

mRNA Vaccination Decreases COVID-19-Associated Morbidity and Mortality Among Organ Transplant Recipients: A Contemporary Cohort Study

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Background. Organ transplant recipients (OTRs) are less protected from vaccination than immunocompetent hosts. Additional vaccine doses have shown increased immunogenicity. Few studies have assessed their clinical efficacy, particularly against Omicron variants, as most included patients from earlier phases of the pandemic, with higher base mortality rates.

Methods. We studied adult OTRs who had coronavirus disease 2019 (COVID-19) between 12/15/21 and 5/25/22. We compared clinical outcomes between those who had received 2 or ≥ 3 doses of an mRNA vaccine and concurrent unvaccinated controls.

Results. Among 103 OTRs, vaccination was associated with lower 90-day mortality (unvaccinated vs 2 vs ≥ 3 doses: 25% vs 7% vs 3%; $P = .003$), hospital (unvaccinated vs 2 vs ≥ 3 doses: 56% vs 37% vs 27%; $P = .018$) and intensive care unit (ICU; unvaccinated vs 2 vs ≥ 3 doses: 25% vs 15% vs 3%; $P = .001$) admission rates, and peak O_2 requirements (ordinal scale Kendall's tau $b = -0.309$ [lower scores, ie, O_2 requirements with more vaccine doses]; $P = .003$). Age (age > 60 years: adjusted hazard ratio [aHR], 7.73; $P = .016$; administration of antispikes monoclonal antibody: aHR, 0.17; $P = .042$) and vaccination, especially with ≥ 3 doses (aHR, 0.105; $P = .01$), were independently associated with 90-day mortality. Black ($P = .021$) and Hispanic ($P = .016$) OTRs were underrepresented among the vaccinated, especially in the ≥ 3 -dose group.

Conclusions. Despite lower mRNA vaccine efficacy in OTRs and against Omicron variants, vaccination protects this vulnerable patient population from severe COVID-19 and death. Ethnic and racial disparities in health care have been exacerbated by the COVID-19 pandemic and warrant better community outreach efforts.

Keywords. COVID-19; SARS-CoV-2; infection; Omicron; transplant; vaccines.

Organ transplant recipients (OTRs) mount weaker immune responses to coronavirus disease 2019 (COVID-19) vaccination compared with immunocompetent patients and are less protected against infection and severe illness [1–3]. In some studies of OTRs, vaccination with 2 doses of an mRNA vaccine was not associated with lower risk of severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) infection [2, 4], critical COVID-19 [4, 5], or mortality [4, 6, 7]. However, others have found that vaccination decreased the risk of symptomatic illness [8], hospitalization [9], use of mechanical ventilation or

extracorporeal membrane oxygenation (ECMO) support [10], and death [10, 11].

Among OTRs, including those with no or low antibody response after 2 doses of an mRNA vaccine, a third [12–16] and fourth [17–19] dose increase protective antibody titers. However, this “boosted” humoral immunity is less effective *in vitro* against Omicron variants compared with wild-type SARS-CoV-2 or Delta strains [20, 21]. Moreover, these observations may not correlate with clinical outcomes, given confounding from the increased transmissibility but attenuated virulence of Omicron strains [22]. For example, one study of vaccinated OTRs during the Omicron surge showed that seronegativity was not associated with risk of breakthrough SARS-CoV-2 infection, which, nonetheless, decreased with increasing antispikes antibody titers [23]. Importantly, most of the aforementioned studies spanned earlier phases of the pandemic [2, 4, 6, 9, 11], with different circulating variants and potentially standards of care, which can affect base mortality rates [24, 25].

There is a relative paucity of recent clinical data on the efficacy of vaccination, and specifically additional mRNA vaccine doses, among OTRs using concurrent unvaccinated controls during the most recent (Omicron) phase of the pandemic. To this end, using real-world, patient-level data from our

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comprehensive institutional registry, we compared clinical outcomes during the current era of Omicron predominance between OTRs who developed COVID-19 after receipt of 2 or ≥ 3 vaccine doses and unvaccinated OTRs who had COVID-19 during the same relatively narrow time period.

