



COVID-19 Vaccine Acceptance Rate and Its Factors among Healthcare Students: A Systematic Review with Meta-Analysis

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Abstract: Healthcare students are clinicians-in-training likely to come into contact with COVID-19 as much as other frontline healthcare professionals. It is therefore necessary to prioritize vaccinations for this group. We conducted a global systematic assessment of COVID-19 vaccine acceptance rates and related factors among healthcare students using the PubMed, Scopus, and Web of Science databases and keyword searches in March of 2022. We found 1779 articles with relevant information and 31 articles that matched our inclusion criteria. We performed a random-effects meta-analysis and quality assessment using the eight-item Joanna Briggs Institute Critical Appraisal test for crosssectional studies. A total of 30,272 individuals from 16 countries were studied. Most of the studies were carried out in the U.S. (n = 6), China (n = 5), Poland (n = 5), India (n = 2), Italy (n = 2), and Israel (n = 2). The prevalence of the COVID-19 vaccine acceptance rate was 68.8% (95% confidence interval [CI]: 60.8–76.3, $I^2 = 100\%$), and the prevalence of the vaccine hesitancy rate was 25.8% (95%) CI: 18.5–33.8, $I^2 = 99\%$). In country-specific analyses, Romania showed the highest acceptance rate (88.0%, 95% CI: 44.5–100%), while Iraq showed the lowest acceptance rate (66.2%, 95% CI: 35.5–90.8%). In time-trend analyses, we found that acceptance rates among healthcare students decreased over time. Students concerned about potentially serious side effects of the vaccine were less willing to accept the vaccine. National and international interventions should be adopted to reduce COVID-19 vaccination hesitancy rates among these important frontline workers.

Keywords: vaccine hesitancy; vaccine acceptance; COVID-19; frontline workers; healthcare students; meta-analysis; SARS-CoV-2; vaccine

1. Introduction

The ongoing COVID-19 pandemic has turned into a global challenge due to its dramatically contagious nature. The virus has led to more than 4.6 million deaths globally between February of 2020 and August of 2021 [1]. Vaccinations are one of the most cost-effective and long-lasting measures in helping to control such a public health disaster [2]. Vaccination rates directly impact herd immunity. Studies reported that when a population's acquired immunity reaches 67%, the prevalence of COVID-19 infections will continue to decline [3]. Multiple biological and chemotherapeutic measures (i.e., plasma therapy, hydroxychloroquine, remdesivir, and tocilizumab) have been used to treat COVID-19 patients, but their curative effects have generally not been recommended or proven for patient treatment [4].



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Copyright: © 2022 by the authors. Licensee MDPI, Basel, Switzerland. This article is an open access article distributed under the terms and conditions of the Creative Commons Attribution (CC BY) license (https:// creativecommons.org/licenses/by/ 4.0/). Many countries have encountered ongoing surges in COVID-19 cases due to relaxed precautionary measures like lockdowns, social distancing, hand washing, and personal hygiene practices [5]. Vaccinations remain the most important tool in combatting the pandemic.

Scientific authorities have undertaken over 300 vaccine invention projects, among which approximately 40 are in the clinical trial stage and a few are available to the general population [6]. Vaccine development was accelerated when two of these vaccines were granted Emergency Use Authorization (EUA) in December of 2020—a process that generally takes several years or more [7]. As of July of 2021, about three billion doses of COVID-19 vaccines have been administered worldwide. More than 11.48 billion doses of vaccines have been approved in at least one country. Clinical trials have shown that some of these have significant promise for real-world use, while others are customized to the unique needs of certain groups (i.e., older adults). Vaccination effectiveness generally varies between 50% and 77%. There is evidence that many vaccines may help minimize the likelihood of severe illness and asymptomatic disease, thus limiting the spread of SARS-CoV-2 [8,9].

Any vaccination program's success depends upon people's willingness to be vaccinated, the demand for vaccines, and positive attitudes toward vaccines [10,11]. Therefore, vaccine hesitancy limits the success of a vaccination program's success; such hesitancy is defined by the indecision, reluctance, or refusal of vaccination [12,13]. The World Health Organization (WHO) has stated that vaccine hesitancy is a serious threat to public health [14]. For example, the 2018 measles outbreak in New York City revealed that vaccine hesitancy resulted in continuous transmission [15]. Vaccine hesitancy has been linked to numerous factors, such as distrust in the government, fear of side effects, and religious convictions [16].

During the COVID-19 pandemic, there has been much discussion about vaccines, especially among healthcare workers (HCWs) and students [17,18]. Healthcare students (HCSs) in medical, dental, nursing, and related programs are future clinical caregivers and important populations who need to be vaccinated against COVID-19. Several governments have chosen to incorporate medical students as volunteers who assist with coronavirus treatment while finishing their residency training [19]. HCSs are likely come into contact with COVID-19-infected patients during training sessions and clinical practice [20,21]. To avoid further infection and increase vaccine acceptance rates, medical students must be taught about the benefits of vaccines as part of their training. Furthermore, their families and friends look to them as competent and trustworthy resources of information, which means their opinions and views have an influence on the general public's vaccine acceptance levels [22].

Numerous studies have investigated vaccine acceptance or hesitancy rates among HCSs. Some of these studies showed surprisingly high rates of vaccine hesitancy [17]. For example, one study in the U.S. found that nearly one-quarter of medical students were reluctant to be vaccinated, even after an approved COVID-19 vaccine was available [18]. Another study among nursing students in Greece, Albania, Cyprus, Spain, Italy, the Czech Republic, and Kosovo found that less than one-half were willing to be vaccinated [23]. In contrast, nearly 90% of medical students in India [24] and nursing students in China [2] were willing to get vaccinated.

