THE EVIDENCE

Meta-Analysis

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Prevalence of cardiac arrest during pregnancy: a systematic review and metaanalysis

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Abstract

Background: Cardiac arrest during pregnancy represents a critical medical emergency, posing substantial risks to the health of the expectant mother as well as the unborn child. To estimate the worldwide prevalence of cardiac arrest during pregnancy this study was planned.

Methods: Multiple databases, including "PubMed, Cochrane, and Embase", were searched until April 7, 2023, exclusively in English, to ensure a comprehensive selection of relevant studies. A stringent two-stage screening process was employed, evaluating titles, abstracts, and full-text reviews based on the research question, "What is the prevalence of cardiac arrest in women during pregnancy?" Two researchers independently gathered pertinent information from the chosen studies. This data included the names of the authors, years of publication, the design of each study, occurrences of cardiac arrest, and the total count of pregnant participants involved. The third author resolved the conflict. The Random Effects Model (REM) computed the pooled cardiac arrest prevalence. Heterogeneity was assessed via the I² statistic.

Results: The systematic search of the database resulted in the identification of 1246 articles. Out of these, 18 studies were eligible for the systematic review, and ultimately, 17 of them were eligible for the meta-analysis. The pooled cardiac arrest prevalence during pregnancy was estimated at 0.08% (95% CI 0.03-0.19). The PI was <0.001-5.03%. However, the heterogeneity remained high (I^2 =100%). Sub-group analysis disclosed varying prevalence across continents, with Asia displaying the highest rate [(2.76%) (95% CI 1.04-7.12].

Conclusions: The study reveals significant cardiac arrest prevalence in pregnancy, with increasing rates over time and variations by continent. Maternal mortality concerns persist, particularly in Asia.

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Evidence in Context

• Cardiac health during pregnancy is critical, significantly contributing to maternal mortality. • Cardiac arrest affects 1 in 12,000 to 1 in 36,000 pregnant women, varying by geography. • Both mother and fetus face severe health risks and high mortality. • Global studies show varying prevalence and outcomes. • Systematic research is needed to address data gaps.

To view Article





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Keywords: cardiac arrest, maternal mortality, meta-analysis, pregnancy, cardiovascular disease, antenatal, intrapartum, myocardial infarction, angina, evidence synthesis

Introduction

Cardiac health in pregnancy is a significant problem since it involves a complicated interplay between obstetric and cardiovascular care of the mother and foetus. Since the symptoms of the disease and typical physiological changes are similar, diagnosing and treating heart disease during pregnancy can be challenging. Cardiovascular disease is the main contributor to indirect maternal mortality, accounting for almost 33% of deaths of pregnant women [1-6]. Furthermore, the risk varies depending on the underlying cardiac condition, with maternal heart disease occurring as a complication in approximately 4% to 16% of pregnancies in women who have previously diagnosed with cardiac disorders [7-9]. According to estimates, managing cardiovascular diseases could prevent up to 68% of pregnancy-related deaths [1,10]. One of the most difficult clinical situations is cardiovascular arrest during pregnancy. There are significant and distinct differences between resuscitating an adult and a pregnant person, even though they share many traits. A key distinction in cases of cardiac arrest during pregnancy is that there are two patients involved: the mother and the fetus [11]. First, resuscitation within the "golden hour" commonly goes unperformed due to subpar pre-hospital care, a lack of experienced professionals, and improper inter-hospital transfers. However, the results of polytrauma in pregnant women are frequently fatal [12]. Cardiac arrest occurring at the time of pregnancy is a grave medical emergency which might result in considerable health complications and fatality risks for both the expectant mother and her unborn child [13]. Cardiac arrest in the third trimester of pregnancy has implications for the neonatal injury and its survival if not immediately intervened [14]. Maternal cardiac arrest is an indication of severe maternal morbidity or unanticipated effects of labour and delivery that have a considerable negative impact on a woman's short- and long-term health. These conditions can result in pregnancy-related deaths if they are not promptly diagnosed and treated [15]. To facilitate the achievement of the Sustainable Development Goal (SDG) target of <30 maternal deaths per 100,000 live births (LBs) by the year 2030, it is essential to estimate and focus on this mostly preventable cause of maternal mortality [16].

