

Impact of emergency point of care ultrasound on time to diagnosis and treatment amongst patients with ectopic pregnancy: A systematic review and meta-analysis

Keywords

Meta-Analysis, Emergency Department, Ectopic Pregnancy, Point-of-care Ultrasound

Abstract

Introduction

Ectopic pregnancy, a significant cause of morbidity in women of reproductive age, requires timely diagnosis and treatment to reduce adverse outcomes. This study examines the impact of point-of-care ultrasound in the emergency department on the time to diagnosis, treatment, and surgical intervention for ectopic pregnancies.

Material and methods

Our review encompassed studies involving patients with ectopic pregnancies who underwent point-of-care ultrasound in emergency settings. The comparator was standard radiology department ultrasound. We included studies of any design and the key outcomes were time to diagnosis, treatment, and operating room. Data synthesis employed a random-effects inverse-variance model. The GRADE approach was used to assess evidence quality, and the PRISMA checklist guided our methodology.

Results

Overall, four studies were included. For time to diagnosis, the pooled Standardized Mean Difference (SMD) was -1.965, indicating a significant reduction. For time to treatment, the pooled SMD was -0.809, showing a modest effect with considerable heterogeneity ($I^2 = 87.5\%$). In surgically treating ectopic pregnancies, the impact was not statistically significant overall, but significant for the subgroup of ruptured ectopic pregnancies. The overall quality of evidence was deemed very low due to moderate risk of bias, high heterogeneity, and publication bias.

Conclusions

Point-of-care ultrasound in emergency departments significantly reduced the time to diagnose ectopic pregnancies. The effect on time to treatment and operating room varied, with notable benefits observed in more severe cases. Despite these promising findings, the quality of evidence and high heterogeneity necessitate cautious interpretation and call for further research.

Impact of emergency point of care ultrasound on time to diagnosis and treatment amongst patients with ectopic pregnancy: A systematic review and meta-analysis

Yuanyuan Ren ¹, Xupeng Shao ^{2,3,#,*}

¹ Gynecology department, Affiliated Hospital of Shandong University of Traditional Chinese Medicine, No. 16369 Jingshi Road, Jinan City, Shandong Province, 250014, China.

² Emergency Department, Affiliated Hospital of Shandong University of Traditional Chinese Medicine, No. 16369 Jingshi Road, Jinan City, Shandong Province, 250014, China.

³ First Clinical Medical College, Shandong University of Traditional Chinese Medicine, No. 16369 Jingshi Road, Jinan City, Shandong Province, 250014, China.

Yuanyuan Ren and Xupeng Shao contributed equally to the study.

***Corresponding author:**

Dr. Xupeng Shao,

Emergency Department, Affiliated Hospital of Shandong University of Traditional Chinese Medicine, No. 16369 Jingshi Road, Jinan City, Shandong Province, 250014, China.

First Clinical Medical College, Shandong University of Traditional Chinese Medicine, No. 16369 Jingshi Road, Jinan City, Shandong Province, 250014, China.

Email: 1021358002@qq.com

Abstract

Background

Ectopic pregnancy, a significant cause of morbidity in women of reproductive age, requires timely diagnosis and treatment to reduce adverse outcomes. This study examines the impact of point-of-care ultrasound in the emergency department on the time to diagnosis, treatment, and surgical intervention for ectopic pregnancies.

Methods

Our review encompassed studies involving patients with ectopic pregnancies who underwent point-of-care ultrasound in emergency settings. The comparator was standard radiology department ultrasound. We included studies of any design and the key outcomes were time to diagnosis, treatment, and operating room. Data synthesis employed a random-effects inverse-variance model. The GRADE approach was used to assess evidence quality, and the PRISMA checklist guided our methodology.

Results

Overall, four studies were included. For time to diagnosis, the pooled Standardized Mean Difference (SMD) was -1.965, indicating a significant reduction. For time to treatment, the pooled SMD was -0.809, showing a modest effect with considerable heterogeneity ($I^2 = 87.5\%$). In surgically treating ectopic pregnancies, the impact was not statistically significant overall, but significant for the subgroup of ruptured ectopic pregnancies. The overall quality of evidence was deemed very low due to moderate risk of bias, high heterogeneity, and publication bias.

Conclusion

Point-of-care ultrasound in emergency departments significantly reduced the time to diagnose ectopic pregnancies. The effect on time to treatment and operating room varied, with notable benefits observed in more severe cases. Despite these promising findings, the quality of

evidence and high heterogeneity necessitate cautious interpretation and call for further research.

Keywords: Ectopic Pregnancy; Emergency Department; Meta-Analysis; Point-of-care Ultrasound

Introduction

Ectopic pregnancy, a potentially life-threatening condition characterized by the implantation of a fertilized ovum outside the uterine cavity, remains a significant concern in obstetrics.[1]

Tubal pregnancies are the most common, but interstitial and cervical ectopic pregnancies can be more difficult to diagnose due to their atypical locations and overlapping symptoms with other conditions.[2] Abdominal ectopic pregnancies are rare and often diagnosed late, posing significant risks due to the potential for severe hemorrhage.[3] The variability in presentation and location of ectopic pregnancies necessitates a high degree of clinical suspicion and the use of advanced diagnostic tools to ensure accurate and timely identification.[4]

Early diagnosis of heterotopic and cornual pregnancies through ultrasonography is crucial for optimal patient management. Heterotopic pregnancies, which involve simultaneous intrauterine and ectopic pregnancies, pose a significant diagnostic challenge and can be life-threatening if not identified promptly.[5] Utilizing advanced ultrasonography techniques, including Point of Care Ultrasound (POCUS), allows for early detection of these complex cases, enabling timely and appropriate clinical interventions that can prevent severe complications and improve patient outcomes.

