## Letters

### **RESEARCH LETTER**

# Thirty-Year Trends in Perioperative Mortality Risk for Living Kidney Donors

Living kidney donor candidates must be accurately informed of the risks of perioperative mortality. The best current estimate, 3.1 deaths within 90 days per 10 000 donations, comes from a study of donors from 1994 to 2009. Open donor ne-



#### Supplemental content

phrectomy, standard of care in the 1990s, has now been replaced almost completely by

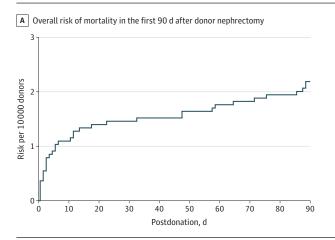
laparoscopic nephrectomy. <sup>2,3</sup> Because of this transition, as well as improvements in donor selection, perioperative care, and surgical technique, <sup>4</sup> prior estimates of perioperative mortality may not accurately represent current risk to donors. We performed a national registry study to characterize temporal trends in perioperative mortality in donors and risk factors associated with this event.

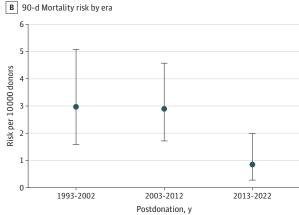
Methods | Using Scientific Registry of Transplant Recipients data on living kidney donors from 1993 to 2022, we calculated mortality ratios within 90 days of donation, stratified across 3 eras: 1993 through 2002, 2003 through 2012, and 2013 through 2022. Death events were captured from Organ Procurement and Transplantation Network (OPTN) living donor follow-up reported by transplant programs; National Technical Information Service Limited Access Death Master File; and deaths made available to the OPTN contractor through an interagency data sharing agreement between the Centers for Medicare & Medicaid Services and the Health Resources and Services Administration, and secondarily verified by the OPTN contractor. Per OPTN policy, all donor

deaths within 2 years of donation must be reported within 72 hours of the hospital's becoming aware of the death. The institutional review board of NYU Langone Health determined that this study of deidentified data did not constitute human subjects research and waived consent. We compared mortality across eras and across subgroups of donors by age, sex, race, ethnicity, procedure type, body mass index (BMI; calculated as weight in kilograms divided by height in meters squared), and hypertension, using Fisher exact tests and reporting Clopper-Pearson exact CIs. Subgroups by procedure type and BMI category were analyzed during 1999-2022 and by history of hypertension during 2004-2022, when these variables were reliably captured in the registry. Owing to the small number of events, we did not conduct multivariable analyses. Analyses were conducted with Stata 17, using 2-sided tests with  $\alpha = .05$ .

Results | There were 164 593 donors in our study, of whom 36 died within 90 days postdonation (2.2 per 10 000 cases); 50% of deaths occurred within the first 7 days (Figure, A). When cause-of-death data were available, the most common cause was hemorrhage (8 deaths of 19 with reported cause of death). Mortality was comparable in 1993-2002 (13 deaths, 3.0 [95% CI, 1.6-5.1] per 10 000) and 2003-2012 (18 deaths, 2.9 [95% CI, 1.7-4.6] per 10 000) and then declined statistically significantly in 2013-2022 (5 deaths, 0.9 [95% CI, 0.3-2.0] per 10 000; P = .01) (Figure, B). There were no statistically significant differences across subgroups of age, race, and ethnicity; compared with that of White donors (2.0 per 10 000), mortality was higher for Black donors (4.2 per 10 000) and lower among donors who were neither Black nor White (1.3 per 10 000; P = .12) (Table). Male donors were at higher risk than

### Figure. Perioperative Mortality and Donor Nephrectomy





A, Eight individuals died due to hemorrhage; 2, due to infection; 2, due to pulmonary embolism; 2, due to cardiovascular causes; 1, due to cerebrovascular accident; 1, due to homicide; and 3, due to other reasons not specified.

B, Vertical lines denote 95% confidence bars. Cause-of-death information was available for 19 of the 28 deaths that occurred in 2000 or later.

