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Association of the Kidney Allocation System with Dialysis Exposure before Deceased Donor Kidney Transplantation by Pre-Emptive Wait-listing Status

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Abstract

Background: It is unknown whether the new kidney transplant allocation system (KAS) has attenuated the advantages of preemptive wait-listing as a strategy to minimize pre-transplant dialysis exposure.

Methods: We performed a retrospective study of adult US deceased donor kidney transplant (DDKT) recipients between December 4, 2011-December 3, 2014 (pre-KAS) and December 4, 2014-December 3, 2017 (post-KAS). We estimated pre-transplant dialysis durations by preemptive listing status in the pre- and post-KAS periods using multivariable gamma regression models.

Results: Among 65,385 DDKT recipients, preemptively listed recipients (21%, n=13,696) were more likely to be white (59% vs 34%, p<0.001) and have private insurance (64% vs 30%, p<0.001). In the pre- and post-KAS periods, average adjusted pre-transplant dialysis durations for

Disclosures

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MNH and MOH designed the study and analyzed the data; MNH, MOH, KR, SMB, LLM, SG, GEM, GX, DJR, and REP interpreted the data and drafted and revised the manuscript; REP provided mentorship on the study design and interpretation; all authors approved the final version of the manuscript.

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preemptively listed recipients were <2 years in all racial groups. Compared to recipients who were listed after starting dialysis, preemptively listed recipients experienced 3.85 (95% Confidence Interval [CI] 3.71–3.99) and 4.53 (95% CI 4.32–4.74) fewer average years of pre-transplant dialysis in the pre- and post-KAS periods, respectively (p<0.001 for all comparisons).

Conclusions: Preemptively wait-listed DDKT recipients continue to experience substantially fewer years of pre-transplant dialysis than recipients listed after dialysis onset. Efforts are needed to improve both socioeconomic and racial disparities in preemptive wait-listing.

Keywords

kidney transplant; waiting time; Kidney; waiting list

Introduction

On December 4, 2014, the Organ Procurement and Transplantation Network implemented major changes to the deceased donor kidney allocation policy in the United States. Under the new kidney allocation system (KAS), deceased donor kidney transplant (DDKT) candidates who are waitlisted after the onset of maintenance dialysis accrue a priority point for each year of pre-listing dialysis exposure, while candidates who are preemptively listed (i.e., listed before dialysis onset) continue to receive a point for each year waiting after reaching the qualifying estimated glomerular filtration rate of 20 milliliters/minute/1.73 meters squared.¹ Early studies on the effects of the KAS have indicated that more individuals with long dialysis durations are receiving DDKT, whereas fewer individuals are receiving preemptive DDKT (i.e., DDKT before the need for maintenance dialysis).^{2–5} Therefore, it is unknown whether DDKT candidates who were preemptively waitlisted as a strategy to minimize pre-transplant dialysis exposure will continue to receive a similar benefit under the new KAS.^{6,7}

Prior to the KAS, DDKT candidates who were preemptively wait-listed with a qualifying eGFR and those who were wait-listed after starting maintenance dialysis accrued waiting time from their listing date onward. In this context, pre-emptively wait-listed individuals were found to have superior transplant outcomes compared to those listed after dialysis onset, a finding attributed to both socioeconomic differences and to the deleterious health impacts of prolonged dialysis exposure.^{6–8} As unequal access to the kidney transplant waiting list among low income and minority candidates contributed to large socioeconomic and racial disparities in DDKT,⁹ one of the primary objectives of the new kidney allocation system (KAS) was to improve equity in organ allocation for candidates who were wait-listed after enduring years of dialysis.¹⁰ Early studies have shown that the KAS has been successful in closing the gap in transplant rates between wait-listed whites and minorities.² However, data are needed on other potential implications of prioritizing dialysis exposure for organ allocation, including its impacts on the known relative benefits of preemptive wait-listing.^{7,8,11,12}

The goal of this study was to examine whether the new KAS was associated with differences in pre-transplant dialysis durations for DDKT recipients with and without preemptive waiting time. We performed a retrospective pre-post cohort study to examine whether

average pre-transplant dialysis durations differed among kidney transplant recipients before and after KAS implementation based on the recipient's preemptive listing status and by race/ ethnicity.

Methods

Study Population

The study population was derived from all individuals who received DDKT in the US between December 4, 2011 and December 3, 2017, as indicated in the United Network for Organ Sharing's (UNOS) standard transplant analytic (STAR) file. Multi-organ recipients were excluded given material differences in organ allocation protocol. To focus on DDKT recipients who did not receive pediatric priority, recipients were excluded if they were <18 years at the time of wait-listing (Appendix Figure 1). Preemptively waitlisted recipients were identified based on dialysis status at the time of wait-listing. Pre-transplant dialysis durations were calculated by subtracting recipient dialysis dates (derived from the STAR file TRR form) from transplant dates. If TRR form dialysis dates were missing, we verified that TCR form dialysis dates were also missing, or used the TCR form dialysis dates if they were non-missing. The study was approved by the Institutional Review Board at the Drexel University College of Medicine and all study procedures were in accordance with the Declaration of Helsinki.