We conducted a rapid systematic review with meta-analysis on HCSs' perception of being vaccinated in response to this growing body of literature on vaccination acceptance and hesitancy among HCSs and the seemingly disparate results. Comparable systematic reviews and meta-analyses have been conducted among general populations [10,25,26] and healthcare workers [27,28], but not HCSs. Our aim was to assess the acceptance and hesitancy of COVID-19 vaccination rates among HCSs globally and identify the factors predicting vaccine acceptance. We expected the findings to help understand the challenges associated with vaccine hesitancy among HCSs, as well as inform strategies for overcoming these challenges.

2. Materials and Methods

We followed the Cochrane Rapid Review guidelines to conduct a rapid systematic review with a streamlined but robust approach. The criteria were searches in English and peer-reviewed studies. Similar approaches have been used to provide time-sensitive information that informs decision-making surrounding COVID-19 immunization programs [29].

2.1. Search Strategy

We systematically searched three databases (PubMed, Web of Science, and Scopus) using the PRISMA checklist (http://www.prisma-statement.org/) on 5 March 2022. We utilized the following Medical Subject Heading (MeSH) terms as well as text words (tw) for COVID-19: "COVID-19", "SARS-CoV-2", "coronavirus", "novel coronavirus", "nCoV", "2019-ncov", "SARS-2", and "severe acute respiratory syndrome coronavirus 2". For vaccines, we used: "vaccines", "vaccination", "COVID-19 vaccines", "vaccina", "vaccine uptake", and "SARS-CoV-2 vaccine". For acceptance/hesitancy, we used: "vaccine hesitancy", "vaccine hesitance", "vaccine acceptance", "vaccine confidence", "vaccine safety", "vaccination attitudes", "vaccine rejection", and "vaccine willingness". We did not specify the population terms to avoid excluding potentially important and relevant articles. Additional articles were identified using the references and citation lists of articles and reviews found in the keyword searches via forward and backward citation tracking in Google Scholar.

2.2. Study Selection

All records were imported to 'Rayyan' (https://www.rayyan.ai/; accessed on 5 March 2022). This is a tool for intelligent systematic reviews. Duplicates were removed using this software. Irrelevant records were excluded through title and abstract screening. Next, the full texts of the remaining articles were screened (Figure 1). Discrepancies were resolved by discussion among the three reviewers (MMP, MB, and MZH) and, if required, consultation with other co-authors for reaching a consensus.

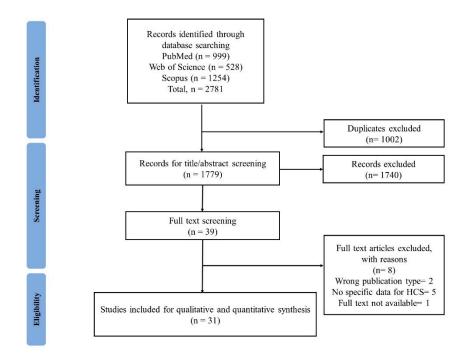


Figure 1. PRISMA flow diagram of the study selection process.

We had six inclusion criteria for the articles. These included: (1) survey studies among HCSs; (2) descriptive and observational studies among HCSs with cross-sectional, experimental, or longitudinal designs; (3) studies focused on evaluating COVID-19 vaccine

acceptance and/or hesitancy; (4) studies published in English with no restriction to country; (5) studies published since January of 2020; and (6) peer-reviewed scientific journal articles.

Six exclusion criteria were included. These were: (1) articles not aiming to evaluate COVID-19 vaccine acceptance or hesitancy; (2) study populations other than HCSs; (3) publication types other than peer-reviewed journal articles, such as literature reviews, systematic reviews, unpublished data, books, conference papers, editorials, commentaries, letters to the editor, and case reports; (4) studies with non-human subjects; (5) studies without available full-texts; and (6) studies other than in English.

2.3. Data Extraction

Data extraction was performed independently by three co-authors. The extracted data included: author-name; publication year; study country; study design; survey method and period; target population; sampling method; sample size; measurement scale of vaccine acceptance; statistical analysis; acceptance rate; hesitancy rate; factors associated with vaccine acceptance, hesitance, or refusal; and summary of results. These data are summarized in Table 1. After independent data extraction, any differences were resolved by consensus among the same three co-authors.

2.4. Assessment of Study Quality

Regarding quality assessment and evaluating the risk of bias, we adopted the Joanna Briggs Institute critical appraisal tools for analytical cross-sectional studies (Table S1) [30]. This allowed us to determine whether certain articles should be included or excluded, or if additional information was required. We used a checklist with eight questions on the study's methods and applicable data analysis for this purpose. The total score for each study was assessed by aggregating the individual scores and categorizing them into a high-or low-quality group following previous studies [31,32].

2.5. Data Analysis

Acceptance and hesitancy rates were pooled using random-effects models. The Higgin's and Thompsons's I^2 statistics determined the heterogeneity [33,34]. Funnel plots and the Egger's tests identified potential publication bias. We considered the survey year and country for subgroup analysis and conducted meta-regression analyses for four predictors: sex, residence, history of prior vaccinations, and concern about serious side effects. All analyses were performed using the 'meta' statistical packages in R software (version 4.2.1).

3. Results

3.1. Search Results

A total of 2781 articles were identified in preliminary searches across three databases including PubMed, Web of Science, and Scopus. Of these, 1002 articles were duplicates. After assessing their eligibility based on the title and abstract, 39 articles were eligible for full-text screening. Ultimately, 31 articles were included in the analyses (Figure 1).

3.2. Characteristics of Included Studies

The characteristics of the included articles are summarized in Table 1. Most used a cross-sectional design and collected data via telephone or online surveys. The majority also relied heavily on snowball sampling (i.e., via social media or email) and convenience sampling for recruitment. Studies were mostly conducted between March of 2020 and March of 2021.

