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Systematic Review / Meta-analysis

Global prevalence and determinants of mortality among patients with COVID-19: A systematic review and meta-analysis



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ARTICLE INFO	A B S T R A C T		
Keywords: Coronavirus COVID-19 Severe acute respiratory syndrome Mortality	 Background: The challenge of COVID-19 is very high globally due to a lack of proven treatment and the complexity of its transmission. The prevalence of in-hospital mortality among patients with COVID-19 was very high which ranged from 1 to 52% of hospital admission. The prevalence of mortality among intensive care patients with COVID-19 was very high which ranged from 6% to 86% of admitted patients. Methods: A three-stage search strategy was conducted on PubMed/Medline; Science direct Cochrane Library. The Heterogeneity among the included studies was checked with forest plot, χ2 test, 12 test, and the p-values. Publication bias was checked with a funnel plot and the objective diagnostic test was conducted with Egger's correlation, Begg's regression tests. Result: The Meta-Analysis revealed that the pooled prevalence of in-hospital mortality in patients with COVID-19 was strongly related to different factors. Patients with Acute respiratory distress syndrome were eight times more likely to die as compared to those who didn't have, RR = 7.99(95% CI: 4.9 to 13). Conclusion: The review revealed that more than fifteen percent of patients admitted to the hospital with coronavirus died. This presages the health care stakeholders to manage morbidity and mortality among patients with coronavirus through the mobilization of adequate resources and skilled health care providers. Registration: This systematic review and meta-analysis was registered in research registry with UIN of reviewregistry1093. 		

1. Introduction

Severe acute respiratory syndrome Coronavirus 2 (SARS-CoV-2) that cause Coronavirus disease 2019 (COVID-19) mainly affects the respiratory, gastrointestinal, liver, and central nervous system of humans, livestock, Bats, mice, and other wild animal [1-4]. The infection mainly affects the respiratory system and present with fever, dry cough, and difficulty of breathing, and lately, the patient may die due to pneumonia and acute respiratory distress syndrome [5-12].

The SARS-CoV-2 novel coronavirus was identified in Wuhan, Hubei province of China in December 2019 by the Chinese Center for Disease and Prevention from the throat swab of a patient and the virus is named severe acute respiratory distress CoV-2 by WHO which causes Coronaviruses disease 2019 (COVID-19) [13,14]. The clinical manifestation of the current coronavirus infection is similar to the one that occurred in

China in 2002 by the name severe acute respiratory distress syndrome [15–19].

Approximately, 5 million confirmed cases and more than 300 thousand deaths were reported by the World Health Organization (WHO) as of May 22, 2020 [20]. The American region accounted for the highest number of cases and deaths which was 2.2 million and 132 thousand respectively. The European region accounted for the second-highest confirmed cases and death which was approximately 2 million confirmed cases and 170 thousand deaths. The total number of confirmed cases and death in the Eastern Mediterranean region accounted for approximately 400 thousand and 11 thousand respectively [20].

Though the COVID-19 pandemic has emerged in the Western Pacific region, China, Wuhan city, the number of infected cases, and deaths was the lowest as compared to the American and European regions. The

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number of laboratory-confirmed cases and deaths in the African region was the lowest for the last couple of months but the rate of spreading in this region is increasing at an alarming rate and is expected to be very high in the next couple of months if it continues as this rate [20–22].

The COVID-19 report in Ethiopia was very small which is 2500 confirmed cases and 27 deaths but there were many cases in short periods which is more than 150 cases per day. It is estimated that the number even may be very high because the diagnosis is limited only in the capital [23,24].

The challenge of COVID-19 is very high globally due to a lack of proven treatment and the complexity of its transmission [13,25–31]. However, it will be more catastrophic for low and middle-income countries because of very poor health care system, high illiteracy and low awareness of the disease and its prevention, lack of skilled health personnel, scarce Intensive Care Unit, a limited number of mechanical ventilators, and prevalence of co-morbidities/infection along with malnutrition.

The severity of the disease is depending on several factors. Studies showed that patients with co-morbidities including (Asthma, COPD, Tuberculosis, Pneumonia, Acute respiratory distress syndrome (ARDS), Diabetes mellitus, hypertension, renal disease, hepatic disease, and cardiac disease), history of smoking, and history of substance use, male gender and age greater than 60 years were more likely to die or develop undesirable outcomes [6,28,31–37].