Covariates

Variables were collected from the UNOS STAR file at the time of transplant and included recipient age at DDKT (as a continuous and squared term to account for non-linear associations), race/ethnicity (white/black/Hispanic/other), sex, diabetes history, body mass index category (categorized as <18.5 kg/m², 18.5 to <25 kg/m², 25 to <30 kg/m², 30 to <40 kg/m², and _40 kg/m²), human immunodeficiency virus (HIV) serostatus, hepatitis C serostatus, prior living organ donor history, prior organ transplant history, calculated panel reactive antibody (as a continuous and squared term to account for non-linear associations), blood group; waiting time from date of wait-listing to transplant, kidney donor profile index (categorized as 0 to _20, >20 to _85, >85),¹³ the donor's public health service (PHS) increased risk status, zero antigen human leukocyte antigen mismatch, private versus other insurance status, educational attainment (< High School, _High School Graduate & < College, _College Graduate, Unknown), and UNOS organ procurement organization region, for which there are 11.

Statistical Analysis

All analyses were performed using STATA/MP version 14 for Mac (College Station, TX, USA). Categorical variables (e.g., sex, ethnicity) were described by their frequencies. Continuous variables (e.g., age) were described by their medians and ranges. Binary variables were compared between groups using chi-square tests. To compare continuous variables between groups, we used Wilcoxon rank-sum tests and Kruskal-Wallis tests, as appropriate.

Multivariable Modeling Strategy

First, we examined pre-transplant dialysis exposure among recipients with and without preemptive wait-listing over time using a multivariable generalized linear model with a gamma family and log link. We estimated average pre-transplant dialysis durations by race/ ethnicity and preemptive listing status during each quarter (three-month period) of our study period. Next, to examine whether the KAS was associated with differences in pre-transplant dialysis durations among DDKT recipients with preemptive listing, we estimated a multivariable generalized linear model with a gamma family and log link for years of pre-transplant dialysis among DDKT recipients with preemptive listing that included an interaction term for KAS period (pre/post) and race/ethnicity. We then compared pre-transplant dialysis durations among recipients with preemptive wait-listing in the pre- and post-KAS periods, by race/ethnicity, with a difference-in-differences approach.¹⁴

To determine whether the KAS was associated with changes in the relative difference in pretransplant dialysis durations between DDKT recipients without and without preemptive wait-listing, we estimated a multivariable generalized linear model for years of pretransplant dialysis exposure that included a three-way interaction term for KAS period, race/ ethnicity, and preemptive wait-listing status. We then compared the adjusted average difference in duration of pre-transplant dialysis among recipients with and without preemptive wait-listing, by race/ethnicity, in the pre-and post-KAS periods. We used marginal standardization (i.e., predicted probabilities summed to a weighted average of the distribution of confounders in the cohort) to calculate adjusted dialysis durations by preemptive wait-listing status and race/ethnicity group in each time frame of the study period.¹⁵ We used the 'margins' package in STATA, with confidence intervals estimated using the delta method.¹⁶ To account for potential clustering of waiting time by transplant center we calculated cluster-robust standard errors.¹⁷

Sensitivity Analyses

Given prior evidence of a "bolus effect" of DDKT among newly listed individuals with long dialysis durations in the immediate post-KAS period,^{1–3} we performed sensitivity analysis in which we estimated dialysis durations after partitioning the post-KAS period into the first 12-month period post-KAS (period 1) and the following 24 months post-KAS years (period 2). We also performed sensitivity analyses in which we compared dialysis durations in the post-KAS period to the pre-KAS period after excluding the first six months after the KAS was implemented. Finally, we considered a model in which we also excluded prior living donors, given the prioritization of these individuals for transplantation,¹⁸ to determine if this exclusion impacted our results.

Missing Data

With the exception of HIV serostatus (missing in less than 7% of the cohort), data were missing in less than 3% of the cohort on any covariates, and all primary analyses were performed on complete cases.¹⁹ In sensitivity analyses, we imputed data for missing observations of HIV serostatus using multiple imputation with 10 iterations.²⁰ (supplementary Table 4 and 5).

