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Clinical Epidemiology and Global Health

journal homepage: www.elsevier.com/locate/cegh



The risk factors and pregnant women's willingness toward the SARS-CoV-2 vaccination in various countries: A systematic review and meta-analysis

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ARTICLE INFO

Keywords: Pregnant women Risk factors SARS-CoV-2 Vaccine Willingness

ABSTRACT

Introduction: Pregnant women will benefit from research on immunization during pregnancy because they will have more accurate information on the SARS-CoV-2 vaccine. The purpose of this study was to determine the risk factors and pregnant women's desire to get the SARS-CoV-2 vaccine in various countries. *Methods:* A search of PubMed, ProQuest, and EBSCO for related publications published (January and December 2021) on risk factors and pregnant women's desire to get the SARS-CoV-2 vaccine in various countries. The Pooled Odds Ratio (POR) were calculated using fixed and random-effect analysis. The I-squared formula was used to calculate the heterogeneity. Egger's and Begg's tests were used to identify study bias. STATA 16.0 was used for data analysis. *Results:* This study revealed good practice has the highest POR (8.99), followed by received influenza vaccine last year (2.72), high perception of SARS-CoV-2 vaccine (2.70), >35 years (2.01), sufficient information about the SARS-COV-2 vaccine (1.94), higher school education (1.84), and third trimester (1.35) with pregnant women's desire to ward the SARS-CoV-2 vaccination. The heterogeneity analysis revealed homogenous among risk factors in >35 years, high perception of SARS-CoV-2 vaccine, good practice, and third trimester (1² ≤ 50%). In the articles combined in this study, there was no indication of study bias.

Conclusion: The insights of this study might help the authorities in determining the most effective strategy to deploy SARS-CoV-2 mass immunization campaigns for pregnant women.

1. Introduction

Many people have died as a condition of the severe acute respiratory syndrome coronavirus 2 (SARS-CoV-2) outbreak.¹ Massive human death has resulted in public health issues, overwhelmed health systems, interrupted distribution chains and the economy.^{2,3} Pregnant women may be more vulnerable to SARS-CoV-2 infection than non-pregnant women.^{4–6} Severe illness is defined as a condition that necessitates hospitalization, intensive care, the use of a ventilator, special breathing equipment, and/or death. According to certain studies, infection with the SARS-CoV-2 in pregnant mothers is related to a higher chance of a

severe illness course, requiring invasive ventilation or ecmo, and/or death. $^{5,6}_{\rm \cdot,6}$

In another research, a link between SARS-CoV-2 and the likelihood of preterm and cesarean births was discovered.⁷ Vertical transmission of the virus, which can induce hydrops fetalis and fetal death.^{8,9} Children are more vulnerable to asymptomatic illnesses, but they also carry the SARS-COV-2 virus, which they may pass on to others, especially pregnant women.^{10,11} Pregnant women who are infected with SARS-CoV-2 have a higher risk of preterm delivery and other poor pregnancy outcomes than pregnant women who are not infected with SARS-CoV-2.¹ It's also important to note that the SARS-CoV-2 pandemic leads pregnant

https://doi.org/10.1016/j.cegh.2022.100982

Received 18 December 2021; Received in revised form 29 January 2022; Accepted 3 February 2022

Available online 10 February 2022

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women to be concerned about their fetus's and personal health, which has a substantial impact on their well-being. 12

Immunization during pregnancy research will give information on vaccine safety and efficacy concerns. It has been produced information for healthcare practitioners and patients on how to utilize it in counseling.¹³ Unfortunately, insufficient study has been conducted to establish the global population's sentiments regarding vaccination among pregnant women. To our knowledge, no previously published study has been subjected to meta-analysis.

2. Materials and methods

2.1. Study design and research sample

A meta-analysis studies were undertaken to review current studies related to risk factors and pregnant women's desire to get the SARS-CoV-2 vaccine in various countries. This study follows the preferred reporting items for systematic reviews and meta-analysis (PRISMA) guidelines.¹⁴

2.2. Eligibility criteria

Original articles having a cross-sectional study design, English language, and human participants as study subjects were only considered for inclusion. Exclusion criteria for the study included the unavailability of a full text version, irrelevant topics, and data from articles that could not be used for further evaluation.

2.3. Search approach and study collection

A search of PubMed, ProQuest, and EBSCO for related publications published (January and December 2021) with keywords ("pregnant women" AND "COVID-19" OR "coronavirus" OR "SARS-CoV-2" AND "vaccine" AND "acceptance"). In this study, pregnant women's desire toward the SARS-CoV-2 vaccination was the dependent variable. The independent variables were the risk factors of SARS-CoV-2 vaccine willingness. The literature quest was carried out by two independent investigators. After the initial search, the duplicates were manually deleted, and the title/abstracts were screened for possible relevance. Following that, the full-texts of possible papers were evaluated using the criterion.