The outcomes of patients with coronavirus infection are very variable. Studies revealed that in-hospital mortality of patients with COVID-19 was very high which varied from 1% to 52% of hospitalized patients [25,27–29,36,38–41]. Studies also showed that the rate of ICU admission among coronavirus infected patients was higher which ranged from 3% to 100% of confirmed cases [16,27,29,31,42–46]. Studies also showed that the prevalence of mortality among intensive care patients with coronavirus infection was very high which ranged from 6% to 86% of admitted patients [16,27,29,31,42–46].

The global prevalence of mortality among hospitalized patients, number of cases requiring ICU care, number of cases need a mechanical ventilator, the prevalence of death in ICU, length of stay and independent risk factors for in-hospital mortality are very important variables to be determined to reduce patient mortality and morbidity through varies strategies including but not limited to increasing number of ICU beds, mechanical ventilator, skilled professionals, integrated monitors and reducing possible risk factors. Therefore, this systematic review aimed to provide global evidence on the prevalence and risk factors including age, gender, comorbidity, and substance use on hospital mortality among hospitalized patients with COVID-19.

2. Methods

2.1. Protocol and registration

The systematic review and meta-analysis were conducted based on the Preferred Reporting Items for Systematic and Meta-analysis (PRISMA) protocols [47]. This systematic review and meta-analysis was registered in research registry with UIN of reviewregistry1093 and available at: https://www.researchregistry.com/browse-the-re gistry#registryofsystematicreviewsmeta-analyses/

2.2. Eligibility criteria

2.2.1. Types of studies

All observational (case series, cross-sectional, cohort, and casecontrol) studies reporting the prevalence of mortality and its determinants among hospitalized patients with coronavirus disease (COVID-19) will be incorporated.

2.2.2. Types of participants

All participants with confirmed Coronaviruses admitted to the

hospital for any kind of care will be included.

2.2.3. Outcomes of interest

The primary outcomes of interest were the global prevalence of mortality, morbidity, and complications among patients with Coronaviruses worldwide. Prevalence of ICU mortality Lengths of hospital stay and the number of days on a mechanical ventilator were secondary outcomes.

2.2.4. Context

This systematic review incorporated studies conducted worldwide to assess the prevalence of mortality and associated risk factors among hospitalized patients with COVID-19.

2.2.5. Inclusion criteria

The review included all cross-sectional studies and a single cohort conducted among adult patients hospitalized with COVID-19 to assess mortality in patients with coronavirus infection.

2.2.6. Exclusion criteria

Studies that didn't report the prevalence of mortality among hospitalized patients with COVID-19, articles that didn't report full information for data extraction, articles with different outcomes of interest, studies with a methodological score less than fifty percent, studies with randomized controlled trials, case reports, and reviews were excluded.

2.3. Search strategy

The search strategy was intended to explore all available published and unpublished studies among COVID-19 patients admitted to the hospital from December 2019 to May 2020 without language restrictions. A three steps search strategy was employed in this review. An initial search on PubMed/Medline, Science Direct, and Cochrane Library was carried out followed by an analysis of the text words contained in Title/Abstract and indexed terms. A second search was undertaken by combining free text words and indexed terms with Boolean operators. The third search was conducted with the reference lists of all identified reports and articles for additional studies. Finally, an additional and grey literature search was conducted on Google scholars up to ten pages. The PubMed was searched with the following search terms using PICO strategy as: (coronavirus) or (coronavirus disease 2019)) or (SARS-CoV-2)) or (COVID-19)) and (mortality)) or (fatality)) or (morbidity)) or (comorbidity)) or (complications)) and (risk factors)) or (determinants)) and (prevalence)) and (global). The final search results were shown with the Prisma flow diagram (Fig. 1).

2.4. Data extraction

The data from each study were extracted with two independent authors with a customized format excel sheet. The disagreements between the two independent authors were resolved by the other two authors. The extracted data included: Author names, country, date of publication, sample size, events mortality, need of mechanical ventilator, the number of days on a mechanical ventilator, presence of co-morbidities, and complication. Finally, the data were then imported for analysis in R software version 3.6.1 and STATA 14.

