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LETTER TO THE EDITOR

COVID-19 infection in solid organ transplant recipients after SARS-CoV-2 vaccination

To the Editor:

The prevalence and mortality of COVID-19 are higher in solid organ transplant recipients (SOTs) compared to the general population. ¹⁻³ Two SARS-CoV-2 messenger RNA (mRNA) vaccines have been approved by the FDA; both are 95% efficient in preventing COVID-19 in the general population. The efficacy of these vaccines in SOTs remains to be unknown as immunocompromised patients have been excluded from the vaccine studies. Initial reports indicate low immunogenicity in SOTs with only 11%–17% having detectable antispike antibody 20–28 days after one vaccine dose. ^{4,5} This finding concerned the transplant community but there is hope that the second vaccine dose will be more efficacious.

After obtaining Mayo Institutional Review Board (IRB) approval, we reviewed the records of 7 SOTs (2 heart, 1 lung, 1 heart/kidney, 1 kidney/pancreas, and 2 kidney alone) who received either 1 (n = 2, 28%) or 2 (n = 5, 71%) doses of the BNT162b2 (Pfizer-BioNTech) or the mRNA-1273 (Moderna) SARS-CoV-2 mRNA vaccines and developed COVID-19 after a median of 28 (6-44) days of their last dose. Demographics of these patients are summarized in Table 1. Five of the 7 (71%) patients had blood type A, 1 had AB, and 1 had O blood type. All patients were symptomatic. Fever developed in 4 (57%), 4 (57%) had hypoxia/dyspnea, and 2 (28%) had diarrhea. Diagnosis was confirmed in all patients with polymerase chain reaction (PCR) of nasal swabs. Six of the patients had antibodies to COVID-19 tested at presentation. Of these, five patients had undetectable antispike antibodies and one patient, who had received his second mRNA-1273 vaccine dose 44 days prior, had low titer antispike antibody (1.4 U/ml, reference range < 0.8 U/ml). None of the six tested had detectable nucleocapsid antibody. Five patients required hospitalization, four due to hypoxia and lung infiltrates that required supplemental oxygen but no intubation, while one patient was hospitalized with acute kidney injury from severe vomiting and diarrhea. All hospitalized patients received remdesivir, three received dexamethasone, four received convalescent plasma, and two received tocilizumab. Two patients had received monoclonal antibody treatment. Antimetabolites were discontinued in three of five hospitalized patients. All five patients were discharged, three on supplemental oxygen. Clinical presentation, management, and outcome of these seven patients are summarized in Table 2.

Of the 1624 SOT recipients transplanted in our center over the last 6 years who are Florida residents, 629 (39%) received two doses and 163 (10%) have received one dose of the BNT162b2 (Pfizer-BioNTech) or the mRNA-1273 (Moderna) SARS-CoV-2 mRNA vaccine. Five out of the seven patients in this report were Florida residents suggesting a post-vaccination infection rate of approximately 0.6% which is much higher than the rate of 0.05% reported in the general population, but this needs to be confirmed with more complete vaccination data.

In conclusion, we report seven SOTs with undetectable or low titer antispike antibodies who developed COVID-19 infection after receiving one or two doses of the SARS-CoV-2 mRNA vaccine. The clinical presentation and course of these patients were comparable to those of SOTs who had COVID-19 infection and have not been vaccinated.² This finding suggests that SOTs are still at risk of acquiring COVID-19 infection even after vaccination and calls to continue measures to prevent COVID-19 infection including masking, social distancing, and regular hand hygiene in these patients even after receiving the required doses of the SARS-CoV-2 vaccine. Our findings also call for further research to study the efficacy of vaccination, to examine the post-vaccination infection rate, and to identify methods to boost the vaccine-related immune response in these immunocompromised patients.

