### Features associated with SARS-COV-2 positivity among people presenting with acute respiratory tract infections to public Hospitals in Harari region, Ethiopia

SAGE Open Medicine Volume 9: 1-7 © The Author(s) 2021 Article reuse guidelines: sagepub.com/journals-permissions DOI: 10.1177/20503121211062793 journals.sagepub.com/home/smo



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### Abstract

Background: Despite investigating coronavirus among respiratory tract infected cases is a top priority to prevent further transmission, severe acute respiratory syndrome coronavirus 2 positivity among this group of patients remains unexplored in resource-limited settings. Therefore, this study intended to assess the severe acute respiratory syndrome coronavirus 2 positivity among patients presenting with acute respiratory tract infection from 1 July to 31 December 2020 in Harar Region, Ethiopia, from 15 February to 10 March 2021.

Methods: A facility-based cross-sectional study design was used. Severe acute respiratory syndrome coronavirus 2 was tested by assaying oropharyngeal swabs using reverse transcriptase-polymerase chain reaction among patients presenting with acute respiratory tract infection in Harari Public Hospitals. A binary logistic regression was used to identify factors associated with severe acute respiratory syndrome coronavirus 2 positivity with an adjusted odds ratio at a 95% confidence interval.

Results: Out of a total of 1692 study participants, 388 (22.9%) of them tested positive for severe acute respiratory syndrome coronavirus 2. Of these severe acute respiratory syndrome coronavirus 2 positive patients, 364 (21.6%) patients presented with lower respiratory tract infection, while the rest only 24 (1.4%) presented with upper respiratory tract infection. Independent variables included separated/divorced in marital status (AOR = 0.53, 95% CI: 0.29–0.95), presenting with cough, fever, and difficulty of breathing (AOR = 2.5, 95% CI: 1.22-4.7), age group of 30-39 years (AOR = 0.35, 95% CI: 0.15-0.79), 40-49 years (AOR = 0.37, 95% CI: 0.14-0.94), and 50-59 years (AOR = 0.31, 95% CI: 0.13-0.76) compared to patients with the age of  $\geq$  60 years, had statistically significant association with severe acute respiratory syndrome coronavirus 2 positivity. **Conclusion:** Severe acute respiratory syndrome coronavirus 2 was positive among 388 (22.9%) acute respiratory tract infected people. Elder age, particular symptoms, such as cough, fever, and difficulty of breathing, and married marital status were associated with a severe acute respiratory syndrome coronavirus 2 positive test. In resource-limited setups, where a shortage of testing equipment is common, these findings could contribute to boosting targeted symptom-oriented screening schemes. Moreover, this study could have paramount clinical importance for further studies in the country.

### **Keywords**

Associated factors, SARS-COV-2, COVID-19, RT-PCR, Ethiopia

Date received: 17 June 2021; accepted: 8 November 2021

### Introduction

Coronaviruses belong to a large family of single-stranded RNA (ssRNA) viruses that are causative agents of the common cold and severe respiratory infections, which include severe acute respiratory syndrome (SARS) and Middle East respiratory syndrome (MERS).<sup>1-5</sup> On 31 December 2019, an outbreak of pneumonia caused by an unknown agent was <sup>1</sup>School of Medicine, College of Health and Medical Sciences, Haramaya University, Harar, Ethiopia

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reported in Wuhan, China, later named severe acute respiratory syndrome coronavirus 2 (SARS-COV-2).6,7 The outbreak of Coronavirus disease 2019 (COVID-19) has become a global most challenging health crisis.<sup>8</sup> Several assessments reported that because of the incident of this pandemic, there is the social, economic, and physiological damage to the society in both developed and developing countries.<sup>9</sup> According to the World Health Organization, reports starting from the first-day existence of coronavirus in both the number of deaths and the confirmed cases are increasing.<sup>10</sup> A study showed that the progression of the coronavirus could result in severe respiratory tract infection.<sup>11</sup> Even though countless efforts are being made, there are excess deaths and impacts on the life expectancy of the world's population due to COVID-19.12 In Ethiopia, before the virus reached the country, the government had designed directions to undertake preventive measures for the passengers and service providers. However, Ethiopia formally announced the first coronavirus case on 13 March 2020 and had increased to 2019 cases with 27 deaths as of 7 July 2020.<sup>13</sup> Besides, as of 10 April 2021, there were 225,516 confirmed COVID-19 cases and 3111 deaths recorded in Ethiopia.<sup>14</sup>

