



Diagnostic accuracy of automated breast volume scanning, hand-held ultrasound and molybdenum-target mammography for breast lesions: a systematic review and meta-analysis

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Background: Given the high incidence and increasing burden of breast cancer, more approaches are needed to improve the early diagnosis of breast cancer. The three mainstream diagnostic methods, automated breast volume scanning (ABVS), hand-held ultrasound (HHUS) and mammography, are still controversial in their diagnostic accuracy. The aim of this study is to systematically evaluate the accuracy of three diagnostic methods.

Methods: PubMed, Embase, Web of Science, The Cochrane Library, Wanfang Data, China National Knowledge Infrastructure (CNKI), VIP and SinoMed databases were searched by computer. Studies on the accuracy of ABVS, HHUS and mammography in the diagnosis of benign and malignant breast lesions were collected, and the search time limit was from the establishment of the database to August 2022. The Chi-square test was then performed using Meta-Disc software, and the Quality Assessment of Diagnostic Accuracy Studies-2 (QUADAS-2) tool was used for bias and quality assessment.

Results: A total of 31 studies involving 8,107 benign or malignant lesion were included in the meta-analysis. The pooled sensitivity, specificity, positive likelihood ratio, negative likelihood ratio, diagnostic odds ratio, and area under curve for HHUS were 0.86 (0.84, 0.87), 0.80 (0.78, 0.81), 4.20 (3.53, 4.99), 0.20 (0.16, 0.24), 22.88 (16.84, 31.08) and 0.898, respectively. And those for ABVS were 0.90 (0.89, 0.91), 0.87 (0.86, 0.88), 7.93 (5.05, 12.45), 0.11 (0.09, 0.15), 74.63 (45.37, 122.76) and 0.956, respectively. And those for molybdenum-target mammography were 0.81 (0.78, 0.84), 0.90 (0.88, 0.91), 6.94 (4.32, 11.17), 0.23 (0.18, 0.29), 31.41 (17.01, 57.98) and 0.887, respectively. Indicators related to patient selection and reference standards suggested a high risk of bias in several included studies.

Conclusions: Meta-analysis found a higher diagnostic accuracy of ABVS in benign and malignant breast lesions. These results provide a reference for clinical practitioners in the selection of diagnostic methods, but considering the possible bias of the included studies, the results need to be treated with caution and further verified.

Keywords: Breast cancer; automated breast volume scanning (ABVS); hand-held ultrasound (HHUS); molybdenum-target mammography; meta-analysis

Submitted Apr 25, 2024. Accepted for publication Feb 11, 2025. Published online Mar 26, 2025.

doi: 10.21037/gs-24-135

View this article at: <https://dx.doi.org/10.21037/gs-24-135>

Introduction

Breast cancer is one of the most common malignancies and the fifth leading cause of death in women worldwide (1). The incidence of breast cancer is increasing year by year and shows younger trend, which threatens women's health (2). The 5-year survival rate of breast cancer reaches only 85%, and the rate is ever worse for breast cancer patients with advanced stage (3). Therefore, the advanced diagnostic methods are urgently required to effectively improve the prognosis for women with breast cancer.

However, the ideal breast cancer diagnostic system is yet to be undetermined. Molybdenum-target mammography is the main screening method for breast cancer with low diagnostic performance, of which the sensitivity is only reported to be 69–90% (4,5). Moreover, the efficacy of molybdenum-target mammography screening in reducing mortality in women younger than 40 years of age has declined because of concerns about radiation and generally thicker breasts (6). The magnetic resonance imaging (MRI) also has some limitations, such as the inadequate particularity and inadmissible accessibility to non-wealthy

regions (7). Hand-held ultrasound (HHUS) is another traditional imaging methods for the diagnosis of breast cancer. It is safe and sensitive at distinguishing the echo of gland tissue and fat, and is good at defining the boundary and morphology of lesions (6). However, the duration of the whole-breast examination and operator dependence make it inconvenient in the clinical use. Automated breast volume scanning (ABVS) is a volumetric sonographic technique in which the whole breast volume is acquired with almost isotropic voxels, providing multiple reconstruction of the breast. Compared with HHUS, ABVS has a large number of advantages, such as shorter duration of image acquisition, higher reproducibility, less operator dependence and providing reproducible and high-resolution images as well as extra information on the diagnosis of the coronal plane (8). In 2012, ABVS was approved for use by the U.S. Food and Drug Administration. Previous studies have confirmed that tumor size diagnosed by ABVS is associated with the diagnostic of clinical T1-T2 breast cancer [odds ratio (OR) =1.033; P=0.002] and sclerosing adenosis (sensitivity, specificity, and accuracy: 75%, 86.76%, 73.53%) (9,10). Currently, ABVS is only been recommended as a promising auxiliary diagnostic method for mammography (11). Considering the need for simple, rapid, and cost-effective screening methods, evaluating the accuracy of a single screening tool can provide the necessary data to support clinical decision making. However, the current research results have not reached a unanimous conclusion on this issue. Several studies have shown that ABVS has many advantages over HHUS, such as shorter image acquisition time, higher repeatability, and less dependence on the operator (12,13) While other studies did not provide the evidence which showed the difference between ABVS and other methods in the diagnostic performance for breast cancer (14). Due to the inconsistent results, this research was conducted to systematically compare the efficacy of ABVS, HHUS, and mammography for breast cancer diagnosis. We present this article in accordance with the PRISMA-DTA reporting checklist (available at <https://gs.amegroups.com/article/view/10.21037/gc-24-135/rc>).

