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Geographic disparities in liver supply/demand ratio within fixeddistance and fixed-population circles

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Abstract

Recent OPTN proposals to address geographic disparity in liver allocation have involved circular boundaries: the policy selected 12/17 allocated to 150-mile circles in addition to DSAs/regions, and the policy selected 12/18 allocates to 150-mile circles eliminating DSA/region boundaries. However, methods to reduce geographic disparity remain controversial, within the OPTN and the transplant community. To inform ongoing discussions, we studied center-level supply/demand ratios using SRTR data (07/2013–06/2017) for 27,334 transplanted deceased donor livers and 44,652 incident waitlist candidates. Supply was the number of donors from an allocation unit (DSA or circle), allocated proportionally (by waitlist size) to the centers drawing on these donors. We measured geographic disparity as variance in log-transformed supply/demand ratio, comparing allocation based on DSAs, fixed-distance circles (150- or 400-mile radius), and fixed-population (12- or 50-million) circles. The recently proposed 150-mile radius circles (variance=0.11, p=0.9) or 12-million-population circles (variance=0.08, p=0.1) did not reduce geographic disparity compared to DSA-based allocation (variance=0.11). However, geographic disparity decreased substantially to 0.02 in both larger fixed-distance (400-mile, p<0.001) and larger fixed-population (50-million, p<0.001) circles (p=0.9 comparing fixed-distance and fixed-population). For allocation circles to reduce geographic disparities, they must be larger than a 150-mile radius; additionally, fixed-population circles are not superior to fixed-distance circles.

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Author Contributions

Disclosure

Data Availability Statement

All authors contributed to the conception and design or acquisition of data or analysis and interpretation of the data, drafting the article or revising it critically for important intellectual content, and final approval of the version to be published.

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The data that support these findings are available from the Scientific Registry of Transplant Recipients March 2018 external release.

INTRODUCTION

Although the Model for End-stage Liver Disease (MELD) based allocation system prioritizes the sickest patients, geographic disparities in organ supply and demand mean that the sickest patients are frequently not at the top of the priority list when an organ becomes available (1–6). Geographic disparity can be illustrated by the large range in median allocation MELD scores in different donor service areas (DSAs), from 20 in Indianapolis, Indiana, to 39 in Los Angeles, California (7), by the variability in liver transplant rates (8), and by the wide DSA variation in liver supply/demand ratios and mortality rates (1, 9, 10).

To address these disparities, the Organ Procurement and Transplantation Network (OPTN) approved policy changes in December 2017 that included: adding to the region a 150-mile circle surrounding the donor hospital, lowering the MELD threshold for regional sharing to 32, and adding proximity points to MELD for candidates within the DSA or within 150 nautical miles of the donor hospital (11, 12). However, before the December 2017 changes were implemented, the OPTN board decided that organ allocation policies must respect new geographical frameworks, which allow circular allocation and which disallow the use of DSA and region boundaries (13, 14). The liver committee wanted to test allocation circles based on population density, but the short timeline for policy change did not allow that (15). The liver committee, after a very close vote, advocated a broader 2-circle distribution policy that treated candidates from MELD 15 up to MELD 31 with identical priority, but in December 2018 the board selected a different policy, acuity circles, to share at several MELD tiers across 150-mile, 250-mile, and 500-mile circles, eliminating DSA or region boundaries (16, 17). None of the aforementioned changes to liver allocation have been implemented. The goal of our study is to inform this ongoing discussion, and it remains possible that more changes to policy are forthcoming.

Fixed-distance circular areas might or might not reduce geographic disparities in transplantation. First, size matters: the smaller the circles, the less likely organs will reach the highest MELD candidates in the country. With a US surface area of 3.8 million square miles divided across 58 DSAs, the average radius of each DSA is 144 miles; therefore, circles larger than 150 miles might be needed to improve on current geographic disparities. Second, fixed-distance circles will have different effects on centers based on each center's physical location (in coastal versus land-locked cities, near oceans or national borders, in densely versus sparsely populated areas), so fixed-population circles might be more equitable than fixed-distance circles.