METHODS

Study Design and Data Collection

We included OTRs followed at Brown University-affiliated hospitals who had polymerase chain reaction (PCR)-confirmed SARS-CoV-2 infection between 12/15/2021 (date of the first OTR case after the Omicron variant was detected in Rhode Island) and 5/25/2022. Baseline was defined as the day of symptom onset or the day of first positive SARS-CoV-2 PCR if asymptomatic. Patients were excluded if they were <18 years old, if they received <2 doses of an mRNA vaccine, or if their baseline was within <2 weeks of receiving their most recent mRNA vaccine dose. Exclusion criteria for vaccination status were determined based on the (a) low efficacy of the adenoviral vector vaccine or of “partial” vaccination with 1 dose of an mRNA vaccine (or <2 weeks from the last dose), and (b) small number of patients in each of these categories. Relevant baseline demographic and clinical information was extracted retrospectively from electronic medical records (EMRs) (Table 1). The study was approved by the Lifespan Institutional Review Board.

The primary outcome was 90-day mortality given substantial delayed mortality among OTRs with COVID-19 [10, 26]. Secondary outcomes were 30-day mortality, hospitalization, intensive care unit (ICU) admission, length of stay among admitted OTRs who survived at least 90 days, and peak (worst) oxygen (O_2) requirement on a modified ordinal scale: 0, outpatient only; 1, admitted to the hospital but without supplemental O_2 requirement; 2, low-flow O_2 requirement; 3, high-flow O_2 requirement; 4, noninvasive mechanical ventilation (BiPAP, CPAP); 5, invasive mechanical ventilation.

Statistical Analyses

The normality of distribution was assessed with the Kolmogorov-Smirnoff test. Data for continuous variables are presented as median (interquartile range [IQR]) and compared with the Kruskal-Wallis test or Mann-Whitney U criterion, the latter for pairwise comparisons. Categorical variables are presented as number (%) and compared with χ^2 for linear trend (Mantel-Haenszel) or the Fisher exact (for 2×2 comparisons) test.

Ninety-day survival was also analyzed by Kaplan-Meier curves (log-rank test for trend) and univariable and multivariable Cox regression models. The proportional hazards assumption was confirmed by visual assessment of Schoenfeld residuals and by building time-dependent variables. Factors with a P value of <.2 on univariate analyses were entered into

the multivariate models and retained if the P value for variable removal was <.05.

The association between vaccination status and peak O_2 requirements was assessed by calculation of the Kendall's tau b correlation coefficient. Additional sensitivity analyses were performed by analyzing age as a continuous variable, after excluding patients diagnosed in December 2021 (for Omicron and Delta overlap), or with history of prior COVID-19 infection, by using binary logistic instead of Cox regression, by entering variables that differed substantially between the 3 groups, and by using ordinal logistic regression instead of ranks correlation to examine the association between vaccination status and peak O_2 requirement.

All analyses were performed with SPSS statistical software, version 24.0 (IBM Corporation, Armonk, NY, USA). Statistical significance was set at a 2-tailed P value of .05, unless otherwise indicated above.

RESULTS

One hundred eleven OTRs acquired COVID-19 during the study period. Six received an adenoviral vector vaccine, and 2 were <18 years old. One hundred three OTRs met the inclusion criteria. Of these, 49 (48%) were female, 13 (13%) Hispanic, and 12 (12%) Black. Most (96, 93%) had received a kidney transplant. The median age at the time of COVID-19 diagnosis (IQR) was 58 (43–68) years. Sixteen (16%) patients were unvaccinated, 27 (26%) received 2 vaccine doses, and 60 (58%) received ≥ 3 COVID-19 mRNA vaccine doses (3 of whom received 4 doses), all >2 weeks from baseline.

