.57)	< 0.0001
Scottand	0.71 (0.51–1.00)	0.04/	0.72 (0.50-1.03)	0.073

Т



FIGURE 2: Multivariable logistic regression analysis of factors associated with LDKT versus DDKT. N. Ireland, Northern Ireland.

disparities in the utilization of LDKT versus DDKT. Older age; Black and Asian ethnicity; being divorced, separated or widowed; lower educational attainment and measures of greater socio-economic deprivation (non-car and non-home ownership) were significantly and independently associated with a reduced likelihood of LDKT versus DDKT. For the period of the study, geographic differences were also noted, with patients in NI having a greater probability of LDKT versus DDKT compared with patients in the rest of the UK. Furthermore, the study demonstrated that among those who do undergo LDKT, ethnic and socio-economic disparities persist in determining whether LDKT is received pre-emptively. Asian ethnicity, unemployment and greater socio-economic deprivation were associated with a lower likelihood of pre-emptive LDKT versus LDKT after the initiation of dialysis.

A major strength of the present study is that we recruited all patients prospectively and collected accurate, reliable and comprehensive data. A large proportion (72%) of the national adult kidney transplant population was included in the study. Nevertheless, as it was not possible to recruit the entire kidney transplant population, it must be recognized that the study is limited by a risk of selection bias. Reassuringly, the age, gender and ethnicity of study participants were not significantly different from the national adult kidney transplant population [11]. Furthermore, the study cohort included patients from all 23 UK renal transplant centres as well as nationally comparable proportions of LDKT, DDKT and pre-emptive recipients, thereby reducing the potential for bias. However, differences in other unmeasured characteristics between study participants and non-participants cannot be ruled out. Another limitation of the study is that we were unable to account for the fact that some patients may not have had a medically suitable living donor. This could be a potential explanation for the observed lower utilization of LDKT for certain patient groups. It is known that ethnic minorities have a higher prevalence of hypertension and diabetes with associated ESRD, thus precluding kidney donation [16, 17]. Similarly, greater socio-economic deprivation is linked to poorer health [18], potentially limiting the pool of living donors available to more deprived patients. Furthermore, due to the observational nature of the study, the results can only describe associations and thus the causality of the observed relationships cannot be inferred.

In recent years, a great deal of attention has been directed towards disparities in access to DDKT in the UK. Individuals who are older, more socially deprived, from ethnic minority backgrounds or treated in certain transplant centres are less likely to be listed for and subsequently receive DDKT [19-23]. Despite LDKT providing optimal clinical outcomes for patients with ESRD, there have been limited data on whether patients experience disparities in utilizing this treatment. Udayaraj *et al.* [24], reported a lower probability of LDKT for patients with greater socio-economic deprivation and patients from Black and South Asian backgrounds in the UK. However, this study analysed the rates of LDKT among patients starting RRT, therefore a major confounding factor is the poorer health among more socio-economically deprived and ethnic minority populations, leading to a higher proportion of patients being medically unsuitable for transplantation. The present study adds new knowledge about the factors associated with receiving LDKT as opposed to DDKT among a cohort of patients deemed suitable to undergo transplantation. This is a select population of patients who have already successfully navigated the process of

Table 5. Multivariable logistic regression analysis of factors associated

 with LDKT versus DDKT among White patients only

Recipient variables	OR (95% CI)	P-value
Age group (years)		
18-34	1 (reference)	
35-49	0.31 (0.22-0.44)	< 0.0001
50-64	0.17 (0.12-0.25)	< 0.0001
65–75	0.11 (0.07-0.17)	< 0.0001
Civil status		
Married/living with partner	1 (reference)	
Divorced/separated/widowed	0.60 (0.42-0.86)	0.006
Single	0.70 (0.51-0.96)	0.028
Qualifications		
Higher education	1 (reference)	
Secondary education	0.73 (0.55-0.96)	0.027
No qualifications	0.53 (0.38-0.74)	0.0001
Car ownership		
Yes	1 (reference)	
No	0.50 (0.35-0.73)	0.0003
Home ownership		
Yes	1 (reference)	
No	0.68 (0.50-0.91)	0.01
Country		
England	1 (reference)	
Wales	0.91 (0.56-1.47)	0.693
Northern Ireland	3.43 (1.98-5.95)	< 0.0001
Scotland	0.71 (0.49–1.04)	0.076