Despite that a survey conducted in Addis Ababa suggested that the large majority of the people are susceptible to COVID-19,<sup>15</sup> no data revealed the magnitude of this disease at the community and facility level in the country, even in Africa, with adequate sample size. In addition, a resourcelimited country, such as Ethiopia, might not afford to test all respiratory tract infected patients for SARS-COV-2.<sup>16</sup> Thus, prioritizing resources and knowing the people at high risk of getting SARS-COV-2 is crucial to contain the pandemic by policymakers, planners, and health facilities in the resourcepoor setups. Even if inclusively identifying all possible SARS-COV-2 positive cases using PCR has public health benefits for the screening and providing immediate response through prevention and control measures against COVID-19, SARS-COV-2 positivity among acute respiratory tract infections (ARTI) remains unknown. Therefore, this study aimed to assess the positivity of SARS-COV-2 and its associated factors among patients presented with ARTI.

### **Methods and materials**

### Study area, period, and design

A facility-based cross-sectional study design was implemented in the public hospitals that found in Harari Regional State. Harari region is one of the 10 regional administrations in Ethiopia. The region is located 522 km away from Addis Ababa, the capital city of Ethiopia due east. The region contains two public hospitals (Hiwot Fana Specialized University Hospital and Jogul General Hospital), one Federal Police Hospital, one Fistula Center, Private Hospitals, and eight health centers. This study was conducted in both Hiwot Fana Specialized University Hospital and Jugol General Hospital, the public hospitals in the region. The two hospitals are the public health facilities owned by the government where everybody who seeks medical care could get service. Hiwot Fana Specialized University Hospital is the largest referral and teaching hospital in Eastern Ethiopia and receives tertiary referrals from the Harari region, Eastern Oromia, Somali region, and Dire Dawa City. During COVID-19 pandemic, the hospital is one of the 10 regional centers designated by the Ethiopia Ministry of Health to manage the COVID-19 cases as a center in Eastern Ethiopia.

### Study population and eligibility criteria

The source population was all patients who presented with ARTI in Hiwot Fana Specialized University Hospital and Jugol Hospital. And the study population was all patients who presented with ARTI in Hiwot Fana Specialized University Hospital and Jugol Hospital from 1 July to 31 December 2020. All patients who presented with respiratory tract infections in the Hiwot Fana Specialized University Hospital and Jugol Hospital were retrospectively included in the study. However, a patient who had no documented SARS-COV-2 result (positive or negative) was excluded.

# Sample size determination and sampling procedure

Initially, a minimum sample size was determined using single proportion population with p=0.5 (no prior data from the study area), Z=95% confidence interval, margin of error (d)=0.05,  $n=(z\alpha/2)^2$ pq/d<sup>2</sup>=384. The final sample was determined to be 422 with 10% of non-response rate. However, we included all patients presenting with ARTI at both public hospitals. A total of 1256 and 436 study participants were included from Hiwot Fana Specialized University Hospital and Jugol Hospital, respectively. Finally, a total of 1692 study participants were involved in this study (Figure 1).

### Study variables

The dependent variable was the SARS-COV-2 positivity. The independent variables included socio-demographics, such as age, sex, residence, marital status, education status, and occupation. COVID-19-related risk factors, signs, and symptoms included chills, rigors, cough, sputum, malaise, myalgia, diarrhea, rhinorrhea, fever, travel history, contact history, smoking, medical history, history of medication, and SARS-COV-2 testing result. Other independent variables included underlying diseases, such as diabetes, hypertension, chronic lung disease, ischemic heart disease, chronic renal disease, thyroid disease, and neoplastic disease.

### Operational definitions

Acute respiratory infection is an infection that may interfere with the normal breathing system that can be commonly

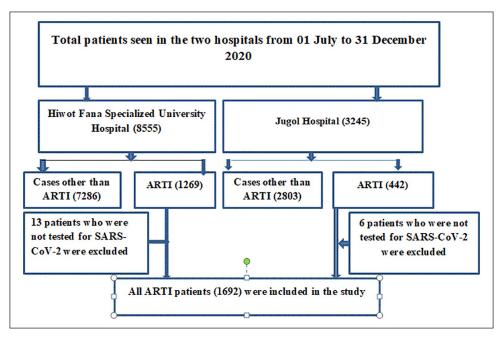


Figure 1. Schematic diagrammatic presentation of sampling procedure.

presented with cough, fever, shortness of breathing, runny nose, sneezing, and chest pain for short period.<sup>17</sup> The classifications of the respiratory tract infections, lower and upper types of infection, were diagnosed by physicians depending on the anatomical sites involved by the infection and the symptoms that the patients presented with.

