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LETTER TO THE EDITOR

SARS-CoV-2 infection in chronic kidney disease patients vaccinated with Oxford/AstraZeneca COVID-19 vaccine: initial Indian experience

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Severe acute respiratory distress syndrome coronavirus 2 (SARS-CoV-2) has grossly unsettled all aspects of humanity across the globe. As of May 2021, India is battling with the coronavirus diease 2019 (COVID-19) crisis and topping the world with the highest number of daily cases [1]. Additionally, the burden of chronic kidney disease (CKD) is among the highest in the world and has worsened further in the pandemic [2]. COVID-19associated mortality has been reported higher in CKD compared with the general population [3, 4]. The vaccination campaign has progressed in the developed world relative to India, which is still up against the herculean task of vaccinating an enormous population amid the COVID-19 surge. Indian advisories, through May 2021, have approved two vaccines, the Oxford-AstraZeneca vaccine (ChAdOx1 nCov-19; Covishield) and BBV152 (Covaxin). The fear of suboptimal antibody response to COVID-19 vaccines in CKD was relieved by recent studies showing adequate immunogenicity of vaccines [5, 6]. However, with various vaccines available across the world and with diverse ethnicity, the antibody response and protection is expected to vary. Herein we report our experience of SARS-CoV-2 in CKD patients admitted in Institute of Kidney Diseases and Research Center, Institute of Transplantation Sciences, Ahmedabad, Gujarat, India, who received either a single or two doses of the Oxford-AstraZeneca COVID-19 vaccine. To the best of our knowledge, this is the first such report.

Overall, during the study period from 3 May 2021 to 10 May 2021, we detected 10 vaccinated CKD patients (6 with two doses and 4 with one dose) who contracted COVID-19 (Table 1). The

median age of the case series was 55 years [interquartile range (IQR) 50-64], with the majority being males (70%). Six patients were on maintenance hemodialysis. All of the patients had hypertension as a common comorbidity. The SARS-CoV-2 severity of the eight surviving patients ranged from mild (n=3) to moderate (n=1) to severe (n=3). Most of the laboratory parameters in the study were out of the normal range, except in Patient 4, who was not investigated further and was managed at home. Patients were managed mostly with oxygen support (n = 7), anticoagulation (n=8), remdesivir (n=7) and steroids (n=5). Two patients died and eight were discharged. Patient 2 had prior a COVID-19 infection 8 months earlier and had a mild illness in both episodes. The median duration from the last dose of vaccine to the onset of COVID-19 symptoms was 29 days (IQR 23-34). SARS-CoV-2 antibody levels were >40 AU/mL in most cases, except for the two patients who died and were on immunosuppression.

The first noteworthy finding from our report is that the CKD patients on immunosuppression may have an inadequate response with the Oxford-AstraZeneca vaccine, which makes them more prone to acquiring severe COVID-19. The second finding is that CKD patients are still susceptible to COVID-19 even with adequate antibody response. The different strains circulating are possibly responsible for this finding, but due to resource limitations, genomic sequencing was not completed. We suggest continued research in the field of vaccine development and the impact of the vaccine on variants to assess the real-world impacts of the vaccination. The CKD group, even though

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Table 1. Summary of the 10 cases