The total number of healthcare students included in the studies was 30,272. Sample sizes ranged from 104 in Israel [35] to 6639 in one study across 22 countries [36]. Approximately 19,425 students (64% of total sample) were female. Most of the studies were conducted in the U.S. (n = 6), China (n = 5), Poland (n = 2), India (n = 2), Italy (n = 2), and Israel (n = 2). The largest share of HCSs were medical students, followed by nursing and dental students.

SL	Author	Study Country	Type of Healthcare Students	Study Design	Survey Method	Survey Period	Sampling Method	Sample Size, N	Gender, Female (%)	Vaccine Acceptance Rate (%)
1	Al Janabi et al. [37]	USA	Osteopathic medical	Cross-sectional	Online	October 2020	NR	197	57.9	45
2	Bălan et al. [38]	Romania	General Medicine, Dentistry, Pharmacy and Nursing and Midwifery	Cross-sectional	Online	12 January until 3 March 2021	NR	1581	74.5	88
3	Belingheri et al. [39]	Italy	Nursing	Cross-sectional	Online	21–27 December 2020	NR	422	82.9	80.9
4	Bolatov et al. [40]	Kazakhstan	Medical	Cross-sectional	Online	March 2021	NR	888	76.5	22.4
5	De Sousa Chaves et al. [41]	Brazil	Medical	Cross-sectional	Online	18 December 2020 to 8 January 2021	Snowball sampling	250	58.5	84
6	Gao et al. [42]	China	Medical	Cross-sectional	Online	February– March 2021	Convenience sampling	612	63.2	NR
7	Gotlib et al. [43]	Poland	Nursing undergraduate students	Cross-sectional	Online	March–April 2021	NR	793	90.8	38
8	Grochowska et al. [44]	Poland	Medical	Cross-sectional	Online/Off line	4 September-5 November 2020	NR	419	<i>n</i> = 331	70.7
9	Jain et al. [45]	India	Medical	Cross-sectional	Online	2 February–7 March 2021	Respondent- driven sampling strategy	1068	48.6	89.4
10	Jiang et al. [2]	China	Nursing	Cross-sectional	Online	February–April 2021	Convenience	1488	84.27	1256
11	Kanyike et al. [4]	Uganda	Medical	Cross-sectional	Online	15–21 March 2021	Convenience	600	37.2	224
12	Katz et al. [35]	Israel	Medical	Cross-sectional	Online	December 2020	NR	104	61.5	91.35
13	Kelekar et al. [17]	USA	Medical	Cross-sectional	Online	November– December	NR	167	- NR -	126
15	Relekal et al. [17]	USA	Dental	Cross-sectional	Onime	2019	INIX	248	INK	135
14	L. Jain et al. [46]	India	Healthcare student	Cross-sectional	Online	November 2020–January 2021	Snowball sampling	655	61.98	63.82
15	Li et al. [47]	China	Medical	Cross-sectional	Online	15 March–30 March 2021	NR	2196	81.7	1291
16	Lindner- Pawłowicz et al. [48]	Poland	Medical	Cross-sectional	Online	8–31 December 2020	NR	350	NR	76.9

Table 1. Characteristics of included studies.

Table 1. Cont.

SL	Author	Study Country	Type of Healthcare Students	Study Design	Survey Method	Survey Period	Sampling Method	Sample Size, N	Gender, Female (%)	Vaccine Acceptance Rate (%)
17	Lucia et al. [18]	USA	Medical	Cross-sectional	Online	NR	NR	167	57	126
18	Lo Moro et al. [49]	Italy	Medical	Cross-sectional	Online	20 November 2020–2 February 2021	NR	838	63.5	93.3
19	Mahdi [50]	Iraq	Medical	Cross-sectional	Online	2021	NR	810	60.2	33.83
20	Manning et al. [21]	USA	Nursing	Cross-sectional	Online	10 August–14 September 2020	NR	1029	87.7	466
21	Mascarenhas et al. [20]	USA	Dental	Cross-sectional	Online	2020	NR	248	58	136
22	Mayan et al. [51]	USA	Medical	Cross-sectional	Online	9 February–15 March 2021	NR	1899	64.3	93.31
23	Mose et al. [52]	Ethiopia	Medical and health science	Cross-sectional	NR	1–30 March 2021	Simple random sampling	420	41.7	58.8
24	Petravic et al. [53]	Slovenia	Medical & Healthcare students	Cross-sectional	Online	December 2020	NR	624	79.49	Medical: 82, Healthcare: 51
25	Riad et al. [36]	22 countries	Dental	Cross-sectional	Online	6–28 February 2021	NR	6639	70.5	63.6
26	Rosental and Shmueli [54]	Israel	Medical and nursing	Cross-sectional	Online	27 August–28 September 2020	NR	628	66.6	Medical: 282Nursing: 234
27	Saied et al. [6]	Egypt	Medical	Cross-sectional	Online	8–15 January 2021	Convenience sampling	2133	NR	34.9
28	Szmyd et al. [55]	Poland	Medical	Cross-sectional	Online	22–25 December 2020	NR	687	64.77	632
29	Talarek et al. [56]	Poland	Medical	Cross-sectional	Online	March and April 2020	NR	411	68.4	94.6
30	Zhang et al. [57]	China	Healthcare students	Cross-sectional	Online	16–20 August 2021	NR	631	79.71	77.81
31	Zhou et al. [58]	China	Nursing	Cross-sectional	Online	4–20 January 2021	NR	1070	82.1	51.9

Notes: NR, Not Reported.

3.3. Prevalence of Vaccine Acceptance and Hesitancy

The estimated total COVID-19 vaccination acceptance rate among HCSs was 68.8% (95% CI: 60.8–76.3% $I^2 = 100\%$) (Figure 2). Talarek et al. [56] observed the highest acceptance rate (95.6%, 95% CI: 92.0–96.6%) in a study in Poland. The study in Kazakhstan by Bolatov et al. [40] reported the lowest vaccination acceptance rate of 22.4% (95% CI: 19.7–25.3%).

