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COVID-19 mortality and its associated factors in Nepal: A cross-sectional study



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ARTICLE INFO	A B S T R A C T
<i>Keywords:</i> Comorbidities Coronavirus disease COVID-19 pneumonia Mortality	Objectives: Reports from other countries have indicated that severe forms and fatal cases of COVID-19 in older adults and people with underlying comorbidities. The aim of this study was to assess the risk factors associated with COVID-19 mortality in Nepal. Methods: A cross-sectional study was conducted from April 12 to July 23, 2021 to identify the underlying factors associated with COVID-19 deaths. Our sample included all cases diagnosed and registered as COVID-19–related

Results: A total of 1459 COVID-19 hospital-based death records were collected from 30 hospitals. Mean age at death was 60.2 (±15.6) years. One-third of cases were admitted with fever, cough, and shortness of breath. The computerized tomography Severity Score showed that 7.3% of the individuals who underwent high-resolution computerized tomography chest had a severe form of lung involvement, and 3.6% had mild to moderate involvement. The most common comorbidities were hypertension (43.7%) followed by diabetes mellitus (25.8%). Among the deceased, 37.7% were diagnosed as cases of COVID-19 pneumonia. The most common recorded causes

Conclusions: Individuals with comorbidities including hypertension and diabetes mellitus were at greater risk of developing complications and had a higher rate of mortality.

related mortality.

the deceased were explored.

Methods

Introduction

COVID-19 was first reported in Wuhan, China, in December 2019 and spread rapidly to become a pandemic as declared by WHO on January 23, 2020 [1]. According to the WHO estimates, as of May 20, 2022, the pandemic witnessed 521,920,560 cases and 6,274,323 deaths globally and 58,056,402 cases and 787,989 deaths in South East Asia [2]. The first COVID-19 case was identified in Nepal on January 23, 2020 [3]. According to the Ministry of Health and Population of Nepal, as of April 3, 2023, the total number of confirmed COVID-19 cases reached up to 1,001,470 and 12,020 deaths [4]. Previous studies have shown that the older people and patients with comorbidities including chronic lung diseases, liver disease, diabetes mellitus, chronic kidney disease, immunosuppressed, and overweight are likely to have a more severe illness [5]. In Nepal, the majority of the COVID-19 deaths were associated with people having comorbid conditions including hypertension, diabetes, respiratory illness, and chronic kidney patients [6-8]. The objective of this study was to identify the risk factors associated with COVID-

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deaths at 30 hospitals of Nepal.

of death were respiratory failure followed by cardio-pulmonary arrest.

19 deaths in Nepal. The results of this study would assist in developing evidence-based policies and prevention strategies to reduce COVID-19-

A cross-sectional study was conducted to identify the underlying risk

factors associated with COVID-19 deaths. We collected data of poly-

merase chain reaction (PCR)-diagnosed cases that were registered as

COVID-19-related deaths between May 2020 and July 2021. Of the total

hospitals requested to participate in this study only 30 hospitals agreed

for data collection. The in-hospital mortality data was provided by the

Ministry of Population and Health of Nepal. Those hospitals which were

not willing to share their data were excluded from the study. Data were

collected from hospital's medical records and from the death certificates

of the patients. The socio-demographic profile and clinical details such

as clinical presentation, risk factors, treatment history, and outcomes of

ulation [9]. Hence, the final sample size was calculated as 351. All the

The sample size was calculated using the formula, N/ (1+Ne2), where "e" (margin of error) is taken at 0.05, and "N" refers to the pop-



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Figure 1. Age distribution of COVID-19 deaths (n = 1459).

Figure 2. Oxygen saturation of COVID-19–infected individuals on admission (n = 1459). BiPAP, Bilevel Positive Airway Pressure

data were entered in a Microsoft Excel spreadsheet from the paper-based hospital records. The collected data were analyzed using SPSS (Statistical Package for Social Science), version 16. Various descriptive and inferential data analyses were used.

