Supplementary material and methods

Our tertiary care hospital of a medical university in South India had been ordained as the State COVID Hospital on 13 March 2020. From then, till now we had admitted and managed COVID-19 disease patients from several districts of our state and neighbour states.

On 3 April 2020 we had commissioned a haemodialysis and a peritoneal dialysis units exclusively for COVID-19 disease patients. A dialysis water treatment plant with the capacity of 1000 L/hour had been initiated.

We shifted six haemodialysis machines to the state COVID hospital. We earmarked one machine each for hepatitis B, hepatitis C and human immunodeficiency virus infected patients. We also shifted two automated peritoneal dialysis machines.

Patients in these wards are managed by teams of doctors. The nephrologists of the institute had always been part of these teams. The continuity of care for nephrology patients was established by a team of nephrologists, out of which one doctor provided care on all days.

All patients had serum creatinine, blood urea, serum sodium and serum potassium, complete haemogram, liver function tests, prothrombin time and partial thromboplastin time, serum procalcitonin, serum ferritin, C-reactive protein, lactate dehydrogenase and serum D-dimer sent on the first day of admission. We requisitioned for serum IL-6 and later considered to prescribe injection tocilizumab when the patients had features that suggested cytokine storm.

We collected the data of end stage renal disease (ESRD) patients on maintenance haemodialysis (MHD) contemporaneously in a register. From this register and based on the information in this register the data were retrieved from patients files—a retrospective observational study. The data from admission to the outcomes were recorded on a computerised proforma. We included the demography, clinical features, laboratory data and treatment schedules of haemodialysis and COVID-19 disease in this proforma.

Revised Guidelines on Clinical Management of COVID – 19 by Ministry of Health & Family Welfare, Government of India published on 31 March 2020 classified the COVID-19 disease patients as patients of uncomplicated illness, mild pneumonia, severe pneumonia and adult respiratory distress syndrome. In clinical practice most patients of mild pneumonia required oxygen either by non-rebreather mask or by simple mask, severe pneumonia and adult respiratory distress syndrome (ARDS) required non-invasive ventilation or mechanical ventilation. Similar classification was enunciated by the World Health Organisation (WHO). The clinical management of COVID-19 guidelines of WHO published on 27 March 2020 (WHO/2019-nCoV/clinical/2020.5) classified the COVID-19 disease patients as patients of mild, moderate, severe and critical disease. In practice the patients, after admission had migrated from the category of mild pneumonia to severe pneumonia to ARDS and also viceversa. Therefore, for the purpose of this study we followed these definitions.

Non-invasive ventilation at admission (NIVa): Patients who required non-invasive ventilation at admission.

Non-invasive ventilation patients in hospital (NIVh): Patients who required non-invasive ventilation at any time in hospital. This group included the patients who required NIV at admission. Oxygen dependent: Requirement of oxygen either by non-rebreather mask or by simple mask. These patients did not require NIV.

Our institute has a standard treatment protocol for COVID 19 disease patients. We initiated patients from admission on injection remedisivir 200 mg on the day 1, followed by 100 mg from day 2 to day 5 extended up to day 10 as per the clinical condition, injection dexamethasone 0.1 mg/kg/day iv and low molecular weight heparin till discharge. Almost all patients of maintenance haemodialysis were given this protocol of treatment.

Patients based on the requirement, received antihypertensives, insulin, 1, 25, dihydroxy vitamin D, phosphate binders, carnitine and erythropoietin.

We prescribed injection tocilizumab (Cipla Ltd.) when the serum IL-6 was elevated 10 times the reference range of our lab. We prescribed injection tocilizumab at 8 mg/kg per dose; up to a maximum dose of 400 mg. Likewise total two doses were prescribed. A thorough history and examination including chest radiograph was taken to exclude the current tuberculosis and any other bacterial or fungal infection. We also ensured that the following were not present: platelet count <100,000/mm³, neutrophil count <2000/mm³, and alanine aminotransferase (ALT) or aspartate aminotransferase (AST) more than five times the upper limit of the normal range.

