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LETTER TO THE EDITOR

COVID-19 after two doses of mRNA vaccines in kidney transplant recipients

To the Editor:

Mortality due to severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) infection and the resulting coronavirus disease 2019 (COVID-19) among transplant recipients is high. In December 2020 the BNT162b2 (Pfizer-BioNTech) and the mRNA-1273 (Moderna) vaccines received emergency use authorization in the United States. These mRNA vaccines administered in a two-dose series were more than 94% effective in preventing COVID-19 in clinical trials, without safety concerns identified. Transplant recipients were excluded from the clinical trials. Although decreased efficacy and immunogenicity were expected for transplant recipients, transplant recipients were included in the early vaccination prioritization groups and transplant societies, including the American Society of Transplantation, urged transplant recipients to get vaccinated as soon as vaccine was available.

Recently, a study showed that only 54% of organ transplant recipients developed a positive antibody response after two doses of SARS-CoV-2 mRNA vaccine and, among kidney transplant recipients, only 48% had an antibody response.⁴ At our center the seropositivity rate in organ transplant recipients was 37.2%.⁵ This suggests that a significant proportion of transplant recipients remain at risk for COVID-19 despite completing the vaccination series.

Our center encourages COVID-19 vaccination for all transplant recipients. We estimate an active follow-up of approximately 1680 fully vaccinated kidney transplant recipients. We had eight kidney transplant recipients who developed COVID-19 more than 14 days after receipt of two doses of SARS-CoV-2 mRNA vaccine, administered at the recommended dosing interval (Table 1). All patients were symptomatic, most commonly with fatigue and cough. The median age was 65 years old (range 36-73), 50% had received a previous transplant, the median time from the most recent transplant to the first dose of mRNA vaccine was 31 months (range 3-248), and three of eight patients had received thymoglobulin for induction in the 6 months preceding vaccination. All patients were on calcineurin inhibitor (CNI)-based immunosuppression and seven of the eight were on mycophenolate mofetil. CNI level goals were not changed during illness, anti-metabolite doses were reduced or held based on symptoms and severity, at the discretion of the nephrologist. Symptom onset and SARS-CoV-2 testing occurred at a median of 56 days (range 18-77) and 59 days (range 20-77) after the second dose of the vaccine respectively. Three patients received

casirivimab/imdevimab as outpatient and did not require hospital admission. Three patients were hospitalized and had a length of stay of 3–5 days, without requiring ICU admission or mechanical ventilation. All were discharged to home without oxygen. None of the patients had anti-SARS-CoV-2 spike protein antibody checked after vaccination or disease and the viruses were not sequenced.

Data on the COVID-19 vaccine effectiveness is emerging, however, vaccine effectiveness in the immunocompromised population is lacking. It is encouraging that our breakthrough cases of COVID-19 after vaccination were mild and associated with good outcomes. However, we should continue to encourage our patients to be vaccinated and also encourage vaccination of household members and close contacts. Recent guidance allowing fully vaccinated individuals to resume pre-pandemic activities probably should not apply to transplant recipients.

KEYWORDS

clinical research/practice, infection and infectious agents - viral, infectious disease, kidney transplantation/nephrology, vaccine

DISCLOSURE

The authors of this manuscript have no conflicts of interest to disclose as described by the *American Journal of Transplantation*.

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TABLE 1 Characteristics of eight kidney transplant recipients who developed COVID-19 after two doses of mRNA vaccines

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Status ,, at last f/u	Alive	Dead ^b	Alive	Alive	Alive	Alive	Alive	Alive
Duration of follow-up t d	26	30	100	71	79	20	63	56
D of Previous fo transplant d	° Z	Š	Yes	Yes	Yes	Yes	°Z	°Z
Duration of O2 supplement Previous follow-up, on discharge transplant d		0			0	o Z		
_		Š			Š	Z		
Ler of ition star		5			4	М		
II Disposi		Home			Home	Home		
Mechanical ventilation		° Z			°Z	o Z		
DOI		°Z			Š	°Z		
Casirivimab and Mechanical of imdevimab Hospitalization ICU ventilation Disposition stay, d	0 N	Yes	° Z	° Z	Yes	Yes	° Z	o _N
	Yes	Yes ^a	o _N	Yes	°N	°Z	Yes	°Z
Days from second dose to SARS- COV-2 test	55	63	37	37	62	50	77	62
Days from second dose to symptoms	54	62	34	29	58	18	77	09
Symptoms	Loss of taste	Shortness of breath, fatigue	Cough, fatigue	Fatigue, cough	Weakness, cough	Dyspnea, weakness, fatigue, cough	Sinus pressure, throat irritation	Cough, congestion, fever
MMF/ Aza stopped or reduced	Yes	o Z	°Z	o Z	o Z	Yes	Yes	° Z
	Tacrolimus, MMF,	CsA, MMF	Tacrolimus, I MMF, pred	Tacrolimus, I MMF, pred	Tacrolimus, I pred	Tacrolimus, Aza	Tacrolimus, MMF	Tacrolimus, I MMF, pred
Thymoglobulin 6 months prior to first vaccine Maintenance dose	Ľ	Ö	<u> </u>	Ë	Ĕ	<u> </u>	<u> </u>	Ë
Thymo 6 mon to firs dose	3 Yes	S S	N N	Yes	°Z	Yes	S S	°Z
Type of vaccine	mRNA-1273 Yes	mRNA-1273 No	mRNA-1273 No	BNT162b2	BNT162b2	BNT162b2	mRNA-1273 No	BNT162b2
Time from transplant to first vaccine dose,	м	28	34	м	248	ო	87	151
Gender	Σ	Σ	ட	Σ	Σ	ш	ட	Σ
Age	1 69	2 66	3 42	4 36	5 71	6 73	7 63	8 57

Abbreviations: Aza, azathioprine, pred, prednisone, IS, immunosuppression; MMF, mycophenolate mofetil.

^aAdministered during hospitalization.

^bPatient recovered from COVID-19, death unrelated.

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