Despite advances in diagnostic modalities, the timely and accurate diagnosis of ectopic pregnancy remains challenging, often leading to delays in treatment and increased morbidity.[6] In recent years, the advent of POCUS has revolutionized emergency medicine, offering rapid, bedside diagnostic capabilities.[7] POCUS, with its real-time imaging, provides

an invaluable tool in the early detection of ectopic pregnancies, particularly in emergency settings where prompt diagnosis is crucial.[8] Its role in reducing time to diagnosis and subsequent treatment initiation is increasingly being recognized.[8]

While both POCUS and traditional transvaginal ultrasound (TVUS) are pivotal in diagnosing ectopic pregnancies, they differ significantly in terms of execution, accessibility, and application. Traditional TVUS, typically conducted by a radiologist or a specialized gynecologist, involves detailed imaging of the pelvic region and requires more sophisticated equipment and longer examination times.[8,9] In contrast, POCUS can be performed at the bedside by emergency physicians or other trained healthcare providers, offering immediate, real-time imaging that facilitates rapid clinical decision-making (**Table 1**).

Experienced operators and high-quality equipment generally yield lower false positive rates, while resource-limited settings and less experienced users may see higher rates. Thus, while POCUS is a valuable tool for rapid diagnosis, its accuracy can be affected by these variables, underscoring the importance of proper training and equipment quality.[9,10] In POCUS diagnosis, handheld or portable ultrasound devices are typically used, differing from the stationary, high-resolution machines found in traditional ultrasound settings. Portable devices are designed for quick, bedside assessments and can be operated by various healthcare providers, while traditional devices require more advanced imaging capabilities and are often managed by specialists. These differences can affect results, as portable devices may have lower image resolution and limited functionality compared to their traditional counterparts, potentially impacting diagnostic accuracy, especially in complex cases.[8-13]

Numerous studies have investigated the efficacy of POCUS in the emergency diagnosis of ectopic pregnancy. A study by Stone et al. (2021) demonstrated the effectiveness of POCUS in reducing the time to treatment and operating room in emergency departments.[14] However,

the literature is varied, with some studies presenting contrasting findings on the accuracy and reliability of POCUS in different clinical settings.

Early detection and treatment can prevent complications such as tubal rupture, which is associated with severe haemorrhage and can be life-threatening.[15-17] Moreover, the psychological impact on patients undergoing a suspected ectopic pregnancy cannot be understated. The anxiety and distress associated with prolonged diagnostic processes can be alleviated through the expedited clarity that POCUS provides.[18,19]

The significance of this study lies in its potential to inform clinical practice and policy-making in obstetrics and emergency medicine. By providing evidence-based insights, it aims to contribute towards improving patient outcomes in ectopic pregnancy management, potentially guiding future research and clinical protocols. Hence, this study aims to synthesize existing literature to provide a clearer understanding of the impact of POCUS on the time to diagnosis and treatment in patients with ectopic pregnancy.

Methods

Eligibility Criteria

Inclusion criteria: The review included the studies satisfying all the following characteristics:

- (1) Population: patients with ectopic pregnancy;
- (2) Intervention: POCUS at the emergency department;
- (3) Comparison: standard or usual ordered radiology department ultrasound;
- (4) Outcome: time to diagnosis (from the time of arrival to the hospital till the point of diagnosis), time to treatment (from the time of arrival to the hospital till time of initiation of the treatment) and time to operating room (from the time of arrival to the hospital till the time of arrival to operating room).

(5) Study design: Studies of all the designs i.e., randomized controlled trials (RCTs), observational studies like case-control, cohort or cross-sectional studies) were considered.

Exclusion criteria: Studies were excluded if they did not provide sufficient data for the calculation of effect size or were published as abstracts only.

Search Strategy

We conducted comprehensive searches in various electronic databases, including PubMed, Scopus, the Cochrane Central Register of Controlled Trials (CENTRAL), ScienceDirect and Google Scholar. To enhance our search, we manually examined the reference lists of relevant reviews and studies that we included. In cases where additional unpublished data or clarifications were needed, we reached out to the authors of these studies. Our search focused on terms related to "ectopic pregnancy", "point of care ultrasound", and "emergency department", utilizing both medical subject headings (MeSH) and corresponding keywords suitable for each specific database. Our search strategy did not impose any restrictions regarding the language of publication or the date of publication.

Study Selection Process

The initial screening of titles and abstracts from the collected studies was carried out independently by two reviewers. Following this, they acquired and thoroughly assessed the full texts of studies that appeared to meet the eligibility criteria. In instances of disagreement between the reviewers, they resolved these through discussion, or by consulting a third reviewer when needed.

Data Collection Process

For the process of data collection, two reviewers extracted information separately using a pre-defined data extraction template. This included extracting details such as the study's author, the journal, the year of the study and its publication, and the study design. Additionally, they gathered data on the characteristics of the participants, such as the total sample size and sample

sizes in each study group, the average or median age, the distribution of genders, and whether the ectopic pregnancies were ruptured or unruptured. Further data on the specifics of the POCUS and its comparator group, the time taken for diagnosis, treatment, or transfer to the operating room, and information vital for evaluating the risk of bias were also collected. Finally, any data regarding the funding sources for the studies and any potential conflicts of interest were noted.

Risk of Bias in Individual Studies

Two investigators took on the task of evaluating the quality of the studies included. They used the Newcastle Ottawa scale (NOS) for observational studies,[20] which includes three criteria: selection of study groups (rated from 0 to 4 stars), comparability of the groups (0 to 2 stars), and the determination of exposure or outcome (0 to 3 stars). Studies scoring seven stars or more were deemed high quality. The researchers resolved any differences in opinion through discussion to reach a consensus. In cases where consensus was not possible, a third opinion was sought for a decision.

Data Synthesis

The meta-analysis was performed for pooling the data between the studies. Random-effects model with inverse variance method was used to account for the heterogeneity.[21] The mean, standard deviation (SD) and sample size was entered for patients in both the groups. The measures of effect utilized for interpretation was standardized mean difference (SMD) as all the outcomes are continuous. Statistical heterogeneity was assessed using the chi square test and I^2 statistic. However, these are relative measures of heterogeneity and hence, prediction interval was reported along with the confidence interval for the pooled estimate. This interval provides a range in which the true effect size for individual settings or populations is expected to fall, considering the heterogeneity among the studies.

Subgroup analyses were performed based on the rupture status of ectopic pregnancies. Sensitivity analyses were done to assess the robustness of the results. Doi plot and Luis Furuya-Kanamori (LFK) index was utilized to check the publication bias for all the outcomes.[22] The Doi Plot visualizes asymmetry in meta-analyses, while the LFK Index quantifies this asymmetry. A high LFK index may indicate potential publication bias. We also checked for selective reporting within studies by comparing the reported outcomes with those listed in the study protocols or trial registries.