2.5. Assessment of methodological quality

Articles identified for retrieval were assessed by two independent Authors for methodological quality before inclusion in the review using a standardized critical appraisal Tool adapted from the Joanna Briggs Institute [48] (Supplemental Table 1) and a critical appraisal tool for systematic reviews that include randomized or non-randomized studies of healthcare interventions(AMSTAR2) [49]. The disagreements between the Authors appraising the articles were resolved through

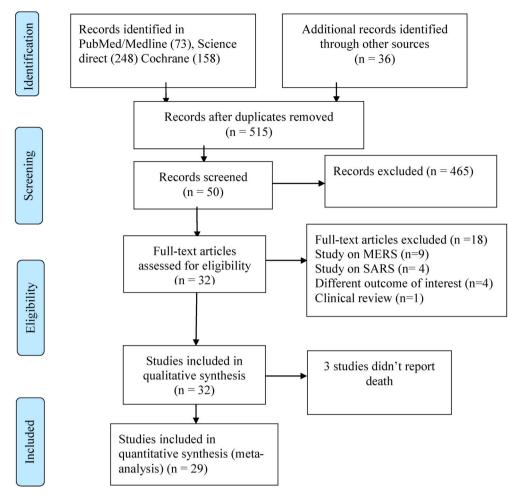


Fig. 1. Prisma flow chart.

discussion with the other two Authors. Articles with average scores greater than fifty percent were included for data extraction.

2.6. Data analysis

Data analysis was carried out in R statistical software version 3.6.1 and STATA 14. The pooled global prevalence of mortality, comorbidity, and complication among hospitalized patients with COVID-19 was determined with a random effect model as there was substantial heterogeneity. The Heterogeneity among the included studies was checked with forest plot, χ^2 test, I² test, and the p-values. Substantial heterogeneity among the included studies was investigated with subgroup analysis.

Publication bias was checked with a funnel plot and the objective diagnostic test was conducted with Egger's correlation, Begg's regression tests, and Trim and fill method. Furthermore, moderator analysis was carried out to identify the independent predictors of mortality among corona cases. The results were presented based on the Preferred Reporting Items for Systemic Reviews and Meta-Analysis (PRISMA) [47].

3. Results

3.1. Selection of studies

A total of 515 articles were identified from different databases and 50 articles were selected for evaluation after the successive screening. Thirty-two Articles with 23082 participants were included and the rest

3.2. Description of included studies

were excluded with reasons (Fig. 1).

Thirty-two Articles with 23082 participants were included in the review while twenty-nine studies were included in the Meta-Analysis for the prevalence of mortality. Studies with the prevalence of mortality and/or prevalence of comorbidity and prevalence of complications among hospitalized patients with COVID-19 were included and the characteristics of each included studies were described in (Table 1) and the rest were excluded with reasons [15,17,19,27,42,50–52,54–57].

The included studies were published from December 16, 2019, to April 7, 2020, with sample sizes, ranged from 18 to 8910. The mean (SD) ages of the included studies varied from 38 to 59.7 years. The majority of the included studies, Twenty-three from thirty-two were conducted in China [12,13,16,31,37,58–75]. One study was conducted among 169 hospitals of Asia, Europe, and North America [76] while six studies were conducted in America [44,77–81] and the remaining two were conducted in Singapore and France [82,83].

Twenty-nine of the included studies reported the prevalence of mortality among hospitalized patients with COVID-19 while three of the included studies didn't report the prevalence of mortality among COVID-19 patients in the hospital. The prevalence of mortality in hospitalized patients with COVID-19 from the included studies varied from 1% to 52%.

Twenty-six studies with 22216 participants reported the prevalence of comorbidity including hypertension, diabetes mellitus, cardiovascular disease, COPD, obesity as the major comorbidity among patients

Table 1

Description of included studies.