Highlight box

Key findings

- Automated breast volume scanning (ABVS) has the highest diagnostic accuracy for differentiating between benign and malignant breast lesions, compared to hand-held ultrasound (HHUS) and molybdenum-target mammography. ABVS shows superior sensitivity, specificity, positive and negative likelihood ratios, and diagnostic odds ratio.

What is known and what is new?

- Molybdenum-target mammography is traditionally used for breast cancer screening but has limitations in sensitivity, especially among younger women. HHUS offers a safe alternative but is operator-dependent and time-consuming.
- This meta-analysis confirms ABVS as a more effective and less operator-dependent method compared to HHUS and molybdenum-target mammography, offering higher reproducibility and accuracy in the diagnosis of breast lesions.

What is the implication, and what should change now?

- The superior diagnostic performance of ABVS suggests it should be considered more prominently in clinical settings for early breast cancer detection. The adoption of ABVS can potentially improve diagnostic workflows and patient outcomes, particularly in settings where quick and reliable diagnosis is critical. Future policies should support broader access to ABVS and further research to solidify its advantages over traditional methods.

Methods

Search strategy and selection criteria

Based on the predetermined protocol, we searched PubMed, Web of Science, Embase, the Cochrane Database, China National Knowledge Infrastructure (CNKI), VIP,

Wanfang and SinoMed from inception to August, 2022, with the terms “breast cancer”, “automated breast volume scanner”, “Hand-held Ultrasound” and “Mammography”. Additionally, the reference lists of reviews were screened for qualifying studies. Two authors independently confirmed the eligibility of studies, collated the data from the qualifying studies and extracted the data. Discrepancies were resolved through discussion. Study quality was assessed as recommended in the Cochrane Handbook. The study language was restricted to English or Chinese.

Selection criteria

The inclusion criteria were listed as follows:

- (I) Population: studies targeted on excisional biopsy confirmed benign or malignant breast lesions patients;
- (II) Intervention and comparison: one of the following diagnostic method should be included, ABVS, HHUS or mammography;
- (III) Outcomes: data on diagnostic accuracy need to be provided, including true positive, false negative, positive predictive value and negative predictive value;
- (IV) Study types: diagnostic test.

The exclusion criteria were listed as follows:

- (I) Repeated publications;
- (II) Literature whose data cannot be used;
- (III) Literature with incomplete information;
- (IV) Reviews or case reports;
- (V) Non-Chinese or English literature.

Data collection was made regarding the name of the first author, year of publication, the country in which the examination was performed, the objectives, study design, number of participants, screening methods of assessment, patients' mean age, and the number of lesions.

Quality assessment

Conference papers, original research articles, and review papers became the basis of this study. A thorough check was made on all duplications in maintaining the nature of the review. For the quality assessment, the QUADAS-2 tool was used which involved four domains including ‘index test’, ‘reference standard’, ‘patient selection’, and ‘flow and timing’. Risk of bias in each domain is assessed, and the first 3 domains are also assessed in terms of concerns regarding applicability (15). Signalling questions are included to help

judge risk of bias. QUADAS-2 tabular and graphical display can be retrieved from the Web page, <http://www.bris.ac.uk/quadas/quadas-2>. Dispute between the two reviewers in assessing the quality of the study was settled through discussion.