Results

Study Population

Among 65,385 DDKT recipients included in the study (Appendix Figure 1), 46% (n=30,126) received DDKT in the pre-KAS period and 53% (n=35,259) received DDKT in the post-KAS period (Table 1). Compared to the pre-KAS period, DDKT recipients in the post-KAS period were younger (median 54 vs 56 years, p<0.001), fewer were white (36% vs 43%, p<0.001), and more were female (40% vs 39%, p=0.003). In the pre-KAS period, 24% of recipients (n=7,282) had been preemptively wait-listed, compared to 14.6% of recipients within the first 12 months after the KAS, and 19.8% of recipients between December 4, 2015 to December 3, 2017 (Appendix Table 1 and Appendix Figure 2). Table 2 compares characteristics between recipients with and without preemptive wait-listing in the pre- and post-KAS periods, respectively. Overall, compared to recipients without preemptive waitlisting, preemptively listed recipients were more likely to be white (59% vs 34%, p<0.001) have private insurance (64% vs 30%, p<0.001), and be college graduates (31.2% vs 20.2%, p<0.001). The proportion of organs allocated with KDPI<20 (i.e., the highest quality allografts) was similar between recipients with and without preemptive listing in the pre-KAS period (20% vs 19%, p=0.26), and was higher among individuals with preemptive listing in the post-KAS period (24% vs 19%, p<0.001).

Trends in Pre-Transplant Dialysis Duration Among DDKT Recipients Who Were and Were Not Preemptively Waitlisted

Among recipients listed after dialysis, average pre-transplant dialysis durations rose steeply in the last quarter (three-month period) of 2014 and peaked in 2015, with a subsequent decline in 2016 and 2017 (Figure 1). In the fourth quarter of 2017, average pre-transplant dialysis duration among recipients without preemptive listing was 5.18 years (95% Confidence Interval [CI]: 4.85–5.51 years) among whites, 6.30 years (95% CI: 5.91–6.69 years) among blacks, 5.72 years (95% CI: 5.35–6.09 years) among Hispanics, and 5.43 years (95% CI 4.87–5.99 years) among other race/ethnicities. Average pre-transplant dialysis durations remained similar over the study period within all racial groups of preemptively listed recipients. Among preemptively listed recipients in the last quarter of 2017, average pre-transplant dialysis durations were 1.12 years (95% CI: 0.96–1.29 years) among preemptively-listed white recipients, 1.79 years (95% CI: 1.40–2.18 years) among preemptively-listed black recipients, 1.53 years among preemptively-listed Hispanic recipients (95% CI: 1.12–1.96 years), and 1.51 years among preemptively listed recipients of other races/ethnicities (95% CI 1.05–1.97 years).

Association of the KAS and Pre-Transplant Dialysis Duration Among Recipients with Preemptive Wait-listing by Race/Ethnicity

In a multivariable adjusted generalized linear difference-in-differences model for the outcome of pre-transplant dialysis duration, the KAS was associated with non-statistically significant differences in pre-transplant dialysis duration among preemptively listed white recipients (1.19 vs 1.24 years, p=0.29), preemptively listed black recipients (1.64 vs 1.74 years, p=0.14), and preemptively listed recipients of other races/ethnicities (1.61 vs 1.70, p=0.37). The KAS was associated with a statistically significant 0.2-year increase in dialysis

exposure among preemptively listed Hispanic recipients (1.43 vs 1.63 years, p=0.01) (Table 3, Appendix Figure 3).

Association of the KAS and the Difference in Pre-Transplant Dialysis Duration Among Recipients with and without Preemptive Wait-listing by Race/Ethnicity

Compared to recipients without preemptive wait-listing, recipients with preemptive waitlisting received DDKT with 3.85 fewer average years of pre-transplant dialysis than those listed after dialysis in the pre-KAS period (95% CI 3.71–3.99 years, p<0.001), compared to 4.53 fewer years of pre-transplant dialysis in the post-KAS period (95% CI 4.32–4.74 years, p<0.001) (difference-in-differences of 0.66 additional years, p<0.001). The post-KAS gap in pre-transplant dialysis duration between preemptively listed and non-preemptively listed recipients became wider within all race/ethnicity groups, with the widest gap in dialysis duration between black recipients with and without preemptive listing (Table 4).

Sensitivity Analyses

Results were similar when comparing pre-transplant dialysis durations among preemptively listed recipients between the pre-KAS period to the early and late post-KAS periods, respectively, and after excluding the first six months of the post-KAS period (Appendix Tables 2 and 3). The post-KAS period was associated with similar gaps in pre-transplant dialysis duration between recipients with and without preemptive wait-listing following multiple imputation for missing HIV serostatus (Appendix Tables 4 and 5). Our results were also robust to the exclusion of prior living donors (Appendix Table 6).