Author	Year	Study period	Sample	Country	Follow up(days)	Prevalence of mortality(95% CI)
Young et al. [83]	2020	Jan 23 to Feb 3, 2020	18	Singapore	10	-
Arentz et al. [44]	2020	Feb 20 to March 5, 2020	21	USA	15	52[30,74]
Bhatraju et al. [77]	2020	March 23, 2020	24	USA	1	50[29,71]
Huang et al. [62]	2020	Dec 16, 2019, to Jan 2, 2020	41	China	17	32[18,48]
Yang et al. [31]	2020	Dec 24, 2019 to Jan 26, 2020	52	China	31	62[47,75]
Xu et al.(12)	2020	Jan 10 to 26, 2020	62	China	16	-
Tang et al. [67]	2020	Dec 24, 2019 to Feb 7, 2020	73	China	41	36[25,48]
Zhao et al. [73]	2020	Jan 21 to Feb 8, 2020	77	China	17	6[2,15]
Liu et al. [65]	2020	Dec 30, 2019 to Jan 15, 2020	78	China	16	_
Wu et al. [70]	2020	Jan 22 to Feb 14, 2020	80	China	22	43[33,55]
Chen et al.(13)	2020	Jan 1 to Jan 20, 2020	99	China	19	11[6,19]
Cao et al. [58]	2020	Jan 3 to Feb 1, 2020	102	China	29	17[10,25]
Wu et al. [69]	2020	Dec 25, 2019 to Jan 26, 2020	201	China	32	3[0,9]
Bialek et al. [78]	2020	March 18, 2020	121	USA	1	45[36,55]
Simonnet et al. [82]	2020	Feb 27 to April 5, 2020	124	France	7	15[9,22]
Wang et al. [68]	2020	Jan 1 to 28, 2020	138	China	27	4[2,9]
McMichael et al. [79]	2020	Feb 28 to March 18, 2020	167	USA	30	21[15,28]
Guo et al. [75]	2020	Jan 23 to Feb 23, 2020	187	China	30	23[17,30]
Zhou et al. [74]	2020	January 2020	191	China	1	28[22,35]
Guan et al. [61]	2020	January 2019	1099	China	1	3[2,4]
Pan et al. [66]	2020	Jan 18 to Feb 28, 2020	204	China	15	18[13,24]
Chen et al.(16)	2020	Jan 20 to Feb 25, 2020	249	China	35	1[0,3]
Zhang et al. [72]	2020	Jan29 -Feb 12, 2020	258	China	13	6[3,9]
Zeng et al. [71]	2020	Jan 11 to Feb 29, 2020	338	China	49	1[0,3]
Li et al. [64]	2020	Feb 3 to March 3, 2020	548	China	30	16[13,20]
Cheng et al. [59]	2020	Feb 20, 2020	701	China	1	16[13,19]
Cheng et al. [59]	2020	Jan 28 to February 2020	710	china	3	13[10,15]
Jin et al. [63]	2020	Jan29 -Febr15,2020	1019	China	18	4[3,5]
Guan et al. [60]	2020	Dec 11, 2019 to Jan 31, 2020	1590	China	51	1[1,2]
Petrilli et al. [80]	2020	March 1 to April 7, 2020	1999	USA	34	15[13,16]
Richardson et al. [81]	2020	March 1 to April 4, 2020	3700	USA	31	15[13,17]
Mehra et al. [76]	2020	Dec 2019 to March 28, 2020	8910	Multi- Country	-	6[5,6]

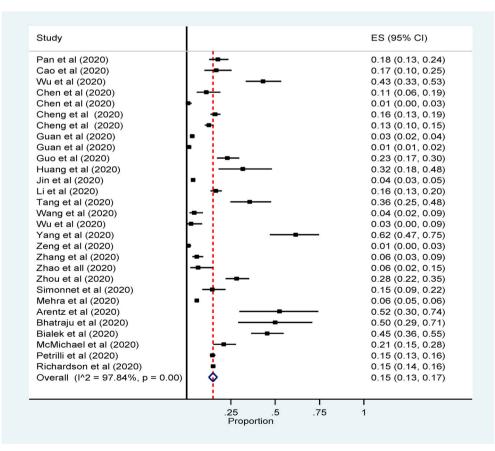


Fig. 2. Forest plot for the prevalence of mortality among hospitalized patients with COVID: The midpoint of each line illustrates the prevalence; the horizontal line indicates the confidence interval, and the diamond shows the pooled prevalence.

hospitalized with COVID-19 while sixteen studies with 8280 participants reporting the prevalence of complications including ARDS, acute kidney injury, liver dysfunction, sepsis and arrhythmia as the major complications. The prevalence of mortality among hospitalized patients with the newly emerging coronavirus was very high which varied from 3 to 100% of the intensive care unit admissions.