Statistical analysis

Meta-analysis was performed by Meta-Disc, and outcome measurements are the pooled sensitivity (SEN), specificity (SPE), negatively likelihood ratio (NLR), diagnostic odds ratio (DOR), area under curve (AUC) and their 95% confidence intervals (CIs) (http://www.hrc.es/investigacion/metadisc_en.htm). The heterogeneity between the results of the study was analyzed by the Cochran Q-test, and the I^2 index ($I^2 > 50\%$ and Q-test $P > 0.10$ indicated high heterogeneity). If there was no statistical heterogeneity among the results, the fixed effect model was adopted for meta-analysis; If there was a statistical heterogeneity among the research results, the sources of heterogeneity would be needing further analysis, and after excluding the influence of obvious clinical heterogeneity, the random effects model was used for meta-analysis. P values < 0.05 were considered statistically significant. We used Stata statistical software (STATA, version 15.1) to obtain a quantitative analysis of all the publication bias according to funnel plots and the Deek's test. We did subgroup analysis according to the year of publication, study design, sample size and the mean age of participants.

Results

Literature screening and study quality

Literature search yielded a total of 619 publications, including 12 from PubMed, 45 from Embase, 2 from Cochrane, 70 from Web of Science, 45 from CNKI, 83 from VIP, 321 from Wanfang and 41 from SinoMed. After removing duplications, 589 publications remained. After subsequent title and abstract screening, 546 were excluded (533 publications were non-relevant to the topic and 13 publications were reviews). Then the full-text content of the remaining 43 papers was screened. Twelve papers were excluded due to the insufficient calculated data. A total of 31 studies were included in this meta-analysis, comprising 19 retrospective studies and 12 prospective studies (Table 1). A flow diagram of the literature screening selection was outlined in Figure 1. And the characteristics and details

Table 1 Characteristics of included studies reporting the diagnostic estimates of ABVS for diagnosis of breast cancer

No.	Author	Study design	Age, years	No. of lesion	No. of malignant lesion	No. of benign lesion	Instrument		
							HHUS	ABVS	Molybdenum-target mammography
1	Zhou, 2010 (16)	Retrospective	48	171	78	93	GE logiq	GE logiq	–
2	Chen, 2011 (17)	Perspective	42.3	101	45	56	GE logiq	GE logiq	–
3	Wang, 2012 (14)	Perspective	43	239	85	154	GE logiq	Acuson	–
4	Wang, 2012 (11)	Retrospective	43.1	165	62	103	Acuson	Acuson	Senograph
5	Zhu, 2013 (18)	Retrospective	43.4	294	91	203	Acuson	Acuson	–
6	Chen, 2013 (19)	Retrospective	41.7	219	67	152	Acuson	Acuson	–
7	Li, 2014 (20)	Perspective	45.2	134	68	66	Acuson	Acuson	–
8	Cheng, 2014 (21)	Retrospective	42.3	104	39	65	Acuson	Acuson	–
9	Yang, 2014 (22)	Retrospective	48.8	130	74	56	Acuson	Acuson	–
10	Shen, 2015 (23)	Retrospective	46.1	470	70	400	Acuson	Acuson	–
11	Min, 2015 (24)	Retrospective	44.1	156	51	105	Acuson	Acuson	–
12	Zhong, 2016 (25)	Retrospective	46.9	179	Not reported	Not reported	Acuson	Acuson	–
13	Cheng, 2016 (26)	Perspective	43.7	71	31	40	Acuson	Acuson	Siemens
14	Kuang, 2016 (27)	Perspective	46.8	136	56	80	Philips	Philips	–
15	He, 2016 (28)	Retrospective	50.8	170	102	68	Acuson	Acuson	–
16	Dai, 2016 (29)	Retrospective	50.5	118	68	50	Philips	Acuson	–
17	Zhu, 2017 (30)	Perspective	50.6	130	72	58	Acuson	Acuson	–
18	Wei, 2017 (31)	Retrospective	44.2	311	141	170	Acuson	Acuson	–
19	Huang, 2017 (32)	Perspective	47.7	156	83	73	Acuson	Acuson	–
20	Yan, 2018 (33)	Retrospective	53.2	168	100	68	Acuson	Acuson	–
21	Yan, 2018 (34)	Retrospective	50.5	210	127	83	Acuson	Acuson	–
22	Du, 2018 (35)	Retrospective	46	272	120	152	Acuson	Acuson	–
23	Zeng, 2018 (36)	Retrospective	47.6	393	200	193	Acuson	Acuson	–
24	Huang, 2018 (37)	Perspective	50	76	18	58	Acuson	Acuson	Philips
25	Xu, 2018 (38)	Retrospective	45.7	267	18	249	Acuson	Acuson	Senograph
26	Zhang, 2018 (39)	Perspective	45.4	1,973	417	1,556	Not reported	Not reported	–
27	Xu, 2019 (40)	Retrospective	48.5	130	56	74	Acuson	Acuson	–
28	Pan, 2019 (41)	Retrospective	49.5	120	43	77	Not reported	Acuson	–
29	Zhang, 2019 (42)	Perspective	50.1	594	75	519	Acuson	GE logiq	GE logiq
30	Wu, 2020 (43)	Perspective	47.4	200	81	119	–	Acuson	Siemens
31	Wang, 2022 (44)	Retrospective	49.2	250	143	107	Philips	GE logiq	Siemens

ABVS, automated breast volume scanning; HHUS, hand-held ultrasound.