			E	vents	per 10	0			
Study	Events	Total			ations		Events	95%-CI	Weight
Janabi et al., 2021	89	197			- :		15.2	[38.1; 52.4]	3.2%
Bălan et al., 2021	1391	1581						[86.3; 89.5]	3.2%
Belingheri et al., 2021	341	422						[76.7; 84.5]	3.2%
	199	888						[19.7; 25.3]	3.2%
Bolatov et al., 2021	210	250				100		[78.9; 88.3]	3.2%
Chaves et al., 2021	301	793				1		[70.9, 00.3]	3.2%
Gotlib et al., 2021	296	419						[66.0; 75.0]	3.2%
Grochowska et al., 2021		1068							3.2%
Kumar, et al., 2021	955							[87.4; 91.2]	
Jiang et al., 2021	1256	1488		_				[82.5; 86.2]	3.2%
Kanyike et al., 2021	224	600				12		[33.5; 41.3]	3.2%
Katz et al., 2021	95	104						[84.2; 96.0]	3.1%
Kelekar et al., 2021	126	167			_ 1			[68.2; 81.8]	3.2%
Kelekar et al., 2021	135	248						[48.0; 60.7]	3.2%
Jain et al., 2021	418	655						[60.0; 67.5]	3.2%
Zheng, et al., 2021	1291	2196			•			[56.7; 60.9]	3.3%
Lindner-Pawłowicz et al., 2021		350			1			[72.1; 81.2]	3.2%
Lucia et al., 2020	126	167			÷	•		[68.2; 81.8]	3.2%
Lo Moro et al., 2022	782	838			1			[91.4; 94.9]	3.2%
Mahdi, 2021	274	810		-				[30.6; 37.2]	3.2%
Manning et al., 2021	466	1029		-				[42.2; 48.4]	3.2%
Mascarenhas et al., 2021	136	248		-	•			[48.4; 61.1]	3.2%
Mayan et al., 2021	1772	1899						[92.1; 94.4]	3.3%
Mose et al., 2022	247	420			-			[53.9; 63.6]	3.2%
Petravić et al., 2021	414	624			-			[62.5; 70.0]	3.2%
Riad et al., 2021	4222	6639						[62.4; 64.8]	3.3%
Rosental et al.,, 2021	516	628						[78.9; 85.1]	3.2%
Saied et al., 2021	744	2133		-			34.9	[32.9; 36.9]	3.3%
Szmyd et al., 2021	632	687				-	92.0	[89.7; 93.9]	3.2%
Talarek et al., 2021	389	411				-	94.6	[92.0; 96.6]	3.2%
Zhang et al., 2022	491	631				-	77.8	[74.4; 81.0]	3.2%
Zhou et al., 2021	555	1070		-	•		51.9	[48.8; 54.9]	3.2%
-									
Random effects model		29660			-	-	68.8	[60.8; 76.3]	100.0%
Heterogeneity: $I^2 = 100\%$, $\tau^2 = 0.0$	0563, p = 1		1	1	1	1	1		
		0	20	40	60	80 1	00		

Figure 2. COVID-19 vaccine acceptance rates among healthcare students by study.

The total estimated COVID-19 vaccination hesitancy rates among HCSs was 25.8% (95%CI: 18.5–33.8% I^2 = 99%) (Figure 3). Mahdi [50] reported the highest hesitancy rates in Iraq (66.2%, 95%CI: 62.8–69.4%), and the lowest rate of hesitancy was found in Poland (3.9%, 95% CI: 2.6–5.7%) by Szmyd et al. [55].

Study	Events	Total		Events observ			Events	95%-CI	Weight
Bălan et al., 2021	126	1581					8.0	[6.7; 9.4]	5.0%
Belingheri et al., 2021	63	422	-				14.9	[11.7; 18.7]	5.0%
Chaves et al., 2021	35	250					14.0	[9.9; 18.9]	4.9%
Gao et al., 2021	356	612			-		58.2	[54.1; 62.1]	5.0%
Gotlib et al., 2021	155	793	+				19.5	[16.8; 22.5]	5.0%
Grochowska et al., 2021	83	419	-				19.8	[16.1; 24.0]	5.0%
Kumar, et al., 2021	70	1068		_			6.6	[5.1; 8.2]	5.0%
Kanyike et al., 2021	184	600					30.7	[27.0; 34.5]	5.0%
Kelekar et al., 2021	38	167		-			22.8	[16.6; 29.9]	4.9%
Kelekar et al., 2021	112	248					45.2	[38.9; 51.6]	4.9%
Zheng, et al., 2021	905	2196					41.2	[39.1; 43.3]	5.0%
Lo Moro et al., 2022	56	838					6.7	[5.1; 8.6]	5.0%
Mahdi, 2021	536	810			-		66.2	[62.8; 69.4]	5.0%
Manning et al., 2021	341	1029		-			33.1	[30.3; 36.1]	5.0%
Mose et al., 2022	173	420					41.2	[36.4; 46.1]	5.0%
Riad et al., 2021	1494	6639					22.5	[21.5; 23.5]	5.1%
Saied et al., 2021	975	2133					45.7	[43.6; 47.9]	5.0%
Szmyd et al., 2021	27	687					3.9	[2.6; 5.7]	5.0%
Zhang et al., 2022	69	631	-				10.9	[8.6; 13.6]	5.0%
Zhou et al., 2021	464	1070		+			43.4	[40.4; 46.4]	5.0%
Random effects model Heterogeneity: I^2 = 99%, τ		22613 , p = 0 (20	40	60	80	25.8	[18.5; 33.8]	100.0%

Figure 3. COVID-19 vaccine hesitancy rates among healthcare students by study.