We used the Grading of Recommendations Assessment, Development and Evaluation (GRADE) approach to assess the quality of the evidence for each outcome.[23] GRADE (Grading of Recommendations Assessment, Development and Evaluation) is a systematic approach to grading the quality of evidence and the strength of recommendations. It considers several factors:

Risk of Bias: Evaluates the reliability of the study design.

Consistency: Looks for similarity in the study results.

Directness: Assesses the direct application of the study results to the research question.

Precision: Examines the certainty or the degree of variability in the effect estimates.[24]

Publication Bias: Identifies the likelihood of selective publication of studies.

This entire review and analysis was done in accordance with the PRISMA checklist.[25]

Results

Search results

Overall, after searching all the databases, 932 studies were obtained, and they underwent duplicate removal. Then, primary screening of title and abstract was done and then 51 full-texts were retrieved. After screening these full-texts with eligibility criteria, four studies satisfied all the aspects and included in the review and then meta-analysis (**Figure 1**).[14,26-28]

Characteristics of the included studies

Table 2 contains the characteristics of the included studies. All the studies were conducted in United States of America and were retrospective review of records. Sample size ranged from 10 to 38 in POCUS group and 21 to 73 in control group. Two studies were conducted exclusively amongst ruptured ectopic pregnancy patients. Mean age of the patients ranged from 28.6 to 31.2 years. All the studies (except Stone 2021 – low risk of bias)[10] had moderate risk of bias (**Table 3**).

Impact of POCUS on time to diagnosis of ectopic pregnancy in emergency setting

In this meta-analysis of two studies with a total of 69 participants, the pooled SMD using the random-effects inverse-variance model was -1.965 (95% CI: -2.562 to -1.368), indicating a significant overall effect ($z = -6.453$, $p < 0.001$) (**Figure 2**). The heterogeneity was negligible with an I^2 of 0.0% (Cochran's $Q = 0.18$, $p = 0.674$), suggesting consistency among the included studies. These results demonstrate a substantial and uniform effect of reduction in time to diagnosis due to use of POCUS in emergency setting across the studies analysed. Additional subgroup analysis, prediction interval estimation, sensitivity analysis and publication bias assessment could not be performed for this outcome due to inclusion of only two studies for analysis.

Impact of POCUS on time to treatment of ectopic pregnancy in emergency setting

About four studies with a total of 266 participants have studied the impact of POCUS on time to treatment of ectopic pregnancy in emergency setting. The pooled SMD was -0.809 (95% CI: -1.602 to -0.017), indicating a modest overall effect ($z = -2.002$, $p = 0.045$) (**Figure 3**). However, there was significant heterogeneity among the studies ($I^2 = 87.5\%$; Cochran's $Q = 24.08$, $p < 0.001$), suggesting substantial variability in the effect sizes across studies. The prediction interval ranged from -4.45 to 2.84, highlighting a wide range of possible true effects.

Subgroup analysis was made to see the impact of POCUS on time to treatment of ruptured ectopic pregnancy patients alone. Three studies have reported estimates separately for ruptured ectopic pregnancy. The pooled SMD was -1.252 (95%CI: -1.834 to -0.669) with significant overall effect ($p < 0.001$) (Figure 4). Doi plot (Figure 5) showed a major asymmetry, which was further confirmed by LFK index of -2.12. This indicates the presence of publication bias. However, leave-one-out sensitivity analysis revealed that the study estimates are robust and credible (Figure 6).

Impact of POCUS on time to operating room for surgically treating the ectopic pregnancy in emergency setting

About two studies with 141 participants have reported the impact of POCUS on time to operating room for surgically treating the ectopic pregnancy in emergency setting, the pooled SMD was -1.038, but this did not reach statistical significance (95% CI: -2.276 to 0.200, $z = -1.644$, $p = 0.100$) (Figure 7). There was considerable heterogeneity observed among the studies ($I^2 = 85.2\%$; Cochran's $Q = 6.77$, $p = 0.009$), indicating notable variation in the effect sizes between the included studies.

Separate analysis including only ruptured ectopic pregnancy patients revealed that the pooled SMD was -1.110 (95%CI: -2.203 to -0.017) with significant overall effect of $p = 0.046$ (Figure 8). This indicates that the POCUS is beneficial in terms of time to operating room for at least ruptured ectopic pregnancy patients. Additional prediction interval estimation, sensitivity analysis and publication bias assessment could not be performed for this outcome due to inclusion of only two studies for analysis.

As per GRADE evidence profile, risk of bias was moderate, there was no indirectness (as separate analysis based on ruptured ectopic pregnancy was provided), but imprecision is present as the CI of SMD crossed 0.5 on either side of CI for all the outcomes, moderate to

high heterogeneity found across almost all the outcomes and publication bias was present for the only outcome it was assessed, indicating that the overall quality of evidence is very low.

GRADE evidence:

According to the GRADE evidence profile, the risk of bias was deemed moderate. There was no evidence of indirectness, as analyses were appropriately stratified based on ruptured ectopic pregnancies. However, the quality of evidence was compromised by imprecision, as indicated by the confidence intervals of the Standardized Mean Differences (SMD) extending beyond 0.5 on either side for all outcomes. Additionally, moderate to high heterogeneity was observed across nearly all outcomes. Publication bias was also detected in the one outcome where it was assessed. Collectively, these factors lead to the conclusion that the overall quality of evidence is very low.

Discussion

Our meta-analysis presents significant insights into the use of POCUS in the diagnosis and management of ectopic pregnancy in emergency settings. The analysis across various outcomes reveals a complex picture. The use of POCUS significantly reduced the time to diagnosis of ectopic pregnancies, with a pooled SMD of -1.965.

A modest but significant effect on time to treatment was observed (pooled SMD of -0.809). The notable heterogeneity ($I^2 = 87.5\%$) in these results suggests variability in implementation or effectiveness across different settings. For the overall cohort, the reduction in time to operating room was not statistically significant. However, in the subgroup of ruptured ectopic pregnancies, a significant reduction was noted. This indicates that POCUS may particularly benefit patients with more severe presentations.