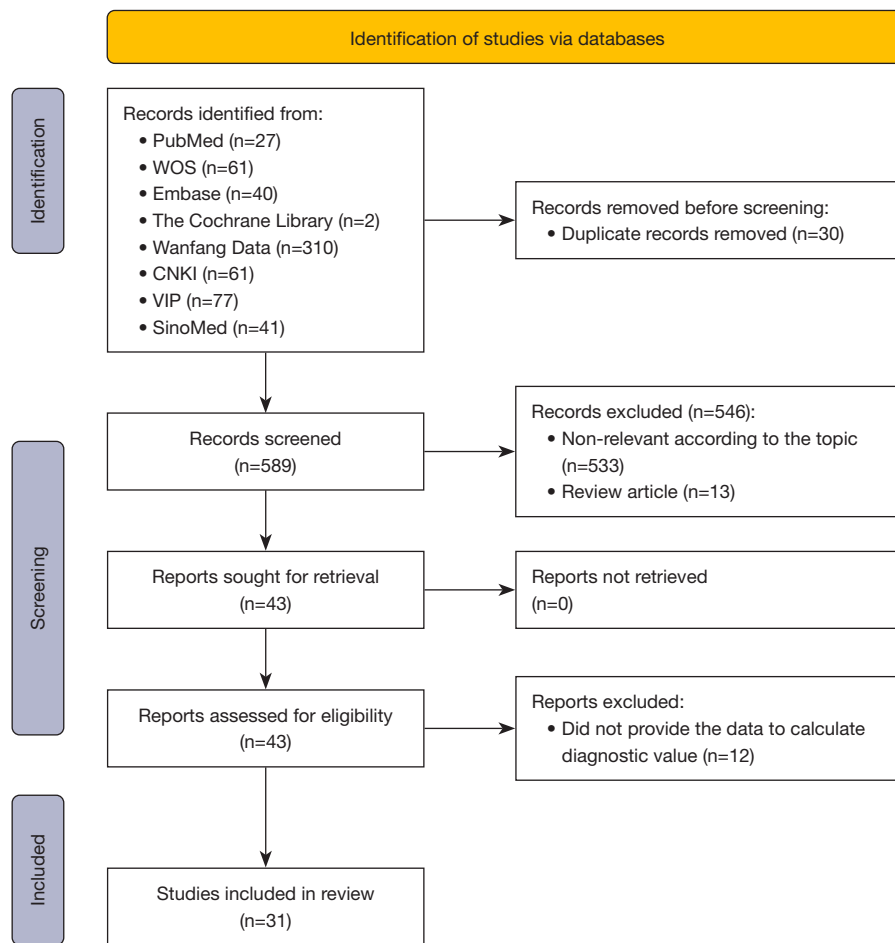


Figure 1 Study flow diagram. A diagram summarizing the search and screening process of the included studies.

concerning the literature type of the included studies were summarized in *Table 1*. The 31 studies underwent quality assessment using the QUADAS criteria for diagnostic studies. The results of the quality assessment after consensus were displayed graphically in *Figure S1* and *Table S1*.

Test of heterogeneity

We first explored heterogeneity through threshold analysis by finding the logarithm of true-positive rates and the logarithm of false-positive rates. The Spearman correlation coefficient for HHUS, ABVS and molybdenum-target mammography, performed as a test for a threshold effect, was 0.091 ($P=0.63$), -0.052 ($P=0.78$) and -0.300 ($P=0.62$), respectively, indicating that the heterogeneity was not caused by a threshold.

Meta-analysis of diagnostic performance for breast cancer

Thirty studies provided data on the diagnostic values of HHUS for the diagnosis of breast cancer. The pooled SEN, SPE, PLR and NLR were 0.86 (0.84, 0.87), 0.80 (0.78, 0.81), 4.20 (3.53, 4.99) and 0.20 (0.16, 0.24), respectively (*Table 2*). And the DOR and AUC of HHUS were 22.88 (16.84, 31.08) and 0.898, respectively (*Figure 2* and *Figure S2*). The Cochrane's Q tests and I^2 tests revealed significant heterogeneity among studies.