4. Meta-analysis

4.1. Global prevalence of mortality

Twenty-nine studies reported prevalence and associated risk factors of mortality among hospitalized patients with coronavirus. The pooled prevalence of mortality was 15% (95% CI: 13 to 17, 29 studies, and 22924 participants) (Fig. 2). The subgroup analysis by country revealed that the mortality of patients with COVID-19 admitted in the hospital was the highest in USA followed by France 25% (95% CI: 19 to 30, 6 studies, 6032 participants) and 15% (95% CI: 9 to 22, one study, 124 participants) respectively (Supplemental Fig. 1).

4.2. Prevalence of ICU mortality

The prevalence of mortality among hospitalized patients with COVID-19 in the Intensive Care Unit was 29% (95% CI: 20 to 38, 16 studies, 2227 participants) (Fig. 3).

4.3. Prevalence of comorbidity

The Meta-Analysis revealed that the prevalence of comorbidity among hospitalized patients with COVID-19 was 48% (95% confidence interval (CI):35 to 62, 26 studies, 22528 participants) (Fig. 4).

The subgroup analysis by the commonest comorbidities showed that diabetes mellitus was 48% (95% confidence interval (CI): 29 to 67, 3 studies, 117 participants) followed by hypertension and cardiovascular diseases 15% (95% confidence interval (CI):11 to 20, 14 studies, 8970 participants and 15% (95% confidence interval (CI): 8 to 21, 6 studies, 2505 participants) respectively (Fig. 5).

4.4. Prevalence of complications

Plenty of complications were mentioned in included studies including Acute Respiratory distress syndrome, acute kidney injury, sepsis, liver dysfunction, arrhythmia was the amongst reported as the common complications. The overall pooled prevalence of complications among hospitalized with COVID-19 was 36% (95% confidence interval (CI): 28 to 43, 16 studies, 8280 participants (Fig. 6).

The Meta-Analysis revealed that sepsis was the most prevalent complication, 55% (95% confidence interval (CI):49 to 61, 2 studies, 246 participants) followed by Liver dysfunction and ARDS, 32% (95% confidence interval (CI): 23 to 44, one study, 77 participants and 26% (95% confidence interval (CI): 9 studies, 2524 participants) respectively (Fig. 7) (see Fig. 8).

4.5. Determinants of mortality

The prevalence of mortality of patients with COVID-19 is greatly affected by several factors including but not limited to the presence of co-morbidities, history of smoking, male gender, older age groups, ICU admission, nosocomial infection, and others. Moderator analysis was conducted to identify the independent predictor of mortality among

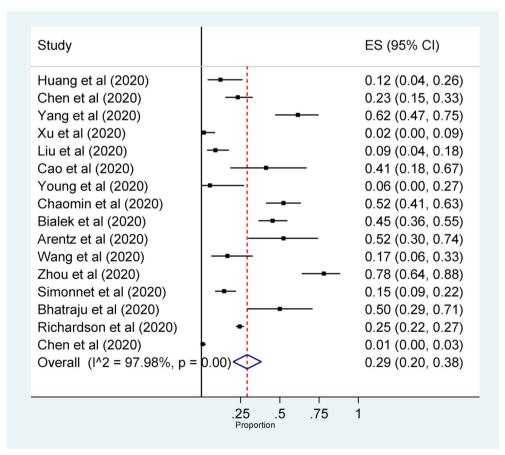


Fig. 3. Forest plot for the prevalence of mortality among patients with Coronaviruses admitted in Intensive Care Unit: The midpoint of each line illustrates the prevalence; the horizontal line indicates the confidence interval, and the diamond shows the pooled prevalence.

Study	ES (95% CI)
Pan et al (2020)	0.50 (0.43, 0.57)
Cao et al (2020)	← 0.17 (0.10, 0.25)
Chen et al (2020)	0.36 (0.30, 0.42)
Chen et al (2020)	0.51 (0.40, 0.61)
Cheng et al (2020)	- 0.42 (0.39, 0.46)
Cheng et al (2020)	■ 0.13 (0.10, 0.15)
Guan et al (2020)	• 0.32 (0.30, 0.35)
Guan et al (2020)	
Guo et al (2020)	— 0.89 (0.83, 0.93)
Huang et al (2020)	0.32 (0.18, 0.48)
Jin et al (2020)	0.47 (0.31, 0.62)
Li et al (2020)	- 0.64 (0.60, 0.68)
Petrilli et al (2020)	 0.57 (0.55, 0.59)
Richardson et al (2020)	 0.91 (0.90, 0.91)
Simonnet et al (2020)	- 0.98 (0.94, 1.00)
Tang et al (2020)	0.36 (0.25, 0.48)
Wang et al (2020)	0.46 (0.38, 0.55)
Wu et al (2020)	0.47 (0.36, 0.59)
Xu et al (2020)	—— 0.32 (0.21, 0.45)
Wu et al (2020)	
Yang et al (2020)	0.62 (0.47, 0.75)
Zhang et al (2020)	—— 0.71 (0.65, 0.76)
Zhao et all (2020)	0.31 (0.21, 0.43)
Zhou et al (2020)	0.48 (0.40, 0.55)
Overall (I ² = 99.71%, p = 0.	00) 0.48 (0.35, 0.62)
	.25 .5 .75 1
	Proportion