To examine the feasibility of reducing geographic disparities through use of circular allocation boundaries, we used national transplant registry and ZIP code data to compare supply/demand variance between DSA-based allocation to smaller versus larger fixed-distance circles (150- or 400-mile radius) and comparably-drawn smaller versus larger fixed-population circles (12- or 50-million-population).

METHODS

Data source

This study used data from the Scientific Registry of Transplant Recipients (SRTR) external release made available in March 2018. The SRTR data system includes data on all donors, waitlisted candidates, and transplant recipients in the US submitted by members of the Organ Procurement and Transplantation Network (OPTN) and has been described elsewhere (18). The Health Resources and Services Administration, United States Department of Health and Human Services provides oversight to the activities of the OPTN and SRTR contractors.

Study population

We studied 27,334 transplanted deceased liver donors and 44,652 waitlist candidates who first achieved an allocation MELD/PELD 15 during the study period (i.e. incident candidates), from 142 liver transplant centers from 7/2013–06/2017, after excluding donors and waitlist candidates from Hawaii and Puerto Rico. We did not include candidates who were already listed and achieved an allocation MELD/PELD 15 before the study period (i.e. prevalent candidates).

Fixed-distance and fixed-population circles

The distance between transplant center and each ZIP code was calculated in Python 3.6.1 using Python's geopy module (version 1.11.0) and Vincenity distance. ZIP code longitude, latitude, and population data were taken from U.S. Census Bureau 2015 Gazetteer Files and U.S. Census Bureau 2015 American Community Survey 5 Year Estimates. ZIP codes were sorted by their centroid distance to transplant center. All ZIP codes with distances less than or equal to the specified fixed radius (150 or 400 miles) were added to the fixed-distance circle around the transplant center. The population within each ZIP code was added to fixed-population circle, in order of their distance from the transplant center, until the population within the circle became greater than or equal to the specified fixed-population (12 or 50 million). Populations of 12 million and 50 million were chosen to designate areas with populations comparable to the average population within fixed-radius circles of 150 and 400 miles, respectively.

Supply/demand ratio under current DSA allocation

We calculated the supply/demand ratio for each of the 50 DSAs with a liver transplant center. For each DSA, we calculated the number of deceased donor livers recovered in the study period from all ZIP codes within the DSA (ZIP_{donors}). Six DSAs do not have a liver transplant center, so supply from these DSAs was distributed according to the number of deceased donor livers from these DSAs that were transplanted in each of the other 50 DSAs during the study period (DSA_{donors}). The supply for each of the 50 DSAs with a transplant center was the sum of these numbers (supply_{DSA} = $ZIP_{donors} + DSA_{donors}$). Demand was the number of active waitlist candidates from all transplant centers within that particular DSA (demand_{DSA} = $tx_{candidates}$) who first achieved MELD 15 during the study period.

Supply/demand ratios for each DSA were calculated by dividing the supply by the demand for that particular DSA (SD_{DSA} = supply_{DSA}/demand_{DSA}).

Supply/demand ratio under fixed-distance and fixed-population circles

Supply for each transplant center was a proportionally allocated fraction of the estimated number of deceased donor livers from all ZIP codes (ZIP_{donors}) within that transplant center's circle. Because the circles overlap, multiple transplant centers might have a claim to donor livers from each ZIP code, so we allocated each ZIP code's supply as proportional to length of the overlapping centers' waitlists. First, we added the number of waitlist candidates from all transplant centers who claim donor livers for each ZIP code (tx_{candidates}), and calculated the fractional claim on donor organs for each transplant center by dividing that center's waitlist size by $tx_{candidates}$ from each ZIP code that the center could claim $(tx_{candidates}/ tx_{candidates})$. Then, we estimated the number of deceased donor livers for a transplant center from that particular ZIP code. For example, 29 deceased donor livers were recovered from ZIP code 50309, and the ZIP code was in the overlapping circles of 6 transplant centers with waitlist size of 21, 451, 668, 553, 141, 510 (Figure 1). We added the numbers of waitlist candidates from these 6 transplant centers ($tx_{candidates} = 2344$). Then, we calculated the fractional claim on donor organs for each transplant center: SDMK (21/2344), MNUM (451/2344), MNMC (668/2344), IAIV (141/2344), NEUN (553/2344), and KSUK (510/2344). Third, we estimated the number of deceased donor livers for each transplant center from ZIP code 50309 for transplant center SDMK (21*29/2344), MNUM (451*29/2344), MNMC (668*29/2344), IAIV (141*29/2344), NEUN (553*29/2344) and KSUK (510*29/2344) (Figure 1). The tranplant center's supply was the sum of proportionally allocated deceased donor livers from all ZIP codes within the transplant center's circle. Some ZIP codes did not fall under any circle. We distributed those donors among the transplant centers within same DSA proportionally according to the centers' waitlist size. Demand was the number of active waitlist candidates from all transplant centers within the particular fixed-distance or fixed-population circle who first achieved MELD 15 during the study period period. The supply/demand ratio for each transplant center was calculated by dividing the supply by the demand for that particular transplant center.