Previous studies have also highlighted the utility of POCUS in emergency settings, especially for conditions requiring rapid decision-making.[11-13,29,30] Our findings align with these

studies but go a step further by quantifying the impact specifically for ectopic pregnancies.[29,30] The immediacy of POCUS likely contributes to quicker decision-making, bypassing the usual delays associated with standard radiology department scans. The emphasis of this study on the specific application of POCUS for ectopic pregnancies in emergency settings is a critical addition to the existing body of literature. Historically, the integration of ultrasound technology in emergency medicine has been a transformative development. Prior research predominantly focused on the utility of ultrasound for a range of urgent conditions, but our study provides a targeted analysis of its effectiveness specifically for ectopic pregnancy.[31] This specificity offers valuable insights into the nuanced application of ultrasound technology in emergency gynaecological care, contributing to a more comprehensive understanding of its role in different emergency scenarios.

The observed variability in the effectiveness of POCUS can be attributed to several factors. First, the emergency department workflow variations i.e., Differences in the protocols, staffing, and resources of emergency departments can significantly impact the efficiency of ultrasound use.[32] In environments where staff are trained and protocols are optimized for ultrasound use, the time to diagnosis and treatment can be notably reduced. Second, the operator expertise i.e., the skill level and experience of the ultrasound operator play a crucial role. Studies have shown that operator proficiency can significantly influence diagnostic accuracy and efficiency, impacting patient outcomes.[33] Finally, the patient demographics and presentation i.e., the factors such as the patient's age, medical history, and severity at presentation might also influence the effectiveness of POCUS.[14,34] For instance, in more acute presentations like ruptured ectopic pregnancies, the rapid diagnosis afforded by ultrasound can be particularly beneficial.

This review has certain strengths. Our study provides a comprehensive analysis across multiple relevant outcomes. The inclusion of both overall and subgroup analyses offers a nuanced view

of the utility of POCUS in varying clinical scenarios. We have also reported the prediction interval, publication bias assessment and sensitivity analysis, wherever possible. However, the study has certain limitations. The presence of heterogeneity and publication bias, particularly in the time to treatment analysis, warrants caution in interpreting these findings. The limited number of studies included, especially for certain outcomes, restricts the generalizability of our conclusions. **Small sample sizes in the included studies limit generalizability and hence, future studies should aim for larger, multi-centre trials.**

While integrating POCUS into emergency protocols may entail additional costs, these should be weighed against the potential benefits of reduced time to diagnosis and treatment, which can lead to improved patient outcomes and potentially lower overall healthcare costs. The results support the need for cross-disciplinary collaboration, ensuring that obstetrics teams are promptly informed and involved in cases of ectopic pregnancy identified in the emergency department.

The findings of this study have significant clinical implications, particularly for the management of ectopic pregnancies in emergency settings. The demonstrated reduction in time to diagnosis and initiation of treatment with the use of POCUS underscores its potential to improve patient outcomes by facilitating timely clinical decisions and interventions. This study contributes to the existing literature by providing a quantified analysis of the benefits of POCUS, particularly highlighting its efficacy in reducing diagnostic and treatment delays. These insights support the integration of POCUS into standard emergency department protocols for suspected ectopic pregnancies. Future studies should focus on optimizing POCUS training programs for emergency physicians, exploring long-term patient outcomes, and addressing the identified heterogeneity to further refine its application in clinical practice. Future studies should also aim to address the heterogeneity observed in our findings through

larger, more standardized studies. Future studies should also explore the impact of operator skill and experience on the effectiveness of POCUS.

Conclusion

The main finding of this study highlights the substantial benefit of POCUS in reducing both diagnostic and treatment delays for patients with ectopic pregnancy. By significantly shortening the time to diagnosis and initiation of treatment, POCUS not only improves clinical outcomes but also alleviates the psychological burden on patients. These benefits emphasize the need for widespread adoption of POCUS in emergency settings, reinforcing its value as a rapid, reliable, and non-invasive diagnostic tool. While the findings indicate potential improvements in time to treatment and operating room readiness, especially in more severe cases, the quality of evidence and heterogeneity suggest a need for cautious interpretation.

Declarations

Ethical approval

This study was approved by the Ethical Committee of Affiliated Hospital of Shandong University of Traditional Chinese Medicine (Approval No. 2023-077-KY).

Acknowledgments

None.

Conflict of interest disclosure

The authors declare that they have no competing interests.

Consent for publication

Not applicable.

Availability of data and materials

The datasets used and analysed during the current study are available from the corresponding author on reasonable request.

Funding

The study was supported by the Development Plan of Shandong Medical and Health Technology (Grant Number. 202117010252) and Shandong Province Medical and Health Science and Technology Project (Grant Number. 202410000997).

Author contribution statement

Yuanyuan Ren & Xupeng Shao were involved in conception and design of the study, conduct of the study, analysis and interpretation of the data, writing first draft, review and critical review of the manuscript. Both the authors approved the final version of the article.

References:

1. Mummert T, Gnugnoli DM. Ectopic Pregnancy. 2023 Aug 8. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2023 Jan-. PMID: 30969682.
2. Feng H, Zheng Y, Ke Y. Ultrasound vs. histologic findings in 40 patients with special types of ectopic pregnancy. *Am J Transl Res.* 2021 Jul 15;13(7):7829-7838. PMID: 34377260;
3. Mraihi F, Buzzaccarini G, D'Amato A, Laganà AS, Basly J, Mejri C, et al. Cornual Pregnancy: Results of a Single-Center Retrospective Experience and Systematic Review on Reproductive Outcomes. *Medicina (Kaunas).* 2024 Jan 21;60(1):186. doi: 10.3390/medicina60010186.
4. Sadłecki P, Grabiec M, Walentowicz-Sadłecka M. Broad ligament pregnancy - A rare and challenging diagnosis. *Clin Case Rep.* 2021; 9(9):e04823. doi: 10.1002/ccr3.4823.
5. Cucinella G, Gullo G, Etrusco A, Dolce E, Culmone S, Buzzaccarini G. Early diagnosis and surgical management of heterotopic pregnancy allows us to save the intrauterine pregnancy. *Prz Menopauzalny.* 2021; 20(4):222-225. doi: 10.5114/pm.2021.111277
6. Mullany K, Minneci M, Monjazebe R, C Coiado O. Overview of ectopic pregnancy diagnosis, management, and innovation. *Womens Health (Lond).* 2023; 19:17455057231160349. doi: 10.1177/17455057231160349.
7. Faruqi I, Siddiqi M, Buhumaid R. Point-of-Care Ultrasound in the Emergency Department [Internet]. *Essentials of Accident and Emergency Medicine.* IntechOpen; 2019. Available from: <http://dx.doi.org/10.5772/intechopen.74123>
8. Moake MM, Price AB, Titus MO, Barnes RM. Point-of-Care Ultrasound Facilitates Management of Ruptured Ectopic Pregnancy. *Pediatr Emerg Care.* 2021; 37(5):282-285. doi: 10.1097/PEC.0000000000002375