Fig. 4. Forest plot for the prevalence of comorbidity among hospitalized patients with COVID-19: The midpoint of each line illustrates the prevalence; the horizontal line indicates the confidence interval, and the diamond shows the pooled prevalence.

Coronavirus patients.

Mortality among hospitalized patients with COVID-19 was two times more likely in patients with any co-morbidity as compared to those who didn't have co-morbidities, RR = 2.20(95% CI:1.75 to 2.77). Acute respiratory distress syndrome was the most likely independent predictor of in-hospital mortality in patients with COVID-19. Those Patients with Acute respiratory distress syndrome were eight times more likely to die as compared to patients with no acute respiratory distress syndrome, RR = 7.99(95% CI: 4.9 to 13). Mortality among hospitalized patients with COVID-19 was two times more likely in a patient with a history of smoking as compared to nonsmokers, RR = 2.19(95% CI: 1.48 to 3.22). Mortality of patients among males with COVID-19 increased the risk by thirty-seven percent when compared to the female counterparts, RR = 1.63(95% CI: 1.33 to 1.99(Supplemental Fig. 2).

4.6. Sensitivity analysis and publication bias

Sensitivity analysis was conducted to identify the most influential study on the pooled summary effect and we didn't find a significant influencing summary effect. The funnel plot didn't show significant publication bias. Besides, egger's regression and Begg's correlation rank correlation failed to show significant difference (p < 0.05).

5. Discussion

The total laboratory confirmed infected cases and the death of patients with SARS-CoV-2 virus is unpredictably high as compared to the previous two outbreaks [15,17,18,27,29,54,56,84].

This Meta-Analysis revealed that the pooled global prevalence of mortality among hospitalized patients with COVID-19 was very high, 15% (95% CI: 13 to 17). This finding is in line with individual studies conducted among Coronavirus confirmed cases during the first outbreak in 2002, China [15,17,18,27,29,54,56,84]. The possible explanation for a high number of deaths among hospitalized patients with COVID-19 may be explained in terms of disease severity, presence of co-morbidities, inadequate laboratory investigation, complications, and some others. However, the finding of this review is higher than other systematic reviews conducted among patients with COVID-19 and this discrepancy might be due to the inclusion of plenty of studies [37, 85–87].

The subgroup analysis showed that the prevalence of in-hospital mortality among COVID-19 patients was higher in the USA followed by France and China. This higher prevalence of in-hospital mortality in the USA compared to China might be due to the inclusion of a small number of studies from America as compared to china.

The Meta-Analysis revealed that the overall prevalence of comorbidities among patients hospitalized with COVID-19 was very high as compared to other systematic reviews and Meta-Analysis conducted to investigate the prevalence of comorbidity among COVID-19 patients [37,86–88]. This dissimilarity might be due to the inclusion of plenty of studies in this systematic review. The Meta-analysis also revealed that the prevalence of hypertension, diabetes mellitus, cardiovascular disease, and respiratory disease which is comparable to other Meta-Analysis [37,86–88].

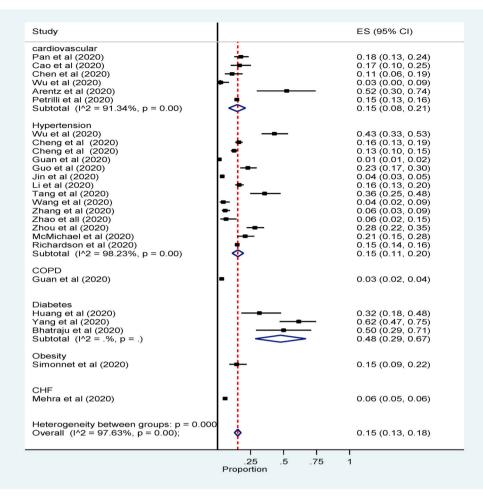


Fig. 5. Forest plot for subgroup analysis of the prevalence of comorbidity among patients in hospital by major comorbidities. The midpoint of each line illustrates the prevalence; the horizontal line indicates the confidence interval, and the diamond shows the pooled prevalence.