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Table. Donor Characteristics 1993-2022 and Risk of Mortality

Overall         164593         36 (2.2)         NA           Era		No. of donors	No. (per 10 000) who died within 90 d	P value
1993-2002 43752 13 (3) 2003-2012 62 185 18 (2.9) 2013-2022 58 656 5 (0.9)  Age, y³ ≤40 74 421 14 (1.9) 40-49 47 308 15 (3.2) ≥50 58 656 7 (1.6)  Sex³  Male 64 703 26 (4) Female 99 890 10 (1)  Race³  Black 19 011 8 (4.2) Other or multiracial⁵ 22 994 3 (1.3) White 122 525 25 (2)  Ethnicity (any race)³  Hispanic/Latino 22 379 4 (1.8) Non-Hispanic/non-Latino 142 214 32 (2.3)  Among donors 1999-2022  Procedure type³  Open 16 447 7 (4.3) Laparoscopic 124 771 24 (1.9)  BMI category³ Underweight or healthy weight 46 338 6 (1.3) Overweight 54 775 12 (2.2) Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension³ No 106 967 15 (1.4)  03	Overall	164 593	36 (2.2)	NA
2003-2012 62 185 18 (2.9)	Era			
2013-2022	1993-2002	43 752	13 (3)	.01
Age, y³  \$40 74 421 40 -49 40 -49 40 -49 250 58 656 7 (1.6)  Sex³  Male 64 703 Female 99 890 10 (1)  Race³  Black 19 011 8 (4.2) Other or multiracialb 22 994 3 (1.3) White 122 525 25 (2)  Ethnicity (any race)³  Hispanic/Latino 22 379 Non-Hispanic/non-Latino 142 214 32 (2.3)  Among donors 1999-2022  Procedure type³  Open 16 447 16 447 Open 16 447 17 (4.3) Laparoscopic 124 771 24 (1.9)  BMI category³ Underweight or healthy weight 54 775 12 (2.2) Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension³ No 106 967 15 (1.4)  03	2003-2012	62 185	18 (2.9)	
\$\frac{\$40}{40-49} & 47 308 & 15 (3.2) \\ \frac{250}{250} & 58 656 & 7 (1.6)\$  \$\frac{\$58x^a\$}{Male} & 64 703 & 26 (4) \\ Female & 99 890 & 10 (1)\$  \$\frac{\$64703}{\$10} & 8 (4.2)\$  Other or multiracial \( \frac{1}{2} & 22 994 \)  White & 122 525 & 25 (2)  \$\frac{\$250}{\$10} & 8 (4.2)\$  Other or multiracial \( \frac{1}{2} & 22 994 \)  White & 122 525 & 25 (2)  \$\frac{\$250}{\$25} & 25 (2)\$  \$\f	2013-2022	58 656	5 (0.9)	
40-49 47 308 15 (3.2) .25  250 58 656 7 (1.6)  Sex³  Male 64703 26 (4) Female 99 890 10 (1)  Race³  Black 19 011 8 (4.2) Other or multiracialb 22 994 3 (1.3) .12  White 122 525 25 (2)  Ethnicity (any race)³  Hispanic/Latino 22 379 4 (1.8) Non-Hispanic/non-Latino 142 214 32 (2.3)  Among donors 1999-2022  Procedure type³  Open 16 447 7 (4.3) Laparoscopic 124 771 24 (1.9)  BMI category³ Underweight or healthy weight 46 338 6 (1.3) Overweight 54 775 12 (2.2) .59  Among donors 2004-2022  Predonation hypertension³ No 106 967 15 (1.4)	Age, y <sup>a</sup>			
250 58 656 7 (1.6)  Sex³  Male 64703 26 (4) Female 99 890 10 (1)  Race³  Black 19 011 8 (4.2) Other or multiracialb 22 994 3 (1.3) White 122 525 25 (2)  Ethnicity (any race)³  Hispanic/Latino 22 379 4 (1.8) Non-Hispanic/non-Latino 142 214 32 (2.3)  Among donors 1999-2022  Procedure type³  Open 16 447 7 (4.3) Laparoscopic 124771 24 (1.9)  BMI category³ Underweight or healthy weight 46 338 6 (1.3) Overweight 54 775 12 (2.2) Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension³ No 106 967 15 (1.4)	≤40	74 421	14 (1.9)	.25
Male	40-49	47 308	15 (3.2)	
Male       64703       26 (4)       <.001	≥50	58 656	7 (1.6)	
Female 99 890 10 (1)  Race <sup>a</sup> Black 19 011 8 (4.2) Other or multiracial <sup>b</sup> 22 994 3 (1.3) .12 White 122 525 25 (2)  Ethnicity (any race) <sup>a</sup> Hispanic/Latino 22 379 4 (1.8) Non-Hispanic/non-Latino 142 214 32 (2.3)  Among donors 1999-2022  Procedure type <sup>a</sup> Open 16 447 7 (4.3) Laparoscopic 124 771 24 (1.9)  BMI category <sup>a</sup> Underweight or healthy weight 46 338 6 (1.3) Overweight 54 775 12 (2.2) .59 Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)	Sex <sup>a</sup>			
Female 99 890 10 (1)  Racea  Black 19 011 8 (4.2)  Other or multiracial <sup>b</sup> 22 994 3 (1.3)  White 122 525 25 (2)  Ethnicity (any race)a  Hispanic/Latino 22 379 4 (1.8)  Non-Hispanic/non-Latino 142 214 32 (2.3)  Among donors 1999-2022  Procedure typea  Open 16 447 7 (4.3)  Laparoscopic 124 771 24 (1.9)  BMI categorya  Underweight or healthy weight 46 338 6 (1.3)  Overweight 54 775 12 (2.2)  Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertensiona  No 106 967 15 (1.4)	Male	64 703	26 (4)	<.001
