



# The added value of S-detect in the diagnostic accuracy of breast masses by senior and junior radiologist groups: a systematic review and meta-analysis

Peijun Chen<sup>1,2</sup>, Jiahui Tong<sup>1</sup>, Ting Lin<sup>1</sup>, Ying Wang<sup>3</sup>, Yuehui Yu<sup>3</sup>, Menghan Chen<sup>3</sup>, Gaoyi Yang<sup>2</sup>

<sup>1</sup>Department of Ultrasonography, The Second Clinical Medical College, Zhejiang Chinese Medicine University, Hangzhou, China; <sup>2</sup>Department of Ultrasonography, Affiliated Hangzhou Chest Hospital, Zhejiang University School of Medicine, Chinese and Western Hospital of Zhejiang Province (Hangzhou Red Cross Hospital), Hangzhou, China; <sup>3</sup>Department of Ultrasonography, Hangzhou Normal University Division of Health Sciences, Hangzhou, China

**Contributions:** (I) Conception and design: P Chen; (II) Administrative support: G Yang; (III) Provision of study materials or patients: J Tong; (IV) Collection and assembly of data: T Lin, Y Wang; (V) Data analysis and interpretation: Y Yu, M Chen; (VI) Manuscript writing: All authors; (VII) Final approval of manuscript: All authors.

**Correspondence to:** Gaoyi Yang. Department of Ultrasonography, Affiliated Hangzhou Chest Hospital, Zhejiang University School of Medicine, Chinese and Western Hospital of Zhejiang Province (Hangzhou Red Cross Hospital), No. 208 Huancheng East Road, Hangzhou 310003, China. Email: yanggaoyi8@163.com.

**Background:** S-detect is an emerging computer-aided diagnosis (CAD) technique that provides a reference for radiologists to identify breast cancer. Some studies have shown that US (ultrasound) + S-detect can improve the diagnostic accuracy of junior radiologists more than senior radiologists, but the results are inconsistent in various studies. Therefore, this meta-analysis aimed to assess the value of S-detect combined with the US outcomes from senior and junior radiologists for the diagnosis of breast cancer.

**Methods:** We searched the PubMed, Cochrane Library, Embase, Web of Science, and Wanfang databases, China Biology Medicine disc, China National Knowledge Infrastructure (CNKI), and VIP database for trials on the diagnostic accuracy of US + S-detect for the diagnosis of breast masses. The search time frame was from the date of establishment of the database to August 20, 2022. Two researchers independently screened the literature, extracted the information, and evaluated the quality of the included literature using the Quality Assessment of Diagnostic Accuracy Studies-2 (QUADAS-2) scale. StataSE 15.1 software was utilized to assess pooled metrics, including sensitivity, specificity, and the area under the curve (AUC).

**Results:** A total of 19 articles with 3,349 patients and 3,895 breast masses were included in this meta-analysis. Of these, seventeen articles evaluated the diagnostic performance of senior radiologists' US + S-detect for breast cancer, while twelve articles reported junior radiologists' diagnostic performance. The risk of bias was primarily attributed to patient selection, flow and timing. In the senior radiologist group, the pooled sensitivity and specificity of US + S-detect were 0.93 [95% confidence interval (CI): 0.89–0.95] and 0.86 (95% CI: 0.80–0.90), respectively, with an AUC of 0.96. As for the junior radiologist group, the pooled sensitivity and specificity of US + S-detect were 0.89 (95% CI: 0.83–0.93) and 0.79 (95% CI: 0.72–0.84), respectively, and the AUC was 0.91.

**Conclusions:** The results of this meta-analysis showed that the pooled sensitivity and the AUC of both the senior and junior radiologist groups were high, with good diagnostic efficacy and high clinical application. However, the results of this study are highly heterogeneous and need to be validated by collecting more high-quality studies and accumulating a larger sample size.

**Keywords:** S-detect; ultrasound; breast; senior radiologist; junior radiologist

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## Introduction

The incidence and mortality of breast cancer among women worldwide have been increasing in recent years, and in 2020, breast cancer has surpassed lung cancer as the most common cancer worldwide. Breast cancer mortality accounts for 6.9% of all tumor-related deaths (1), which seriously endangers the health of women. Early diagnosis and treatment of breast cancer are key to reducing the mortality rate and improving the survival quality of patients (2,3). Ultrasound (US) is an important imaging tool for breast cancer screening. However, the diagnostic results are easily affected by the operation and experience level of radiologists, and there is a risk of misdiagnosis and missed diagnosis (4-7).