SARS-COV-2 positive is the proportion of positively tested SARS-COV-2 cases using reverse transcriptase–polymerase chain reaction (RT-PCR) detection from oropharyngeal swabs.<sup>18</sup>

Contact history was defined as having a close or causal contact with COVID-19 confirmed/probable cases in the last 14 days prior to attending the health facilities.

Travel history was defined as having a history of travel to COVID-19 reporting areas in the past 14 days before attending the health facilities.

Comorbidity is any disease that co-exists with COVID-19.

History of medication was reviewed for chronic health conditions.

### Equipment used and preparation made on the SARS-COV-2 assay using RT-PCR

Viral transport medium (VTM), swab sticks, ice-box, icepads, tongue depressor, marker, requisition form, and personal protective equipment (PPE) were prepared and used beforehand. The VTM that contains 3 mL fluids composed of gelatin and antimicrobial agents in a buffered salt solution was used. This VTM was used to prevent the sample collected from drying, maintains the viability of the virus, and avoids the growth of contaminants. The used swabs are made up of rayon with a plastic shaft. Ice-box and ice-pads are used for maintaining a cold chain during sample transportation from the sample collection area to the laboratory center.<sup>19</sup> The ice-pad was filled with water and stored in the freezer  $(-20^{\circ}C)$  before and after use.

The sample contamination from the nasal vestibule was avoided by sterilely opening the outer case of the swab, inserting the swab into the mouth by slightly elevating the tip of the nose, and keeping the tip of the swab in the oropharynx for a few seconds, then rotated to achieve the highest absorption of oropharyngeal secretions.<sup>20,21</sup>

### Data quality and procedures

Data were retrospectively extracted from the patients' medical record by six BSc health professionals. Two public health professionals were involved in the data collection as supervisors. Before data collection, they took 2 days of training on the objective and relevance. Before actual data collection, 5% of the sample size was pre-tested on the patients presented with ARTI before 1 July 2020, in Hiwot Fana Specialized University Hospital. All patients who presented with ARTI were screened by RT-PCR for SARS-COV-2 in oropharyngeal. SARS-COV-2 loads were estimated with cycle threshold (Ct) values. SARS-COV-2 viral load is estimated with Ct values with the cutoff value of the mean value plus 3 standard deviations of the negative control.<sup>22</sup>

### Statistical analysis

The quality of the data was maintained by checking for consistency, completeness, and accuracy manually during data

Variables	Category	Ν	%	
Sex	Male	816	48.2	
	Female	876	51.8	
Age (years)	Mean = 32.6 ( $\pm$ 2SD)			
Age (years)	0–9	375	22.2	
	10–19	112	6.6	
	20–29	557	32.9	
	30–39	265	15.6	
	4049	148	8.7	
	50–59	112	6.6	
	≥60	123	7.3	
Religion	Muslim	1292	77.5	
	Orthodox	280	16.8	
	Protestant	94	5.5	
	Others	4	0.24	
Marital status	Single	224	13.4	
	Married	932	55.6	
	Separated/divorced	520	31	
Educational status	Unable to read and write	529	31.3	
	Able to read and write	1163	68.7	
Region	Harari	754	44.6	
	Oromia	938	55.4	
Contact history	Yes	244	14.4	
	No	1448	85.6	
Travel history	Yes	28	1.6	
-	No	1664	98.4	

**Table I.** Socio-demographic characteristics of the study participants (*N* = 1692).

collection time. Data were entered using Epi-data 3.1 and exported to STATA 14.2 for analysis. Descriptive measures, such as mean with standard deviation, percentages, and frequencies, were used to characterize the study population. The binary logistic regression was used to identify the association between outcome variables (SARS-COV-2 positivity) and independent variables with an adjusted odds ratio (AOR) at a 95% confidence interval. Statistically, a significance level was declared at a *p*-value of less than 0.05.