2.4. Data extraction

Data was retrieved by two separate authors using structured extraction forms. The quality of the publications was evaluated using the Newcastle-Ottawa Quality Assessment Scale (NOS). Articles were categorized into low, medium, and high quality groups using the numbers 0-3, 4-6, and 7-9.¹⁵ PRISMA flowcharts were used to show the stages required in looking for study publications (Fig. 1).

2.5. Data analysis

The Pooled Odds Ratio (POR) from the obtained data was calculated with a 95% confidence interval for further data analysis. I^2 indicates that there was heterogeneity between publications if it was greater than 50%. The random effect analysis was used if the outcome was heterogeneous, and the fixed effect analysis was used if it was homogenous. Furthermore, the results were summarized as forest plots, and Egger's and Begg's tests were used to examine study bias. There was no publication bias among the studies, according to the p > 0.05 findings of the two tests. The role of covariate in lower middle income countries (LMICs) was investigated using restricted-maximum likelihood random effects meta-regression. Processing and analysis of all data, STATA 16.0 was used.

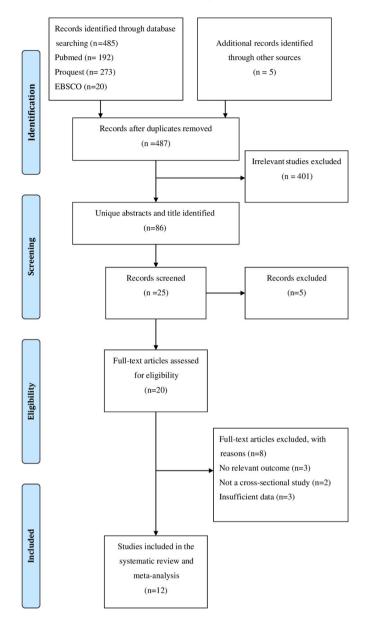


Fig. 1. PRISMA flow diagram.

3. Results

A total of 12 recent studies were considered in this systematic review (Table 1). There were 15,444 participants in all that participated in the study. $^{16\text{-}27}$

Table 1 is based on a review of 12 cross-sectional studies that looked at risk factors and pregnant women's desire to have the SARS-CoV-2 vaccine in various countries. This research revealed variables related to willingness of SARS-CoV-2 vaccination among pregnant women >35 years, higher school education, sufficient information about the SARS-COV-2 vaccine, high perception, good practice, received influenza vaccine last year and third trimester.

Meta-estimate of willingness of pregnant women toward the SARS-CoV-2 vaccination and associated factors (Table 2 and Fig. 2).

Table 2 and Fig. 2 revealed good practice has the highest POR (95% CI) (8.99, 7.42–10.56), followed by received influenza vaccine last year (2.72, 2.09–3.35), high perception of SARS-CoV-2 vaccine (2.70, 2.13–3.27), >35 years (2.01, 1.10–2.93), sufficient information about the SARS-COV-2 vaccine (1.94, 0.94–2.95), higher school education (1.84, 1.40–2.28), and third trimester (1.35, 1.10–1.60) with desire of

Table 1

Systematic review of pregnant women's willingness toward the SARS-CoV-2 vaccination and associated factors.