Demographic and clinical data are summarized in Table 1 and were largely comparable. However, those who were vaccinated, especially with ≥ 3 doses, were less likely to have chronic kidney disease (unvaccinated vs 2 vs ≥ 3 doses: 69% vs 59% vs 45%; $P = .046$) or hypertension (unvaccinated vs 2 vs ≥ 3 doses: 100% vs 93% vs 80%; $P = .034$). Patients identified in the EMR as Black (unvaccinated vs 2 vs ≥ 3 doses: 19% vs 18.5% vs 7%; $P = .021$) or Hispanic (unvaccinated vs 2 vs ≥ 3 doses: 25% vs 19% vs 7%; $P = .016$) were underrepresented among the vaccinated, especially in the ≥ 3 -dose group (Table 1).

Vaccinated patients were less likely to be treated with remdesivir, which at our institution is not administered to outpatients, as they were less likely to require hospital admission (unvaccinated vs 2 vs ≥ 3 doses: 50% vs 19% vs 17%; $P = .008$); this difference did not reach statistical significance among hospitalized patients, where remdesivir is administered to most OTRs, especially those requiring supplemental O_2 (unvaccinated vs 2 vs ≥ 3 doses: 89% vs 50% vs 63%) (Table 1).

Clinical outcomes are summarized in Table 2. Vaccinated patients were less likely to be admitted to the hospital (unvaccinated vs 2 vs ≥ 3 doses: 56% vs 37% vs 27%; $P = .018$) or ICU (unvaccinated vs 2 vs ≥ 3 doses: 25% vs 15% vs 3%; $P = .001$) or

Table 1. Demographic and Clinical Characteristics

Parameter	Unvaccinated	2 Doses	≥3 Doses	P Value
No. of patients	16	27	60	
Age, y	55 (43.5–66)	57 (39–65)	58 (45.8–69.3)	.344
Female sex	8 (50.0)	11 (40.7)	30 (50.0)	.811
Ethnicity and race				
Hispanic	4 (25.0)	5 (18.5)	4 (6.7)	.016
Black	3 (18.75)	5 (18.5)	4 (6.7)	.021
White	8 (50.0)	17 (63.0)	46 (76.7)	.042
Other	1 (6.25)	0 (0)	6 (10.0)	.430
BMI, kg/m ²	29.7 (22.9–33.3)	27.1 (23.7–33.8)	29.0 (25.0–32.4)	.919
Smoking status				
Never	10 (62.5)	16 (59.3)	38 (63.3)	.965
Current or former	6 (37.5)	11 (40.7)	22 (36.7)	
Comorbidities ^a				
Hypertension	16 (100)	25 (92.6)	48 (80.0)	.034
Diabetes	4 (25.0)	11 (40.7)	18 (30.0)	.866
Cardiac	2 (12.5)	4 (14.8)	7 (11.7)	.676
Chronic kidney disease	11 (68.8)	16 (59.3)	27 (45.0)	.046
Pulmonary	2 (12.5)	4 (14.8)	6 (10.0)	.573
Time since transplant, mo	57 (34–111)	61 (15–138)	67 (32.3–130)	.754
Transplanted organ ^a				
Kidney	16 (100)	24 (88.9)	56 (93.3)	.430
Heart	2 (12.5)	3 (11.1)	3 (5.0)	.152
Pancreas	1 (6.25)	1 (3.7)	2 (3.3)	.410
Lung	0 (0)	0 (0)	1 (1.7)	.922
Liver	0 (0)	0 (0)	1 (1.7)	.922
>1	3 (18.8)	1 (3.7)	3 (5.0)	.068
Maintenance immunosuppressive drug regimen				
3-drug regimen	12 (75.0)	21 (77.8)	42 (70.0)	.453
2-drug regimen	4 (25.0)	6 (22.2)	18 (30.0)	.649
Calcineurin/mTOR inhibitor or costimulatory blocker				
Tacrolimus	12 (75.0)	21 (77.8)	51 (85.0)	.374
Sirolimus	1 (6.25)	2 (7.4)	8 (13.3)	.443
Tacrolimus & sirolimus	1 (6.25)	0 (0)	0 (0)	.010
Cyclosporine	1 (6.25)	3 (11.1)	1 (1.7)	.107
Belatacept	1 (6.25)	0 (0)	0 (0)	.010
None	0 (0)	1 (3.7)	0 (0)	.214
Antimetabolite				
Azathioprine	2 (12.5)	3 (11.1)	8 (13.3)	.983
Mycophenolic acid (Myfortic)	10 (62.5)	14 (51.9)	29 (48.3)	.277
Mycophenolate mofetil (CellCept)	1 (6.25)	4 (14.8)	10 (16.7)	.435
None	3 (18.75)	6 (22.2)	13 (21.7)	.974
Held or decreased ^b	10 (76.9)	17 (80.95)	30 (63.8)	.200
Prednisone	15 (93.8)	27 (100)	53 (88.3)	.130
Vaccine type				
BNT162b2 (Pfizer-BioNTech)	NA	20 (74.1)	37 (61.7)	.379
mRNA-1273 (Moderna)	NA	7 (25.9)	23 (38.3)	
Mixed (Pfizer-BioNTech/Moderna)	NA	0	1 (1.7)	
Antiviral medications				
mAb	5 (31.3)	19 (70.4)	34 (56.7)	.323
Remdesivir (all, hospitalized only)	8 (50.0, 88.8)	5 (18.5, 50)	10 (16.7, 62.5)	.008 , .185
Nirmatrelvir/ritonavir (Paxlovid)	0	1 (3.7)	3 (5.0)	.590
COVID-19 baseline characteristics				
Year 2021	3 (18.8)	6 (22.2)	14 (23.3)	.831
Reinfection	2 (12.5)	2 (7.4)	4 (6.7)	.346
Days of symptoms	4 (1–8.5)	2.5 (1–8.5)	3 (1–5)	.488