Table 6. Multivariable logistic regression analysis of factors associated with pre-emptive LDKT

Recipient variables	OR (95% CI)	P-value
Ethnicity		
White	1 (reference)	
Asian	0.45 (0.23-0.86)	0.016
Black	1.19 (0.53-2.65)	0.672
Other	1.17 (0.17-7.79)	0.874
Employment status		
Employed	1 (reference)	
Unemployed	0.44 (0.21-0.92)	0.029
Long-term sick/disability	0.44 (0.28-0.68)	0.0002
Retired	0.47 (0.29-0.75)	0.002
Other	1.41 (0.80-2.50)	0.240
Car ownership		
Yes	1 (reference)	
No	0.41 (0.19-0.86)	0.018
Home ownership		
Yes	1 (reference)	
No	0.65 (0.44–0.96)	0.029

transplant referral, evaluation and listing. Therefore, it is concerning that the striking disparities observed appear to occur over and above the well-recognized inequities that patients face before even reaching this stage. These findings are not confined to the UK. Our results are consistent with those of a USA study by Gore *et al.* [25], which reported lower odds of LDKT relative to DDKT for patients who were older, from ethnic minority groups, with lower socio-economic status and with lower levels of education. Roodnat *et al.* [26], showed the same factors reduced the likelihood of LDKT versus DDKT in The Netherlands. It is interesting that similar results have been demonstrated both within publicly funded as well as private health care systems, suggesting factors other than financial disadvantage play an important role.

The well-recognized markers of socio-economic deprivation (car ownership and home ownership) were strongly associated with a reduced likelihood of LDKT versus DDKT in this study. A subgroup analysis of only White patients confirmed that the effects of socio-economic deprivation were independent of ethnicity. Lower rates of LDKT in socio-economically deprived patients have also been reported in Australia [27] and the USA [28, 29]. The reasons behind this finding are unclear. It is known that living donor-recipient pairs usually come from the same socio-economic group [30]. In the UK, kidney transplantation including medication and aftercare are provided free of charge. However, it is possible that other costs such as transportation, childcare and lost income from time off work could play a role in deterring potential living donors or deterring those in need of a kidney from approaching potential donors [31]. A financial reimbursement policy for expenses incurred by living donors does exist in the UK, but it is not implemented consistently by transplant centres. A recent qualitative study of DDKT recipients found that many were unaware of the living donor reimbursement policy [32]. Despite this, socio-economically deprived patients did not perceive financial concerns to be a major barrier to LDKT and described passivity and disempowerment in treatment decisions, short-term focus and lack of social support as more significant obstacles to LDKT [32].

It is well recognized that ethnic minority patients wait longer for DDKT in the UK, due to the mismatch between the HLA types of minority patients and those of the predominantly White donor pool [33]. One might, therefore, expect a higher uptake of LDKT in ethnic minority patients. Our study found the opposite, with patients from Black and Asian backgrounds having lower odds of LDKT than DDKT compared with White patients. Similar disparities have been reported in the USA [15, 34] and Canada [35]. These disparities have worsened over time and are likely contributing to differences in outcomes between White and non-White patients [36]. The reasons for these disparities are not well understood. Possible explanations cited include cultural and religious beliefs [37, 38], reluctance to engage with the medical system [39, 40], institutional prejudice [41, 42], language barriers [43] and concern over a higher risk for living donors from minority ethnic backgrounds [44–46].

We have demonstrated that a patient's level of educational attainment is independently associated with their likelihood of LDKT versus DDKT. Educational attainment is related to health literacy, which has been shown to be an important factor for both potential kidney transplant recipients as well as potential living donors in successfully navigating the living donation and transplantation process [47, 48]. Higher academic achievement may be linked to a better ability to understand the benefits of LDKT or to take part in informed and shared decision making.

The finding that patients who were married or living with a partner had better access to LDKT is likely to be related to the opportunity for spousal donation. Spouses represented a considerable proportion (23.3%) of living donors in this study, and the

majority were female (61.7%). Being married or living with a partner may also confer other benefits, such as having a better social support network or access to more unrelated or child donors.

Older age was associated with dramatically reduced odds of LDKT versus DDKT. Previous research has demonstrated that older age is associated with a lower probability of attempted donor recruitment [49]. Older patients have reported an unwill-ingness to put younger donors at risk, particularly their children [50]. In our study, 18.2% of the living donors were parents while only 9.3% were children.