National-level population-based research from the Euorpean and American countries reveals that between 1 in 12,000 and 1 in 36,000 pregnant women experience cardiac arrest [17-20]. In a recent cohort study in the United States, cardiac arrest occurred in one among every 9,000 hospital deliveries from 2017 to 2019, and nearly seven out of ten women survived the incident [21]. The cardiac arrest rate from 2017 to 2019 was 13.4 per 100,000 delivery hospitalisations. Among them, 68.6% of the patients who experienced cardiac arrest made it out of the hospital. In another study from Thailand, 23 women experienced cardiac arrest out of 89,368 deliveries from 2006 to 2015 (1:3885), with three of the arrests taking place prior to delivery (1:29,789), [22,23]. According to this study, pregnancy-related hypertension and cardiovascular conditions were the most frequent causes of cardiac arrests. Pregnancy-related cardiac arrests were more frequent than previously reported [19,24,25]. Several factors, such as those related to anaesthesia, haemorrhage, cardiovascular illness, embolism, uterine atony, and hypertension/pre-eclampsia/eclampsia, are frequently responsible for maternal cardiac arrest and mortality [24,25]. Furthermore, in a study conducted between 2015 and 2019, no cardiac arrest event was reported in the Philippine General Hospital, with 30,053 women admitted for deliveries, of whom 293 had cardiovascular disease [26]. No cardiac arrests were noted in another cohort study from the Swedish population that included 6630 pregnant women [27].

In India, 60% of the heart diseases identified for the first time during pregnancy were later determined to be rheumatic heart disease in 42% of cases and pulmonary hypertension in 33% of cases, 15% of pregnancies experienced maternal cardiac problems, with heart failure being the most prevalent (66%) of these [28].

Overall, the collective evidence from studies regarding cardiac arrest during pregnancy is limited and contradictory. This emphasises the importance of doing a systematic inquiry to determine the frequency of incidents. The contradiction in the findings of the studies makes it more essential to investigate and quantify the issue of the threat of cardiac arrest during pregnancy. This will enable better resource allocation by policymakers towards the problem of cardiac arrest in pregnancy. However, there have been previous systematic reviews (SR) and meta-analyses (MA) focusing on the management of cardiac arrest at the time of pregnancy [24,29,30]. In a systematic search undertaken by the authors in the Cochrane Library and the "International Prospective Register of Systematic Reviews (PROSPERO)", no systematic reviews were in progress or planned on the frequency of cardiac arrest in pregnant women. Therefore, the index study systematically reviewed and presented the pooled prevalence of cardiac arrest among pregnant women, globally.

Materials and Methods

The "Preferred Reporting Items for Systematic Review and Meta-analysis" (PRISMA) reporting requirements were followed during the conduct of this study [Table S1]. The study was registered in PROSPERO under registration number CRD42023410718. Informed consent and ethical approval do not apply to the index study because it did not collect data from human participants.

Data source

The research question for the present study was "What is the prevalence of cardiac arrest in pregnant women?", which was elaborated as per the criteria listed in Table S2. Three databases were utilised for the systematic search: Embase, Cochrane, and PubMed [Table S3]. "cardiac arrest," "cardiopulmonary arrest," and "pregnant women" were among the search terms. The search approach and the exact search strings used for each database are listed in Table S3. To handle citations, eliminate duplications, and coordinate the review process, we used Mendeley Desktop V1.19.5 software. After removing duplicates, the search results were exported into a Microsoft Excel spreadsheet.

Data Extraction in the index review

Two reviewers (SBS and GS) assessed the titles and abstracts of studies retrieved from the systematic search, independently. The reviewers discussed amongst themselves to come to an understanding when there was any disagreement over the study's inclusion criteria for the review of full-text articles. In instances where there was a persisting discrepancy in opinions about the eligibility of a publication, the third author (PA) was consulted to review the title and abstract for a decisive evaluation. PA decided whether to submit the article for full-text review. The same procedure was adopted for the full-text review of the studies, and the final list of studies eligible for data extraction was identified. In order to extract the data, the two reviewers (NK and DK) read the full texts of the eligible studies. The two reviewers extracted the data independently. A conference was held to discuss the disagreement among the reviewers at the end of the data extraction. The third reviewer (SBS) sorted the unresolvable disagreement. Using the PRISMA flowchart [Figure 1], the screening process and its results at various stages have been depicted.



Figure 1. PRISMA diagram indicating the systematic inclusion of studies for evidence synthesis

Data Management

For analysis, a data extraction table was created using a Microsoft Excel spreadsheet. From each article deemed eligible after the final selection process, specific details were extracted, including the first author's name, year of publication, the design of the study, the number of pregnant women who experienced cardiac arrest (the numerator), the total number of pregnant participants in the study (the denominator), and the name of the country where the study was conducted. Following the exclusion of studies that were not available, all eligible studies were methodically cataloged into an Excel spreadsheet for further analysis.