Thirty-one studies provided data on the diagnostic values of ABVS for the diagnosis of breast cancer. The pooled SEN, SPE, PLR and NLR were 0.90 (0.89, 0.91), 0.87 (0.86, 0.88), 7.93 (5.05, 12.45) and 0.11 (0.09, 0.15), respectively (*Table 2*). And the DOR and AUC of ABVS were 74.63 (45.37, 122.76) and 0.956, respectively (*Figure 3* and *Figure S3*). The Cochrane's Q tests and I^2 tests revealed

Table 2 The diagnostic performance of HHUS, ABVS and molybdenum-target mammography for breast cancer

Diagnostic performance indicators	HHUS	ABVS	Molybdenum-target mammography
SEN (95% CI)	0.86 (0.84, 0.87)	0.90 (0.89, 0.91)	0.81 (0.78, 0.84)
SPE (95% CI)	0.80 (0.78, 0.81)	0.87 (0.86, 0.88)	0.90 (0.88, 0.91)
PLR (95% CI)	4.20 (3.53, 4.99)	7.93 (5.05, 12.45)	6.94 (4.32, 11.17)
NLR (95% CI)	0.20 (0.16, 0.24)	0.11 (0.09, 0.15)	0.23 (0.18, 0.29)
DOR (95% CI)	22.88 (16.84, 31.08)	74.63 (45.37, 122.76)	31.41 (17.01, 57.98)
AUC	0.898	0.956	0.887

ABVS, automated breast volume scanning; AUC, area under curve; DOR, diagnostic odds ratio; HHUS, hand-held ultrasound; NLR, negative likely ratio; SEN, sensitivity; SPE, specificity; PLR, positive likely ratio; 95% CI, 95% confidence interval.