The diagnosis of mucormycosis of the paranasal sinuses, orbit and brain were done with the aid of computerised tomography and/or MRI followed by endoscopic biopsy. The patients were treated with injection liposomal amphotericin and posoconazole and their combination. Surgery was decided by the departments of surgery.

Statistics: The data were entered into Microsoft Excel. The frequency and percentage was calculated for qualitative variables. The mean and standard deviation or median and interquartile range (IQR) was calculated for quantitative variables. Independent sample t test was used to test the significant difference between two means. Chi-square test was used to test the significant difference between proportions. Odds ratio with 95% confidence intervals was calculated. Multivariate binary logistic regression was used to predict the occurrence. IBM SPSS 26th version was used. P value less than 0.05 was considered as statistically significant. We obtained the clearance from the institutional ethics committee, number: 1247.

Supplementary references

- S1. Wang R, Liao C, He H, Hu C, Wei Z, Hong Z et al. COVID-19 in Hemodialysis Patients: A Report of 5 Cases. Am J Kidney Dis. 2020;76(1):141-143
- S2. Chan L, Jaladanki SK, Somani S, Paranjpe I, Kumar A, Zhao S et al. Mount Sinai COVID Informatics Center (MSCIC). Outcomes of Patients on Maintenance Dialysis Hospitalized with COVID-19. Clin J Am SocNephrol. 2021;16(3):452-455.
- S3. Ghonimi TAL, Alkad MM, AbuhelaiqaEA,Othman MM, Elgaali MA, Ibrahim RAM, et al.Mortality and associated risk factors of COVID-19 infection in dialysis patients in Qatar: Anationwide cohort study. PLoS ONE 2021; 16(7): e0254246.
- S4. Creput C, Toledano D, Diaconita M, Izzedine H. COVID-19 in patients undergoing hemodialysis: prevalence and asymptomatic screening during a period of high community prevalence in a large Paris center. Kidney Med. 2(6):716-723.
- S5. Smolander J, Bruchfeld A. The COVID-19 epidemic: management and outcomes of hemodialysis and peritoneal dialysis patients in Stockholm, Sweden. Kidney Blood Press Res 2021;46:250–256.
- S6. Kazmi S, Alam A, Salman B, Saeed F, Memon S, Chughtai J et al. Clinical course and outcome of esrd patients on maintenance hemodialysis infected with COVID-19: A single-center study. Int J NephrolRenovasc Dis, 2021;14:193-199.
- S7. Valeri AM, Robbins-Juarez SY, Stevens JS, Ahn W, Rao MK, Radhakrishnan J et al. Presentation and outcomes of patients with ESKD and COVID-19. J Am SocNephrol. 2020;31(7):1409-1415.
- S8.Ng JH, Hirsch JS, Wanchoo R, Sachdeva M, Sakhiya V, Hong S, et al.Outcomes of patients with end-stage kidney disease hospitalized with COVID-19. Kidney Int. 2020;98(6):1530–9.
- S9. Vardhan H, Kumar A, Shyama S, Chaudhary N, Pandey S, Rai DK et al. Clinical Profile and Outcome of Haemodialysis in Patients With COVID-19 A Single Centre Experience. Cureus. 2021 Aug 14;13(8):e17170.
- S10.Prasad N, Behera MR, Bhatt M, Agarwal SK, Gopalakrishnan N, Fernando E et al. Outcomes of symptomatic coronavirus disease 19 in maintenance hemodialysis patient in India. Semin Dial. 2021;34(5):360-367.
- S11.Banerjee S, Patel HV, Engineer DP, Gupta V, Patel H, Gupta A, Shah PR, Kute VB. COVID-19 in Hemodialysis Patients: Experience from a Western Indian Center. Indian J Nephrol. 2022;32(3):216-222

S12.Trivedi M, Shingada A, Shah M, Khanna U, Karnik ND, Ramachandran R. Impact of COVID-19 on maintenance haemodialysis patients: The Indian scenario. Nephrology (Carlton). 2020 Dec;25(12):929-932