The proposed breast imaging reporting and data system (BI-RADS) has standardized breast US examinations (8), but the influence of inter-observer variability on US diagnostic results still exists, especially in primary hospitals, due to the large number of junior radiologists, the accuracy of BI-RADS classification in the diagnosis of the nature of breast lesions and the final pathological results is low. In recent years, with the rise of artificial intelligence and the rapid development of digital healthcare, the application of computer-aided diagnosis (CAD) in medical imaging has become a research hotspot (9-11). S-detect is a CAD system based on deep learning algorithms grafted onto US instruments, which is based on the BI-RADS interpretation of the benignity and malignancy of lesions, which indicates that the breast lesion is “probably benign” or “probably malignant”. This technique can help to improve diagnostic

efficiency and reduce subjective variability and has broad application prospects (12). Some studies have shown that US + S-detect can improve the diagnostic accuracy of junior radiologists more than senior radiologists, but the results are inconsistent in various studies (13,14). Because the results of previous studies were variable, there are no relevant meta-analyses examining the results of this technique in combination with senior and junior radiologists. The present meta-analysis included several domestic and international articles for quantitative synthesis, aiming to evaluate the usefulness of applying S-detect to breast US and examining whether there are differences between senior and junior radiologists, exploring how the S-detect technique can help radiologists of different levels of seniority in the clinical setting. We present the following article in accordance with the PRISMA-DTA reporting checklist (available at <https://gs.amegroups.com/article/view/10.21037/gS-22-643/rc>).

## Methods

### *Data sources and search strategy*

On August 20, 2022, we searched the PubMed, Cochrane Library, Embase, Web of Science, Wanfang databases, China Biology Medicine disc, China National Knowledge Infrastructure (CNKI), and VIP database for studies evaluating the accuracy of US + S-detect in breast cancer. Taking PubMed as an example, the search strategy used the following formula: “(ultrasound) AND (S-detect) AND (breast)”. Similar search formulas were used for the Cochrane Library, Embase, Web of Science, Wanfang databases, China Biology Medicine disc, China National Knowledge Infrastructure (CNKI), and VIP database.

### *Eligibility criteria*

We relied on the population, intervention, comparison, outcome and study design principle (PICOS) to define study eligibility.

Population: patients with S-detect diagnosis of breast masses were included.

Intervention: S-detect diagnosis combined with the diagnosis of senior and junior radiologists.

Comparison: the gold diagnosis of the breast masses (final histopathological examination)

Outcome: diagnostic accuracy of breast lesions by S-detect combined with senior and junior radiologists (sensitivity, specificity, accuracy).

### Highlight box

#### Key findings

- The pooled sensitivity, pooled specificity, and the AUC of senior radiologists + S-detect were all higher than that of junior radiologists + S-detect.

#### What is known and what is new?

- A previous meta-analysis reported that the pooled sensitivity and specificity of S-detect in diagnosing breast cancer were 0.82 and 0.86
- We assessed the value of S-detect combined with the US outcomes from senior and junior radiologists for the diagnosis of breast cancer.

#### What is the implication, and what should change now?

- S-detect can assist radiologists to diagnose breast cancer and reduce the subjectivity between different radiologists.

Study design: any type of study design, such as retrospective, prospective, or case-control, were eligible for inclusion provided the study assessed the diagnostic accuracy of breast lesions by S-detect combined with senior and junior radiologists.

Exclusion criteria: (I) case reports, conference proceedings, reviews, animal studies, etc., for which the original data were unavailable or lacking; (II) articles with small sample sizes (<20 cases); (III) articles written using the same data; and (IV) repeated articles from different databases.

### *Literature screening and data extraction*

Two researchers (Chen PJ and Tong JH) independently completed the literature screening and data extraction. In cases of disagreement and those in which no consensus could be reached after discussion, a third researcher determined the data for inclusion. The following data were extracted from the literature: (I) basic information including the first author, year of article publication, author's country, reference standard, study design, age of patients, number of patients and lesions, tumor diameter, type of US equipment, BI-RADS edition, radiologist's seniority level, BI-RADS diagnostic threshold, and method of US + S-detection; and (II) 2×2 table information including true positive (TP), false positive (FP), false negative (FN), and true negative (TN). If there are multiple sets of data in one article, take the average. If the 2×2 table information was not directly obtained from the original literature, it could be calculated based on the sample size, sensitivity, and specificity provided in the article.