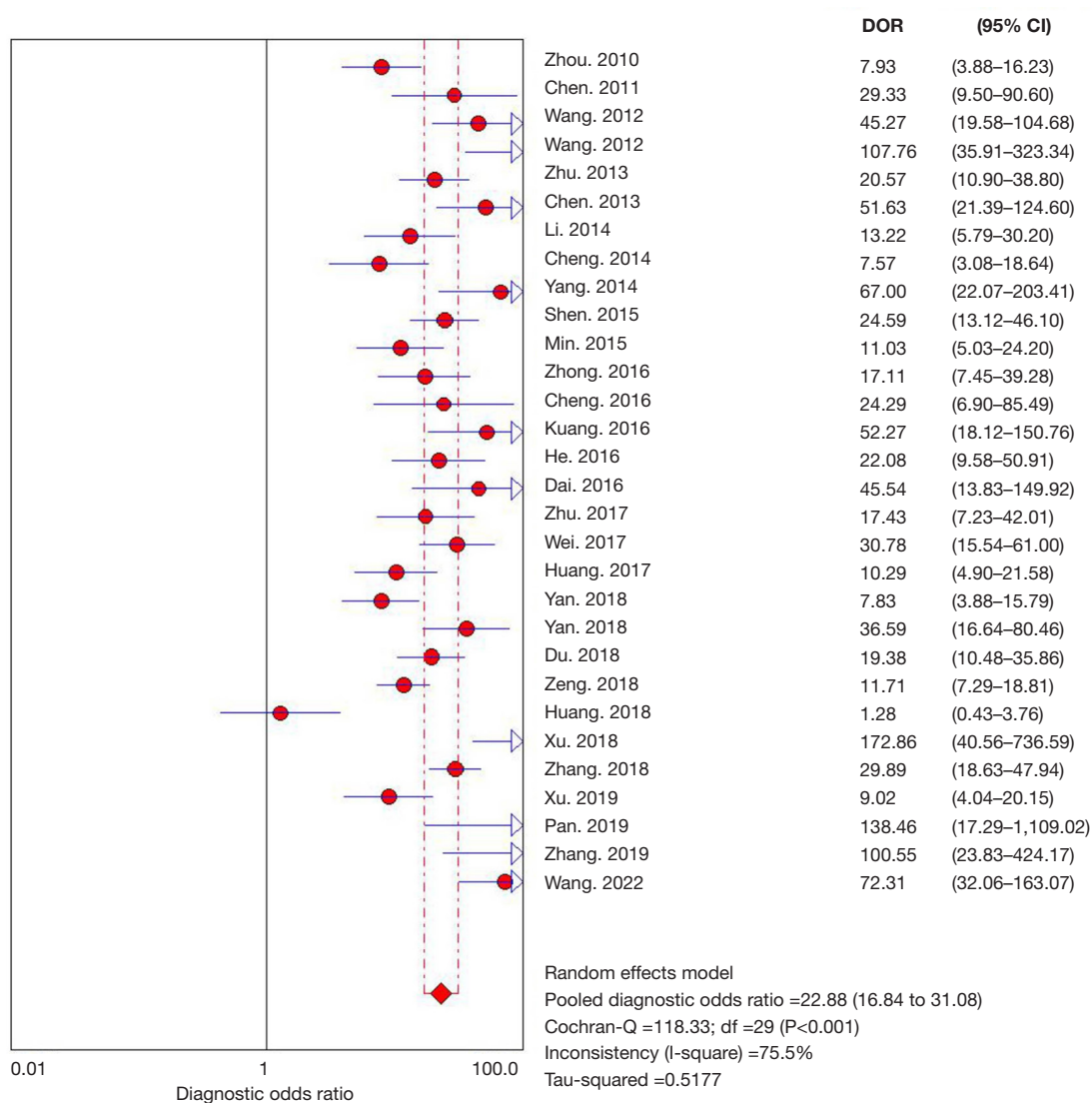


Figure 2 Forest plot of pooled DOR for HHUS in the diagnosis of breast cancer. DOR, diagnostic odds ratio; HHUS, hand-held ultrasound; 95% CI, 95% confidence interval.

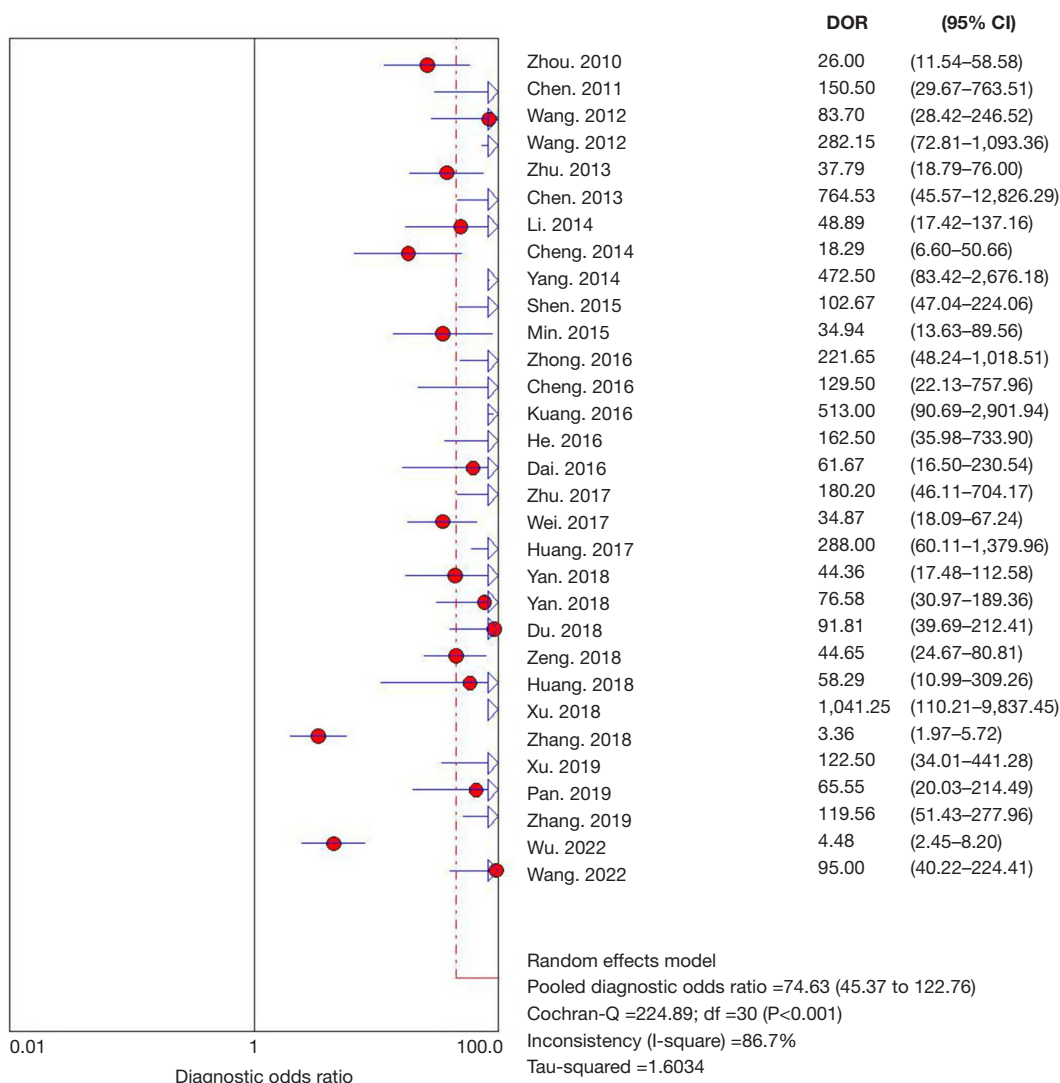


Figure 3 Forest plot of pooled DOR for ABVS in the diagnosis of breast cancer. ABVS, automated breast volume scanning; DOR, diagnostic odds ratio; 95% CI, 95% confidence interval.

significant heterogeneity among studies.