Conclusion

This nationally representative study is the first to examine whether the KAS, in prioritizing DDKT for candidates with long dialysis durations, was associated with differences in the pre-transplant dialysis durations of recipients with and without preemptive wait-listing for DDKT. Our results demonstrate that DDKT recipients with preemptive wait-listing continue to receive DDKT with substantially fewer years of pre-transplant dialysis than those without preemptive wait-listing under the KAS, underscoring the importance of efforts to improve preemptive access to the DDKT waiting list.

Individuals who receive DDKT after prolonged dialysis exposure have higher risks of graft loss and death than individuals with preemptive and early DDKT.^{11,21,22} As expected given the prioritization of pre-listing dialysis time, preemptive transplantation rates have declined under the new KAS.^{3–5} However, our findings suggest that preemptive wait-listing remains advantageous to minimize pre-transplant dialysis durations under the new KAS, independent of recipients' types of insurance coverage and their races/ethnicities. We found that DDKT recipients with preemptive wait-listing in the post-KAS period received transplant after dialysis durations of less than two years on average, compared to recipients who were listed after starting dialysis, who had average pre-transplant dialysis durations of five to six years. Numerous factors may be maintaining the wide gap in pre-transplant dialysis durations between DDKT recipients with and without preemptive listing under the new KAS. For example, individuals who begin maintenance dialysis before being wait-listed may

encounter numerous additional health burdens that delay transplant referral and prolong waiting time, including increased risks of functional dependence, vascular disease, and hospitalizations.^{23–32} Delayed transplant referral after dialysis onset may also be a reflect variation in dialysis center transplant referral practices,^{33,34} a factor that is coming under increasing scrutiny as a potential quality-of-care indicator.^{35,36}

In addition to promoting shorter pre-transplant dialysis durations, preemptive wait-listing may also increase the likelihood of receiving the highest quality allografts under the new KAS. Acknowledging the deleterious effects of long dialysis durations on health and transplant outcomes, the KAS incorporated dialysis duration into a new longevity matching paradigm, that allocates the highest quality kidneys for recipients who are expected to live longest.³⁷ This policy change may help to explain our finding that recipients who were listed after dialysis were equally likely as preemptively listed recipients to receive the highest quality allografts before the KAS, and less likely to receive these kidneys after the KAS. In aggregate, our results suggest that efforts to further improve equitable outcomes in DDKT under the new KAS may require added focus on improving unequal access to preemptive wait-listing.^{9,38–40} Unequal access may be driven in part by lack of health insurance among many US individuals with non-dialysis dependent chronic kidney disease, 40,41 a disparity which may be narrowing with the national gains in insurance coverage under the Affordable Care Act (ACA).^{42–44} In addition to providing affordable health insurance coverage options, efforts to improve access to preemptive wait-listing should also include educational interventions targeted to those most likely to lack CKD awareness and pre-dialysis health care, including low income individuals with CKD. 45-50

Historically, racial and ethnic minority DDKT candidates have been less likely to be preemptively waitlisted than white candidates.^{9,40,51,52} Racial disparities in DDKT rates have improved under the new KAS,^{6,53} but concerns have been raised that the improvements in racial disparities in DDKT may only be temporary.^{2–4} Experts have posited that early gains in DDKT among racial and ethnic minorities may have represented a "bolus effect" in which a relatively small proportion of individuals with very long dialysis durations receive DDKT, followed by a return to prior patterns of inequitable organ access.^{3,4} Consistent with this hypothesis, a recent study by Melanson and colleagues showed that dialysis durations among new DDKT recipients peaked in all racial groups immediately after implementation of the KAS, though white DDKT recipients continued to experience substantially shorter dialysis durations prior to DDKT than black and Hispanic recipients in the post-KAS period. ² Our study results suggest that this latter finding may be explained in part by racial differences in preemptive wait-listing. For example, though the absolute and relative benefits of preemptive wait-listing with respect to pre-transplant dialysis minimization were raceindependent, the majority of recipients with preemptive listing were white in both the preand post-KAS periods. Therefore, increasing rates of preemptive listing among racial and ethnic minorities with advanced CKD is a high priority to improve equity in DDKT outcomes under the new KAS.^{7,11,54–56} Efforts are needed to address numerous factors that may contribute to delays in transplant referral among minorities with CKD, including geographic and provider-related variation in referral for transplant evaluation, ^{33,57–62} perceived discrimination,⁶³ excess financial burdens,^{9,51,64,65} and lack of supportive social networks.66,67