Black 19 011 8 (4.2) Other or multiracial <sup>b</sup> 22 994 3 (1.3) White 122 525 25 (2)  Ethnicity (any race) <sup>a</sup> Hispanic/Latino 22 379 4 (1.8) Non-Hispanic/non-Latino 142 214 32 (2.3)  Among donors 1999-2022  Procedure type <sup>a</sup> Open 16 447 7 (4.3) Laparoscopic 124 771 24 (1.9)  BMI category <sup>a</sup> Underweight or healthy weight 46 338 6 (1.3) Overweight 54 775 12 (2.2) Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)	Female	99 890	10 (1)	
Other or multiracial <sup>b</sup> 22 994 3 (1.3)	Race <sup>a</sup>			
White 122 525 25 (2)  Ethnicity (any race) <sup>a</sup> Hispanic/Latino 22 379 4 (1.8)  Non-Hispanic/non-Latino 142 214 32 (2.3)  Among donors 1999-2022  Procedure type <sup>a</sup> Open 16 447 7 (4.3)  Laparoscopic 124 771 24 (1.9)  BMI category <sup>a</sup> Underweight or healthy weight 46 338 6 (1.3)  Overweight 54 775 12 (2.2)  Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)	Black	19011	8 (4.2)	.12
Ethnicity (any race) <sup>a</sup> Hispanic/Latino Non-Hispanic/non-Latino 142 214 32 (2.3)  Among donors 1999-2022  Procedure type <sup>a</sup> Open 16 447 17 (4.3) Laparoscopic 124 771 24 (1.9)  BMI category <sup>a</sup> Underweight or healthy weight 46 338 Overweight 54 775 12 (2.2) Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)  81  81  81  81  81  81  81  81  81  8	Other or multiracial <sup>b</sup>	22 994	3 (1.3)	
Hispanic/Latino 22 379 4 (1.8) Non-Hispanic/non-Latino 142 214 32 (2.3)  Among donors 1999-2022  Procedure type <sup>a</sup> Open 16 447 7 (4.3) Laparoscopic 124 771 24 (1.9)  BMI category <sup>a</sup> Underweight or healthy weight 46 338 6 (1.3)  Overweight 54 775 12 (2.2) Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)	White	122 525	25 (2)	
Non-Hispanic/non-Latino   142 214   32 (2.3)   .81	Ethnicity (any race) <sup>a</sup>			
Non-Hispanic/non-Latino       142 214       32 (2.3)         Among donors 1999-2022       Procedure type <sup>a</sup> Open       16 447       7 (4.3)         Laparoscopic       124 771       24 (1.9)         BMI category <sup>a</sup> Underweight or healthy weight       46 338       6 (1.3)         Overweight       54 775       12 (2.2)       .59         Obese       32 417       6 (1.9)         Among donors 2004-2022       Predonation hypertension <sup>a</sup> No       106 967       15 (1.4)	Hispanic/Latino	22 379	4 (1.8)	.81
Procedure type <sup>a</sup> Open 16 447 7 (4.3) .08  Laparoscopic 124 771 24 (1.9)  BMI category <sup>a</sup> Underweight or healthy weight 46 338 6 (1.3)  Overweight 54 775 12 (2.2) .59  Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)	Non-Hispanic/non-Latino	142 214	32 (2.3)	
Open       16 447       7 (4.3)       .08         Laparoscopic       124 771       24 (1.9)       .08         BMI category³       Underweight or healthy weight       46 338       6 (1.3)       .59         Overweight       54 775       12 (2.2)       .59         Obese       32 417       6 (1.9)         Among donors 2004-2022       Predonation hypertension³         No       106 967       15 (1.4)	Among donors 1999-2022			
Laparoscopic   124771   24 (1.9)   .08	Procedure type <sup>a</sup>			
Laparoscopic 124771 24 (1.9)  BMI category <sup>a</sup> Underweight or healthy weight 46 338 6 (1.3)  Overweight 54775 12 (2.2)  Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)	Open	16 447	7 (4.3)	.08
Underweight or healthy weight 46 338 6 (1.3)  Overweight 54 775 12 (2.2)  Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)	Laparoscopic	124771	24 (1.9)	
Overweight     54 775     12 (2.2)     .59       Obese     32 417     6 (1.9)       Among donors 2004-2022       Predonation hypertension <sup>a</sup> No     106 967     15 (1.4)       03	BMI category <sup>a</sup>			
Obese 32 417 6 (1.9)  Among donors 2004-2022  Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)	Underweight or healthy weight	46 338	6 (1.3)	.59
Among donors 2004-2022 Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)	Overweight	54 775	12 (2.2)	
Predonation hypertension <sup>a</sup> No 106 967 15 (1.4)	Obese	32 417	6 (1.9)	
No 106 967 15 (1.4)	Among donors 2004-2022			
03	Predonation hypertension <sup>a</sup>			
Yes 4006 3 (7.5)	No	106 967	15 (1.4)	.03
	Yes	4006	3 (7.5)	