The systematic review showed that the prevalence of mortality among ICU admitted cases with Coronavirus was very high which is in line with systematic reviews conducted among hospitalized patients with COVID-19 [37,85–87].

The overall pooled prevalence of complications among hospitalized with COVID-19 was 36% (95% confidence interval (CI): 28 to 43, 16 studies. The subgroup analysis showed that sepsis was the highest prevalent followed by liver dysfunction and ARDS, unlike other systematic reviews where ARDS was the most prevalent complication. This discrepancy might be due to the number of included studies and sample size contribution.

The prevalence of mortality among patients with co-morbidities, history of smoking, gender, advanced age, and others was very high. Acute respiratory distress syndrome was the most likely independent predictor of in-hospital mortality.

5.1. Quality of evidence

The systematic review and meta-analysis included plenty of studies with adequate sample size. The methodological quality of included studies was moderate to high quality as depicted with Joanna Briggs Institute assessment tool for meta-analysis of cross-sectional studies. However, substantial heterogeneity associated with dissimilarities of included studies in sample size, study setting, study design may limit generalization of this finding to the global community.

5.2. Limitation of the review

The review incorporated plenty of studies with a large number of participants but some of the included studies in this review didn't report risk factors, comorbidities, and complications for factor analysis. The included studies were conducted in a different setting with different sample sizes, population, and study design which caused substantial heterogeneity. Besides, there were a limited number of studies in some countries and it would be difficult to provide conclusive evidence with results pooled from fewer studies.

5.3. Implication for practice

The current COVID-19 pandemic is spreading swiftly around the world due to uncertain mode of transmission, lack of proven treatment and vaccination, incompliance of people with preventive measures. Body of evidence revealed that the global prevalence of mortality, morbidity, and complications among hospitalized patients with COVID-19 was very high. The magnitude of this problem will be worse than this particularly in low and middle-income countries with weak health care systems and lack of well-equipped hospitals, ICU, skilled health care providers, quarantine centers, and laboratory centers. Therefore, global unity is highly required than ever to combat this deadly pandemic from the globe.

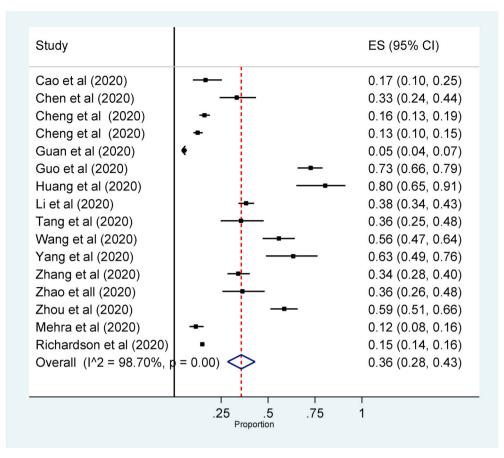


Fig. 6. Forest plot for the prevalence of complication among hospitalized patients with COVID-19: The midpoint of each line illustrates the prevalence; the horizontal line indicates the confidence interval, and the diamond shows the pooled prevalence.

5.4. The implication for further research

The Meta-analysis revealed that the global prevalence of mortality, morbidity, and complications among hospitalized patients with COVID-19 was very high and the major independent predictors were identified. However, the included studies were too heterogeneous, and crosssectional studies also don't show temporal relationship outcomes and their determinants. Therefore, further observational and randomized controlled trials are in demand for specific comorbidity by stratifying the possible independent predictors.

6. Conclusion

The systematic review and Meta-Analysis revealed that the prevalence of mortality among hospitalized patients with COVID-19 was very high. The systematic review also showed that approximately fifty percent of patients hospitalized COVID-19 had one or more comorbidities and among which hypertension, cardiovascular disease, diabetes, and obesity were the most prevalent comorbidities. The prevalence of complications during hospitalization in patients with COVID-19 was as high as thirty-six percent. Sepsis, Acute liver failure, and ARDS were the most prevalent complications during hospitalization. Therefore, the body of evidence warns the health care stakeholders to give attention to hospitalized patients with COVID-19 through accessing mechanical ventilators, integrated patient monitors, skilled ICU staffs, creation of awareness about infection prevention and more others.