Our study has several strengths, including a large, national study sample and focus on the potential implications of policy changes on existing disparities in transplantation. However, our study must be considered with respect to its limitations, particularly concerning the possible biases associated with retrospective, observational analyses of registry data. For example, the retrospective study design may be vulnerable to selection bias due to the lack of data on CKD patients who may be eligible for preemptive wait-listing. Our analysis is also limited by a lack of granular data on recipient socioeconomic status and community-level health indicators, which may have important implications for transplant candidacy.⁶⁸ Further, our study does not provide insight on other unmeasured confounders, such as social support or health literacy, that may also impact transplant referral and waiting time.⁶⁹

In summary, this study found that among kidney transplant recipients of all races and ethnicities, preemptive wait-listing continues to confer a large benefit with respect to minimizing pre-transplant dialysis duration compared to listing after dialysis under the new KAS. Future studies should be directed at mitigating persistent drivers of disparate access to preemptive wait-listing.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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Abbreviations:

BMI	Body Mass Index
CI	Confidence Interval
cPRA	Calculated Panel Reactive Antibody
DDKT	Deceased Donor Kidney Transplantation
HIV	Human Immunodeficiency Virus
KAS	Kidney Allocation System
KDPI	Kidney Donor Profile Index
UNOS	United Network for Organ Sharing
US	United States

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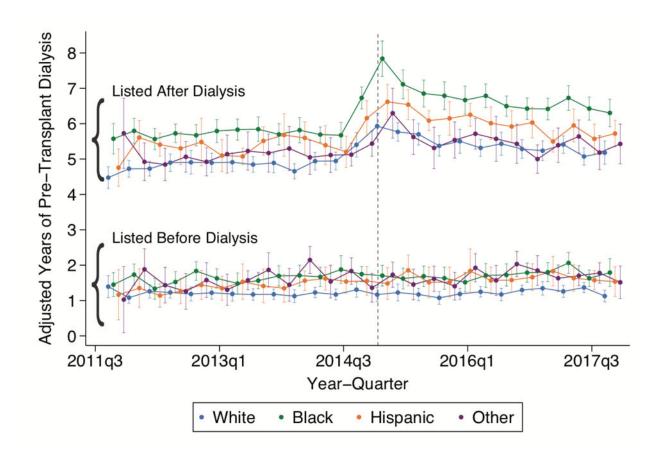


Figure 1: Average Pre-Transplant Dialysis Durations of Deceased Donor Kidney Transplant Recipients in the United States by Race/Ethnicity and Preemptive Wait-Listing Status.

This figure displays results of multivariable generalized linear model for average pretransplant dialysis duration, in years, by recipient race/ethnicity and preemptive wait-listing status. The dashed line indicates the onset of the new kidney allocation system on December 4, 2014. Each year/quarter point estimate (with 95% Confidence Interval) represents the predicted marginal mean dialysis duration within race/ethnicity group and preemptive listing category. Green, orange, blue, and purple circles indicate black, Hispanic, white, and other race/ethnicity recipients, respectively.

Table 1:

Kidney Transplant Recipient Characteristics, Stratified by Transplant in Pre- or Post- Kidney Allocation System Period

	Pre-KAS December 4, 2011-December 3, 2014	Post-KAS December 4, 2014-December 3, 2017	p-value
	N=30126	N=35259	
Race/Ethnicity			< 0.001
White	12860 (42.7%)	12793 (36.3%)	
Black	9671 (32.1%)	12492 (35.4%)	
Hispanic	4865 (16.1%)	6594 (18.7%)	
Other	2730 (9.1%)	3380 (9.6%)	
Preemptively Listed	7282 (24.2%)	6414 (18.2%)	< 0.001
Preemptively Transplanted	2689 (8.9%)	2368 (6.7%)	< 0.001
KDPI Category			< 0.001
≤20	5805 (19.3%)	6855 (19.4%)	
>20 - <85	21219 (70.4%)	25346 (71.9%)	
>85	3071 (10.2%)	3057 (8.7%)	
missing	31 (0.1%)	1 (<1%)	
Age (years)	56.0 (45.0, 64.0)	54.0 (43.0, 63.0)	< 0.001
Sex			0.003
Female	11835 (39.3%)	14250 (40.4%)	
Male	18291 (60.7%)	21009 (59.6%)	
Prior Living Donor			0.53
No	30017 (99.6%)	35136 (99.7%)	
Yes	107 (0.4%)	115 (0.3%)	
Missing	2 (<1%)	8 (<1%)	
Private Insurance			< 0.001
No	17633 (58.5%)	23242 (65.9%)	
Yes	12489 (41.5%)	12012 (34.1%)	
Missing	4 (<1%)	5 (<1%)	
Educational Attainment			0.28
Less than High School	2197 (7.3%)	2683 (7.6%)	
High School Graduate	20208 (67.1%)	23667 (67.1%)	
College Graduate or Higher	6813 (22.6%)	7901 (22.4%)	
Missing/Unknown	908 (3.0%)	1008 (2.9%)	
Diabetic			< 0.001
No	18825 (62.5%)	23023 (65.3%)	
Yes	11223 (37.3%)	12200 (34.6%)	
Missing	78 (0.3%)	36 (0.1%)	
BMI Category (kg/m ²)			< 0.001
18.5–24.9	8212 (27.3%)	10279 (29.2%)	
<18.5	479 (1.6%)	608 (1.7%)	
25-29.9	10273 (34.1%)	11807 (33.5%)	