Data are presented as No. (%) for categorical variables and median (IQR) for continuous variables. All patients were coded as either female or male sex in the hospital EMR; none were listed as intersex. Ethnicity and race data were taken from the hospital EMR and may not reflect patient self-identification. Bolded *P*-values are those ≤ .05 (considered statistically significant).

Abbreviations: BMI, body mass index; EMR, electronic medical record; IQR, interquartile range; mAb, antispikes monoclonal antibody; OTRs, organ transplant recipients.

^aTotal will be greater than the total number of patients due to row overlap.

^bPercentage of OTRs who were receiving an antimetabolite at baseline.

Table 2. Clinical Outcomes

Outcome	Unvaccinated	2 Doses	≥3 Doses	<i>P</i> Value
No. of patients	16	27	60	
90-d mortality	4 (25)	2 (7.4)	2 (3.3)	.003
30-d mortality	3 (18.8)	2 (7.4)	2 (3.3)	.019
Hospitalization	9 (56.25)	10 (37.0)	16 (26.7)	.018
ICU admission	4 (25)	4 (14.8)	2 (3.3)	.001
Length of hospital stay, d	7 (3–12)	7.5 (1.3–11)	3.5 (2–5.5)	.130

Length of hospital stay was isolated to admitted OTRs who survived at least 90 days and is presented as median (IQR). All other variables are presented as No. (%). Bolded *P*-values are those $\leq .05$ (considered statistically significant).

Abbreviations: ICU, intensive care unit; IQR, interquartile range; OTRs, organ transplant recipients.

to die by day 30 (unvaccinated vs 2 vs ≥ 3 doses: 19% vs 7% vs 3%; $P = .019$) or day 90 (unvaccinated vs 2 vs ≥ 3 doses: 25% vs 7% vs 3%; $P = .003$; log-rank $P = .006$) (Figure 1). There was no significant difference in 90-day mortality between subjects who received either 2 or ≥ 3 doses of BNT162b2 (Pfizer: 3/57, 5.3%) and those who received 2 or ≥ 3 doses of mRNA-1273 (Moderna: 1/30, 3.3%) vaccines.