Statistical analysis

We used skewness and the kurtosis test to check the normality of the supply/demand ratios distribution. Since the distribution of supply/demand ratios was skewed, we log-transformed the supply/demand ratios for subsequent analyses. The variance ratio test was used to test whether the variance of transplant center log supply/demand ratios was different when comparing DSAs, fixed-distance circles, and fixed-population circles. Additionally, we conducted modified Leven's test (based on median for skewed data) to test whether the variance of transplant center supply/demand ratios was different when comparing DSAs, fixed-distance circles, and fixed-population circles. Additionally, set was a p-value of 0.05 was considered statistically significant. We used a histogram to show the distribution of supply/demand ratio under DSAs, fixed-distance circles, and fixed-population circles. We plotted the average supply/demand ratios for the transplant centers in 50 DSAs under fixed-distance and fixed-population circles in color-scaled maps using R version 3.0.1.

Darker colors represented higher supply/demand ratio, while lighter colors represented lower supply/demand ratio. All other statistical analyses were performed using Stata 14.2/SE for Linux (Stata Corp., College Station, TX).

Sensitivity analysis

To check whether variation in listing practices for low-MELD candidates distorted our measure of demand, we ran the supply/demand analyses with two other measures of demand, first considering only waitlist candidates who first achieved an allocation MELD/PELD 20 during the study period, and second considering only waitlist candidates who first achieved an allocation MELD/PELD 22 during the study period. Additionally, we estimated the national supply/demand ratio with no geographic restriction.

RESULTS

Study population

Among the 27,334 deceased liver donors, the median (interquartile range [IQR]) age was 38 (25–53), 65.3% were Caucasian, and 40% were female. Among the 44,652 active incident waitlist candidates who first achieved a MELD 15 during the study period, the median (IQR) age at listing was 57 (48–62), 68.87% were Caucasian, 36.85% were female, 3.63% were status 1 candidates, and the median (IQR) MELD score was 18 (13–25) at registration. Additionally, 27.3% had alcoholic cirrhosis, 24% had Hepatitis C virus, 20.6% had hepatocellular carcinoma, and 15.8% had non-alcoholic steatohepatitis as the indication for liver transplant.

Fixed-distance and fixed-population circles

The population sizes (median [IQR], millions) within the small 150-mile and large 400miles fixed-distance circles were 11.49 (6.32–18.88) million and 50.00 (32.76–71.48) million. The circle radius sizes (median [IQR], miles) for the small 12-million and large 50million fixed-population circles were 155 (97–218) miles and 404 (305–590) miles. Fixedpopulation circles in the Northeast were generally smaller than in the Midwest or West coast (Figure 2).

Supply/demand ratio in DSAs

Within the 50 DSAs that have liver transplant programs, with historically proportional allocation of the supply from the remaining DSAs, the median (IQR) supply/demand ratio was 0.66 (0.48–0.87) with a range of 0.34–2.03 (Table 1) (Figure 3). The calculated variance in supply/demand ratio was 0.11 (Table 1).