Results

Between May 2020 and July 2021, there were 1459 COVID-19related deaths in 30 hospitals of seven provinces in Nepal. These cases were tested positive for COVID-19 by reverse transcriptase (RT)-PCR. Almost half of the deaths (735 cases or 50.4%) occurred in the hospitals in Bagmati province, whereas Karnali province had the least number of deaths (52 cases or 3.6%). The mean age was 60.2 ±15.6 years, and the median age was 60 years with interguartile range (17-102). The age distribution of the cases showed that highest rate of in-hospital COVID-19-related mortality (23.9%) occurred in the age group 56-65 years (Figure 1). Among all COVID-19 deaths, 952 (65.3%) were male while 507 (34.7%) were female. Majority of the COVID-19-infected individuals 1217 (83.4%) who presented to the healthcare center had shortness of breath as the common presenting symptom followed by cough in 1077 (73.8%) patients. The mean (\pm SD) duration of hospital stay before the event (death) was 6.8 (\pm 6 days). The mean (\pm SD) duration of illness on presentation was 5.2 (\pm 4.3) days and median days were 4 (interquartile range 1-39). Among the total COVID-19-related deaths, 826 (56.6%) were smokers.

Oxygen saturation status on admission to the hospital

At the time of admission, out of all individuals who had COVID-19– related death, 139 (24.5%) had oxygen saturation of <90% in room air, 450 (31.5%) had oxygen saturation of <90% even with oxygen supplementation, and 18 (1.2%) had oxygen saturation of <90% even under Bilevel Positive Airway Pressure support (Figure 2).

Of the 311 patients who had done chest X-ray, 71 (4.9%) individuals showed homogenous patchy opacities with lower zone infiltration in bilateral lungs. Similarly, among the 203 patients who had HRCT (High Resolution Computed Tomography) chest, 70 (4.8%) showed diffuse ground glass opacities in bilateral lungs as the most common finding. The computerized tomography Severity Score showed that 107 (7.3%) of the individuals who underwent an HRCT chest had a severe form of lung involvement and 52 (3.6%) had a mild to moderate form of the disease.

Comorbidities on admission to the hospital

The most commonly known comorbidities on admission to the hospital among the COVID-19 mortality cases were found to be hypertension (638 cases or 43.7%) followed by diabetes mellitus (376 cases or 25.8%), acute and chronic renal disorders, (180 cases or 12.3%) and chronic obstructive pulmonary disease (100 cases or 6.9%).

Diagnosis upon hospital admission

Among all deceased, the most common diagnosis at admission to the hospital were COVID-19 pneumonia (619 cases or 42.4%) without comorbidities followed by COVID-19 pneumonia with hypertension (210 cases or 14.4%). In addition to chronic liver disease and coronary artery disease, interestingly, hypothyroidism was also seen in a significant number of cases (27 or 1.9%) who had COVID-19–related deaths.

Table 1

Diagnosis upon admission to the hospital in the order of frequency (n = 1459).

Diagnosis	Frequency (N)	Percent (%)
COVID-19 PCR positive with Pneumonia		
Without comorbidities	619	42.4
With comorbidities	521	35.8
HTN	210	14.4
HTN and DM	90	6.2
CKD	58	4.0
AKI	28	2.0
DM/ DKA	42	2.9
CLD	10	0.6
Other Comorbidities		
Hypothyroidism	27	1.9
Coronary Artery Disease	21	1.4
Carcinoma	35	2.4
COVID-19 Pneumonia with complications		
ARDS	56	3.9
Sepsis	9	0.6
COVID-19 PCR positive without Pneumonia		
RT-PCR positive for COVID-19 only	136	9.3
Acute exacerbation of COPD	72	4.9
MI	16	1.1
Ischemic stroke	13	0.9
RTA	5	0.3
Fall injury with multiple fractures	3	0.2
OP poisoning	1	0.1
Other infections with RT-PCR positive		
PTB	3	0.2
Scrub Typhus	3	0.2
Dengue fever	2	0.1

AKI, Acute Kidney Injury; ARDS, Acute Respiratory Distress Syndrome; CKD, Chronic Kidney Disease; CLD, Chronic Liver Disease; COPD, Chronic Obstructive Pulmonary Disease; DKA, Diabetic Ketoacidosis; DM, Diabetes Mellitus; HTN, Hypertension; MI, Myocardial Infarction; OP, Organophosphorus Poisoning; PCR, Polymerase Chain Reaction, PTB, Pulmonary Tuberculosis; RTA, Road Traffic Accident.