### Ethical statement

Haramaya University College of Health and Medical Sciences Institutional Health Research Ethical Review Committee (IHRERC) ethically cleared the paper (Ref. no: IHRERC/ 019/2021, with the approval date of 10 February 2021). A letter of permission was sent from the College of Health and Medical Sciences to Hiwot Fana Specialized University Hospital and Jugol Hospital. Due to difficulty to reach patients to take informed consent, written informed consent was obtained from legally authorized representatives of the facilities before the extraction of data. The data were obtained from patients' medical records anonymously guaranteeing information confidentiality, and all data extraction processes were conducted as per the declaration of Helsinki. The collected data were used only for the intended purpose.

### Results

### Socio-demographic characteristics of the study participants

Out of a total of 1692 patients, 816 (48.2%) were males, while 876 (51.8%) were females. The mean age of the participants was 32.6 years, with standard deviation of 2 years. The majority (77.5%) of the study participants were Muslim, followed by 16.8% Orthodox religion. Among the total study participants, single, married, and separated/divorced were 13.4%, 55.6%, and 31%, respectively. Among the total study participants, 1163 (68.7%) of them had formal education, while the rest 529 (31.3%) had no formal education. Out of a total of 1692 study participants, 754 (44.6%) of them were Harari region residents, while the rest 938 (55.4%) were from the Oromia region. Among the total participants, only 14.4% and 1.6% of them had a contact history with suspected/confirmed COVID-19 cases and travel history to the pandemic areas, respectively (Table 1).

# Clinical characteristics and comorbidities of study participants

Out of a total of 1692 study participants, 388 (22.9%) of them tested positive for SARS-COV-2. Out of a total of 1692 study participants, a majority (78.3%) of them presented with cough, fever, and shortness of breathing. The mean duration of COVID-19 symptoms presentation was 14.6 days  $(\pm 2.6$ SD). About 182 (10.8%) of the study participants had comorbidities at the time of visit. Of these comorbidities, hypertension, diabetes mellitus (DM), cardiac diseases, tumor, and HIV/AIDS diagnosed among 165 (9.8%), 63 (3.7%), 87 (5.1%), 8 (0.5%), and 24 (1.4%), respectively. A majority of the study participants, 220 (13%) and 64 (3.8%), had history of taking chronic case medication and smoking, respectively. Regarding the types of respiratory tract infection, 510 (30.1%) participants presented with lower respiratory tract infection, while the rest 1182 (69.9%) presented with upper respiratory tract infection (Table 2).

### Factors affecting SARS-COV-2 positivity

Variables that had a *p*-value less than 0.3 were transformed to multivariable logistic regression. A separated/divorced marital status, age above, and symptoms with the patients presented had statistically significant association with SARS-COV-2 infection (p < 0.05). The odd of SARS-COV-2 positivity was reduced by 47% among separated/ divorced in marital status compared to married individuals (AOR=0.53, 95% CI: 0.29–0.95). Besides, patients who were in the age group of 30–39 years became positive for SARS-COV-2 65% less likely than the patients with the age of 60 years and above (AOR=0.35, 95% CI: 0.15–0.79). Moreover, patients who were in the age group of 40–49 years became positive for SARS-COV-2 63% less likely compared

Variables	Category	N	%	
COVID-19 symptoms on time	Sore throat, headache, sneezing	141	8.4	
of visit	Cough, fever, and SOB	1325	78.3	
	Chest pain, diarrhea	226	13.3	
Mean duration of the symptoms	14.6 days (±2.6SD)			
Existence of	Yes	182	10.8	
comorbidities on presentation	No	1510	89.2	
Hypertension	Yes	165	9.8	
	No	1524	90.2	
Diabetes mellitus	Yes	63	3.7	
	No	1629	96.3	
Cardiac disease	Yes	87	5.1	
	No	1603	94.9	
Malignant/tumors	Yes	8	0.5	
	No	1684	99.5	
HIV/AIDS	Yes	24	1.4	
	No	1668	98.6	
History of taking any	Yes	220	13	
medication	No	1472	87	
Smoking history	Yes	64	3.8	
	No	1628	96.2	
Types of respiratory	Lower	510	30.I	
tract infection	Upper	1182	69.9	

Table 2.	Clinical	characteristics	of the	study	participants
(N=1692)	).				