Supplementary table 1: Age and Mortality

Quartile-Age (years)	Number of patients	Mortality
		N (%)
<45	158	36 (22.8)
46-54	144	55 (38.2)
55-62	146	51 (34.9)
>62	147	61 (41.5)
Total	595	203 (100)

Supplementary table 2: Risk factors for mortality: Univariate analysis of continuous variables

Variable	Mortality		P
	Yes	No	
	Mean±SD	Mean±SD	
Duration of stay (days)	8.24±8.01	8.15±4.10	0.868
Age (years)	56.12±12.42	51.90±13.16	<0.001
Computerised	14.29±7.67	14.93±5.05	0.815
tomography severity			
score			
SPO ₂ at admission	88.49±11.83	94.42±6.09	<0.001
(percentage)			
SBP at admission (mm	133.21±25.67	134.11±21.85	0.653

Hg)			
DBP at admission	82.06±15.32	82.55±12.92	0.676
(mm Hg)			
Number .of dialysis	2.92±2.47	3.43±2.00	<0.001
sessions			
Haemoglobin (g/dL)	9.73±2.63	9.33±2.87	0.124
Total leucocyte count	13193.68±8424.44	8441.90±5570.57	<0.001
(/cu mm)			
Neutrophils	84.49±11.39	74.51±13.07	<0.001
Lymphocytes	9.39±8.41	17.50±11.71	<0.001
ESR (mm at end of	69.62±32.41	71.69±31.74	0.587
first hour)			
Platelet count (/cu	2.11±1.24	2.00±0.86	0.257
mm)			
Blood urea (mg/dL)	161.54±67.89	120.28±58.54	<0.001
Serum creatinine	9.61±22.75	8.27±4.06	0.355
(mg/dL)			
Serum potassium	5.03±1.096	4.77±2.56	0.208
(mEq/L)			
Serum sodium	145.81±112.36	135.21±12.18	0.806
(mEq/L)			

Serum bilirubin	0.66±0.66	0.58±0.91	0.326
(mg/dL)			
AST (IU/L)	73.82±108.35	44.72±89.65	<0.001
ALT (IU/L)	48.21±83.75	39.03±105.12	<0.001
Serum alkaline	137.91±96.54	130.54±99.52	0.161
phosphatase (IU/L)			
Serum total protein	5.16±2.36	4.79±2.82	0.163
(g/dL)			
Serum albumin (g/dL)	3.32±2.36	3.42±1.85	0.614
C reactive protein	106.63±89.57	61.92±71.33	<0.001
(mg/L) (reference			
range: 0-10mg/L)			
Procalcitonin (ng/mL)	46.50±115.95	9.82±18.94	0.066
(reference range: 0.1			
ng/mL)			
Serum ferritin	896.49±1138.97	661.64±613.45	0.003
(ng/mL) (reference			
range: 20 to 250			
ng/mL)			
D-dimer (μg/mL))	3.89±6.96	8.789±38.62	0.499
(reference range: 0.5			
μg/mL)			

Serum IL-6 (pg/mL)	106.84±107.81	57.27±100.43	0.299
(reference range:			
between 0 and 43.5			
pg/mL)			
LDH (U/L) (reference	746.79±1178.74	419.16±263.99	0.000
range: 140 to 280			
U/L)			

SPO₂ Oxygen saturation of blood, SBP: systolic blood pressure, DBP: diastolic blood pressure, AST: aspartate aminotransferase, AST: alanine transaminase, IL-6: interlukein-6, LDH: lactate dehydrogenase

Supplementary table 3: Risk factors for mortality: Univariate analysis of categorical variables