	Pre-KAS December 4, 2011-December 3, 2014	Post-KAS December 4, 2014-December 3, 2017	p-value
	N=30126	N=35259	
30–39.9	10591 (35.2%)	12048 (34.2%)	
≥40	571 (1.9%)	517 (1.5%)	
HIV Positive			< 0.001
No	27850 (92.4%)	33687 (95.5%)	
Yes	251 (0.8%)	528 (1.5%)	
Missing	2025 (6.7%)	1044 (3.0%)	
HCV Positive			< 0.001
No	28483 (94.5%)	32901 (93.3%)	
Yes	1502 (5.0%)	2253 (6.4%)	
Missing	141 (0.5%)	105 (0.3%)	
Blood Group			0.018
А	10580 (35.1%)	12117 (34.4%)	
A1	338 (1.1%)	349 (1.0%)	
A1B	28 (0.1%)	46 (0.1%)	
A2	57 (0.2%)	43 (0.1%)	
A2B	6 (<1%)	13 (<1%)	
AB	1570 (5.2%)	1833 (5.2%)	
В	3922 (13.0%)	4744 (13.5%)	
0	13625 (45.2%)	16114 (45.7%)	
cPRA	0.0 (0.0, 44.0)	0.0 (0.0, 62.0)	< 0.001
Zero Antigen Mismatch	2527 (8.4%)	1726 (4.9%)	< 0.001
PHS Increased Risk Allograft			< 0.001
No	25673 (85.2%)	26884 (76.2%)	
Yes	4440 (14.7%)	8366 (23.7%)	
Missing	13 (<1%)	9 (<1%)	
Prior Organ Transplant	3870 (12.8%)	5215 (14.8%)	< 0.001
Days Inactive on Waiting List	18.0 (0.0, 258.0)	0.0 (0.0, 219.0)	< 0.001
Waiting Time			< 0.001
<1 Year	6721 (22.3%)	12047 (34.2%)	
1-3 Years	10923 (36.3%)	10674 (30.3%)	
≥3 Years	12482 (41.4%)	12538 (35.6%)	

Abbreviations: BMI—Body Mass Index; KAS—kidney allocation system; cPRA—calculated panel reactive antibody; KDPI—kidney donor profile index; kg—kilograms; m—meters; PHS—Public Health Service

Values presented as median (interquartile range) and n (%)

 I p-values comparing non-missing data from Chi-Square Tests and Wilcoxon Rank Sum Tests, as appropriate

Table 2.

Kidney Transplant Recipient and Allograft Characteristics, Stratified by Preemptive Listing Status and Transplant in Pre- or Post- Kidney Allocation System Period

	Pre-KAS December 4, 2011-December 3, 2014	AS ecember 3, 2014		Post-KAS December 4, 2014-December 3, 2017	AS ecember 3, 2017	
	Not Preemptively Listed Preemptively Listed	Preemptively Listed	p-value ^a	Not Preemptively Listed	Preemptively Listed	p -value b
	22844	7282		28845	6414	
Race/Ethnicity			<0.001			<0.001
White	8455 (37.0%)	4405 (60.5%)		9109 (31.6%)	3684 (57.4%)	
Black	8112 (35.5%)	1559 (21.4%)		11094 (38.5%)	1398 (21.8%)	
Hispanic	4212 (18.4%)	653 (9.0%)		5885 (20.4%)	709 (11.1%)	
Other	2065 (9.0%)	665 (9.1%)		2757 (9.6%)	623 (9.7%)	
Age (years)	55.0 (44.0, 63.0)	58.0(49.0,65.0)	<0.001	53.0 (42.0, 62.0)	57.0 (46.0, 65.0)	< 0.001
Sex			<0.001			< 0.001
Female	8608 (37.7%)	3227 (44.3%)		11117 (38.5%)	3133 (48.8%)	
Male	14236 (62.3%)	4055 (55.7%)		17728 (61.5%)	3281 (51.2%)	
Waiting Time (from Listing to Transplant)			<0.001			< 0.001
<1 Year	5627 (24.6%)	1094 (15.0%)		10708 (37.1%)	1339 (20.9%)	
1–2.9 Years	8208 (35.9%)	2715 (37.3%)		8809 (30.5%)	1865 (29.1%)	
≥ 3 Years	9009 (39.4%)	3473 (47.7%)		9328 (32.3%)	3210 (50.0%)	
Prior Donor			0.022			<0.001
No	22772 (99.7%)	7245 (99.5%)		28781 (99.8%)	6355 (99.1%)	
Yes	71 (0.3%)	36 (0.5%)		58 (0.2%)	57 (0.9%)	
Missing	1 (<1%)	1 (<1%)		6 (<1%)	2 (<1%)	
Private Insurance			<0.001			<0.001
No	15059 (65.9%)	2574 (35.3%)		20923 (72.5%)	2319 (36.2%)	
Yes	7784 (34.1%)	4705 (64.6%)		7919 (27.5%)	4093 (63.8%)	
Missing	1 (<1%)	3 (<1%)		3 (<1%)	2 (<1%)	
Education			<0.001			<0.001
Less than High School	1923 (8.4%)	274 (3.8%)		2448 (8.5%)	235 (3.7%)	
High School Graduate	15676 (68.6%)	4532 (62.2%)		19732 (68.4%)	3935 (61.4%)	
College Graduate or Higher	4576 (20.0%)	2237 (30.7%)		5854 (20.3%)	2047 (31.9%)	