1st author	Year	Study location	Study design	Sample size	Risk factors (OR, 95% CI)	NOS
Goncu Ayhan et al. ¹⁶	2021	Ankara, Turkey	Cross-sectional	300	Sufficient information about the SARS- CoV-2 vaccine (2.07, 1.22–3.51)	7
Battarbee et al. ¹⁷	2021	U.S.	Cross-sectional multicenter	915	Higher school education (2.40, 1.30–4.70) Received influenza vaccine last year	8
Levy et al. ¹⁸	2021	New York	Survey study	653	(2.60, 1.90–3.60) Sufficient information about the SARS- CoV-2 vaccine (30.95, 9.55–100.33)	7
Geoghegan et al. ¹⁹	2021	Dublin, Ireland	Online survey	300	>35 years old (1.36, 0.80–2.32) Higher school education (1.78, 1.09–2.92)	6
Hailemariam et al. ²⁰	2021	Southwest Ethiopia	Cross-sectional	412	 >35 years old (6.73, 3.84–11.79) Higher school education (5.87, 3.14–10.97) High perception of SARS-CoV-2 vaccine 	7
Mose et al. ²¹	2021	Southwest Ethiopia	Cross-sectional	396	(4.35, 2.73–6.95) >35 years old (2.55, 1.06–6.08) Higher school education (3.28, 1.92–5.59) Sufficient information about the SARS- CoV-2 vaccine (3.24, 1.78–5.89)	7
Nguyen et al. ²²	2021	Vietnam	Cross-sectional	651	Good practice (9.15, 8.73–12.19) Higher school education (1.98, 1.24–3.14) High perception of SARS-COV-2	7
Skjefte et al. ²³	2021	U.S., U.K., India, Brazil, Russia, Spain, Argentina, Colombia, Mexico, Peru, South Africa, Italy, Chile, Philippines, Australia and New Zealand	Cross-sectional	5,294	vaccine (2.71, 1.93–3.82) Higher school education (1.31, 1.12–1.54) Received influenza vaccine last year (3.29, 2.91–3.72)	8
Stuckelberger et al. ²⁴	2021	Switzerland	Cross-sectional	1,551	 (3.5), 2.31-30.20 >35 years old (2.00, 1.30-3.00) Higher school education (1.70, 1.30-2.20) Received influenza vaccine last year (3.60, 2.80-4.70) Third trimester (1.40,1.00-2.00) 	7
Sutton et al. ²⁵	2021	U.S.	Online survey	1,012	Received influenza vaccine last year (2.25, 1.66–3.05)	7
Tao et al. ²⁶	2021	China	Cross-sectional	1,392	(2.25, 1:00-5:03) Higher school education (2.85, 1.45–5:59) Sufficient information about the SARS- CoV-2 vaccine (1.05, 1.01–1.10) Third trimester (1.49,1.03–2.16)	7
Wang et al. ²⁷	2021	China	Cross-sectional	2,568	Sufficient information about the SARS- CoV-2 vaccine (2.63, 1.38–5.00) Received influenza vaccine last year (1.81, 1.18–2.80) High perception of SARS-CoV-2 vaccine (2.48, 1.83–3.35) Good practice (8.27, 5.35–12.77) Third trimester (1.27, 0.98–1.65)	8
Total samples NOS score				15,444 7.17 \pm 0.58		

Abbreviation: CI = confidence interval; OR = odds ratio; NOS, Newcastle–Ottawa Quality Assessment Scale.

pregnant women toward the SARS-CoV-2 vaccination. The heterogeneity analysis revealed homogenous among risk factors in >35 years, high perception, good practice, and third trimester (I² \leq 50%).

(p=<0.001). This study found that pregnant women's desire toward the SARS-CoV-2 vaccination varies depending on the country type.

The findings of Egger's and Begg's tests to identify study bias (Table 3).

Table 3 revealed that based on the results of Egger's and Begg's tests (p > 0.05), an associated factors of >35 years, higher school education, sufficient information, high perception, good practice, received influenza vaccine last year, and third trimester had no study bias among publications included.

The relationship between LMICs and pregnant women's desire toward the SARS-CoV-2 vaccination based on meta-regression (Fig. 3).

Fig. 3 revealed that the relationship between LMICs and decreased pregnant women's desire toward the SARS-CoV-2 vaccination

4. Discussion

Our findings revealed that pregnant women who received the SARS-CoV-2 immunization had a high level of practice about the vaccine. Vaccine practice rates can support in the planning of activities and initiatives that will assist increase knowledge and reassure people about the safety and advantages of vaccines, which will help prevent the spread of the virus and minimize the negative impacts of this historic pandemic.^{28,29} Identification of practice and vaccination rates for SARS-CoV-2 vaccines can assist in the selection of the most effective communication method for boosting vaccination confidence.³⁰ Because

Table 2

Meta-estimate of pregnant women's willingness toward the SARS-CoV-2 vaccination and associated factors.