Supply/demand ratio under fixed-distance and fixed-population circle

With a 150-mile fixed-distance circle, the median (IQR) supply/demand ratio was 0.61 (0.48–0.74) with a range of 0.33–2.87 (Table 1) (Figure 4a). With a 400-mile fixed radius circle, the median (IQR) supply/demand ratio was 0.61 (0.54–0.69) with a range of 0.39–1.09 (Table 1) (Figure 5a). The calculated variances in supply/demand ratio for 150-mile and 400-mile circles were 0.11 and 0.02 (Table 1).

With 12-million fixed-population circles, the median (IQR) supply/demand ratio was 0.59 (0.47–0.83) with a range of 0.27–2.14 (Table 1) (Figure 4b). With 50-million fixed-population circles, the median (IQR) supply/demand ratio was 0.61 (0.51–0.65) with a range of 0.43–1.01 (Table 1) (Figure 5b). The caculated variances in supply/demand ratio for 12-million-population and 50-million-population circles were 0.08 and 0.02 (Table 1).

Comparision of supply/demand ratios

The 150-mile fixed-distance circles did not significantly reduce variance in supply/demand ratios compared to DSAs (0.11 vs. 0.11, p=0.2) (Table 1). Additionally, 12-million-population circles did not significantly reduce the variance in supply/demand ratios compared to DSAs (0.08 vs. 0.11, p=0.5) (Table 1). However, 400-mile fixed-distance circles and 50-million-population circles substantially reduced the variance in supply/ demand ratio compared to DSAs (400-mile: 0.02 vs. 0.11, p<0.001; 50-million: 0.02 vs. 0.11, p<0.001) (Table 1).

The variance in supply/demand ratio was significantly lower for the 400-mile circle compared to that of the 150-mile circle (0.02 vs. 0.11, p<0.001). Similarly, the variance in supply/demand ratio was significantly lower for the 50-million-population circle compared to that of the 12-million-population circle (0.02 vs. 0.08, p<0.001).

Comparing the 'smaller' circles, the variance in supply/demand ratio was not significantly lower for the 12-million-population circle compared to the 150-mile circle (0.08 vs. 0.11, p=0.3). Comparing the 'larger' circles, the variance between the 50-million-population and 400-mile circles was also similar (0.02 vs. 0.02, p=0.9).

Sensitivity analyses

Including only active incident waitlist candidates with MELD/PELD 20, inferences did not change. We found no significant reduction of variances in supply/demand ratios for 150-mile (p=0.1) and 12-million-population (p=0.3) circles compared to DSA. Similar to our main results, the 400-mile circle (p<0.001) and 50-million-population (p<0.001) circles significantly reduce the variance in supply/demand ratios compared to DSAs. The variance in supply/demand ratio was significantly lower for 'smaller' circles compared to 'larger' circles (150-mile vs. 400-mile p<0.001; 12-million-population vs. 50-million-population p<0.001). Comparing the 'smaller' and 'larger' circles, the variances were similar (150-mile vs. 12-million-population p=0.4; 400-mile vs. 50-million-population p=0.9). Including active incident waitlist candidates with MELD/PELD 22, there was also no change in our conclusions. If livers were distributed nationally, the supply/demand ratio would be 0.61 everywhere; there would be no variation in this availability metric across transplant centers.

DISCUSSION

In this national study of geographic disparities in liver supply/demand, we found large variance in the supply/demand ratio between DSAs (variance=0.11). Geographic disparities with 150-mile fixed-distance circles, as recently proposed by the OPTN liver committee, were no better than the currently existing geographic disparities within DSAs (variance=0.11, p=0.2). Geographic disparities with 12-million-population circles

(variance=0.08, p=0.5) were also no better than disparities within DSAs. Only larger circles sufficed to significantly reduce variation in supply/demand ratios: 400-mile fixed-distance circles (variance=0.02, p<0.001) or 50-million-population circles (variance=0.02, p<0.001) yielded substantially more equitable supply/demand ratios than DSAs or smaller circles.

Our finding of a wide range and large variance in the liver supply/demand ratios across DSAs is consistent with many previous reports of geographic disparities in liver allocation in the United States, including a previous report that showed the large national variation in DSA liver supply/demand has not improved since implementation of MELD and that unequal liver supply/demand led to higher waitlist mortality (10).