COVID-19: coronavirus disease 2019; SD: standard deviation; SOB: shortness of breathing.

to patients with the age of 60 years and above (AOR=0.37, 95% CI: 0.14–0.94). Furthermore, patients who were in the age group of 50–59 years were positive for SARS-COV-2 69% less likely than the patients with the age of  $\geq$  60 years (AOR=0.31, 95% CI: 0.13–0.76). The study subjects who presented with cough, fever, and difficulty of breathing had SARS-COV-2 2.5 times more likely compared to those who presented with symptoms, such as sore throat, headache, and sneezing (AOR=2.5, 95% CI: 1.22–4.7) (Table 3).

### Discussion

This study demonstrated that SARS-COV-2 was positive for 388 (22.9%) acute respiratory infected people. Majority (21.6%) of SARS-COV-2 positive patients presented with lower respiratory tract infection, (p < 0.0001). The factors, such as separated/ divorced marital status, age, and particular symptoms, such as cough, fever, and shortness of breathing, had statistically significant association with SARS-COV-2 positivity.

This study revealed that the SARS-COV-2 positivity was 22.9% among acute respiratory tract infected people. The majority (21.6%) of the patients who were diagnosed with lower respiratory tract infections were positive for SARS-COV-2. This finding was higher than the studies detected SARS-COV-2 using RT-PCR. The SARS-COV-2 was positive

among 13.2% in Thailand<sup>23</sup> and 13.1% in Iran.<sup>24</sup> The discrepancy might be due to the variation in socio-demographic characteristics and potential spatiotemporal variations in viral distribution. The discrepancy may support the importance of generating evidence from both developed and resource-limited setups like the study area of the present study. In addition, the current study's higher finding is likely due to the increasing infectiousness (R0) of SARS-Cov-2 over time. However, this study finding was lower than the report from Iran that was 36.08% cases.<sup>19</sup> This can be because the probability of testing positive for SARS-COV-2 among patients who presented with severe pneumonia is highly expected due to the highest degree of manifestations overlapping of the two infections. In addition, the finding from the current study was lower than the SARS-COV-2 positives (30.4%) reported from Nicaragua, Central America.<sup>20</sup> This can be since healthcare workers are at high risk of getting SARS-COV-2 infection due to their working area than a general population studied in the present study.

The study participants with divorced/separated marital status had 47% less likely to be positive for SARS-Cov-2. The finding was in line with the report from Switzerland that revealed higher SARS-COV-2 positive tests (57·2%) among family members and those who had close contact (19%).<sup>21</sup> A divorced/separated/widowed man/woman may live with a smaller family size/alone compared to married individuals. The smaller family size could decrease familial transmissions of SARS-Cov-2. Thus, living with less family size decreases the probability of unsafe contact with suspected/confirmed COVID-19 patients that result in a lesser viral spread.

This study also revealed that people above the age of 30 years had SARS-COV-2 infection 63% less likely compared to patients with the age of 60 years and above. This study was supported by the finding from Wuhan, China, Iran, and the United Kingdom.<sup>22,24,25</sup> In fact, due to aging and the risk of developing comorbidities, the older (>60 years) population may have hampered immunity that exposes them to SARS-COV-2 infection. According to the survey conducted in Ethiopia, the older population had a 31% lesser probability of engagement in COVID-19 preparedness and response,<sup>26</sup> and this can be due to distinct psychosocial support they need during this critical period for their lives.<sup>27</sup> Therefore, because of their less involvement in the preparedness and response against COVID-19 and a distinct psychosocial need, older population might not fully adhere to SARS-COV-2 containing ways, such as wearing masks, maintaining the physical distance to optimum, frequent hand washing, and practicing safe contact with suspected/confirmed COVID-19 cases.

In addition, the study participants who presented with cough, fever, and difficulty of breathing infected with SARS-COV-2 2.5 times more likely compared to those who presented with symptoms, such as sore throat, headache, and sneezing. In contrary to the present finding, according to the study conducted in Sweden;<sup>28</sup> fever and loss of smell and taste and in Brazil;<sup>29</sup> anosmia and ocular pain were associated with positive SARS-COV-2. Regarding the strength of this study, the study could be the first report in Ethiopia, even