Despite adjustment for demographic and socio-economic factors, we found striking geographic differences in LDKT activity, with patients in NI experiencing higher odds of LDKT versus DDKT compared with patients in England, Wales and Scotland. Our results reflect the actual number of LDKTs pmp, which were around twice as high in NI (31.1) compared with the rest of the UK (England 15.9, Wales 16.6, Scotland 10.9) at the time of the study [51]. Around this time, an initiative was begun in NI to promote LDKT and pre-emptive transplant as the treatment of choice. The key measures included education to promote a change of mindset among nephrologists (particularly nontransplant nephrologists) as well as the entire transplant team, together with improved infrastructure and more streamlined services to enable timely workup and transplantation (e.g. one-stop living donor assessment clinic). Effective leadership, persistence and gaining the support of commissioners and management were critical in achieving these changes [A. Courtney (personal communication, 17 January 2017)]. Our results and the national figures indicate that such a strategy can be very successful in increasing LDKT utilization. The higher LDKT rate in NI led to a lower DDKT rate (NI 15.0, England 24.9, Wales 33.0, Scotland 26.7) [51] and there are now very few long-waiting patients on the waiting list in NI [52]. Moreover, the number of LDKTs in NI has continued to increase (40 pmp in 2016, one of the highest rates in the world), demonstrating that the changes have led to a sustained improvement rather than a temporary peak in activity. This is encouraging when exploring potential avenues to improve LDKT across the UK as a whole.

Our study showed for the first time in the UK that socioeconomic deprivation, unemployment and Asian ethnicity were independently associated with a lower likelihood of pre-emptive LDKT. These findings are consistent with studies from the USA and Australia [5, 25, 27]. The disparity experienced by socioeconomically deprived individuals is likely to be related to an increased likelihood of late referral to specialist renal services in the UK [53]; however, this does not explain the disparity for patients of Asian ethnicity.

LDKT, and in particular pre-emptive LDKT, provides optimal clinical outcomes for patients with ESRD, yet its uptake is variable within the UK. This study has identified specific patient groups with a lower likelihood of undergoing LDKT relative to DDKT. We have demonstrated that demographic, socioeconomic and geographic factors are more strongly associated with the type of transplant received rather than clinical factors, including comorbidity, primary renal diagnosis, HLA sensitization or previous transplantation. Moreover, a remarkable finding is that even among LDKT recipients, disparities persist in receiving pre-emptive transplantation. This demonstrates the strength of social factors in influencing access to health care and may reflect similar inequities across a wide range of health care services. The demonstrated disparities may reflect both barriers in certain patient groups as well as important positive factors in others. Furthermore, these influencing factors are likely to apply to both potential recipients and donors. If particular groups experience avoidable barriers to LDKT receiving or donating, there is a responsibility to provide tailored resources to remove these barriers. Improving access to LDKT will not only benefit individual patients, but will also have favourable effects for the wider ESRD population by effectively increasing the overall pool of available organs. However, both donor and recipient welfare and autonomy undoubtedly remain the primary focus. Some patients may prefer not to pursue LDKT due to concerns about risks to their potential donors, just as some potential donors may be unwilling to donate [50, 54].

Identifying disadvantaged patient groups is essential to directing further research into potentially modifiable factors and appropriate interventions. Several studies in the USA have explored targeted interventions, including culturally sensitive education programmes [55, 56], home-based education [57, 58] and patient advocates [59], with promising results for reducing disparities in LDKT. Similar programmes in the UK may provide a more equitable opportunity for disadvantaged patients to explore the option of LDKT.

SUPPLEMENTARY DATA

Supplementary data are available online at http://ndt.oxfordjournals.org.

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ETHICS APPROVAL

East of England Research Ethics Committee (reference number 11/EE/0120).

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AUTHORS' CONTRIBUTIONS

All authors contributed to the study design. D.W. conducted the literature review and data analysis. M.L.R. and R.J.J. provided statistical input for the data analysis. D.W. and G.C.O. drafted the manuscript. All authors interpreted the data, provided intellectual content, revised the drafts and approved the final version. D.W. and G.C.O. are guarantors for the paper.

CONFLICT OF INTEREST STATEMENT

None declared. The results presented in this article have not been published previously in whole or part, except in abstract form.

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