Nine studies provided data on the diagnostic values of molybdenum-target mammography for the diagnosis of breast cancer. The pooled SEN, SPE, PLR and NLR were 0.81 (0.78, 0.84), 0.90 (0.88, 0.91), 6.94 (4.32, 11.17) and 0.23 (0.18, 0.29), respectively (Table 2). And the DOR and AUC of ABVS were 31.41 (17.01, 57.98) and 0.887, respectively (Figure 4 and Figure S4). The Cochrane’s Q tests and I² tests revealed significant heterogeneity among studies.

Sub-group analysis

Sub-group analysis was performed to explore the potential source of heterogeneity in DOR based on the covariates including year of publication, study design, sample size and the mean age of participants. Table 3 showed the sub-group analysis results. The DOR value of ABVS was lower in the studies which was published after 2016 and in the studies with sample size more than 200 [2016 and before vs. after 2016: 86.74 (52.93, 142.14) vs. 59.21 (26.95, 130.08); sample

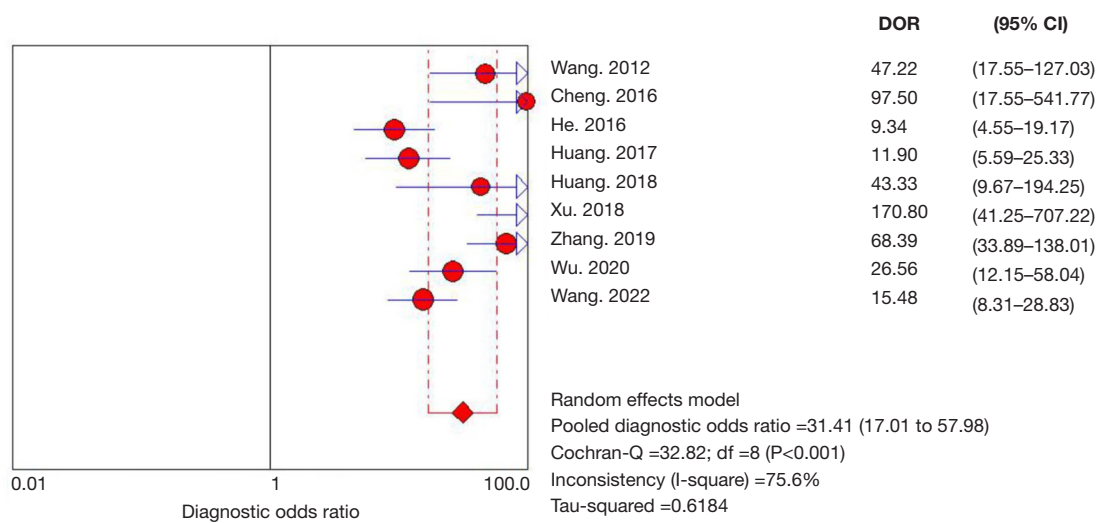


Figure 4 Forest plot of pooled DOR for molybdenum-target mammography in the diagnosis of breast cancer. DOR, diagnostic odds ratio; 95% CI, 95% confidence interval.

Table 3 The sub-group analysis for DOR value (95% CI) of HHUS, ABVS and molybdenum-target mammography for breast cancer

Sub-group	HHUS	ABVS	Molybdenum-target mammography
Year of publication			
2016 and before	24.57 (17.08, 35.34)	86.74 (52.93, 142.14)	30.63 (7.44, 126.20)
After 2016	21.18 (12.74, 35.20)	59.21 (26.95, 130.08)	32.86 (15.76, 68.49)
Average age			
≤45 years	26.80 (16.34, 43.97)	61.72 (34.13, 111.60)	56.59 (24.02, 133.36)
>45 years	21.46 (14.64, 31.47)	77.43 (40.23, 149.02)	27.03 (13.53, 54.01)
Study design			
Retrospective	24.14 (16.87, 34.53)	71.61 (50.07, 102.42)	28.44 (10.18, 79.46)
Perspective	20.25 (10.84, 37.82)	65.39 (20.21, 211.58)	35.03 (15.92, 77.08)
Sample size			
≤200	17.03 (10.99, 26.39)	80.84 (41.63, 156.99)	23.98 (12.36, 46.51)
>200	32.81 (22.79, 47.24)	67.30 (30.34, 149.31)	51.01 (13.40, 194.17)

ABVS, automated breast volume scanning; DOR, diagnostic odds ratio; HHUS, hand-held ultrasound; 95% CI, 95% confidence interval.

size less than 200 vs. sample size more than 200: 80.84 (41.63, 156.99) vs. 67.30 (30.34, 149.31)].