Risk factors	1st author	OR (95% CI)	POR (95% CI)	Heterogeneity	
				I ² (%)	p-value
>35 years			2.01 (1.10-2.93)	45.29	0.35
	Geoghegan et al. ¹⁹	1.36 (0.80-2.32)			
	Hailemariam et al. ²⁰	6.73 (3.84–11.79)			
	Mose et al. ²¹	2.55 (1.06-6.08)			
	Stuckelberger et al. ²⁴	2.00 (1.30-3.00)			
Higher school education	0		1.84 (1.40-2.28)	51.98	0.02
0	Battarbee et al. ¹⁷	2.40 (1.30-4.70)			
	Geoghegan et al. ¹⁹	1.78 (1.09-2.92)			
	Hailemariam et al. ²⁰	5.87 (3.14-10.97)			
	Mose et al. ²¹	3.28 (1.92–5.59)			
	Nguyen et al. ²²	1.98 (1.24–3.14)			
	Skjefte et al. ²³	1.31 (1.12–1.54)			
	Stuckelberger et al. ²⁴	1.70 (1.30–2.20)			
	Tao et al. ²⁶	2.85 (1.45–5.59)			
Sufficient information about the SARS-COV-2 vaccine	nuo et al.	2.05 (1.45-5.55)	1.94 (0.94-2.95)	61.70	0.02
Sumclent information about the SARS-COV-2 Vaccine	Goncu Ayhan et al. ¹⁶	2.07 (1.22-3.51)	1.94 (0.94-2.93)	01.70	0.02
	Levy et al. ¹⁸	30.95 (9.55–100.33)			
	Mose et al. ²¹	3.24 (1.78–5.89)			
	Tao et al. ²⁶	1.05 (1.01–1.10)			
	Wang et al. ²⁷	2.63 (1.38–5.00)			
High perception of SARS-CoV-2 vaccine	wang et al.	2.03 (1.38-3.00)	2.70 (2.13-3.27)	0	0.26
High perception of SARS-Cov-2 vaccine	Hailemariam et al. ²⁰	4.35 (2.73–6.95)	2.70 (2.13-3.27)	0	0.20
	Nguyen et al. ²²	4.35 (2.73-6.95) 2.71 (1.93-3.82)			
	Wang et al. ²⁷				
O a la martína	wang et al.	2.48 (1.83–3.35)		0	0.67
Good practice	Mose et al. ²¹	0.15 (0.70, 10.10)	8.99 (7.42–10.56)	0	0.67
		9.15 (8.73–12.19)			
	Wang et al. ²⁷	8.27 (5.35–12.77)			
Received influenza vaccine last year			2.72 (2.09-3.35)	74.92	<0.00
	Battarbee et al. ¹⁷	2.60 (1.90-3.60)			
	Skjefte et al. ²³	3.29 (2.91–3.72)			
	Stuckelberger et al. ²⁴	3.60 (2.80–4.70)			
	Sutton et al. ²⁵	2.25 (1.66-3.05)			
	Wang et al. ²⁷	1.81 (1.18-2.80)			
Third trimester			1.35 (1.10–1.60)	0	0.78
	Stuckelberger et al. ²⁴	1.40 (1.00–2.00)			
	Tao et al. ²⁶	1.49 (1.03–2.16)			
	Wang et al. ²⁷	1.27 (0.98–1.65)			

Abbreviation: CI = confidence interval; OR = odds ratio; POR = Pooled odds ratio; $I^2 > 50\%$, heterogeneity.

Risk factors		POR (95% CI)
> 35 years		2.10 (1.10, 2.93)
Higher school education		1.84 (1.40, 2.28)
Sufficient information about the SARS-CoV-2 vaccine	— •	1.94 (0.94, 2.95)
High perception of SARS-CoV-2 vaccine	-	2.70 (2.13, 3.27)
Good practice		8.99 (7.42, 10.56)
Received influenza vaccine last year	-	2.72 (2.09, 3.35)
Third trimester	-	1.35 (1.10, 1.60)
.125	1	8

Fig. 2. Forest plots of the risk factors and pregnant women's willingness toward the SARS-CoV-2 vaccination in various countries.

mothers have the biggest effect on whether or not to vaccinate their children and other family members, it is equally critical to assess trust and the most important determinants of pregnant women's vaccination

acceptance.^{23,31}

According to the results of this investigation, there is a link between receiving influenza vaccine last year and pregnant women's acceptance

R.D. Nindrea et al.

Table 3

The findings of Egger's and Begg's tests to identify study bias.

Risk factors	Study bias		
	Egger's test	Begg's test	
>35 years	0.766	0.857	
Higher school education	0.934	0.054	
Sufficient information about the SARS-CoV-2 vaccine	0.349	0.120	
High perception of SARS-CoV-2	0.445	0.065	
Good practice	0.890	0.700	
Received influenza vaccine last year	0.530	0.132	
Third trimester	0.357	0.106	

p > 0.05, no publication bias.

of SARS-CoV-2 immunization. Women who had previously received an influenza vaccination reacted better to the SARS-CoV-2 vaccination. Women who were hesitant to obtain a SARS-CoV-2 vaccine expressed worries about vaccine safety and efficacy. Almost all pregnant women chose their obstetrician/gynecologist as their most trusted source of SARS-CoV-2 facts, with >40% choosing their gynecologist.²³

The desire to get the SARS-CoV-2 vaccine was linked to high perception. This finding is consistent with prior studies in which high perception or sensitivity was linked to acceptance and willingness COVID-19 vaccination.^{32,33} Furthermore, women who saw COVID-19 as a danger among their acquaintances were more likely to get the immunization than others.