3.4. Sub-Group Analysis

Figures 4 and 5 present country-specific COVID-19 vaccine acceptance rates among HCSs. The pooled prevalence of the highest acceptance rate was observed in Romania (88.0%, 95% CI: 44.5–100%), followed by Italy (87.8%, 95% CI: 58.3–100%, $I^2 = 98\%$), Israel (87.0%, 95% CI: 58.7–100%, $I^2 = 84\%$), Brazil (84.0%, 95% CI: 38.4–100%), and India (78.0%, 95% CI: 45.3–98.0%, $I^2 = 99\%$).

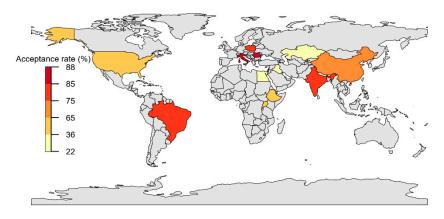


Figure 4. Map of COVID-19 vaccine acceptance rates among healthcare students by country.

Study	Events	Total	Events per 100 observations	Events	95%-CI	Weigl
Country: 22 countries Riad et al., 2021	4222	6639		63.6	[62 4· 64 8]	3.3
Random effects model	4222	6639			[62.4; 64.8] [18.0; 97.4]	3.3
leterogeneity: not applicable		0000		0010	[]	010
Country: Brazil						
Chaves et al., 2021	210	250			[78.9; 88.3]	3.2
Random effects model leterogeneity: not applicable		250		84.0	[38.4; 100.0]	3.2
Country: China						
liang et al., 2021 Zheng, et al., 2021	1256 1291	1488 2196	. •	84.4 58.8	[82.5; 86.2] [56.7; 60.9]	3.2° 3.3°
Zhang et al., 2022	491	631			[74.4; 81.0]	3.2
Zhou et al., 2021	555	1070	-		[48.8; 54.9]	3.2
Random effects model deterogeneity: $I^2 = 99\%$, $\tau^2 = 0.00$	612 n < 0	5385 01		69.0	[45.2; 88.4]	13.0
	o 12, p - 0.					
Sountry: Egypt Saied et al., 2021	744	2133			[32.9; 36.9]	3.3
Random effects model leterogeneity: not applicable		2133		34.9	[2.1; 80.8]	3.3
ountry: Ethiopia						
Mose et al., 2022	247	420			[53.9; 63.6]	3.2
Random effects model deterogeneity: not applicable		420		58.8	[14.1; 95.8]	3.2
Country: India			10.460			
Cumar, et al., 2021 ain et al., 2021	955 418	1068 655			[87.4; 91.2] [60.0; 67.5]	3.2 3.2
Random effects model leterogeneity: $l^2 = 99\%$, $\tau^2 = 0.00$		1723			[45.3; 98.0]	6.5
	<u>.</u> , p ~ 0.					
Country: Iraq Nahdi, 2021	274	810		33.8	[30.6; 37.2]	3.2
Random effects model leterogeneity: not applicable		810		33.8	[1.7; 80.0]	3.2
Country: Israel						
Katz et al., 2021	95	104			[84.2; 96.0]	3.1
Rosental et al.,, 2021	516	628 732	•		[78.9; 85.1]	3.2 6.4
Random effects model leterogeneity: $I^2 = 84\%$, $\tau^2 = 0.00$	612, p = 0.			07.0	[56.7; 100.0]	0.4
Country: Italy	Lange of the	1000000000		Y-100-002101		With a
Belingheri et al., 2021	341 782	422 838	-		[76.7; 84.5]	3.2
ucia et al., 2020 Random effects model		1260			[91.4; 94.9] [58.3; 100.0]	3.2 6.5
Heterogeneity: $I^2 = 98\%$, $\tau^2 = 0.0$	612, p < 0.	.01				
Country: Kazakhstan Bolatov et al., 2021	199	888		22.4	[19.7; 25.3]	3.2
Random effects model	155	888 -			[0.0; 69.0]	3.2
leterogeneity: not applicable						
Country: Poland Sotlib et al., 2021	301	793	4	38.0	[34.6; 41.4]	3.2
Grochowska et al., 2021	296	419	-	70.6		3.2
indner-Pawłowicz et al., 2021		350		76.9	[72.1; 81.2]	3.2
zmyd et al., 2021	632	687	•	92.0		3.2
alarek et al., 2021 tandom effects model	389	411 2660		94.6	[92.0; 96.6] [56.6; 92.3]	3.2 16.1
leterogeneity: $I^2 = 99\%$, $\tau^2 = 0.00$	612, p < 0.			10.3	[00.0, 02.0]	10.1
ountry: Romania	100		6.200		100.0.00	
tälan et al., 2021 tandom effects model	1391	1581 1581	•		[86.3; 89.5] [44.5; 100.0]	3.2 3.2
leterogeneity: not applicable		1001		00.0		9.2
country: Slovenia		001		60 G	100 5. 70	
etravić et al., 2021 tandom effects model	414	624 624			[62.5; 70.0] [20.1; 98.3]	3.2 3.2
leterogeneity: not applicable				0010		w the
country: Uganda	224	600	<u> </u>	27.0	100 E. 44 01	
Canyike et al., 2021 Random effects model	224	600 600			[33.5; 41.3] [2.8; 82.9]	3.2 3.2
leterogeneity: not applicable		000		31.3	[2.0, 02.9]	5.2
ountry: USA	and the second					
anabi et al., 2021 Kelekar et al., 2021	89 126	197 167		45.2 75.4	[38.1; 52.4]	3.2 3.2
kelekar et al., 2021 Kelekar et al., 2021	126	248		75.4 54.4		3.2
Noro et al., 2022	126	167		75.4	[68.2; 81.8]	3.2
lanning et al., 2021	466	1029	-	45.3	[42.2; 48.4]	3.2
Ascarenhas et al., 2021	136	248	-	54.8		3.2
/layan et al., 2021 Random effects model	1772	1899 3955			[92.1; 94.4]	3.3 22.4
leterogeneity: $l^2 = 99\%$, $\tau^2 = 0.0$	612, p < 0.			04.9	[46.6; 81.2]	22.4
andom effects model		29660		68.8	[60.8; 76.3]	100.0

Figure 5. COVID-19 vaccine acceptance rates among healthcare students by country.