Vaccinated patients, especially those who received ≥ 3 doses, had decreased peak supplementary O_2 requirements (ordinal scale Kendall's tau $b = -0.309$ [lower scores, ie, O_2 requirements with more vaccine doses]; $P = .003$) (Figure 2). Among hospitalized 90-day survivors, the difference in length of hospital stay did not reach statistical significance ($P = .13$) (Table 2).

On univariable and multivariable Cox regression analyses (Table 3), older age (>60 years) was associated with increased risk of 90-day mortality (adjusted hazard ratio [aHR], 7.73; $P = .016$). Administration of antispikes mAb (aHR, 0.17; $P = .042$) and vaccination, especially with ≥ 3 doses (aHR, 0.105; $P = .01$), were associated with decreased risk of 90-day mortality. All mAbs were administered under Emergency Use Authorization (EUA), not a clinical trial. The benefits associated with vaccination, especially with ≥ 3 doses, did not change substantially across all sensitivity analyses (data not shown, available upon request).

DISCUSSION

Several studies have yielded mixed results regarding vaccine clinical efficacy among OTRs, especially with 2 doses of an mRNA vaccine or 1 of an adenoviral vector vaccine, during the pre-Omicron waves of the pandemic. Aslam et al. [8] found that vaccination with 2 doses of an mRNA vaccine was protective against symptomatic COVID-19. Their findings were consistent with another report from a multistate registry, which also showed decreased efficacy among immunocompromised patients, especially OTRs, compared with immunocompetent hosts [1]. A recent population-based study among OTRs from Canada showed increasing protection with $3 > 2 > 1$

COVID-19 vaccine doses, against both SARS-CoV-2 infection and severe COVID-19 [3].

On the contrary, a smaller pre-Delta/Delta (January–June 2021) study from Europe showed that kidney transplant recipients who acquired COVID-19 after 1 or 2 doses of the BNT162b2 (Pfizer) mRNA vaccine had clinical outcomes comparable to those of unvaccinated controls from the previous pandemic wave (September–December 2020) [4]. Likewise, we previously found that COVID-19 mortality during the Delta surge among vaccinated OTRs after 2 mRNA vaccine doses or 1 dose of an adenoviral vector vaccine was comparable to that of unvaccinated controls, including controls from the pre-Delta phase [7]. The above findings were consistent with 2 more studies conducted during the Delta wave, which did not show a mortality benefit from vaccination [6, 9].

In our earlier report, vaccinated patients were older than unvaccinated ones and waited longer to seek medical care [7]. After these findings and our results showing benefit from mAb administration [27], we reached out with letters to OTRs followed at our center and emphasized the importance of additional vaccine doses and early treatment of COVID-19. Notably, in the present study, symptom duration at presentation was shorter than before [7], and there were no significant differences among the 3 different vaccination status groups (Table 1), highlighting the importance of patient education.

In this study, we analyzed patient-level data from a rather homogeneous cohort of OTRs, all of whom had COVID-19 in the smallest US state, during a recent time frame without significant variations in health care resources, management protocols, or circulating SARS-CoV-2 variants. Despite lower immunologic [12–14] and clinical [1, 10] efficacy of COVID-19 vaccines among OTRs and against Omicron variants [20, 21], we observed a strong, linear mortality and morbidity benefit with mRNA vaccination. OTRs who had received ≥ 3 doses had the highest survival rate and lowest rates of hospitalization and critical illness (Table 2, Figures 1 and 2).