One strength of this study is the inclusion of a large national cohort using SRTR and ZIP code data to analyze the liver supply/demand ratio within various circular areas that might be used for liver allocation. To isolate the direct impact of sharing circles, our approach considered the supply/demand ratio at a single geographical level. As such, we did not model the changing MELD scores over time of individual candidates, nor the random arrival times of each organ, nor the specific MELD level at which different allocation areas would be enforced. So that our metrics and conclusions would be independent of the above stochastic processes, we did not use variation in MELD at transplant to measure geographic disparity (19). Instead, the range of supply/demand ratios within boundaries is an intuitive indicator of whether using those boundaries in allocation would be likely to result in geographic disparity (9). Also, our measure of demand comprises patients who are waitlisted, but there are documented geographic disparities in listing (20). Disparities in listing are troubling and should be addressed, but not by allocation policy: only listed patients can receive transplants, and therefore the available supply can only be distributed among the listed patients. Because the OPTN proposed circular allocation, we did not address other allocation approaches that might reduce geographic disparity (21-24). We did not correct demand to account for patients who travel to areas with greater organ supply (5, 6, 25) and our findings are orthogonal to the multifactorial causes of or explanations for geographic disparity (20, 26–29). Of course, in parallel with efforts to distribute currently available organs in the most geographically equitable manner to patients in need of them, we always encourage continued efforts to increase supply by increasing deceased donation (30, 31), improving OPO performance, and improving marginal organ utilization by transplant centers (32). Increased competition with larger circles might lead to more people being waitlisted (33); however, previous studies have shown that increased transplant center competition was associated with increased offer acceptance (34). Further, higher utilization of increased risk livers was seen in centers with longer waitlists, higher proportion of higher MELD candidates, higher transplant volume, and in more competitive DSAs (32).

We have demonstrated that 150-mile radius fixed-distance allocation circles will not reduce disparity in supply/demand ratio, compared with the disparity in supply/demand ratio within DSAs. Also, using comparably-drawn fixed-population circles is no more effective at reducing disparity than using fixed-distance circles. However, using larger 400-mile radius or 50-million-population circles would reduce geographic disparity. If, as suggested by the recent report of the OPTN's Ad Hoc Geography Committee (11), DSA boundaries are unacceptable in organ allocation because of the disparity in supply/demand ratios that these

boundaries create, then neither should very small circular geographic boundaries be acceptable in allocation. Neither fixed-distance nor fixed-population circular sharing areas will reduce geographic disparity in liver supply/demand ratios unless these areas are quite large, larger than many proposed circles for liver allocation. Just as broader sharing does not always guarantee reduced geographic disparity (3), neither does sharing within circularshaped units guarantee a reduction in geographic disparity; instead, the devil is in the details of the particular circles constructed.

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Abbreviations:

DSA	donor service area
HRSA	Health Resources and Services Administration
IQR	interquartile range
MELD	Model for End-stage Liver Disease
OPTN	Organ Procurement and Transplantation Network
SRTR	Scientific Registry of Transplant Recipients

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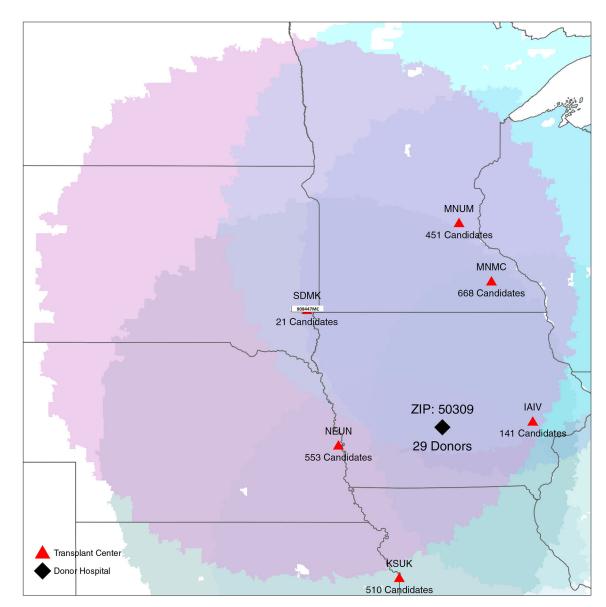


Figure 1.