### *Assessment of study quality*

The quality of the included studies was evaluated using the quality assessment of diagnostic accuracy studies-2 (QUADAS-2) scale after a thorough reading of all selected articles (15), and each of the 14 items was rated according to the actual content of each original study. Determine "yes", "no", "uncertain" according to the relevant signature questions included in each part, the risk of bias and applicability of the scale were assessed as "high", "low", or "unclear". If the answer to the signature question is "yes", the risk of bias is judged to be low. "No" is judged as a high risk of bias. High quality articles are randomized controlled trials (RCTs), high-quality case control or cohort studies with a very low risk of confounding or bias. The quality

evaluation chart of the included literature was obtained using Review Manager 5.4 software (<https://training.cochrane.org/online-learning/core-software-cochrane-reviews/revman>). Quality evaluation was performed independently by two researchers; in cases of inconsistency and those in which no consensus could be reached after discussion, a third researcher made the final decision.

### *Statistical analysis of data*

We inputted TP, FP, FN, and TN into the STATASE 15.1 software (StataCorp, College Station TX, USA) and calculated the estimated pooled sensitivity and specificity of US + S-detect for breast cancer by senior and junior radiologists, respectively. The area under the curve (AUC) was used to evaluate the diagnostic value of US + S-detect for breast cancer by radiologists of different seniority levels. The closer the AUC was to 1, the higher the diagnostic accuracy; however, the closer the AUC was to 0.5, the lower the diagnostic accuracy. Z-test was conducted to compare sensitivity, specificity, and AUC between senior and junior radiologists, and a P value  $\leq 0.05$  was considered to indicate statistical significance.

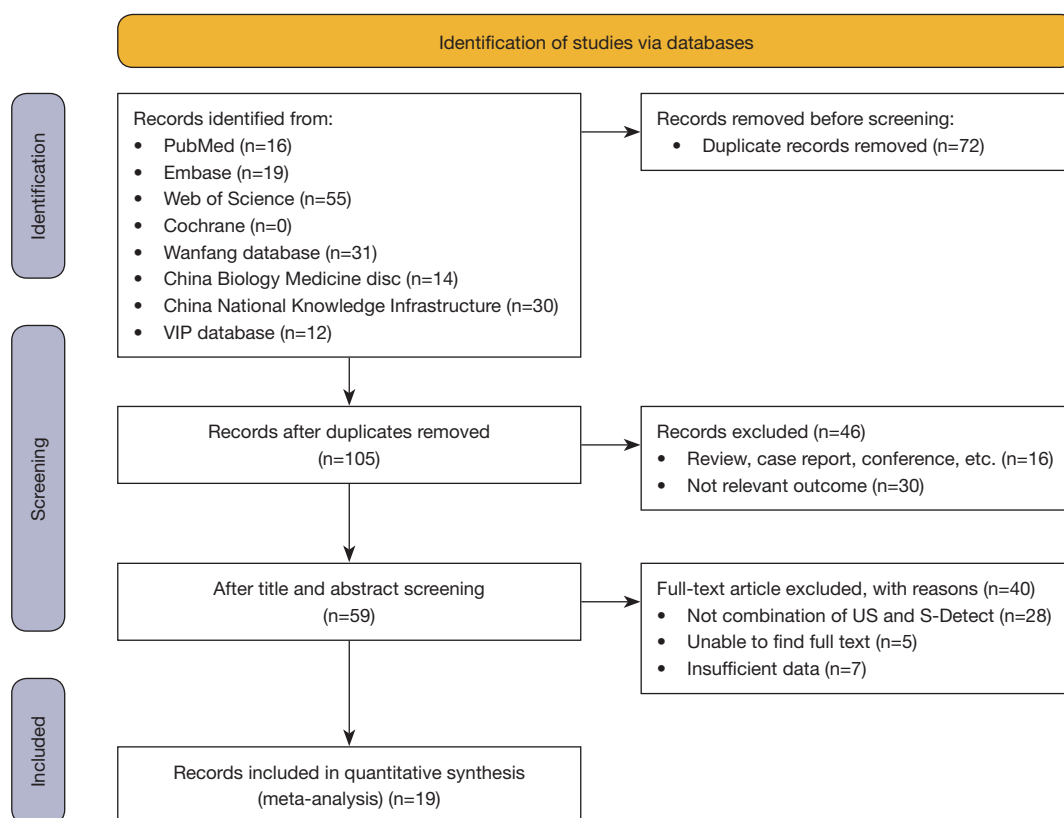
### *Heterogeneity analysis*

Spearman's correlation coefficient was calculated using Meta-Disc 1.4 software (<https://meta-disc.software.informer.com/1.4/>) to test for a threshold effect and assess whether there was heterogeneity due to a threshold effect. A threshold effect was considered to exist when Spearman's correlation coefficient was  $>0$  and  $P < 0.05$ .