Publication bias

Publication bias of all included studies concerning the diagnostic value of HHUS for breast cancer was determined using Deek’s funnel plot test. The result revealed a good

symmetric distribution of the included studies on both sides of the funnel plot, suggesting a small possibility of publication bias (P=0.71, Figure S5).

Discussion

Breast cancer is one of the most common malignancies and has become a serious threat to women’s health worldwide (45).

Perceptions towards breast cancer screening and diagnosis have an important role in the early diagnosis of breast cancer, which is crucial in the reduction of breast cancer mortality rate. As a new imaging technology, ABVS can automatically measure the distance between the lesion, skin and nipple, and position the mass clearly within the breast boundaries, providing accurate location information regarding the mass (8). Moreover, ABVS also makes ultrasonography less operator dependent and facilitates easier interpretation of ultrasonographic images (46). Therefore, it is necessary to explore the diagnostic efficacy of ABVS, which is a cost-effective and convenient method for breast cancer.

Some previous studies have compared the ABVS and HHUS with respect to diagnostic performance in the differential diagnosis of benign and malignant breast lesions. Wang *et al.* included nine studies and found that compared with HHUS, ABVS had higher DOR in the differentiation of benign and malignant breast lesions [61.68 (32.31, 117.76) *vs.* 52.60 (32.06, 86.35)] (47). However, the areas under the SROC curves in the differentiation of benign and malignant breast lesions were 0.93 and 0.94 for ABVS and HHUS, respectively, which were not significantly different ($P=0.853$).

In the current meta-analysis, 31 diagnostic studies were included. We found that ABVS had the numerically higher sensitivity, PLR, DOR and AUC compared with HHUS and molybdenum-target mammography. However, the NLR of ABVS was lower than that of HHUS and molybdenum-target mammography. These results indicated that the diagnostic accuracy of ABVS in differentiating benign from malignant breast lesions was high and ABVS had the higher diagnostic performance than the other two methods. Patients who were diagnosed positive by ABVS had a greater likelihood of breast cancer. Patients who were diagnosed negative by HHUS had a greater likelihood of not developing breast cancer. This result was consistent with the previous study, highlighting the importance of using ABVS in the early diagnosis of breast cancer, which could increase the diagnostic performance of breast cancer.

The results of sub-group analysis showed that the year of publication and sample size might be the source of heterogeneity. The results of studies with larger sample size might tend to be more accurate. In addition, the heterogeneity might be associated with the use of blinding, the low quality of literatures, the lesion mass and the experience of doctors. More studies are worthwhile to be explored in future on a larger number of patients, to enable

more meaningful sub-group analysis.

However, our study still had some limitations. First, most of included studies were from China. That is due to the fact that breast lesion is tended to be diagnosed by molybdenum-target mammography in western countries. Second, we did not consider the lymph node involvement and distant metastasis in the current meta-analysis. Third, the interval between gold standard and examinations may introduce bias, because the disease status may change during the interval.

Conclusions

In conclusion, ABVS has higher diagnostic performance in the diagnosis of benign and malignant breast lesions. However, more studies are still needed to verify the above conclusion.

Acknowledgments

None.

Footnote

Reporting Checklist: The authors have completed the PRISMA-DTA reporting checklist. Available at <https://gs.amegroups.com/article/view/10.21037/gS-24-135/rc>

Peer Review File: Available at <https://gs.amegroups.com/article/view/10.21037/gS-24-135/prf>

Funding: None.

Conflicts of Interest: All authors have completed the ICMJE uniform disclosure form (available at <https://gs.amegroups.com/article/view/10.21037/gS-24-135/coif>). The authors have no conflicts of interest to declare.

Ethical Statement: The authors are accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

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Cite this article as: Liu X, Dai Y, Wu Y, Li F, Liang M, Wu Q. Diagnostic accuracy of automated breast volume scanning, hand-held ultrasound and molybdenum-target mammography for breast lesions: a systematic review and meta-analysis. *Gland Surg* 2025;14(3):294-304. doi: 10.21037/gs-24-135