	Pre-KAS December 4, 2011-December 3, 2014	AS ecember 3, 2014		Post-KAS December 4, 2014-December 3, 2017	AS ecember 3, 2017	
	Not Preemptively Listed 22844	Preemptively Listed 7282	p-value ^a	Not Preemptively Listed 28845	Preemptively Listed 6414	p-value ^b
Missing/Unknown	669 (2.9%)	239 (3.3%)		811 (2.8%)	197 (3.1%)	
KDPI			0.26			<0.001
>20	18454 (80.8%)	5836 (80.1%)		23519 (81.5%)	4884 (76.1%)	
	4369 (19.1%)	1436 (19.7%)		5325 (18.5%)	1530 (23.9%)	
Missing	21 (0.1%)	10~(0.1%)		1 (<1%)	0(0.0%)	
Zero Antigen Mismatch	1796 (7.9%)	731 (10.0%)	<0.001	1177 (4.1%)	549 (8.6%)	<0.001
Prior Transplant	2846 (12.5%)	1024 (14.1%)	<0.001	4197 (14.6%)	1018 (15.9%)	0.007
Diabetic			<0.001			<0.001
No	13832 (60.5%)	4993 (68.6%)		18334 (63.6%)	4689 (73.1%)	
Yes	8962 (39.2%)	2261 (31.0%)		10484 (36.3%)	1716 (26.8%)	
Missing	50 (0.2%)	28 (0.4%)		27 (0.1%)	9 (0.1%)	
BMI Category (kg/m ²)			<0.001			0.003
18.5–24.9	6193 (27.1%)	2019 (27.7%)		8420 (29.2%)	1859 (29.0%)	
<18.5	384 (1.7%)	95 (1.3%)		493 (1.7%)	115 (1.8%)	
25-29.9	7757 (34.0%)	2516 (34.6%)		9589 (33.2%)	2218 (34.6%)	
30–39.9	8039 (35.2%)	2552 (35.0%)		9889 (34.3%)	2159 (33.7%)	
≥40	471 (2.1%)	100 (1.4%)		454 (1.6%)	63 (1.0%)	
HIV Positive			<0.001			<0.001
No	21062 (92.2%)	6788 (93.2%)		27476 (95.3%)	6211 (96.8%)	
Yes	226 (1.0%)	25 (0.3%)		499 (1.7%)	29 (0.5%)	
Unknown	1556 (6.8%)	469 (6.4%)		870 (3.0%)	174 (2.7%)	
HCV Positive			<0.001			<0.001
No	21475 (94.0%)	7008 (96.2%)		26843 (93.1%)	6058 (94.4%)	
Yes	1263 (5.5%)	239 (3.3%)		1920 (6.7%)	333 (5.2%)	
Unknown	106 (0.5%)	35 (0.5%)		82 (0.3%)	23 (0.4%)	
Blood Group			<0.001			<0.001
А	7849 (34.4%)	2731 (37.5%)		9604 (33.3%)	2513 (39.2%)	
AI	251 (1.1%)	87 (1.2%)		287 (1.0%)	62 (1.0%)	