9. Hashim A, Tahir MJ, Ullah I, Asghar MS, Siddiqi H, Yousaf Z. The utility of point of care ultrasonography (POCUS). *Ann Med Surg (Lond)*. 2021; 71:102982. doi: 10.1016/j.amsu.2021.102982.
10. Collins K, Collins C, Kothari A. Point-of-care ultrasound in obstetrics. *Australas J Ultrasound Med*. 2019; 22(1):32-39. doi: 10.1002/ajum.12133.
11. Recker F, Weber E, Strizek B, Gembruch U, Westerway SC, Dietrich CF. Point-of-care ultrasound in obstetrics and gynecology. *Arch Gynecol Obstet*. 2021; 303(4):871-876. doi: 10.1007/s00404-021-05972-5
12. Venkatayogi N, Gupta M, Gupta A, Nallaparaju S, Cheemalamarri N, Gilari K, et al. From seeing to knowing with artificial intelligence: a scoping review of point-of-care ultrasound in low-resource settings. *Appl Sci*. 2023; 13(14):8427. doi: <https://doi.org/10.3390/app13148427>
13. Abrokwa SK, Ruby LC, Heuvelings CC, B elard S. Task shifting for point of care ultrasound in primary healthcare in low- and middle-income countries-a systematic review. *EClinicalMedicine*. 2022;45:101333. doi: 10.1016/j.eclinm.2022.101333.
14. Stone BS, Muruganandan KM, Tonelli MM, Dugas JN, Verriet IE, Pare JR. Impact of point-of-care ultrasound on treatment time for ectopic pregnancy. *Am J Emerg Med*. 2021; 49:226-232. doi: 10.1016/j.ajem.2021.05.071.
15. Lee R, Dupuis C, Chen B, Smith A, Kim YH. Diagnosing ectopic pregnancy in the emergency setting. *Ultrasonography*. 2018; 37(1):78-87. doi: 10.14366/usg.17044.
16. Oluwole AA, Ugwu AO, Omisakin SI, Adaramoye VO. Ectopic Pregnancy: A Life-threatening Gynaecological Emergency Revisited in Lagos, Southwest, Nigeria. *Nigerian J Med*. 2023; 32(2):p 113-116. doi: 10.4103/NJM.NJM_8_23

17. McGurk L, Oliver R, Odejinmi F. Severe morbidity with ectopic pregnancy is associated with late presentation. *J Obstet Gynaecol.* 2019; 39(5):670-674. doi: 10.1080/01443615.2018.1557610.
18. Ren N, Dela Rosa RD, Chen Z, Gao Y, Chang L, Li M, et al. Research Progress on Psychological Distress in Patients with Ectopic Pregnancy in China. *Neuropsychiatr Dis Treat.* 2023; 19:1633-1639. doi: 10.2147/NDT.S410320.
19. Jia L, Li W, Liu Y, Wang L. Psychologic Sequelae in Early Pregnancy Complications. *Int J Womens Health.* 2023;15:51-57. doi: 10.2147/IJWH.S382677
20. Lo CK, Mertz D, Loeb M. Newcastle-Ottawa Scale: comparing reviewers' to authors' assessments. *BMC medical research methodology.* 2014 Dec;14:1-5. doi: 10.1186/1471-2288-14-45
21. Cumpston M, Li T, Page MJ, Chandler J, Welch VA, Higgins JP, et al. Updated guidance for trusted systematic reviews: a new edition of the Cochrane Handbook for Systematic Reviews of Interventions. *Cochrane Database Syst Rev.* 2019;2019(10): ED000142. doi: 10.1002/14651858.ED000142.
22. Furuya-Kanamori L, Barendregt JJ, Doi SAR. A new improved graphical and quantitative method for detecting bias in meta-analysis. *Int J Evid Based Healthc.* 2018 Dec;16(4):195-203. doi: 10.1097/XEB.0000000000000141.
23. Basu A. How to appraise a body of evidence using GRADE system and GRADEPro: a step by step tutorial for the perplexed. *Qeios.* 2020. doi: <https://doi.org/10.32388/2C2WG8>
24. Castellini G, Bruschetti M, Gianola S, Gluud C, Moja L. Assessing imprecision in Cochrane systematic reviews: a comparison of GRADE and Trial Sequential Analysis. *Syst Rev.* 2018; 7(1):110. doi: 10.1186/s13643-018-0770-1.

25. Page MJ, McKenzie JE, Bossuyt PM, Boutron I, Hoffmann TC, Mulrow CD, et al. The PRISMA 2020 statement: an updated guideline for reporting systematic reviews. *BMJ*. 2021;372:n71. doi: 10.1136/bmj.n71
26. Urquhart S, Barnes M, Flannigan M. Comparing Time to Diagnosis and Treatment of Patients with Ruptured Ectopic Pregnancy Based on Type of Ultrasound Performed: A Retrospective Inquiry. *J Emerg Med*. 2022; 62(2):200-206. doi: 10.1016/j.jemermed.2021.07.064.
27. Rodgerson JD, Heegaard WG, Plummer D, Hicks J, Clinton J, Sterner S. Emergency department right upper quadrant ultrasound is associated with a reduced time to diagnosis and treatment of ruptured ectopic pregnancies. *Acad Emerg Med*. 2001; 8(4):331-6. doi: 10.1111/j.1553-2712.2001.tb02110.x.
28. Durston WE, Carl ML, Guerra W, Eaton A, Ackerson LM. Ultrasound availability in the evaluation of ectopic pregnancy in the ED: comparison of quality and cost-effectiveness with different approaches. *Am J Emerg Med*. 2000; 18(4):408-17. doi: 10.1053/ajem.2000.7310.
29. Pouryahya P. The utility of Point-Of-Care Ultrasound (POCUS) in emergency department; an observational study. *Ultrasound Med Biol*. 2019; 45(1):S68. doi: <https://doi.org/10.1016/j.ultrasmedbio.2019.07.639>
30. Smallwood N, Dachsel M. Point-of-care ultrasound (POCUS): unnecessary gadgetry or evidence-based medicine? *Clin Med (Lond)*. 2018; 18(3):219-224. doi: 10.7861/clinmedicine.18-3-219.
31. Baker M, dela Cruz J. Ectopic Pregnancy, Ultrasound. [Updated 2023 Jan 16]. In: StatPearls [Internet]. Treasure Island (FL): StatPearls Publishing; 2023 Jan-. Available from: <https://www.ncbi.nlm.nih.gov/books/NBK482192/>