The Cochran Q test was applied to obtain P values to quantify the heterogeneity between original studies. When the  $I^2$  value was  $<50\%$ , heterogeneity was considered insignificant, and the data were directly combined for analysis. Meanwhile, when the  $I^2$  value was  $>50\%$ , heterogeneity was considered significant, and a random-effects model was used to merge the data and explore the potential factors contributing to heterogeneity (16,17).

### *Subgroup analysis and sensitivity analysis*

Subgroup analysis was performed to search for factors that might contribute to heterogeneity. The original studies were divided into two subgroups based on the possible factors, when the heterogeneity of a group decreased to  $<50\%$  in subgroup analysis, it was considered that the



**Figure 1** Literature retrieval flow chart. The flow chart shows the process of selecting eligible studies and the total number (n=19) of studies included. US, ultrasound.

source of heterogeneity was found through subgroup analysis. In this study, reference standard (one standard *vs.* two standards), number of included lesions ( $\geq 200$  *vs.*  $< 200$ ), BI-RADS diagnostic threshold (BI-RADS 4a and 4b *vs.* BI-RADS 3 and others), the method of US + S-detect (objective binding *vs.* based on radiologist subjective binding or others), the methodological quality of the study (high-quality *vs.* acceptable-quality) were used as independent variables to delineate the subgroups, respectively.

We performed a sensitivity analysis to further explore the sources of heterogeneity and assess the reliability of the combined effect size results, and determine if there were original studies that had a large effect on heterogeneity.

### Clinical application analysis

Fagan plots were created, pre-test probabilities were set, and post-test probabilities were calculated using the combined positive likelihood ratios to analyze the clinical value of using US + S-detect to diagnose breast cancer.

## Results

### Literature screening process

According to the formulated search strategy, eight databases were searched sequentially, and a total of 177 articles were initially obtained. After importing these articles into Endnote software (Clarivate, Philadelphia, PA, USA), 72 identical articles were excluded, and 46 articles that were deemed inconsistent based on their titles and abstracts were further checked. The remaining relevant articles were read and the inclusion and exclusion criteria were strictly implemented. Finally, a total of 19 diagnostic research articles were included in this meta-analysis (Figure 1).

### Basic characteristics of the included literature

This meta-analysis included 19 original studies (18-36), including eight English articles (19-26) and eleven Chinese studies (18,27-36). We extracted and summarized the basic characteristics of the included literature. Four

included studies were prospective (20,21,23,26), seven were retrospective (18,19,25,29-31,33), and the remaining articles were not reported. Also, a total of 3,349 patients with 3,895 lesions were included. The average lesion diameter of the included patients was 1.10–1.93 cm, and the average age of the patients was 42.6–51.0 years. Eleven articles evaluated the diagnostic performance of US + S-detect for breast masses by both senior and junior radiologists', six articles evaluated only senior radiologists, and two articles evaluated only junior radiologists. The basic characteristics of the included literature are shown in *Table 1*.

### **Methodologic quality evaluation**

Review Manager 5.4 software (<https://training.cochrane.org/online-learning/core-software-cochrane-reviews/revman>) was used to evaluate the quality of the included original studies according to the QUADAS-2 scale. The risk of bias was primarily attributed to patient selection and flow and timing. The risk of bias from the index test was low, as was the risk of bias from the reference standard. *Figure 2* displays the quality evaluation of the included literature.

### **Threshold effect**

The 2x2 table data (TP, FP, FN, TN) from nine studies of US + S-detect in the senior radiologist group were imported into the Meta Disc 1.4 statistical software for analysis, yielding a Spearman correlation coefficient of  $-0.213$  ( $P=0.411$ ). The Spearman correlation coefficient from seven studies of US + S-detect in the junior radiologist group was  $0.280$  ( $P=0.379$ ), indicating that there was no threshold effect in either group.