Variables	Category	SARS-COV-2 status		COR	AOR (95% CI)	p-value
		Positive	Negative	-		
Sex	Male	183	633	1.1 (0.84–1.3)	1.1 (0.7–1.6)	0.6
	Female	205	671	, , , , , , , , , , , , , , , , , , ,	l í	
Age (years)	<18	100	387	2.2 (1.4–3.5)	0.54 (0.12-2.3)	0.2
	20–29	113	444	1.9 (1.3–3.1)	0.5 (0.23-1.3)	0.12
	30–39	56	209	1.9 (1.2–3.1)	0.35 (0.15–0.79)	0.01*
	40-49	44	104	1.3 (0.7–2.1)	0.44 (0.14–1.02)	0.05
	50–59	33	79	1.1 (0.7–2.5)	0.39 (0.12–1.04)	0.06
	>60	42	81	l` í	l í	
Marital status	Married	226	706	I	I.	
	Not married	36	188	1.67 (1.13–2.46	1.1 (0.6–2.1)	0.4
	Separated/divorced	123	397	1.1 (0.8–1.3)	0.53 (0.29–0.95)	0.03*
Educational status	Unable to read and write	152	377	l Í	l í	
	Able to read and write	236	927	1.5 (1.2–2.1)	0.86 (0.57-1.3)	0.5
Region	Harari	165	589	l` í	Ì	
-	Oromia	223	715	0.8 (0.7–1.3)	0.7 (0.6–1.4)	0.7
Contact history	Yes	44	200	1.4 (0.9–2.1)	0.86 (0.46-1.59)	0.6
·	No	344	1104	l í	l	
Travel history to areas with	Yes	7	21	0.9 (0.4–2.1)	0.58 (0.13-2.64)	0.5
COVID-19 pandemic	No	381	1283	l Í	Ĺ	
Existence of comorbidities	Yes	50	132	0.7 (0.5–1.1)	1.3 (0.53–3.1)	0.6
	No	338	1172	l í	l Í	
History of medication taking	Yes	63	157	0.7 (0.5–0.9)	0.76 (0.33-1.6)	0.4
	No	323	1145	l í	l	
Smoking history	Yes	21	45	0.6 (0.36–1.1)	1.1 (0.4–2.5)	0.9
	No	366	1528	l í		
COVID-19 symptoms on time of visit	Sore throat, headache, and sneezing	141	502	I	I	
	Cough, fever, and SOB	21	151	2.1 (1.23–3.31)	2.5 (1.22-4.7	0.01
	Fever, chest pain, and diarrhea	226	651	0.81 (0.63–1.2)	0.7 (0.45–1.1	0.5

**Table 3.** Factors associated with the SARS-COV-2 positivity among study participants (N=1692).

AOR; adjusted odds ratio: CI: confidence interval; COVID-19: coronavirus disease 2019; SOB: shortness of breathing; COR: crude odds ratio. Hosmer–Lemeshow  $\chi^2(8)$  = 4.15, Prob >  $\chi^2$  = 0.8435.

\*Significant factors (p < 0.05).

for other developing countries. It could be also the study conducted with a larger sample size inclusion that would make the study representative. As a limitation, false-negativity of RT-PCR could underestimate the positive SARS-COV-2 test. In addition, due to resource constraints, some other laboratory investigations and radiologic investigations were not done for the study participants.

### **Conclusion and recommendations**

The SARS-COV-2 tested positive among 388 (22.9%) acute respiratory infected people. The majority (21.6%) of SARS-COV-2 positive patients presented with lower respiratory tract infections. A divorced/separated, age, presenting with symptoms, such as cough, fever, and difficulty of breathing, had statistically significant associations with of SARS-COV-2 positive test. In resource-limited setups, where a shortage of testing equipment is frequent, these findings could contribute to boosting targeted symptom-oriented screening schemes. Moreover, strengthening the protection and control strategies for older adults is of priority to contain the spread of the virus.

### Acknowledgements

The authors thank their study participants, study facilities staff, data collectors, and supervisors for their commitment throughout the data collection process.

#### Author contributions

A.B. was involved in conception and design of the research idea, analysis, and writing up of the paper. G.M.A., Y.D., A.M., and M.B. participated in the manuscript drafting and editing. Finally, all authors read and approved the final manuscript.

#### **Declaration of conflicting interests**

The author(s) declared no potential conflicts of interest with respect to the research, authorship, and/or publication of this article.

### Funding

The author(s) received no financial support for the research, authorship, and/or publication of this article.

### **Ethical approval**

Ethical approval for this study was obtained from Haramaya University College of Health and Medical Sciences Institutional Health Research Ethical Review Committee (IHRERC) (Ref. no. IHRERC/019/2021, with the approval date of 10 February 2021).

### Informed consent

Written informed consent was obtained from legally authorized representatives before the study.

#### Data availability

The data can be accessed from the corresponding author upon request.

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