32. Abdolrazaghnejad A, Banaie M, Safdari M. Ultrasonography in Emergency Department; a Diagnostic Tool for Better Examination and Decision-Making. *Adv J Emerg Med.* 2017 Dec 11;2(1):e7. doi: 10.22114/AJEM.v0i0.40.
33. Mulder TA, van de Velde T, Dokter E, Boekestijn B, Olgers TJ, Bauer MP, et al. Unravelling the skillset of point-of-care ultrasound: a systematic review. *Ultrasound J.* 2023; 15(1):19. doi: 10.1186/s13089-023-00319-4.
34. Casado-López I, Tung-Chen Y, Torres-Arrese M, Luordo-Tedesco D, Mata-Martínez A, Casas-Rojo JM, et al. Usefulness of Multi-Organ Point-of-Care Ultrasound as a Complement to the Decision-Making Process in Internal Medicine. *J Clin Med.* 2022; 11(8):2256. doi: 10.3390/jcm11082256.

Preprint

Figure captions:

Figure 1: PRISMA flowchart

Figure 2: Forest plot showing the impact of point-of-care ultrasound on time to diagnosis of ectopic pregnancy

Figure 3: Forest plot showing the impact of point-of-care ultrasound on time to treatment of ectopic pregnancy

Figure 4: Forest plot showing the impact of point-of-care ultrasound on time to diagnosis of ruptured ectopic pregnancy

Figure 5: Doi plot showing the asymmetry of the estimates

Figure 6: Sensitivity analysis results

Figure 7: Forest plot showing the impact of point-of-care ultrasound on time to operating room for treating ectopic pregnancy

Figure 8: Forest plot showing the impact of point-of-care ultrasound on time to operating for treating ruptured ectopic pregnancy

Impact of Emergency Point-of-Care Ultrasound on Time to Diagnosis and Treatment of Ectopic Pregnancy



- Time to diagnosis reduced by 1.965 SMD
- Time to treatment reduced by 0.809 SMD
- Significant benefits in ruptured ectopic pregnancies

POCUS significantly improves diagnostic and treatment times, especially in severe cases, but variability in evidence quality suggests need for further research

Preprint

Table 1: Key differences between Point of care ultrasound and traditional Transvaginal Ultrasound in gynecological diagnostics

Aspect	Point of care ultrasound	Traditional Transvaginal Ultrasound
Performed by	Emergency physicians, various trained providers	Radiologists, specialized gynecologists
Location	Bedside, emergency settings	Radiology department, specialized clinics
Equipment	Portable ultrasound devices	Advanced, stationary ultrasound machines
Imaging Time	Immediate, real-time imaging	Longer, detailed examination time
Resource Requirements	Minimal, suitable for resource-limited settings	Requires more resources and specialized facilities
Utility in Emergency	High, ideal for immediate decision-making	Moderate, usually follows initial assessment
Clinical Setting	Emergency departments, field hospitals	Specialized diagnostic centers, outpatient settings

Table 2: Characteristics of the studies included (N=4)

Author	Country	Study design	Study participants	Methodology	Operator skills	Sample size (POCUS)	Sample size (Control)	Outcomes assessed	Mean age (years)
Durston 2000	USA	Retrospective	Both ruptured and unruptured ectopic pregnancy patients	POCUS performed using portable ultrasound devices by emergency physicians with varying levels of training. Data included time to treatment	Emergency physicians with varying levels of training	38	50	Time to treatment	28.6
Rodgerson 2001	USA	Retrospective	Ruptured ectopic pregnancy patients	POCUS conducted using right upper quadrant ultrasound performed by experienced emergency physicians to diagnose hemoperitoneum in ruptured ectopic pregnancies	Experienced emergency physicians	16	21	Time to diagnosis and treatment	29.4

Stone 2021	USA	Retrospective	Both ruptured and unruptured ectopic pregnancy patients	POCUS involved transabdominal scans performed by trained emergency physicians, residents, and fellows. Images were reviewed before any radiology ultrasound results	Trained emergency physicians, residents, and fellows	36	73	Time to treatment and OR	31.2
Urquhart 2021	USA	Retrospective	Ruptured ectopic pregnancy patients	POCUS examinations were performed by resident, fellow, and attending physicians using transabdominal views to detect free fluid in the abdomen and pelvis	Resident, fellow, and attending physicians from novice to expert levels	10	22	Time to diagnosis, treatment and OR	29.5

POCUS- point-of-care ultrasound; NR – Not reported; USA – United States of America; OR – Operating room