Abbreviations: BMI, body mass index (calculated as weight in kilograms divided by height in meters squared); NA, not applicable.

female donors (4.0 vs 1.0 per 10 000). Mortality was higher for donors who had open (vs laparoscopic) procedures, although not statistically significant (4.3 vs 1.9 per 10 000; P = .08). Mortality was consistent across categories of BMI. Mortality was higher for donors with a history of predonation hypertension (7.5 vs 1.4 per 10 000; P = .03).

Discussion | Perioperative mortality after living donation declined substantially in the past decade compared with prior decades, to fewer than 1 event per 10 000 donations. Risk was higher for male donors and donors with a history of hypertension. Current guidelines<sup>5</sup> for donor informed consent, based on 2009 data, should be updated to reflect this information.

Study limitations include that with only 36 perioperative deaths in 30 years, power to estimate relative risks is limited. Additionally, follow-up may be incomplete. However, OPTN requirements for donor follow-up have grown stricter, with 6-month follow-up required for donors since 2013; as such, missingness would bias toward lower mortality in earlier eras, the opposite of what the study found.

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**Author Contributions:** Drs Massie and Segev had full access to all of the data in the study and take responsibility for the integrity of the data and the accuracy of the data analysis.

Concept and design: Massie, Levan, Segev.

Acquisition, analysis, or interpretation of data: All authors.

Drafting of the manuscript: Massie, Levan.

Critical review of the manuscript for important intellectual content: Motter, Snyder, Levan, Segev.

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<sup>&</sup>lt;sup>a</sup> Missingness is as follows: age, 7 donors; sex, 0; race, 63; ethnicity, 0; era, 0; procedure type (among donors 1999-2022), 2121; BMI (among donors 1999-2022), 9808; and predonation hypertension (among donors 2004-2022), 3399.

b Race and ethnicity were assessed to investigate whether risk of perioperative mortality differentially affects members of minority groups. Race and ethnicity were self-reported, with options defined by the Organ Procurement and Transplantation Network. The "other" category includes American Indian or Alaska Native, Asian, multiracial, Native Hawaiian or Other Pacific Islander, and unknown.

Statistical analysis: Massie, Motter.

Obtained funding: Segev.

Administrative, technical, or material support: Motter, Snyder, Levan. Supervision: Segev.

Conflict of Interest Disclosures: Dr Snyder reported serving as director of the Scientific Registry of Transplant Recipients (which supplied the publicly available data), a contract with the Health Resources and Services Administration of the US Department of Health and Human Services. Dr Levan reported fees from PatientsLikeMe and Takeda for research consultancy as a key opinion leader outside the submitted work. Dr Segey reported grants from the National Institutes of Health during the conduct of the study; consulting for AstraZeneca, CareDx, Moderna Therapeutics, Novavax, and Regeneron; and speaker fees and honoraria from Springer Publishing, AstraZeneca, CareDx, Houston Methodist, Northwell Health, Optum Health Education, Sanofi, and WebMD. No other disclosures were reported.

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Registry of Transplant Recipients (SRTR). The SRTR data system includes data on all donor, wait-listed candidates, and transplant recipients in the US, submitted by the members of the Organ Procurement and Transplantation Network (OPTN). The interpretation and reporting of these data are the responsibility of the author(s) and in no way should be seen as an official policy of or interpretation by the SRTR or the US government.

#### Data Sharing Statement: See the Supplement.

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