Recent analyses from the National COVID Cohort Collaborative (NC3; the largest COVID-19 database in the United States) are in agreement with our results: across the pre-Delta, Delta, and Omicron waves, vaccination, especially with 3 vaccine doses, protected OTRs from infection and serious adverse outcomes [10]. The benefits from vaccination were less prominent compared with nonimmunocompromised patients, but the relative risk attenuated significantly over time (pre-Delta $>$ Delta $>$ Omicron), likely indicating inferior standards of care or/and higher strain virulence earlier in the pandemic, disproportionately affecting the immunosuppressed. Nevertheless, the NC3 investigators did not provide detailed outcome results specific to the Omicron era.

To our knowledge, no other study to date has assessed the clinical efficacy of mRNA vaccination among OTRs during

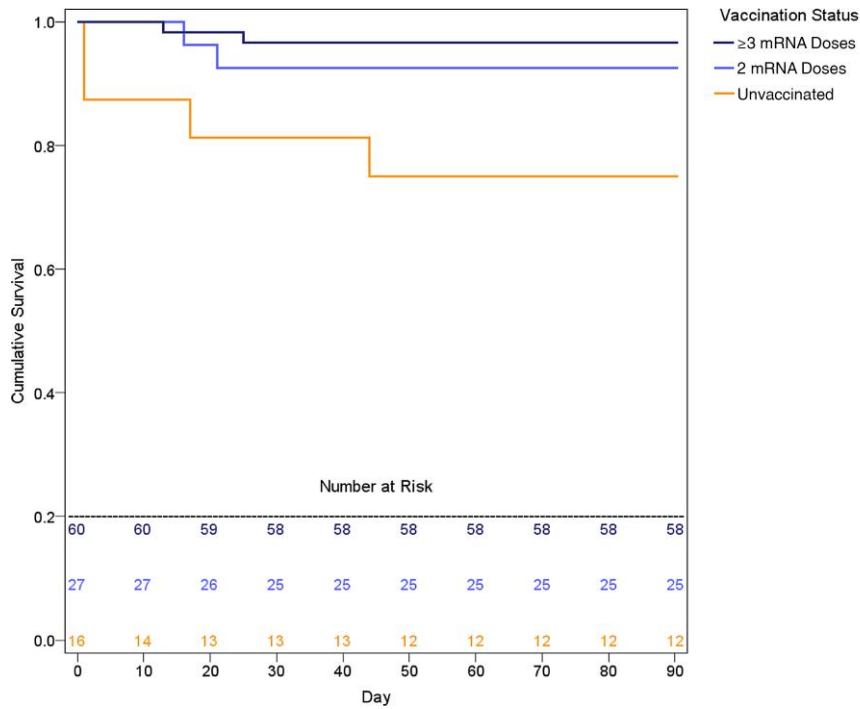


Figure 1. Kaplan-Meier 90-day survival curves. Log-rank $P = .006$.

the Omicron surge using concurrent unvaccinated controls. Our findings and those of the above studies [3, 10] support the recommendation to administer additional vaccine doses to moderately or severely immunosuppressed patients, which was originally based on immunogenicity data alone [12–14].

Vaccination rates, especially with ≥ 3 doses, were significantly lower among OTRs identified as Black or Hispanic in the EMR, compared with non-Hispanic White OTRs (Table 1). Although we cannot rule out type I error due to small numbers, this was a concerning finding. Minority racial and ethnic groups in America, particularly Black, Native American and Alaskan Native, and Latina/o/x groups, are disproportionately burdened with COVID-19 infections [28, 29], hospital admissions [27], and deaths [28, 29]. One comprehensive review found that the nationwide vaccine hesitancy rates for Black (42%) and Hispanic (30%) patients were higher than the nationwide pooled hesitancy rate (26%) [30]. Several factors may be contributing to such observations, namely socioeconomic status [30, 31], geographic proximity and access to transportation, education and health literacy [32], preexisting beliefs and exposure to misinformation [30], language barriers, and medical mistrust, founded in intergenerational trauma from racism and discrimination, provider prejudice, or lack of racial concordance with providers [30, 33].