There were COVID-19 pneumonia with complications like acute respiratory distress syndrome (56 cases or 3.9%). There were few cases of COVID-19 PCR positive with Myocardial Infarction (16 cases or 1.1%) and road traffic accident with the accidental finding of COVID-19 PCR positive (5 cases or 0.3%) (Table 1).

Treatment received by patients for the illness

The majority of deceased, 99.4% who visited the hospital for their symptoms were admitted in the healthcare facilities for further treatments. A total of Nine patients died in the emergency room, 97.3% had oxygen therapy and 85.3% needed intensive care unit care. More than 80% had taken steroids and 40.3% got intubated. Similarly, more than 40% had received heparin and azithromycin for treatment (Table 2).

Final Diagnoses of the COVID-19 deaths

Among a total of 1459 COVID-19–positive deceased cases, 550 (37.6%) had a diagnosis of COVID-19 pneumonia without known comorbidities, 214 (14.6%) with COVID-19 pneumonia with hypertension, and 163 (11.2%) with COVID-19 pneumonia induced acute respiratory distress syndrome and/or sepsis (Table 3).

Cause of death

As per death certificates, the most commonly recorded immediate causes of deaths were respiratory failure, 635 (43.5%) followed by cardio-pulmonary arrest 298, (20.4%). Other common terminal events leading to deaths included acute respiratory distress syndrome, 210 (14.4%), septic shock, 156 (10.7%), and cardiac arrhythmias, 155 (10.6%).

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Table 2

Treatment received	l for	the	illness	(n =	1459).
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	Yes		No	
Treatment	N	%	N	%
Oxygen therapy	1,420	97.3	39	2.7
Symptomatic treatment ^a	1,240	85.5	219	15.0
Steroids (Dexamethasone)	1,186	81.3	273	18.7
Vitamin C	809	55.4	650	44.6
Non- invasive ventilation ^b	732	50.2	727	49.8
Zinc	699	47.9	760	52.1
Heparin	664	45.5	795	54.5
Azithromycin	610	41.8	849	58.2
Invasive ventilation ^c	588	40.3	871	59.7
Antiviral (Remdesivir)	543	37.2	916	62.8
Vitamin D	470	32.2	989	67.8
Ivermectin	241	16.5	1,218	83.5
Hydroxychloroquine	169	11.6	1,290	88.4
Convalescent plasma therapy	152	10.4	1,307	89.6
Warfarin	144	9.9	1,315	90.1

^a Paracetamol, Oral rehydration solution, Ringer's lactate

^b Non-invasive ventilation include CPAP, Bi-PAP

^c Invasive ventilated patients included mechanical ventilator

Table 3

Final Diagnoses (n = 1459).

Diagnoses	Frequency (N)	Percent (%)
COVID-19 pneumonia without known comorbidities	550	37.6
COVID-19 pneumonia with comorbidities	699	48
Hypertension	214	14.6
Kidney disease	115	7.9
Hypertension with Diabetes mellitus	100	6.9
Chronic obstructive pulmonary disease	62	4.2
Diabetic ketoacidosis	57	4
Thyroid disorder	53	3.6
Carcinoma	35	2.6
Coronary Heart Disease	28	1.8
Chronic liver disease	10	0.7
Rheumatoid arthritis	7	0.5
Other Comorbidities ^a	18	1.2
COVID-19 pneumonia with complications		
COVID-19 pneumonia-induced ARDS and/or sepsis	163	11.2
COVID-19 pneumonia with Atrial fibrillation	14	1.0
Other diagnosis		
COVID-19 pneumonia with MI with HTN	22	1.5
COVID-19 pneumonia with Ischemic Stroke with HTN	6	0.4
RTA with SAH with RT-PCR positive for COVID-19	5	0.3

^a Other Comorbidities include: Pulmonary Tuberculosis, Hepatitis C, Crohn's disease, Idiopathic Thrombocytopenic Purpura, Scrub Typhus, Schizophrenia, Cushing's syndrome, Leprosy, Organophosphorus Poisoning.ARDS, Acute Respiratory Distress Syndrome; HTN, Hypertension; MI, Myocardial Infarction; RTA, Road Traffic Accident; RT-PCR, Reverse Transcriptase Polymerase Chain Reaction, SAH, Subarachnoid Hemorrhage.