Quality Assessment

Utilising the JBI quality evaluation tools [31], two authors evaluated the quality of each study separately. A sensitivity analysis was planned following the quality assessment by excluding the studies rated with poor quality.

Data Analysis

All the required analysis for the meta-analysis was undertaken in R Studio. In order to calculate the prevalence of cardiac arrest, the number of cardiac arrest episodes in pregnant women was divided by the total number of research participants. The Random Effects Model (REM) determined the pooled prevalence of cardiac arrest using the Dersimonian-Laird (DL) method. The heterogeneity of studies was evaluated using the I^2 test. I^2 value of more than 50% was declared to have substantially high heterogeneity. A sub-group analysis was planned to explore the heterogeneity based on the continent of the study. The prediction interval (PI) was estimated to present high heterogeneity [32]. Meta-regression was planned to identify other potential reasons for heterogeneity. The Baujot plot, leave-one-out analysis, and "Graphic Display of Heterogeneity (GOSH)" plots were used to discover outliers. Sensitivity analyses were planned after leaving out poor-quality studies and outliers. Doi plot was employed to evaluate publication bias. Publication bias was quantified using the LFK index [32]. Other appropriate graphs, including forest plots and bubble plots, were made.

Results

Screening Process & Eligible Studies

The multiple database searches yielded a total of 1246 studies, among which 134 duplicates were identified and removed. Among the 1112 records screened for title-abstract, 30 were found eligible for full-text review. Seven articles were excluded due to the non-availability of full-text, and five were excluded due to the wrong population, based on the review of full texts.. Finally, 18 studies were included in the systematic review [18,22,33–48], and 17 studies were included in the meta-analysis [18,22,33–43,45–48]. The overall process and results of each stage of screening and eligibility are depicted in Figure 1.

Characteristics of the Eligible Studies

The studies included were undertaken between 2009 and 2022 in countries across the world. The majority of studies were from the USA (8, 44.4%), while three studies were from Canada and one study each from Namibia, Thailand, Poland, India, North African countries, Sweden, and China. The sample sizes of the studies included in the analysis varied widely, ranging from as few as 64 participants to as many as 58,784,013 [Table 1]. There was high heterogeneity between included studies, which is inferred from the I² (100%) [Figure 2a].

Table 1. Basic characteristics of the studies included for analysis (K=18)

S. no.	Author	Year of publication	Study design	Country	Events	Sample size
1	Lopez et.al	2022	Cohort	U.S.A	75	6120

2	Marschner et.al	2022	Cohort	U.S.A	82	74510
3	Nivatpumin et.al	2021	Cohort	Thailand	23	89368
4	Pfalleret.al	2020	Cohort	Canada	3	335
5	Köcheret.al	2020	Cohort	Sweden	0	6812
6	Dongarwar et.al	2020	Cross sectional	U.S.A	117568	58784013
7	Pierce-Williams et.al	2020	Cohort	U.S.A	1	64
8	Easter et.al	2020	Cohort	U.S.A	1	276
9	Balkiet.al	2017	Cohort	Canada	286	3568597
10	Grotegut et.al	2014	Cohort	U.S.A	853	12628746
11	Li et.al	2009	Cohort	U.S.A	8	136422
12	Lipczynska et al.	2021	Cohort	Poland	1	140
13	Kumar et al.	2020	Cohort	India	1	205
14	Ballali et al.	2018	Cohort	North Africa	3	80
15	Lima et al.	2016	Cohort	U.S.A	501	7850381
16	Liu et al.	2012	Cohort	China	2	1741
17	Balint et al.	2010	Cohort	Canada	1	318
18	Heemelaar et al.	2022	Cohort	Namibia	1	65
18	Heemelaar et al.	2022	Cohort	Namibia	1	65



Figure 2. Forest Plots showing overall effect size of the a) proportion of cardiac arrest during pregnancy; b) proportion of cardiac arrest during pregnancy after excluding poorquality studies

Quality Assessment

Of the studies that were deemed appropriate for meta-analysis, 14 were deemed to be of good or fair quality, and three were of poor quality [Table S4] [35,36,40].

Pooled Prevalence of the Cardiac Arrest Among Pregnant Women

There were 83,141,381 pregnant women included in the meta-analysis, among whom 119,410 events of cardiac arrest were reported. The prevalence of cardiac arrest varied from 0.005% to 1.56%. The pooled prevalence of cardiac arrest was estimated to be 0.12%

(95% CI 0.05-0.30). The PI ranged from <0.001 to 8.83% [Figure 2a]. A sensitivity analysis undertaken by excluding the poor-quality studies revealed a slightly lower prevalence of cardiac arrest [0.08% (95% CI 0.03-0.19)] [Figure 2b]. The PI was <0.001-5.03%. However, the heterogeneity remained high (I^2 =100%).