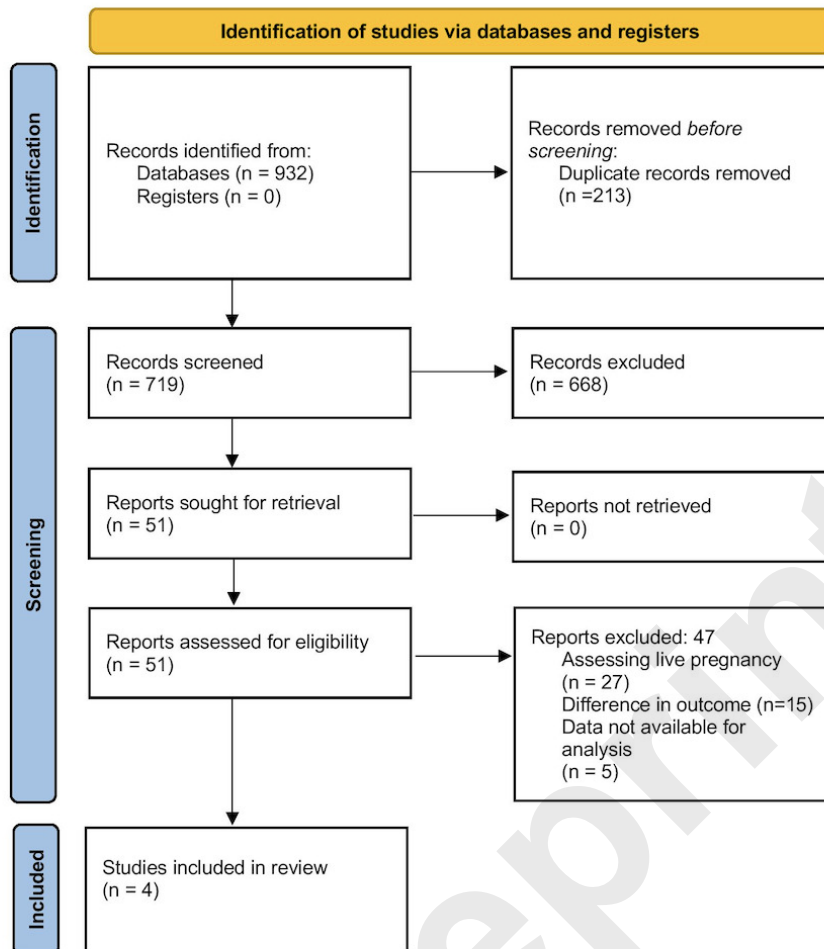
Table 3: Quality assessment of the included studies (N=4)

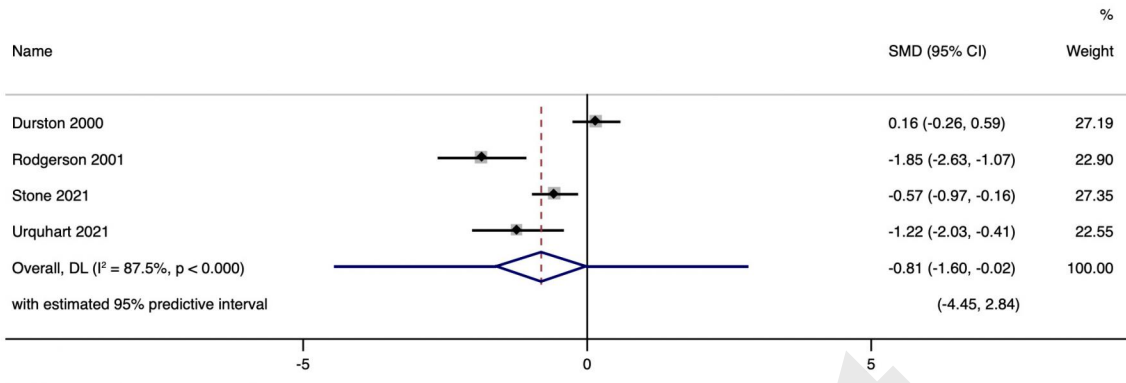
S.N.	Author and year	Representativeness	Sample size justification	Non-response	Ascertainment of exposure	Control for confounding	Assessment of outcome	Statistical tests	Overall Quality
1.	Durston 2000	0 star	0 star	1 star	1 star	0 star	1 star	1 star	4 stars (Moderate)
2.	Rodgerson 2001	0 star	0 star	1 star	1 star	0 star	1 star	1 star	4 stars (Moderate)
3.	Stone 2021	0 star	1 star	1 star	1 star	2 stars	1 star	1 star	7 stars (Low)
4.	Urquhart 2021	0 star	0 star	1 star	1 star	0 star	1 star	1 star	4 stars (Moderate)

Risk of bias rated based on the value of stars: 0-3 – High risk of bias; 4-6 – Moderate risk of bias; 7-9 – Low risk of bias

Preprint

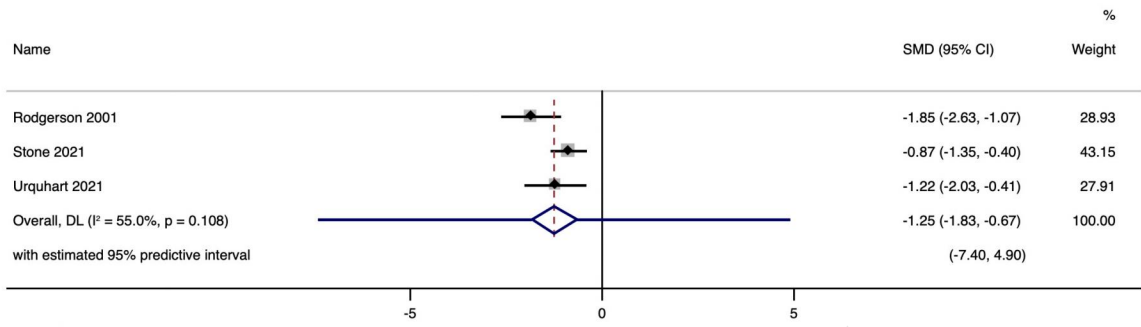
PRISMA 2020 flow diagram for new systematic reviews which included searches of databases and registers only