KEYWORDS

clinical research/practice, editorial/personal viewpoint, infection and infectious agents – viral, organ transplantation in general, patient safety

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 Baseline characteristics of 7 SOT recipients who had COVID-19 infection after SARS-COV-2 mRNA vaccination

Days from last vaccine dose to COVID-19 diagnosis	35	26	44	9	28	9	19
Number of doses	2	2	2	T	1	7	2
Vaccine name	Pfizer/ BioNTech	Pfizer/ BioNTech	Moderna	Pfizer/ BioNTech	Moderna	Pfizer/ BioNTech	Moderna
Years from Tx to COVID-19	7.37	3.21	1.3	0.58	11.35	0.85	0.58
Rejection history	Yes	°Z	No	°Z	Yes	°Z	o N
Maintenance IS	Bela/Pred/MMF	Tac/MMF/Pred	Tac/MMF/Pred	Tac/MMF/Pred	Tac/MMF/Pred	Tac/MMF/Pred	Tac/MMF/Pred
Induction IS	ATG	ATG	Alemtuzumab	ATG	ATG	Basiliximab	Basiliximab
Previous organ Tx	°Z	°Z	No	Yes	o Z	o Z	°N
Cause of organ failure	COPD	ICM/FSGS	DΜ	HIVAN	Σ	ICM	NICM
Blood	⋖	<	⋖	0	⋖	AB	⋖
Race	O	O	AA	¥	O	O	O
Gender	Σ	Σ	Σ	Σ	Σ	Σ	Σ
Age	64	89	09	42	43	69	29
Organ	Double Lung	Heart/ Kidney	Kidney	Kidney	Kidney Pancreas	Heart	Heart
Patient	T	7	က	4	2	9	7

Abbreviations: AA, African American; ATG, antithymocyte globulin; Bela, Belatacept; C, Caucasian; COPD, chronic obstructive pulmonary disease; DM, diabetes mellitus; HIVAN, HIV associated nephropathy; ICM, ischemic cardiomyopathy; IS, immunosuppression; M, male; MMF, mycophenolate mofetil; NICM, nonischemic cardiomyopathy; Pred, prednisone; Tac, Tacrolimus.

TABLE 2 Clinical presentation, serological findings, and outcome of 7 SOT recipients who had developed COVID-19 infection after SARS-COV-2 vaccination

Outcome	DC on RA	DC on 2 L O ₂	Recovered	DC on RA	DC on 2 L O ₂	Recovered	DC on 2 L O ₂
COVID-19 specific treatment	MAB1, Remd, Dexa, CP, Tocilizumab	Remd, Dexa, CP DC on 2 L O ₂	None	Remd	Remd, Dexa, CP, $$ DC on 2 L $$ Tocilizumab $$ O ₂	MAB 2	Remd, CP
IS management	MMF held	MMF dose reduced	No change	MMF held	MMF held	No change	No Change
Antinucleocapside antibody at COVID-19 diagnosis	Negative	Negative	Negative	Negative	Negative	QN	Negative
Antispike antibody at COVID-19 diagnosis	Negative	Negative	Pos (1.4 U/ ml)	Negative	Negative	ΩN	Negative
AKI (Cr >0.3 mg/ dl from baseline)	o N	o Z	o N	Yes	Yes	°Z	Yes
Lymphopenia AKI (absolute (Cr) lymphocytes dl fr <-900/mcL) base	Yes	Yes	Yes	Yes	Yes	°Z	Yes
Intubation	°N	°Z	o Z	No	oN	°Z	oN
Lung infiltrate	Bilateral R>L	Bilateral R>L	None	None	Bilateral	N/A	Bilateral
Hypoxia (O ₂ Sat < 92% on RA)	Yes	Yes	°Z	°N	Yes	°Z	Yes
Hypoxia (O ₂ Hospitalization, Sat < 92% on duration RA)	Yes, 5 days	Yes, 8 days	°N ON	Yes, 3 days	Yes, 11 days	°Z	Yes, 5 days
Presentation	Fever, rigors, SOB	Fever, chills, SOB, cough, N/V	Cough	N/V, diarrhea	Fever, cough, SOB	Cough, runny nose	Cough, chills, weakness
Patient	Т	2	ო	4	2	9	7

Abbreviations: CP, convalescent plasma; DC, discharged; Dexa, dexamethasone; L, left; MAB 1, bamlanivimab; MAB2, casirivimab/imdevimab; N/V, nausea and vomiting; ND, not done; R, right; RA, room air; Remd, remdesivir; SOB, shortness of breath.

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