Our research also found that as people become older, they are more likely to develop chronic conditions including hypertension, renal disease, and heart disease, which can lower a pregnant woman's immunity and raise the risk of SARS-CoV-2 related morbidity and death. As a result, it may instill fear in the elderly population, leading to a greater willingness to receive the SARS-CoV-2 vaccination.²¹

When compared to pregnant women who had limited understanding about SARS-CoV-2, those who had high information were more likely to receive SARS-CoV-2 vaccine. This might be explained by the fact that pregnant women with good understanding of SARS-CoV-2 would be aware of the virus's severity to themselves and their fetus, allowing them to readily receive SARS-CoV-2 vaccination to mitigate the pandemic's effects.³⁴

Higher-educated pregnant women were more likely to wish to

acquire the SARS-CoV-2 vaccine. Previous research have found a significant level of concern against the SARS-CoV-2 immunization among the less educated. This might be because more educated people have easier access to vaccination facts and are better able to interpret facts about the SARS-CoV-2 vaccine's advantages and risks. On the contrary, vaccination misinformation is more likely to affect persons who are less informed.³⁵

We revealed that being pregnant in the second trimester was a negative predictor of SARS-CoV-2 vaccination uptake when compared to the third trimester, indicating a potential concern of caused fetal abnormalities. Fear of any potential detrimental negative consequences of the vaccination on their pregnancy or newborn, as well as worries about safety and advantages, have been identified as important factors for vaccine aversion in various studies.^{17,36}

The essential battle against the pandemic is to figure out what factors influence pregnant women's willingness to get COVID-19 vaccines.²⁰ As more information on the safety and effectiveness of the COVID-19 immunization becomes known, immunization acceptance, perceived motivators, and barriers to acceptance may change among pregnant women.³⁷ Acceptance of the COVID-19 vaccination and its determinants among pregnant women differs throughout the world. As a result, vaccination efforts targeting this community should be tailored to the needs of each country in order to have the most effect.^{38,39}

Our study's strength was performed in various countries. As a result, this is a good setting for investigating potential relationships between pregnant women's willingness toward the SARS-CoV-2 vaccination among LMICs and non-LMICs. There are a few limitations in this metaanalysis study. Three publications appeared to be acceptable in this study, but they lacked sufficient evidence and yielded negligible data estimation findings. The possibility of selection bias will be exacerbated as a result of this issue. Furthermore, we excluded articles written in other languages. This may limit epidemiological data from nations that do not speak English.

The findings suggest that health authorities should establish immediate promotion of health programs and distribute more accurate information. Authorities should take steps to ensure that individuals have access to sufficient facts, adopt positive vibes, and hold high perception about SARS-CoV-2 vaccinations. Another recommendation is for obstetric care specialists to make a clear advice to pregnant women to get

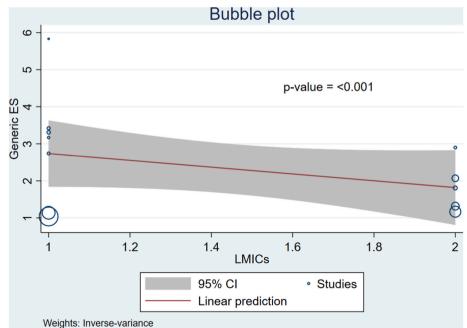


Fig. 3. The relationship between LMICs and pregnant women's desire to get vaccinated against SARS-CoV-2 based on meta-regression.

the vaccination. This will likely enhance COVID-19 vaccine uptake. Apart from that, community-based engagement activities may be necessary to adapt pregnant women education materials and increase communication and shared decision-making in order to accomplish universal health coverage in pregnant immunization.

5. Conclusion

This finding results revealed the risk factors for pregnant women's willingness toward the SARS-CoV-2 vaccination, good practice has the highest risk, followed by received influenza vaccine last year, high perception, >35 years, sufficient information, higher school education and third trimester. The heterogeneity analysis revealed homogenous among risk factors in >35 years, high perception, good practice, and third trimester. The insights of this research might help the authorities in determining the most effective strategy to deploy SARS-CoV-2 mass immunization campaigns for pregnant women.

Funding/financial support

This research received no funding from public, private, or non-profit sources.

Declaration of competing interest

The authors declare that they have no competing interests.

Acknowledgments

The authors would like to thank Riyani Betri Novialita, MA, for her help with data collecting.

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