Country sub-group analyses are presented in Figure 6. Iraq showed the highest rates of vaccine hesitancy (66.2%, 95% CI: 35.5–90.8%), followed by Egypt (45.5%, 95% CI: 17.5–

75.5%), Ethiopia (41.2%, 95% CI: 14.0–71.8%), China (37.1%, 95% CI: 23.0–52.6%, *I*² = 99%), and the U.S. (33.4%, 95% CI: 17.4–51.4%, *I*² = 92%).

Study	Events	Total	Events per 100 observations	Events	95%-CI	Weigh
Country: 22 countries						
Riad et al., 2021	1494	6639	•		[21.5; 23.5]	5.19
Random effects model		6639		22.5	[3.4; 51.8]	5.19
Heterogeneity: not applicat	le					
5. 7 11						
Country: Brazil						
Chaves et al., 2021	35	250		14.0	[9.9; 18.9]	4.9
Random effects model		250			[0.3; 41.6]	
Heterogeneity: not applicat	le					
Country: China						
Gao et al., 2021	356	612		58.2	[54.1; 62.1]	5.0%
Zheng, et al., 2021	905	2196	•	41.2	[39.1; 43.3]	5.0%
Zhang et al., 2022	69	631		10.9	[8.6; 13.6]	5.0%
Zhou et al., 2021	464	1070	-	43.4	[40.4; 46.4]	5.0%
Random effects model		4509			[23.0; 52.6]	
Heterogeneity: $I^2 = 99\%$, τ^2	= 0.0249	, p < 0.0	1			
Country: Egypt						
Saied et al., 2021	975	2133		45.7	[43.6; 47.9]	5.0%
Random effects model		2133			[17.5; 75.5]	5.0%
leterogeneity: not applicat	le	2100		1011	[1110, 1010]	0.07
iotorogeneity. not apprious						
Country: Ethiopia						
Mose et al., 2022	173	420	-	412	[36.4; 46.1]	5.0%
Random effects model	175	420			[14.0; 71.8]	5.0%
Heterogeneity: not applicat	le	420		-7 1 . da	[14.0, 71.0]	0.0
received and the second s	no l					
Country: India						
Kumar, et al., 2021	70	1068	-	6.6	[5.1; 8.2]	5.09
Random effects model	10	1068 -			[0.0; 29.1]	5.0%
	le.	1000		0.0	[0.0, 29.1]	5.0
Heterogeneity: not applicat	Ne					
Country: Iraq						
Mahdi, 2021	526	910	-	66.2	162 9.60 41	5.09
	536	810			[62.8; 69.4]	
Random effects model		810		66.2	[35.5; 90.8]	5.0%
Heterogeneity: not applicat	le					
Countinu Italu						
Country: Italy	63	422	-	14.0	[11 7. 10 7]	5.0%
Belingheri et al., 2021					[11.7; 18.7]	
Lucia et al., 2020	56	838	•		[5.1; 8.6]	
Random effects model	0.0040	1260		10.4	[1.1; 27.3]	10.0%
Heterogeneity: $I^2 = 95\%$, τ^2	= 0.0249	, p < 0.0				
Country: Poland						
	155	793		10 5	[16 0. 00 E]	5.09
Gotlib et al., 2021	83				[16.8; 22.5]	
Grochowska et al., 2021		419			[16.1; 24.0]	
Szmyd et al., 2021	27	687			[2.6; 5.7]	
Random effects model	0.0010	1899		13.3	[3.6; 27.6]	15.09
Heterogeneity: $I^2 = 98\%$, τ^2	= 0.0249	, p < 0.0	1			
Den la Bernie						
Country: Romania	100	4504	_	0.0	107 0 1	F 64
Bălan et al., 2021	126		•	8.0	[6.7; 9.4]	
Random effects model		1581 .		8.0	[0.0; 31.6]	5.0
Heterogeneity: not applicat	le					
Country: Uganda						-
Kanyike et al., 2021	184	600	-		[27.0; 34.5]	5.0%
Random effects model		600		30.7	[7.3; 61.3]	5.0%
leterogeneity: not applicat	le					
Country: USA						
Kelekar et al., 2021	38	167			[16.6; 29.9]	
Kelekar et al., 2021	112	248			[38.9; 51.6]	4.99
Manning et al., 2021	341	1029	-		[30.3; 36.1]	5.09
Random effects model		1444			[17.7; 51.4]	14.89
Heterogeneity: $I^2 = 92\%$, τ^2	= 0.0249	, p < 0.0	1			
Random effects model		22613		25.8	[18.5; 33.8]	100.0

Figure 6. COVID-19 vaccine hesitancy rates among healthcare students by country.

3.5. Time Trends

COVID-19 vaccine acceptance rates decreased with time (Figure 7). During 2020, the pooled acceptance rate was 75.0% (95% CI: 63.5–85.0%, $I^2 = 99\%$). The acceptance rate in 2021 was only 62.8% (95% CI: 51.3–73.6%).