Variable	Number	Presence/absence	Mortali	ty n (%)	Total	P, OR (95%
		of the variable				CI)
			Yes	No		
Oxygen at	595	Yes	117	136	253	<0.001
admission			(46.2)	(53.7)		2.561 (1.809-
		No	86 (25.1)	256	342	3.625)
				(74.8)		
Non-oxygen	595	Yes	32 (11.6)	242	274	<0.001
requirement at				(88.3)		0.116(0.07-
admission						0.178)
		No	171(53.2)	150(46.7)	321	
NIV at	595	Yes	54(79.4)	14(20.5)	68	<0.001
admission						9.785 (5.276-
		No	149(28.2)	378(71.7)	527	18.15)
NIV in	595	Yes	108(74.4)	37(25.5)	145	<0.001
hospital						10.908(7.05-
		No	95(21.1)	355(78.8)	450	16.88)
InjTocilizumab	595	Yes	16 (48.5)	17 (51.5)	33	0.073
		No	187	375	562	1.887;
			(33.3)	(66.7)		95%CI:0.933-
						3.82
HIV	425	Yes	0 (0)	1 (100)	1	0.304

		No	152	272	424	
			(35.8)	(64.2)		
HBs Ag	387	Yes	2 (22.2)	7 (77.8)	9	0.650
		No	164	307	471	
			(34.8)	(65.2)		
Anti HCV	387	Yes	1 (10)	9 (90)	10	0.234
Antibody						
		No	165 (35)	306 (65)	471	

NIV: non-invasive ventilation

Supplementary table 4: Oxygen requirement and Mortality

Parameter	Total	Death
NIV at admission	68	54 (79.4%)
Oxygen at admission	253	117 (46.2%)
Non oxygen at	274	32 (11.6%)
admission		
Total	595	203 (34.1%)

NIV: non-invasive ventilation

Supplementary table 5: NIV patients & patients requiring Oxygen versus patients not requiring oxygen

NIV patients	Definition	Mort	tality	Total	P,
& patients		Yes	No		OR (95%
requiring					CI)
Oxygen					
versus					
patients not					
requiring					
oxygen					
Yes	On NIV & oxygen	171	150	321	< 0.001
	requirement at				8.621
	admission				(5.614-
No	Patients not requiring	32	242	274	13.239)
	oxygen				
Total		203	392	595	

NIV: non-invasive ventilation

Supplementary table 6: Patients requiring Oxygen versus patients not requiring oxygen

Oxygen	Definition	Mor	tality	Total	P,
patients		Yes	No		OR (95%
versus Non-					CI)
oxygen					
patients					
Yes	Oxygen requirement	117	136	253	< 0.001
	at admission, NIV				6.506
	patients not included				(4.174-
No	Patients not requiring	32	242	274	10.141)
	oxygen, NIV patients				
	not included				

Supplementary table 7: Comparison between the patients admitted in 2020 and 2021

Parameter	From March 2020 to 1 March 2021	From March 1, 2021 to till December 31, 2021	Total
Number of patients	269 (37.6%)	445 (62.3%)	714
Number of files available	210 (35.2%)	385 (64.7%)	595
Age (mean ±SD)	52.13 ± 12.32	54.00 ± 13.41	-
(years)			
Sex			
Male	166 (38.0%)	270 (61.9%) 72.9	436
Female	43 (27.0%)	(30.1%)	159
NIV at admission	22 (32.3%)	46 (67.6%)	68
NIV at hospital	46 (31.7%)	99 (68.2%)	145
stay			
Oxygen at admission	83 (32.8%)	170 (67.1%)	253

Non oxygen	114 (41.6%)	160 (58.3%)	274
Mortality	57 (28.0%)	146 (71.9%)	203

Supplementary table 8: International Published Studies

S	Referen	Type of	Duration of	Number of	Mortality	Risk
no	ce	the study	data	ESRD		factors
			collection	patients on		
			and year of	MHD with		
			publication	COVID-19		
				disease		
1	S1 [9]	Single	December	5	0	-
		centre	2019 to			
			February			
			2020/2020			
2	S2 [10]	Single	March 15	122	11 (9%)	None
		centre	and June 7,			
			2020 /2021			
3	2 [11]	Single	March 12th	36	11 (30.5%)	Longer dialysis
		centre	to April			vintage, increased
			10th, 2020			LDH, higher CRP,
			/2020			lower lymphocyte
						count
4	3 [12]	Four	March	94 out of	27 (28.7%)	Fever at disease