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	December 4, 2011-December 3, 2014	ecember 3, 2014		December 4, 2014-December 3, 2017	ecember 3, 2017	
	Not Preemptively Listed	Preemptively Listed	p-value ^a	Not Preemptively Listed Preemptively Listed p-value a Not Preemptively Listed Preemptively Listed p-value b	Preemptively Listed	p-value
	22844	7282		28845	6414	
A1B	22 (0.1%)	6 (0.1%)		32 (0.1%)	14 (0.2%)	
A2	36 (0.2%)	21 (0.3%)		32 (0.1%)	11 (0.2%)	
A2B	6 (<1%)	0 (0.0%)		9 (<1%)	4 (0.1%)	
AB	1165 (5.1%)	405 (5.6%)		1418 (4.9%)	415 (6.5%)	
В	3037 (13.3%)	885 (12.2%)		3908 (13.5%)	836~(13.0%)	
0	10478 (45.9%)	3147 (43.2%)		13555 (47.0%)	2559 (39.9%)	
Calculated PRA	0.0 (0.0, 44.0)	$0.0\ (0.0,\ 41.0)$	0.57	$0.0\ (0.0,\ 59.0)$	0.0 (0.0, 70.0)	<0.001

_____ tibody

Values presented as median (interquartile range) and n (%)

^a P-values comparing non-missing data from Chi-Square Tests and Wilcoxon Rank Sum Tests, as appropriate among recipients in pre-KAS period

b-values comparing non-missing data from Chi-Square Tests and Wilcoxon Rank Sum Tests, as appropriate among recipients in post-KAS period

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Table 3.

Comparison of Adjusted Pre-Transplant Dialysis Durations (in Years) among Recipients with Preemptive Wait-Listing between the Pre- and Post-KAS Periods

	Pre-Transplant Dialys	sis Exposure (in Years)	Difference in Years:	
	Pre-KAS December 4, 2011-December 3, 2014	Post-KAS December 4, 2014-December 3, 2017	Between Periods	In difference
White	1.19 (95% CI 1.10-1.28)	1.24 (95% CI 1.14–1.34)	+0.05 (95% CI -0.04-0.14)	reference
Black	1.64 (95% CI 1.51–1.78)	1.74 (95% CI 1.59–1.89)	+0.10 (95% CI -0.03-0.23)	-0.05 (95% CI -0.18- 0.09)
Hispanic	1.43 (95% CI 1.25–1.61)	1.63 (95% CI 1.48-1.79)	+0.20 [*] (95% CI 0.05–0.36)	-0.15 (95% CI -0.32- 0.02)
Other	1.61 (95% CI 1.42–1.79)	1.70 (95% CI 1.52–1.87)	+0.09 (95% CI -0.11-0.29)	-0.04 (95% CI -0.24- 0.16)

Abbreviations: KAS-Kidney Allocation System; CI-Confidence Interval

* p<0.05

Estimates represent predicted marginal mean dialysis durations that are standardized to the cohort distributions of recipient age (years), sex, diabetes status, hepatitis C status, HIV status, prior living donor status, prior organ transplant status, OPTN region, calculated PRA, blood group, PHS increased risk status, kidney donor profile index category, zero HLA antigen mismatch, education level, private insurance status, and waiting time years)

Table 4.

Comparison of Adjusted Differences in Pre-Transplant Dialysis Durations (in Years) Between Transplant Recipients with and without Preemptive Wait-Listing between the Pre- and Post-KAS Periods

	Difference in Pre-Transplant	Difference in Pre-Transplant Dialysis Exposure (in Years)		Difference in Years:		
	Pre-KAS December 1, 2012-November 30, 2014	Post-KAS December 1, 2014-December 31, 2016	Proportional Difference Between Periods	In difference		
White	3.66 (95% CI 3.52-3.79)	4.22 (95% CI 4.06–4.38)	0.57 (95% CI 0.41–0.72)**	reference		
Black	4.07 (95% CI 3.87-4.27)	5.07 (95% CI 4.78–5.35)	0.99 (95% CI 0.74–1.24)**	-0.43 (95% CI -0.68 - -0.17) *		
Hispanic	3.96 (95% CI 3.69-4.24)	4.41 (95% CI 4.08-4.76)	0.45 (95% CI 0.28–0.93)*	0.11 (95% CI -0.21-0.44)		
Other	3.47 (95% CI 3.23–3.71)	3.86 (95% CI 3.54-4.17)	0.39 (95% CI 0.06–0.71) [*]	0.18 (95% CI -0.14 - 0.50)		

KAS-Kidney Allocation System; CI-Confidence Interval

Estimates represent predicted marginal mean dialysis durations that are standardized to the cohort distributions of recipient age, sex, waiting time, educational attainment, private insurance status, diabetes status, Human Immunodeficiency Virus serostatus, Hepatitis C serostatus, prior living donor, prior organ transplant, UNOS region, calculated PRA, blood group, PHS increased risk status, kidney donor profile index, zero HLA antigen mismatch

p<0.05

*

** p<0.001