95%-CI	Weight	

			Events per 100			
Study	Events	Total	observations E	vents	95%-CI	Weight
Year: 2020			:			
	00	107		45.0	100 4. 50 41	0.00/
Janabi et al., 2021	89	197			[38.1; 52.4]	3.2%
Belingheri et al., 2021	341	422	1		[76.7; 84.5]	3.2%
Grochowska et al., 2021	296	419			[66.0; 75.0]	3.2%
Katz et al., 2021	95	104			[84.2; 96.0]	3.1%
Kelekar et al., 2021	126	167	÷••		[68.2; 81.8]	3.2%
Kelekar et al., 2021	135	248		54.4	[48.0; 60.7]	3.2%
Lindner-Pawłowicz et al., 2021	269	350		76.9	[72.1; 81.2]	3.2%
Lucia et al., 2020	126	167		75.4	[68.2; 81.8]	3.2%
Lo Moro et al., 2022	782	838	÷ •	93.3	[91.4; 94.9]	3.2%
Manning et al., 2021	466	1029	-	45.3	[42.2; 48.4]	3.2%
Mascarenhas et al., 2021	136	248		54.8	[48.4; 61.1]	3.2%
Petravić et al., 2021	414	624	-	66.3	[62.5; 70.0]	3.2%
Rosental et al., 2021	516	628	-		[78.9; 85.1]	3.2%
Szmyd et al., 2021	632	687			[89.7; 93.9]	3.2%
Talarek et al., 2021	389	411			[92.0; 96.6]	3.2%
Random effects model	000	6539			[64.0; 84.7]	48.1%
Heterogeneity: $I^2 = 99\%$, $\tau^2 = 0.09$	557 n < 0			1010	[0 110, 0 111]	1011/0
Hotologonoldy. 7 0078, 7 0.00	501, p · 0	.01				
Year: 2021						
Bălan et al., 2021	1391	1581		88.0	[86.3; 89.5]	3.2%
Bolatov et al., 2021	199	888	+		[19.7; 25.3]	3.2%
Chaves et al., 2021	210	250			[78.9; 88.3]	3.2%
Gotlib et al., 2021	301	793	+		[34.6; 41.4]	3.2%
Kumar, et al., 2021	955	1068			[87.4; 91.2]	3.2%
Jiang et al., 2021	1256	1488			[82.5; 86.2]	3.2%
Kanyike et al., 2021	224	600	-		[33.5; 41.3]	3.2%
Jain et al., 2021	418	655	-		[60.0; 67.5]	3.2%
Zheng, et al., 2021	1291	2196			[56.7; 60.9]	3.3%
Mahdi, 2021	274	810	+		[30.6; 37.2]	3.2%
Mayan et al., 2021	1772	1899			[92.1; 94.4]	3.3%
Mose et al., 2022	247	420	-		[53.9; 63.6]	3.2%
Riad et al., 2021	4222	6639	1000		[62.4; 64.8]	3.3%
Saied et al., 2021	744	2133			[32.9; 36.9]	3.3%
	491	631	1000			3.2%
Zhang et al., 2022	555	1070	225		[74.4; 81.0]	3.2%
Zhou et al., 2021		23121			[48.8; 54.9]	
Random effects model				02.8	[51.3; 73.6]	51.9%
Heterogeneity: $I^2 = 100\%$, $\tau^2 = 0.0$	J557, p =	U				
Random effects model		29660		68.8	[60.8; 76.3]	100.0%
Heterogeneity: $I^2 = 100\%$, $\tau^2 = 0.0$						
Residual heterogeneity: $l^2 = 100\%$	$6.\tau^2 = 0.0$	$557 \ n = 00$	20 40 60 80 100			
Residual field/ogeneity. 7 = 1007	ο, τ = 0.0	οσι, μ =00				

Events per 100

Figure 7. COVID-19 vaccine acceptance rates among healthcare students by year.

3.6. Predictors of Vaccine Acceptance

Figure 8 presents the potential predictors associated with COVID-19 vaccine acceptance among HCSs. Sex, place of residence, previous history of vaccination, and concern about the vaccination side effects were considered. Only one factor-concern about potentially serious side effects of vaccines (n = 3 studies, OR = 0.2, 95% CI: 0.1–0.4)—was significantly associated with lower acceptance rates.

Variables	Categories	Studies	OR				OR (95% CI)	I^2
Sex	Female	6	4				1	
Sex	Male	6					1 [0.6, 1.6]	81.2
Destines	Rural	3	i i				1	
Residence	Urban	3					0.9 [0.6, 1.2]	45.4
Previous history of vaccination	No	3	÷				1	
r revious instory of vaccination	Yes	3					1.6 [0.3, 7.1]	94.4
Concerned about vaccination side effec	No	3	É.				1	
Concerned about vaccination side effect	Yes	3					0.2 [0.1, 0.4]	71.2
			0 2	4 OR	6	8		

Figure 8. Predictors of COVID-19 vaccination acceptance among healthcare students.

3.7. Risk of Bias

All 31 studies were assessed to be of the highest possible quality based on the JBI technique (Table S1). Studies that used ineffective recruitment methods like convenience and snowball sampling via social media were not removed, but their results may not have been representative of the population.

We observed no risk of publication bias. The Egger's tests among studies of vaccine acceptance (p-value = 0.64) and vaccine hesitancy (p-value = 0.97) were not significant (Figures S1 and S2).

4. Discussion

4.1. Summary of the Main Findings

Vaccines have been revolutionary in their the potential to end the COVID-19 pandemic [38]. However, vaccine hesitancy remains high and an important obstacle in many vaccination programs [59,60]. Vaccine skepticism is on the rise among healthcare workers due in part to the rapid development of these vaccines [61]. Healthcare students can act as role models in their communities to increase trust about the safety of vaccinations [38]. Furthermore, healthcare students are frontline workers likely to be exposed to COVID-19 during training and clinical practice. It is necessary to prioritize vaccinations for this group. To our knowledge, no systematic review or meta-analysis had been conducted on vaccination acceptance and hesitancy rates among healthcare students.