The last factor may be pertinent for our small transplant team, as we do not have any Black or Hispanic providers, and our results are discrepant from those observed in the

general population of Rhode Island, where most vaccine-hesitant patients are non-Hispanic White [34]. It should be noted, however, that the underrepresentation of Black and Hispanic OTRs was most prominent in the third dose group. A third mRNA vaccine dose became available later in the pandemic as part of the primary series only for immunocompromised hosts. Therefore, it is possible that our Black or Hispanic OTRs missed that dose not because of vaccine hesitancy, but rather due to incomplete or delayed vaccine education, which highlights the importance of continued outreach to ethnic and racial minorities, especially among the immunosuppressed.

Another important finding from our study was that mAb administration to OTRs with COVID-19 was associated with better survival (Table 3), in agreement with the results of a recent meta-analysis [35]. Vaccinated patients received mAbs more frequently; however, the difference was not statistically significant (Table 2) and is unlikely to have contributed to the independent benefits observed with vaccination (Table 3). In fact, it is possible that because vaccinated patients were less frequently hypoxic or hospitalized due to COVID-19 (both of which preclude mAb use under EUA), they were more likely to receive mAbs than unvaccinated patients.

Regardless, mAbs remain our first-line outpatient treatment for OTRs with COVID-19, as they have been shown to decrease the risk of hospitalization [27] and emergency room visit and death [35]. Nirmatrelvir/ritonavir (Paxlovid), although at least

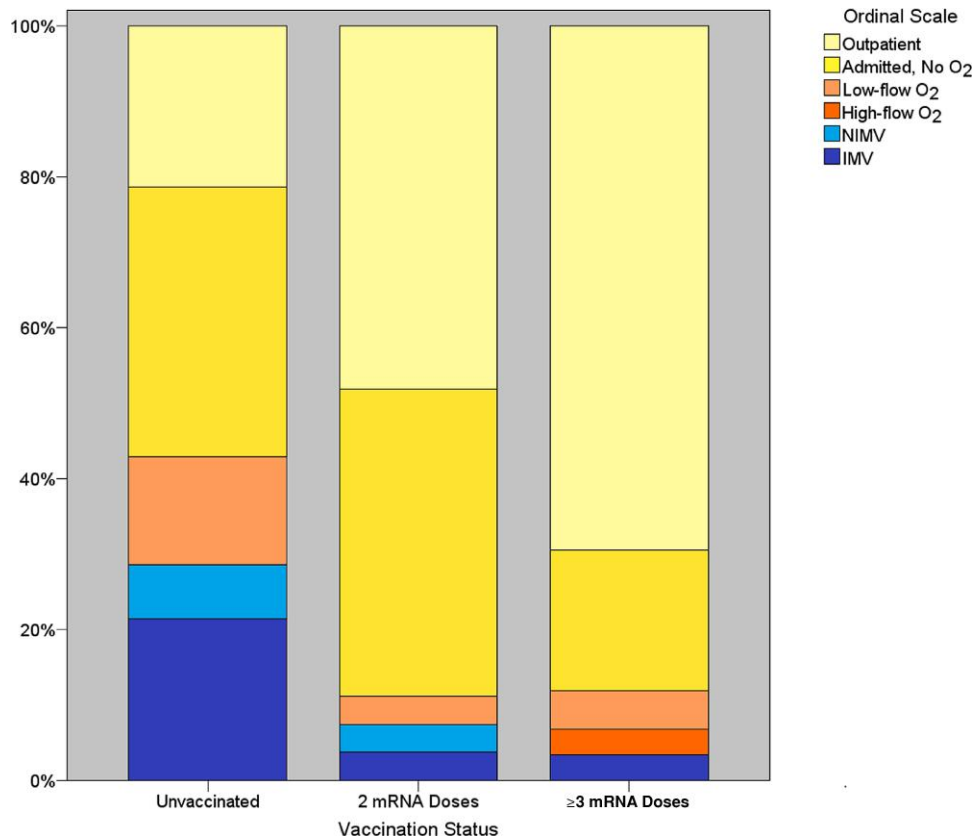


Figure 2. Peak O₂ requirement ordinal scale value distribution by vaccination status. Kendall's tau b = -0.309 (lower scores, ie, O₂ requirements with more vaccine doses; *P* = .003). Abbreviations: IMV, invasive mechanical ventilation; NIMV, noninvasive mechanical ventilation.