Outliers and Sensitivity Analysis

Baujat Plot and influence diagnostics revealed no outliers [Figures S1 and S2]. Leave-one-out analysis also revealed no significant changes in the pooled prevalence of cardiac arrest and the heterogeneity between the studies [Figure S3].

Exploring the Heterogeneity-Meta-regression and Sub-group Analysis

Meta-regression revealed that the sample size of the study did not have any significant impact on the outcome (cardiac arrest) (beta=0, p=0.631) [Figure 3a]. However, the bubble plot between the year of study and the prevalence of cardiac arrest showed a significant association (beta=0.273, p=0.014), indicating an increase in the prevalence of cardiac arrest over the period [Figure 3b]. Sub-group analysis according to geography did not reduce the heterogeneity. However, it revealed a significant difference in the prevalence of cardiac arrest across the continents. Studies from Asia reported the highest cardiac arrest prevalence among pregnant women [2.76% (95% CI 1.04-7.12], while studies from Africa reported the least [0.05% (95% CI 0.01-0.28]. Studies from North America reported a prevalence of 0.08 (95% CI 0.03-0.22) [Table 2].



Bubble plot demonstrating meta-regression based upon sample size b) Bubble plot demonstrating meta-regression based upon year of study

Figure 3. Bubble plot illustrating the influence of a) sample size on the percentage of cardiac arrests during pregnancy b) year of study on prevalence of cardiac arrest during pregnancy

Table 2. Subgroup analysis for cardiac arrest among pregnant women based on geography (continents)

Subgrouping variable	Number of studies	Pooled estimate	Confidence interval	I ²	P - value
Continent					
North America	11	0.08	0.03-0.22	99.9%	
Asia	3	2.76	1.04-7.12	83.0%	P < 0.01

Africa	2	0.05	0.01-0.28	0%
Europe	1	0.71	0.10-4.89	-

Publication bias

Assessment of publication bias revealed an asymmetrical Doi plot [Figure 4]. The LFK index was 1.41, indicating a publication bias towards studies reporting a higher prevalence of cardiac arrest.



Figure 4. Doi plot for assessment of publication bias

Discussion he Evi 2023:01(02)

In the index review, 17 studies were included, in which 119,410 events of cardiac arrest were reported among 83,141,381 pregnant women. Our result shows a high prevalence of cardiac arrest

during pregnancy, with a pooled prevalence of 0.08% (95% CI 0.03-0.19). This reflects the fact that the risk of maternal morbidity due to cardiac arrest is high in pregnant women. The number of high-risk pregnancies and the frequency of serious pregnancy problems, including cardiac arrest, are both on the rise [49-51]. The current analysis shows that the prevalence of cardiac arrest has significantly increased over the period. This approach mirrors that of a previous study which detailed the evolving characteristics of delivery and postpartum hospitalizations in individuals with chronic heart disease, spanning the years 1995 to 2006 [50]. The odds ratios for cardiac arrest complications were 35.7 in 1995-1997, which increased to 82.3 in 2004-2005 for deliveries hospitalised with a chronic heart disease compared to deliveries hospitalised without a chronic heart disease [50]. In a recent study, approximately one in 9,000 women experienced a cardiac arrest during labour and delivery from 2017 to 2019, compared to one in 12,000 from 1998 to 2011 [21]. This suggests that cardiac arrest during labour and delivery has become increasingly frequent in recent years. On the other hand, the overall survival rate increased over time, going from 58.9% after a cardiac arrest during childbirth in a hospital between 1998 and 2011 to 68.6% in the period from 2017 to 2019 [21]. This improvement in the survival rate could be attributed to the advancements in the management of cardiac arrest in pregnancy over the period. In a different study, cardiac arrest accounted for 16.3% of all sudden maternal deaths, while the overall maternal mortality ratio was 1.7 per 100,000 LBs [52]. Similar to this, 6.3 cardiac arrests during pregnancy or right after delivery per 100,000 births (1:16 000) and 2.8 cardiac arrests during pregnancy (1:36 000) per 100,000 natalities were recorded in a study, with nearly 60% of the women surviving [19].