		dialysis	2020/2020	643 (15%)		diagnosis, cough at
		centres				disease diagnosis,
						higher serum CRP at
						disease
						diagnosis
5	4 [13]	47	17th	567	93 (16.3%)	Cox regression
	•	centres	April to 1st			survival analysis:
		in	June 2020.			Age, severe-critical
		Turkey	/2021			disease at the time of
						diagnosis, presence
						of congestive heart
						failure, ferritin
						levels on admission,
						AST (> 2x upper
						limit of normal)
						during
						hospitalization,
						thrombocytopenia
						during
						hospitalization
6	S3 [18]	Nationwi	February 1,	76 out of		Multivariate
		de study	2020, to	1064		analysis: ICU
			July 19,			admission
			2020/2021			Univariate Cox
						regression:age,

						CHF, COPD,
						history of DVT,
						atrial fibrillation,
						hypoxia, ICU
						admission,
						mechanical
						ventilation, using
						inotropes
7	S4 [19]	Two	March 31 to	38 (19%)	8 (21%)	None
		centres	April 4,	out of 200		
			2020			
8	S5 [20]	Single	March 12 to	40	9 (22.5%)	Median age in non-
		centre	April 17,			survivors (78
			2020, /2021			years) was
						significantly higher
						than in survivors,
						median time in
						dialysis (11.5 years)
						significantly
						longer in non-
						survivors, CRP at
						diagnosis
						significantly higher
9	S6 [21]	Single	15 March	43	11 (Multivariate logistic
		centre	2020 to 30		25.6%)	regression: Age >
		<u> </u>				

			September			65 years, high TLC
			2020 /2021			Other significant risk
						factors: lymphocyte
						count, LDH
10	S7 [22]	Single	March 9,	57	18 (31%)	Age >75 years,
		centre	2020 to			higher median
			April 8,			Charlson
			2020 /2020			comorbidity
						index
11	S8[23]	13	March 1,	419/10482	133(31.7%)	Increased age, being
		centres	2020, to	(3.9%)		on a ventilator,
			April 27,			lymphopenia, blood
			2020/2020			urea nitrogen and
						serum ferritin.

TLC: total leucocyte count, LDH: lactate dehydrogenase, C-reactive protein levels, AST: aspartate transaminase, CHF: congestive heart failure, COPD: chronic obstructive pulmonary disease, DVT: deep vein thrombosis

Supplementary table 9: Published studies from India

S	Referen	Type of	Duration of	Number of	Mortality	Risk
no	ce	the study	data	ESRD		factors
			collection	patients on		
			and year of	MHD with		
			publication	COVID-19		
				disease		
1	S9 [14]	Single	Between	40	16 patients	Multiple logistic
		centre	May 1st,		(40%)	regression: Invasive
			2020 and			ventilation, deranged
			March 31st,			TLC,
			2021/ July			hypoalbuminaemia
			2021			
						Other significant
						factors: Age, mean
						SpO ₂ at presentation
						and raised
						inflammatory
						markers were
						significantly
						associated with
						mortality
2	S10	11 public	Between	263	35 (13.3%)	Multivariate
	[15]	and	March 2020			analysis: CVC use,
		private	and July			disease severity,

		centres	2020/July			NIV support
			2021			
						Other significant
						factors: Older,
						diabetic kidney
						disease,
						comorbidities,
						severe COVID-
						19,on twice-weekly
						MHD than thrice-
						weekly, dialysis
						through central
						venous catheter
3	S11 [16	Single	Between	58	22 (37.9%)	Disease severity,
]	centre	May 2020			CRP above 175
			and July			mg/L at admission
			2020			
4	S12	Two	Between 1	37	14 (37.8%)	-
	[17]	dialysis	April and			
		centres	30 April			
			2020			

TLC: total leucocyte count, LDH: lactate dehydrogenase, C-reactive protein levels, AST: aspartate transaminase, CHF: congestive heart failure, COPD: chronic obstructive pulmonary disease, DVT: deep vein thrombosis

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