Discussion

Our review of COVID-19–related deaths among hospitalized patients in Nepal showed that the majority of severe diseases leading to deaths occurred among the older (over 55 years old) and patients with multiple comorbidities. In this study, the mean age of COVID-19 deceased was 60.2 (SD±15.6) years. In comparison to our findings one of the study in Nepal revealed that the mean age of death was 50 years, in contrast to another study by Sanyaolu et al. where the mean age of death was 41 years [10,11]. Moreover, the median age of death in this study was 60.0 years which was almost similar to the publication by Patan Academy of Health Sciences, Nepal [12]. According to our study, nearly one-fourth of the COVID-19 deaths occurring in the hospitals were commonly distributed in the age group of 56-65 years, and almost two-thirds of deaths occurred in patients over 55 years. A systematic review and meta-analysis of 42 studies showed the average age of individuals with COVID-19 deaths occurred in the age range from 48.9-77 years [13]. The majority of mortality occurred among males (65.3% males, 34.7% females) in this study. These results are similar to the findings of a metaanalysis performed in China as well as two other studies done in two different hospitals of Nepal [11,12,14].

In this study, the most common presenting symptoms at the time of hospital admission were shortness of breath (83.4%) and cough (73.8%), which is similar to findings from other studies [12,14,15]. Other studies from Nepal and Bangladesh have also shown fever as the most frequent symptom [10,16]. The median duration of hospital stay in this study before reporting of an event (death) was five days and ranged from 0 to 43 days. Other studies have reported longer duration of hospital admissions prior to death from COVID-19. This outcome was discordant with the findings of Shrestha et al., another retrospective study conducted at Sukraraj Tropical and Infectious Diseases Hospital, Nepal, where the median duration of hospital stay was 10.5 days (range 1-35 days). A German study reported the average length of hospital stay as 16 days before death [12,14,17].

Out of total COVID-19 deaths, 31.5% were hypoxemic (oxygen saturation of <90% in room air) at presentation to the hospital which is similar to other studies [16]. The most common comorbidities among the COVID-19 mortality cases were hypertension, diabetes mellitus, chronic kidney disease, and acute renal insufficiency among patients who presented with COVID-19 pneumonia, whereas exacerbation of chronic obstructive pulmonary disease was the most common problem among those without pneumonia. These findings were similar to various studies conducted in Nepal, Brazil, USA, and Italy and meta-analysis performed by Peishan et al [10,18-21]. Similar findings were reported in various systematic reviews and meta-analysis, and comparable publications from Nepal, Wuhan (China), and France [11-14,17,22-24]. These would help to know the associated comorbidities of COVID-19 infection. In the current study, more than 80% of the patients had received steroids and 40.3% got invasive ventilation. Other medications received by the patients during the course of therapy included heparin, azithromycin, remdesivir, hydroxychloroquine, convalescent plasma therapy and warfarin. The usage of these medications were also reported by Bastola et al. in his study from Nepal, the study done by Koirala et al. and similar other studies from Bangladesh and Germany [14,16,17,25].

The most commonly documented final causes of deaths in the current study were respiratory failure followed by cardio-pulmonary arrest, acute respiratory distress syndrome, and septic shock. This was comparable to the reports from similar other studies from Nepal and other parts of the world including India and USA [14,26–29]. In this study, there were 1.5% cases of Myocardial Infarction with a single case of Organophosphorus poisoning. This might conclude that all of the associated factors may not be the actual cause of COVID-19 death.

Although we had planned to trace all cases of COVID-19–related mortality, we were unable to acquire detailed data because of the inadequate documentation of patient details, inaccessibility to complete data, and unwillingness to share the details of COVID-19 mortality cases by many medical institutions. These could be considered as the limitations of the study.

Conclusions

This analysis of COVID-19 related mortality among hospitalized patients revealed the highest death rates among the age group over 55 years. This study concludes that the patients with comorbidities including hypertension, diabetes mellitus, renal insufficiency, and COPD (Chronic Obstructive Pulmonary Disease) were associated with COVID-19 deaths.

Conflict of interest

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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Ethical approval statement

Ethical clearance was obtained from the Ethical Review Board of Nepal Health Research Council with a reference number of 2915. All the information collected was kept confidential and anonymity of the participants was assured.

Supplementary materials

Supplementary material associated with this article can be found, in the online version, at doi:10.1016/j.ijregi.2023.08.002.

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