### **Diagnostic performance of the combination of S-detect with senior radiologists**

Seventeen of the studies (involving 3,119 lesions) reported on the diagnostic performance of the combination of S-detect with senior radiologists in differentiating benign from malignant breast masses. Combining S-detect with senior radiologists, the pooled sensitivity was  $0.93$  [95% confidence interval (CI):  $0.89-0.95$ ,  $I^2=64.74\%$ ] and the pooled specificity was  $0.86$  (95% CI:  $0.80-0.90$ ,  $I^2=86.11\%$ ). *Figure 3A* shows the Forest plot of the sensitivity and specificity. The AUC of the summary receiver operating characteristic (SROC) curve was  $0.96$  (95% CI:  $0.93-0.97$ ) (*Figure 3B*).

### **Diagnostic performance of the combination of S-detect with junior radiologists**

Twelve of the studies (involving 2,953 lesions) reported on the performance of the combination of S-detect with junior radiologists in diagnosing breast cancer. By combining S-detect with junior radiologists, the pooled sensitivity was  $0.89$  (95% CI:  $0.83-0.93$ ,  $I^2=74.48\%$ ), the pooled specificity was  $0.79$  (95% CI:  $0.72-0.84$ ,  $I^2=92.94\%$ ) (*Figure 4A*), and the AUC of the SROC curve was  $0.91$  (95% CI:  $0.88-0.93$ ) (*Figure 4B*).

### **Comparison of diagnostic accuracy between the combination of S-detect with senior radiologists and the combination of S-detect with junior radiologists**

Combining S-detect with senior radiologists, the pooled sensitivity, pooled specificity, and the AUC in differentiating benign from malignant breast masses were  $0.93$ ,  $0.86$  and  $0.96$ . The pooled sensitivity, pooled specificity, and the AUC of junior radiologists were  $0.89$ ,  $0.79$  and  $0.91$ . The pooled sensitivity, pooled specificity, and the AUC of senior radiologists plus S-detect were all higher than that of junior radiologists plus S-detect. There is significant difference in the AUC ( $P<0.01$ ), however, there were without significant difference in the pooled sensitivity ( $P=0.188$ ) and pooled specificity ( $P=0.086$ ).

### **Heterogeneity detection**

#### **Subgroup analysis**

To identify the potential factors contributing to heterogeneity, the original studies were divided into subgroups according to reference standard, number of included lesions, BI-RADS diagnostic threshold, the method of US + S-detect and the methodological quality of the study for subgroup analysis of the included literature in the senior and junior radiologist groups, respectively. The subgroup analysis results are shown in *Table 2* and *Table 3*.

In the senior radiologist group, the heterogeneity for sensitivity of the BI-RADS 3 and others group, the subjective and others group, and acceptable-quality group were reduced to 50%, indicating that these three factors were among the sources of heterogeneity of sensitivity in the original study. In the junior radiologist group, the heterogeneity for sensitivity of the No. of lesion  $<200$ , subjective and others, and acceptable-quality subgroups were reduced to 50%, indicating that these three factors were a source of heterogeneity of sensitivity in the original study.