The cardiovascular-related outcomes, including cardiac arrest during pregnancy, can occur as a result of a number of medical disorders that either existed prior to pregnancy or developed during pregnancy. The present meta-analysis examined studies involving both pregnant women with and without comorbid conditions. Obstetric causes of cardiac arrest include haemorrhage, pregnancy-induced hypertension, peripartum cardiomyopathy, anaesthesia complications, and amniotic fluid embolism, while non-obstetric causes include pulmonary embolism, sepsis, stroke, myocardial infarction and trauma [17,49,53,54]. A study from India revealed that serious maternal problems

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And mortality are substantially related to hypertensive disorders of pregnancy, particularly preeclampsia and eclampsia [35]. Another study identified a high rate of maternal mortality following cardiac arrests that occurred during hospitalization for childbirth. The most frequently cited causes for these arrests were pregnancy-induced hypertension and cardiovascular complications, accounting for 30.4% and 21.7% of cases, respectively. Maternal fatalities in peripartum cardiac arrests occurred in 20 of 23 cases, accounting for 86.9% of deliveries [22].

The subgroup analysis revealed a significant difference in the prevalence of cardiac arrest across the continents. Studies from Asia reported the highest cardiac arrest prevalence among pregnant women [2.76% (95% CI 1.04-7.12]. The high occurrence of cardiac arrest in Asian countries in this study could be explained by the high maternal mortality ratio in Southern Asia countries, which account for approximately 87% (along with Sub-Saharan Africa) of the expected global maternal fatalities in 2020 [55]. Maternal mortality is notably high in three countries outside of Sub-Saharan Africa: Afghanistan, Yemen, and Haiti. These high-maternal-mortality locations and countries are also known to have a high prevalence of peripartum cardiomyopathy [56], which can relate to the high prevalence of cardiac arrest cases in Asian countries. However, studies from North America reported a prevalence of 0.08 (95% CI 0.03-0.22). Compared to developed nationa, maternal mortality rates are much higher in developing nations [57]. An earlier study discovered a significant relationship between the maternal mortality ratio and socioeconomic characteristics, healthcare system-related factors, morbidity burden, and their complex relationships [58].

Limitations

The findings of the index MA needs to be interpreted with caution owing to the presence of publication bias towards research reporting a higher rate of cardiac arrest. The inherent limitations of both the original studies included in the review must also be taken into consideration. Only three databases ("PubMed, Cochrane, and Embase") could be included in the SR and MA due to resource limitations (accessibility concerns).

Conclusion

The study concludes that the prevalence of cardiac arrest is high among the pregnant women. The prevalence of cardiac arrest has increased over the period, and a spatial variation also exists between the continents. Maternal mortality is a cause of concern, especially in African and Asian countries. Thus, policies and initiatives to improve maternal health and lower maternal mortality, particularly in Asian nations, must focus on the determinants of cardiac arrest among pregnant women, such as population dynamics, socioeconomic influence, and healthcare system characteristics that pose a substantial risk to mothers.

Supporting information

None

Ethical Considerations

None

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Author contribution statement

Shashi B Singh: Conceptualization (Equal); Data Curation (Equal); Methodology; Writing- Original Draft Preparation (Equal). P Aparnavi: Conceptualization (Equal); Data Curation (Equal); Singh et al., (2023): Cardiac arrest during pregnancy meta-analysis

> Methodology; Writing- Original Draft Preparation (Equal). **Geetu Singh:** Data Curation (equal); Formal Analysis (supporting); Project Administration (lead); Writing- Review & Editing (lead). **Dewesh Kumar:** Data Curation (supporting); Formal Analysis (supporting); Writing- Review & Editing (supporting). **Asha Kiran:** Data Curation (supporting); Formal Analysis (supporting); Writing- Review & Editing (supporting). **Nisha Kumari:** Data Curation (supporting); Formal Analysis(supporting); Validation (lead); Writing- Review & Editing (supporting). **Pravin Yannawar:** Data Curation (supporting); Formal Analysis (supporting); Writing- Review & Editing (supporting). **Sandip Kumar:** Data Curation (supporting); Formal Analysis (supporting); Writing- Review & Editing (supporting). **Arvind Kumar:** Data Curation (supporting); Formal Analysis (supporting); Formal Analysis (supporting); Writing- Review & Editing (supporting). **Nitika Keshri:** Data Curation (supporting); Formal Analysis (supporting); Formal Analysis (supporting); Formal Analysis (supporting); Writing- Review & Editing (supporting). **Nitika Keshri:** Data Curation (supporting); Formal Analysis (supporting); Formal Analysis (supporting); Writing- Review & Editing (supporting).

> All authors attest they meet the ICMJE criteria for authorship and gave final approval for submission.

Data availability statement

Data included in article/supp. material/referenced in article.

Additional information

No additional information is available for this paper.

Declaration of competing interest

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

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