sex 55/M 55/F 56/F 56/F 56/F 56/F 56/F 56/F 56/F	62/M Obstructive uropathy Carvedilol, ARB, sevelamer HD <10 <10 LVD No	45/M HTN Calcitriol, nifedipine, levetiracetam HD <10 2 years AVF HTN, CVA	54/F Unknown Anti-HTN aspirin HD <10 2 years Catheter CCVA/IHD No	wn TN CKD CKD artive eter	45/M Post-transplant CKD Steroids HD <10 1 year AVF	46/M Post-transplant CKD Steroids and antimetabolite HD <10	71/M Post-transplant CKD Steroids, antimetabolite, CNI 1.8 34	72/M HTN OHA, anti-HTN
DKD DKD ON OHA On insulin + OHA + CCB + alpha agonist + loop diuretic + aspirin + statin 2.5 HTM diabetes HTM No S months prior, mild COVID-19; 1 cycle of HD required 2 1 42 2 1 48 Positive Positive NRBM Ambient air steroids Fever, cough, COUGH, fever for DOB × 5 days 3 days	Obstructive uropathy Carvedilol, ARB, sevelamer HD Carvedilol, ARB, AVF HTN, diabetes, LVD No				Oost-transplant CKD Steroids HD 10 1 year AVF HTN</td <td>Post-transplant CKD Steroids and antimetabolite HD <10 Thought</td> <td>Post-transplant CKD Steroids, antimetabolite, CNI 1.8 34</td> <td>HTN OHA, anti-HTN</td>	Post-transplant CKD Steroids and antimetabolite HD <10 Thought	Post-transplant CKD Steroids, antimetabolite, CNI 1.8 34	HTN OHA, anti-HTN
On OHA On insulin + OHA + CCB + alpha agonist + loop diuretic + aspirin + statin 2.5 4.3 2.8 1.1 Conservative Conservative HTN, diabetes HTTN No 8 months prior, mild COVID-19; 1 cycle of HD required 2 1 2 2 1 42.2 24 >	Carvedilol, ARB, sevelamer HD <10 2 years AVF HTN, diabetes, LVD No			AMI on CKD <10 Conservative Temporary catheter HTN	Steroids HD <10 AVF HTN	Steroids and antimetabolite HD <10 1 month AVF	Steroids, antimetabolite, CNI 1.8 34	OHA, anti-HTN
durretic + aspirin + statin + statin - 4.3 28 11 Conservative Conservative HTN diabetes HTN No 8 months prior, mild COVID-19; 1 cycle of HD required of HD required 42 2 1 42 2 24 24 8 Positive NRBM Ambient air Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days 3 days	HD <10 2 years AVF HTN, diabetes, LVD No			AKI on CKD <10 Conservative Temporary catheter HTN	HD <10 AVF	HD <10 1 month AVF	1.8 34 Onnsenyative	Ę
2.5 4.3 28 11 Conservative Conservative — HTN, diabetes HTN No 8 months prior, mild COVID-19; 1 cycle of HD required 2 1 42 24 >400 48 Positive Positive Positive NRBM Ambient air steroids Fever, cough, Cough, fever for DOB × 5 days 3 days	HD <10 2 years AVF HTN, diabetes, LVD No			AKI on CKD <10 Conservative Temporary catheter HTN	HD <10 1 year AVF HTN	HD <10	1.8 34 Conservative	Ę
Conservative Conservative HTN, diabetes HTN No 8 months prior, mild COVID-19; 1 cycle of HD required 2 1 42 2 42 Positive Positive NRBM Ambient air steroids Fever, cough, fever for DOB × 5 days 11 3 43 3 48 Positive Positive Positive Ambient air steroids Ambient air steroids Ambient air	<10 2 years AVF HTN, diabetes, LVD No			<10 Conservative Temporary catheter HTN	<10 1 year AVF HTN	<10 1 month AVF	34	H
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HTN, diabetes No 8 months prior, mild COVID-19; 1 cycle of HD required 1 42 2 1 42 24 24 >48 Section Positive NRBM Ambient air Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days 3 days	HTN, diabetes, LVD No	HTN, CVA No	CVA/IHD No	NTH	NTH	ļ	-	3 months AVF
No 8 months prior, mild COVID-19; 1 cycle of HD required 2 1 2 2 2 4 24 2 24 >400 48 Positive Positive Ambient air Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days 3 days	N O	ON O	o N	-14		NTH	Diabetes	HTN, diabetes
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2 1 42 24 24 Solution Positive Positive NRBM Ambient air steroids Fever, cough, Cough, fever for DOB × 5 days 3 days								
24 >400 48 Positive Positive NRBM Ambient air Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days 3 days	2	2	7	2	Τ,	Τ,	2	2
Positive Positive NRBM Ambient air Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days	19	28	30	32	45	23	23	40
Positive Positive NRBM Ambient air Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days								
Positive Positive NRBM Ambient air Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days 3 days	105	110	112	138	3.8	4.6	Non-reactive	Not done
Positive Positive NRBM Ambient air Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days 3 days								
NRBM Ambient air Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days 3 days	Positive	Positive	Negative ^a	Positive ^b	Positive	Positive	Positive	Positive
Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days 3 days	Low flow	Home	Room air	NRBM	Low-flow	NRBM	NRBM	Ambient air
Remdesivir, Favipiravir steroids Fever, cough, Cough, fever for DOB × 5 days 3 days	oxygen 2 days				oxygen			
Fever, cough, DOB × 5 days	Remdesivir	No No	Supportive]	Remdesivir, sternids	Remdesivir, steroids	Remdesivir, steroids	Remdesivir,	Remdesivir
$DOB \times 5 days$	DOB for	Fever cough	DOB for	DOB for	DOB for	Fever, cough,	Fever, cough,	Fever, cough
	2 days	3 days	5 days	5 days	2 days	DOB for 1 day	DOB for 1 day	for 3 days
AKI on CKD Yes (recovered)	I	1	ı		ı	1	ı	1
, No No	MHD	MHD	MHD	H	MHD	MHD	No	MHD
equirement NRBM Ambient air	Low-flow	Home	NRBM	NRBM	Low-flow	Mechanical	Mechanical	Ambient air
	oxygen				oxygen	ventilation	ventilation	
Admitted Discharged	Discharged	Discharged]	Discharged	Discharged	Discharged	Died	Died	Discharged
Radiological abnormalities Yes Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes	Yes
Laboratory abnormalities (worst reported) TLC (\times 10 3 /mm 3) 6.89	5.3	1	16.2	9.5	3.58	15.9	27	9