Fixed-distance circles surrounding each transplant center (red triangles) and donor hospital (black triangle) showing the weighted portion of donor liver supply from the donor hospital based the transplant center waitlist size (demand).

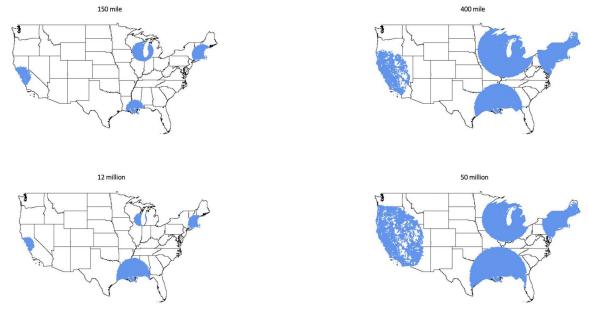


Figure 2.

Fixed-distance (150-mile, 400-mile) and fixed-population (12-million, 50-million) circles around transplant centers CAPC (Stanford), MABI (Beth Israel), LATU (Tulane), WICH (University of Wisconsin).

DSA

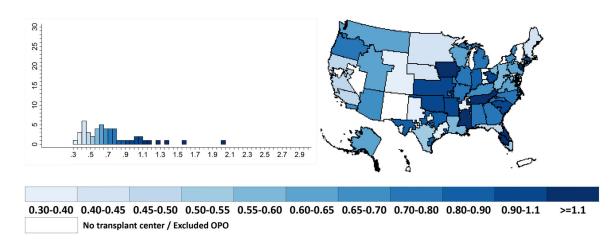


Figure 3.

Supply/demand ratio under current donor service area (DSA) allocation.

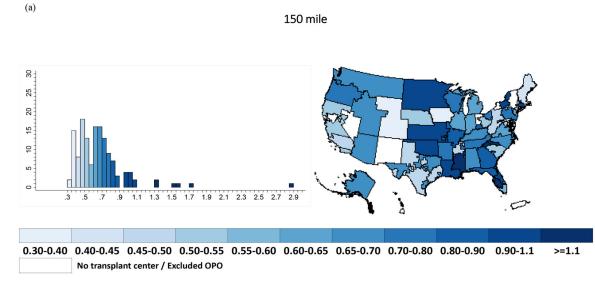


Figure 4.

Supply/demand ratio for (a) 150-mile fixed-distance and (b) 12-million fixed-population circle.

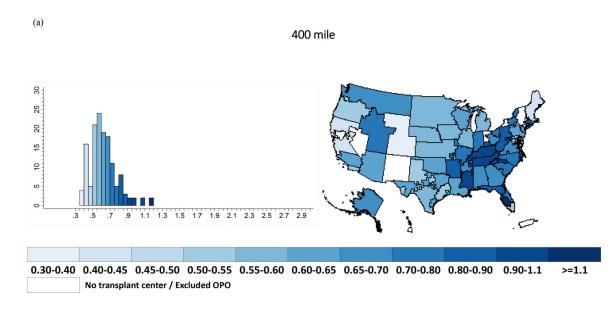


Figure 5.

Supply/demand ratio within (a) 400-mile fixed-distance and (b) 50-million fixed-population circles.

Table 1.

Supply/demand ratios between each sharing area and current donor service area (DSA) allocation.

	Range	Median (IQR)	Mean (SD)	Variance	p-value*	Leven's test
DSA	0.34-2.03	0.66 (0.48 - 0.87)	0.74 (0.34)	0.11	-	
150 mile	0.33-2.87	0.61 (0.48 - 0.74)	0.66 (0.33)	0.11	0.2	0.2
400 mile	0.39-1.09	0.61 (0.54 - 0.69)	0.62 (0.12)	0.02	< 0.001	< 0.001
12 million	0.27-2.14	0.59 (0.47 – 0.83)	0.67 (0.28)	0.08	0.5	0.3
50 million	0.43-1.01	0.61 (0.51 - 0.65)	0.62 (0.14)	0.02	< 0.001	< 0.001

* comparision to DSA