as effective as mAbs, inhibits irreversibly the catabolism of calcineurin and mTOR inhibitors, which are the backbone of maintenance immunosuppression among OTRs. Therefore, its administration in this patient population requires close laboratory monitoring of immunosuppressant levels and has the potential for iatrogenic toxicities [36, 37].

Our study has several limitations, which, nonetheless, are unlikely to have affected our core conclusions. First, it is a single-center report; therefore, the results may not be generalizable, although they are well aligned with those of many

pre-Omicron studies [1, 3, 8, 10]. Second, our cohort included mostly kidney transplant recipients; therefore, our conclusions may not necessarily apply to other OTRs. Third, the number of patients was relatively small, but the differences in clinical outcomes were convincing (Table 2, Figures 1 and 2). Moreover, the 3 patient groups were relatively well balanced not only in their baseline characteristics (Table 1), but also in geography and Omicron phase of the pandemic, unlike older reports [2, 4, 6, 9, 11]. Fourth, data were retrospectively collected, but all key independent and outcome variables were objective and easy to extract from EMRs. Fifth, we did not have patient-level SARS-CoV-2 sequences, and there may have been some overlap between Omicron and Delta variants early on. Nonetheless, by January 2022, >95% of SARS-CoV-2 infections in our state were caused by Omicron [38], the distribution of cases in late 2021 vs 2022 was similar across the 3 vaccination status groups (Table 1), and exclusion of OTRs from 2021 did not significantly change our conclusions.

Also, the size of our cohort did not allow us to match patients who received 2 vs ≥3 doses for time from last dose, to clarify if the benefit from the additional dose was independent of the fact that patients had higher antibody titers at the time of COVID-19, because the most recent vaccine dose was closer

Table 3. Ninety-Day Mortality Univariable and Multivariable Cox Regression Analyses

Variables	Univariable Analysis			Multivariable Analysis		
	HR	95% CI	<i>P</i> Value	aHR	95% CI	<i>P</i> Value
mAb	0.24	0.05–1.20	.083	0.17	0.03–0.94	.042
Age >60 y	4.02	0.81–19.9	.089	7.73	1.47–40.58	.016
Vaccination			.037			.037
2 doses	0.27	0.05–1.47	.129 ^a	0.41	0.07–2.37	.319
≥3 doses	0.12	0.02–0.65	.014^a	0.105	0.02–0.59	.010

Bolded *P*-values are those ≤ .05 (considered statistically significant). Abbreviations: aHR, adjusted hazard ratio; HR, hazard ratio; ICU, intensive care unit; mAb, antispike monoclonal antibody.

^aReference category: unvaccinated.

to infection. However, previous studies have shown that even nonresponders to 2 mRNA vaccine doses can mount an immune response to additional doses, suggesting increased immunogenicity [12–14]. Last, clinicians may have a lower threshold to admit unvaccinated OTRs to the hospital, but this should not affect mortality or peak O₂ requirements.

In conclusion, mRNA vaccination protects OTRs from severe COVID-19 and death. Health care disparities in the prevention and treatment of COVID-19 call for better outreach initiatives and patient education. Every effort should be made toward high enrollment of OTRs in vaccine clinical trials, with strong representation of ethnic and racial minorities.

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Patient consent. This retrospective study was approved by the Lifespan IRB with a waiver of informed consent.

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