Table 1. (continued)

	Patient	Patient	Patient	Patient	Patient	Patient	Patient	Patient	Patient	Patient
Characteristics	1	2	3	4	2	9	7	8	6	10
Platelets $(\times 10^3/\text{mm}^3)$	292	197	06	I	120	225	102	216	238	88
Neutrophil (%)	68	54	71	ı	78	85	75	93	06	06
Lymphocytes (%)	10	42	26	I	20	13	23	2	∞	10
ALC $(\times 10^3/\text{mm}^3)$	1.4	2.8	1.3		3.2	1.2	0.8	7.9	2.1	9.0
NLR	8.9	1.2	2.7	I	3.9	6.5	3.2	18.6	11	6
D-dimer (ng/mL)	630	930	1260	I	>4000	1700	2530	1230	ı	1720
IL-6 (pg/mL)	19	14.22	30	ı	806	ı	228	642	1146	154
Ferritin (ng/mL)	Not done	307		ı	1420	126	1120	915	1000	1000
Serum creatinine (mg/dL)	2.54	6.05	MHD	MHID	MHID	1.8	MHID	MHD	2.46	MHD
hs-CRP (mg/L)	17.3	3.93	18.7	ı	79	70	45	84	81	140
LDH (IU/L)	353	247	203	I	298	272	1733	243	275	253

Outside hospital reported COVID-19 positive.

eGFR, estimated glomerular filtration rate; DKD, diabetic kidney disease; HTN, hypertension, HD, hemodialysis; MHD, maintenance hemodialysis; RT-PCR, reverse transcription polymerase chain reaction, CVA, cerebrovascular accident; GLIA, Clinical total leukocyte count; IL-6, interleukin-6; hs-CRP, high-sensitiv ty C-reactive protein, LDH, lactate dehydrogenase; NLR, neutrophil:lymphocyte ratio; ALC, absolute lymphocyte count; OHA, oral hypoglycemic drugs; CCB, calcium channel blocker; CPC, convalescent plasma component aboratory Improvement Amendments; AKI, acute kidney injury; LVD, left ventricular dysfunction; IHD, ischemic heart disease; DB, difficulty breathing; NRBM, non-rebreather mask; TLC,

they mounted a reasonable antibody response, still acquired COVID-19. The protective cut-off antibody level is unknown in different ethnicities, as the immune composition may vary in individuals of various geographic regions, as is the impact of variants [7]. There have been concerning reports of attenuated antibody response to messenger RNA COVID-19 vaccines in organ transplant recipients [8]. Similar to a previous report [9], the three patients who were CKD and post-renal transplant status did not mount antibody response. In conclusion, we report the first study of COVID-19 in CKD patients vaccinated with the Oxford-AstraZeneca vaccine, emphasizing the need for expanded research with various vaccines and variants in this high-risk population.

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CONFLICT OF INTEREST STATEMENT

None declared.

DATA AVAILABILITY STATEMENT

Data will be available from the corresponding author on reasonable request.

REFERENCES

- 1. The Hindustan Times. Photos: India's daily Covid-19 cases cross 400,000. https://www.hindustantimes.com/photos/ news/photos-india-s-daily-covid-19-cases-cross-400000-101619857250444.html (last accessed on 17 May 2021)
- 2. Prasad N, Bhatt M, Agarwal SK et al. The adverse effect of COVID pandemic on the care of patients with kidney diseases in India. Kidney Int Rep 2020; 5: 1545-1550
- 3. Gansevoort RT, Hilbrands LB. CKD is a key risk factor for COVID-19 mortality. Nat Rev Nephrol 2020; 16: 705-706
- 4. Cai R, Zhang J, Zhu Y et al. Mortality in chronic kidney disease patients with COVID-19: a systematic review and meta-analysis. Int Urol Nephrol 2021; doi: 10.1007/s11255-020-02740-3
- 5. Grupper A, Sharon N, Finn T et al. Humoral response to the Pfizer BNT162b2 vaccine in patients undergoing maintenance hemodialysis. Clin J Am Soc Nephrol 2021; 10.2215/CJN.03500321
- 6. Glenn DA, Hegde A, Kotzen E et al. Systematic review of safety and efficacy of COVID-19 vaccines in patients with kidney disease. Kidney Int Rep 2021; 6: 1407-1410
- 7. Windpessl M, Bruchfeld A, Anders HJ et al. COVID-19 vaccines and kidney disease. Nat Rev Nephrol 2021; 17: 291-293
- 8. Boyarsky BJ, Werbel WA, Avery RK et al. Immunogenicity of a single dose of SARS-CoV-2 messenger RNA vaccine in solid organ transplant recipients. JAMA 2021; 325: 1784-1786
- 9. Alfano G, Fontana F, Mori G et al. Seroconversion after COVID-19 vaccine in a patient on dialysis on immune suppressants. Clin Kidney J 2021; 14: 1983-1984