The current study systematically reviewed and analyzed the data from 30,272 healthcare students across the world. Our pooled estimations showed that approximately twothirds of healthcare students were willing to accept a COVID-19 vaccine. Meanwhile, approximately one-quarter were hesitant about accepting a COVID-19 vaccine. Such rates are similar to those observed in general populations [10] and healthcare workers [62]. One potential explanation for these findings is that healthcare students may be exposed to large amounts of health-related information, which may make them more aware of the vaccine's serious side effects and thus influence their decision to be vaccinated [18].

Country-wise, pooled results found that healthcare students from comparatively highincome countries like Romania, Poland, Italy, and the U.S. were more likely to accept a COVID-19 vaccine than students in other countries. One possible explanation is that vaccines were more prevalent in higher-income countries, making it easier for students to receive vaccinations. A recent study reported that, among 25 countries, 10 high-income countries received a median of 51.7% more vaccine doses than their low-income counterparts (31–14.9%) despite high rates of authorization [63]. Furthermore, most of these studies were conducted during the early stages of the pandemic, when countries were experiencing increasing rates of COVID-19-related mortality. Fear of becoming infected could have influenced vaccine acceptance levels.

Low vaccine acceptance and high vaccine hesitancy were observed in Middle Eastern and African countries (e.g., Kazakhstan, Egypt, and Iraq). Middle Eastern results may be attributed to high belief rates in conspiracy theories and high dependence on social media platforms to obtain vaccine-related information [64]. Lower COVID-19 mortality rates might have influenced vaccine acceptance rates in African countries [65]. In addition, people in Africa have a history of vaccination skepticism, which may have contributed to low acceptance rates [66]. Traditionally, many African groups have shown poor healthseeking behaviors because of spiritual considerations that limit vaccination uptake [67].

Our study found that vaccine acceptance among healthcare students decreased over time. Earlier studies have also found that vaccine acceptance varies over time [68]. For example, a global systematic review on vaccine acceptancy rates reported a decline from 79% in March–May to 60% in June–October of 2020 [10]. This finding could be explained by the fact that students during the early stages of the pandemic were more fearful of being infected, which motivated them to receive a vaccine. Similar findings were observed among Egyptian medical students [6]. Additionally, Wong et al. [69] reported that individuals who were more fearful of COVID-19 demonstrated greater willingness to receive a vaccine due to the perceived benefit of immunization reducing the risk of infection. With time, healthcare students were exposed to more professional information, which likely influenced their decisions. Recent research shows that the observed decreases in vaccination intentions may be caused by COVID-19-related misinformation, as well as public worries about vaccine safety [70,71].

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Finally, we found that concerns about serious side effects of COVID-19 predicted vaccine acceptance. A similar finding was observed in Egypt, where 74% of medical students reported that side effects were major barriers of vaccine acceptance [6]. Another study conducted among Egyptian healthcare professionals (HCWs) found that 57% of HCWs were unwilling to accept a vaccine due to their belief that vaccines were unsafe [72]. Such findings could be explained by students being doubtful of vaccine efficacy due to its rapid development. However, it is worth mentioning that different countries and regions often use different types of vaccines, and potential side effects vary, which may also influence vaccine hesitancy from study to study.

4.2. Implications

COVID-19 vaccinations should be prioritized for frontline workers since they are critical to COVID-19 responses and are at high risk of infection. Given the low degree of intention to vaccinate against COVID-19 among healthcare students, it is necessary to boost vaccine acceptability rates in this population. The majority of countries agree that frontline workers should be immunized against COVID-19 [27]. Our systematic review could be an initial step, as it estimated country-wise vaccine acceptancy and hesitancy coverage among healthcare students. This information could help decision-makers determine where and how to prioritize vaccine distribution. It is critical to focus on establishing confidence in COVID-19 vaccinations among this population. Governments of each country could mandate vaccination policies for not only healthcare workers but also healthcare students.

4.3. Strengths and Limitations

This study has a number of strengths. It is the first comprehensive meta-analysis study on vaccination acceptability among healthcare students that we are aware of. All of the publications considered in this review were judged as high-quality observational studies. Our evaluation considered the most recent study findings when calculating the final vaccination acceptance rate.

Our review also has limitations. First, our search was confined to three databases (Scopus, PubMed, and Web of Science). Other databases, such as Embase, PsycINFO, CINAHL, PMC, or NCBI were not searched. Secondly, we excluded preprints and unpublished grey literature. Given the spike in COVID-19 papers throughout our research period, we may have reached a different outcome if preprints or unpublished grey literature were included. Third, the data collection period for the included studies was from 2020 to early 2022, which may have influenced the findings due to the fact that public sentiments regarding vaccination change over time. Fourth, most of the reviewed research was cross-sectional and performed through online surveys. Conclusions from online research are prone to clarity and self-selection bias [73]. Finally, we were unable to investigate some potential determinants of vaccine acceptance owing to data constraints.

5. Conclusions

Healthcare students were moderately willing to accept a COVID-19 vaccine as of March of 2022. Romania and Kazakhstan showed the highest and lowest vaccine acceptance rates, respectively. Vaccination acceptance rates among healthcare students decreased from 2020 to 2021. Healthcare students who expressed concerns about the potential side effects of the vaccine were less likely to accept a vaccine.

Governments should prioritize vaccine distribution to frontline healthcare workers, including students, as soon as safe vaccines are available. These efforts should be coupled with comprehensive educational programs that reinforce the safety of vaccines to healthcare students. Previous studies indicate that vaccine-exposed medical students have positive attitudes toward vaccines. If more healthcare students are vaccinated, they can relate their positive experiences to their patients and increase vaccine uptake in the general public.

Supplementary Materials: The following supporting information can be downloaded at: https://www.mdpi.com/article/10.3390/vaccines10050806/s1, Figure S1: Funnel plot of vaccine acceptance among healthcare students. Figure S2: Funnel plot of vaccine hesitancy among healthcare students.; Table S1: Quality assessment of included studies.

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