NOTE: Weights are from random-effects model

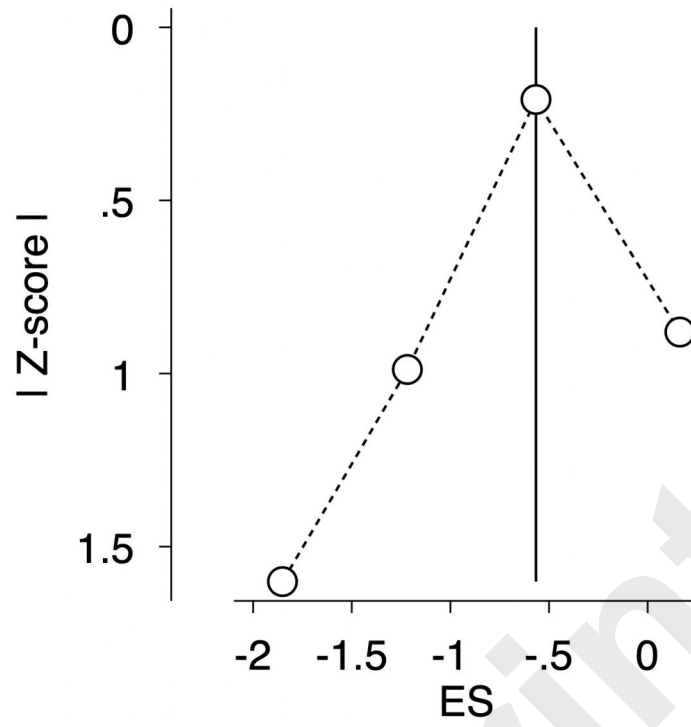
Preprint

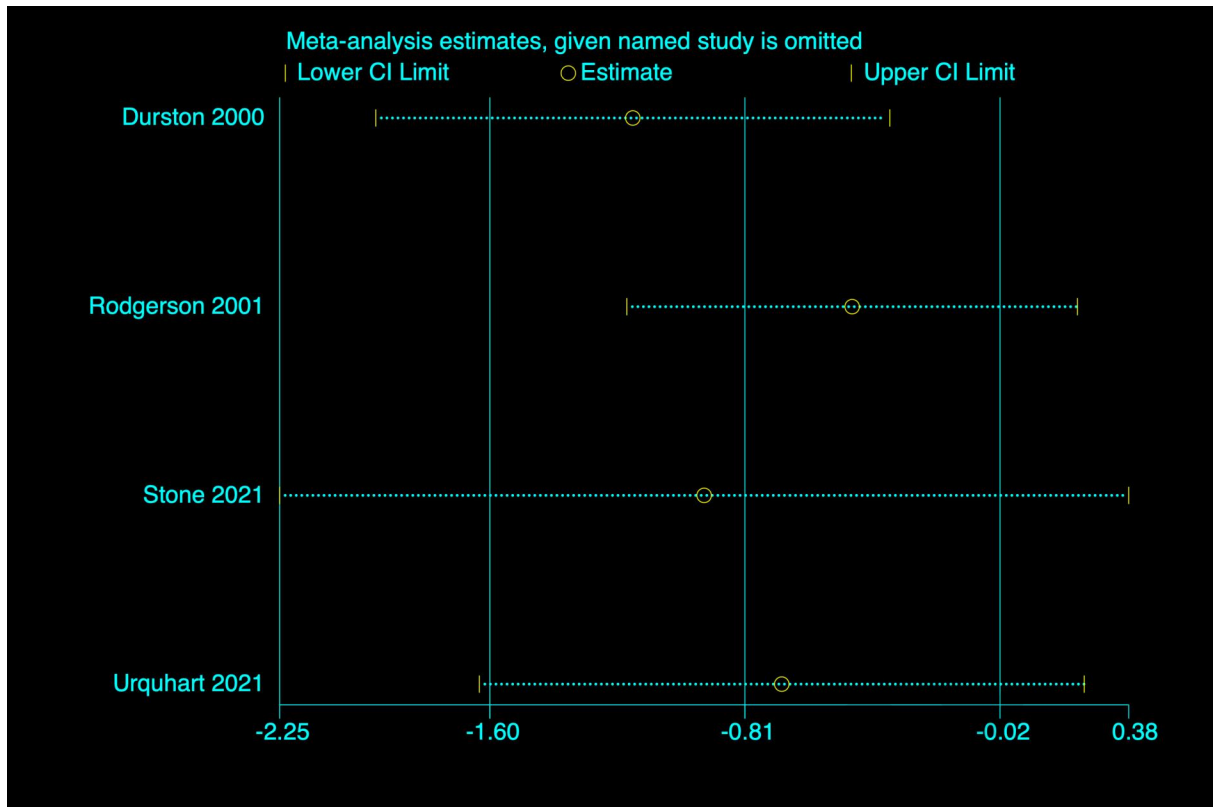


NOTE: Weights are from random-effects model

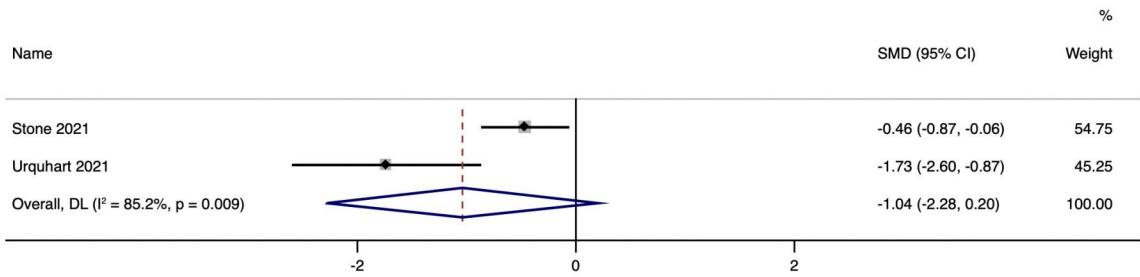
Preprint

LFK index = -2.12 (major asymmetry)



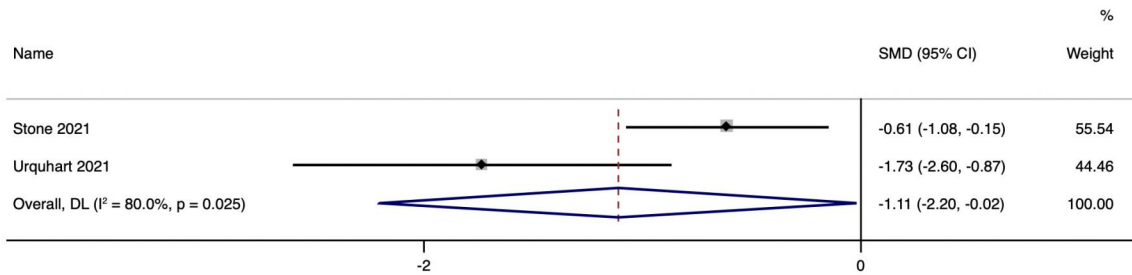


Preprint



NOTE: Weights are from random-effects model

Preprint



NOTE: Weights are from random-effects model

Preprint