Ethical approval

Ethical approval was obtained from Dilla University.

Sources of funding

The authors didn't receive any sources of funding for this systematic review and meta-analysis.

Author contribution

Semagn Mekonnen Abate and Yigrem Ali Chekol conceived the idea and design of the project. Semagn Mekonnen Abate, Yigrem Ali Chekol, Bivash Basu, and Bahiru Mantefardo involved in searching strategy, data extraction, quality assessment, analysis and manuscript preparation. All authors approve the final manuscript.

Registration of research studies

1. Name of the registry: Prospero's international prospective register of systematic reviews.

2. Unique Identifying number or registration ID: CRD42020187721.

3. Hyperlink to your specific registration (must be publicly accessible and will be checked): https://www.crd.york.ac.uk/prospero/#recordDetails.

Guarantor

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Study		ES (95% CI)
AKI	_	0.05 (0.04, 0.07)
Cheng et al (2020)		(, , ,
Cheng et al (2020)		0.03 (0.02, 0.05)
Richardson et al (2020)	\sim	0.14 (0.13, 0.15)
Subtotal (I^2 = .%, p = .)	~	0.07 (0.00, 0.15)
ARDS		
Cao et al (2020)		0.44 (0.34, 0.54)
Chen et al (2020)		0.33 (0.24, 0.44)
Guan et al (2020)		0.03 (0.02, 0.05)
Guo et al (2020)	_ _	0.25 (0.19, 0.31)
luang et al (2020)		0.29 (0.16, 0.46)
i et al (2020)		0.38 (0.34, 0.43)
Nang et al (2020)		0.20 (0.13, 0.27)
rang et al (2020)		0.17 (0.08, 0.30)
Zhang et al (2020)	- t -	0.24 (0.19, 0.30)
Subtotal (I^2 = 98.25%, p = 0.00)	$\langle \rangle$	0.26 (0.13, 0.38)
Arrythmia		
Mehra et al (2020)	+	0.12 (0.08, 0.16)
iver dysfunction Zhao et all (2020)	<u> </u>	0.32 (0.22, 0.44)
		0.32 (0.22, 0.44)
sepsis		0.45 (0.04.0.57)
Tang et al (2020)		0.45 (0.34, 0.57)
Zhou et al (2020)	-	0.59 (0.51, 0.66)
Subtotal (I^2 = .%, p = .)	\sim	0.55 (0.49, 0.61)
leterogeneity between groups: p = 0.00)	
Overall (I^2 = 98.33%, p = 0.00);	\diamond	0.24 (0.19, 0.29)
	.25 .5 .7	

Fig. 7. Forest plot for subgroup analysis of the prevalence of complication among patients in hospital by major complication.: The midpoint of each line illustrates the prevalence; the horizontal line indicates the confidence interval, and the diamond shows the pooled prevalence.

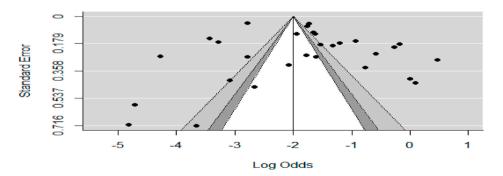


Fig. 8. Funnel plot to assess publication bias. The vertical line indicates the effect size whereas the diagonal line indicates precision of individual studies with 95% confidence interval.

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Consent

Consent was not applicable as we had collected data from previously published articles.

Declaration of competing interest

The authors declared that there is no conflict of interest.

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Abbreviations

ARDS	Acute Respiratory Distress Syndrome
CI	confidence interval
COPD	Chronic Obstructive Pulmonary Disease
COVID-19	Coronavirus Disease 2019
ICU	Intensive Care Unit
PRISMA	Preferred Reporting Items for Systematic Review and Meta-
	Analysis
RR	Relative Risk
SARS-CoV	7-2 Severe Acute Respiratory Distress 2
WHO	World Health Organization

Appendix A. Supplementary data

Supplementary data to this article can be found online at https://doi.org/10.1016/j.amsu.2021.102204.

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