**Table 1** Characteristics of the included studies

First author	Year	Country	Reference standard	Study design	Age (years) (mean ± SD)	No. of patients	No. of masses (benign/malignant)	Size (cm)	US equipment	BI-RADS edition	Seniority of doctor	BI-RADS threshold	Method of US + S-detect
Boyuan Xing (27)	2022	China	Biopsy, excision	NA	44.0±11.5	151	190 (136/54)	1.2±0.4	RS80A	5th	Senior	BI-RADS 4a	S-detect malignant, BI-RADS upgrades, S-detect benign, BI-RADS degrades
Xin-Yi Wang (22)	2021	China	Biopsy, excision	NA	50.8±15.5	167	173 (95/78)	1.6±0.9	RS80A	5th	Senior and junior	BI-RADS 3	S-detect benign, BI-RADS 4 and 5 degrade, if not, the original BI-RADS maintain
Yongping Liang (26)	2020	China	Excision	Prospective	43.11±12.55	261	398 (248/150)	1.92±1.26	RS80A	5th	Senior and junior	BI-RADS 4a	One side of S-detect malignant, BI-RADS upgrades, both sides of S-detect benign, BI-RADS degrades
Min Young Kim (21)	2021	Korea	Biopsy	Prospective	46±10	146	156 (146/10)	1.1±0.5	RS85	NA	Senior and junior	BI-RADS 3	Subjective judgment
Chenyang Zhao (25)	2020	China	Biopsy, excision	Retrospective	45.7	195	195 (113/82)	NA	RS85	NA	Junior	BI-RADS 3	S-detect benign, BI-RADS 4a, indicates benign, S-detect malignant, BI-RADS 3 indicates benign; others maintain original BI-RADS of doctor
Qun Xia (24)	2021	China	Biopsy	NA	50.9±13.9	40	40 (16/24)	NA	RS80A	5th	Senior and junior	BI-RADS 4a	Subjective judgment
Qi Wei (23)	2022	China	Biopsy, excision	Prospective	43±12	248	266 (197/69)	1.48±0.92	RS80A	5th	Senior and junior	Clinical experience	Subjective judgment
Eun Cho (19)	2018	Korea	Biopsy, excision	Retrospective	48.5±12.2	116	119 (65/54)	1.69±1.07	RS80A	5th	Senior and junior	BI-RADS 3	Subjective judgment
Fang He (18)	2018	China	Excision	Retrospective	42.6±6.2	42	54 (30/24)	1.423±0.447	RS80A	NA	Senior	BI-RADS 4b	S-detect benign, BI-RADS ≥4b indicates malignant, S-detect malignant, BI-RADS 3 indicates benign
Ji-Hye Choi (20)	2018	Korea	Biopsy, 2 years of follow-up	Prospective	49.5±11.8	200	200 (188/12)	1.2±0.8	RS80A	NA	Senior and junior	BI-RADS 3	Subjective, conjunctive, disjunctive
Wenjun Xu (28)	2020	China	Biopsy, excision	NA	46.15±12.61	136	145 (87/58)	1.71±0.78	RS80A	5th	Senior	BI-RADS 4a	S-detect malignant, BI-RADS upgrades, S-detect benign, BI-RADS degrades
Hong Yan (33)	2020	China	Excision	Retrospective	43.4±12.2	453	581 (411/170)	1.57±0.85	RS80A	5th	Junior	BI-RADS 4a	Subjective judgment
Jiazheng Pan (32)	2021	China	Excision	NA	46.64±13.88	169	175 (87/88)	1.87±0.93	RS80A	NA	Senior and junior	BI-RADS 4a	S-detect malignant, BI-RADS upgrades, S-detect benign, BI-RADS degrades

**Table 1** (continued)

Table 1 (continued)

First author	Year	Country	Reference standard	Study design	Age (years) (mean ± SD)	No. of patients	No. of masses (benign/malignant)	Size (cm)	US equipment	BI-RADS edition	Seniority of doctor	BI-RADS threshold	Method of US + S-detect
Xue Ge (35)	2020	China	Biopsy, excision	NA	51±15	168	168 (152/16)	NA	SW80A	NA	Senior and junior	BI-RADS 4b	Either S-detect and BI-RADS method is malignant, then malignant, both are benign, then benign
Xiang Li (36)	2019	China	Biopsy, excision	NA	44.02±12.13	435	563 (403/160)	1.93±0.93	RS80A	5th	Senior and junior	BI-RADS 4a	S-detect malignant, BI-RADS upgrades, S-detect benign, BI-RADS degrades
Yanyan Ge (34)	2020	China	Biopsy, excision	NA	44.2±12.0	88	98 (66/42)	1.77±1.14	RS80A	NA	Senior	BI-RADS 4a	Either S-detect and BI-RADS method is malignant, then malignant, both are benign, then benign
Feng Zhao (30)	2021	China	Biopsy, excision	Retrospective	44.2±12.0	120	120 (70/50)	1.25±0.53	RS80A	5th	Senior	BI-RADS 4a	Subjective, conjunctive, disjunctive
Jinhong Wang (29)	2019	China	Excision	Retrospective	46.02±12.53	79	92 (69/33)	1.73±0.88	RS80A	NA	Senior and junior	BI-RADS 4a	S-detect malignant, BI-RADS upgrades, S-detect benign, BI-RADS degrades
Lan Shen (31)	2022	China	Excision	Retrospective	42.69±12.34	135	162 (107/55)	1.87±0.51	RS80A	5th	Senior	NA	Either S-detect and BI-RADS method is malignant, then malignant, both are benign, then benign

US, ultrasound; BI-RADS, breast imaging reporting and data system; NA, not available.



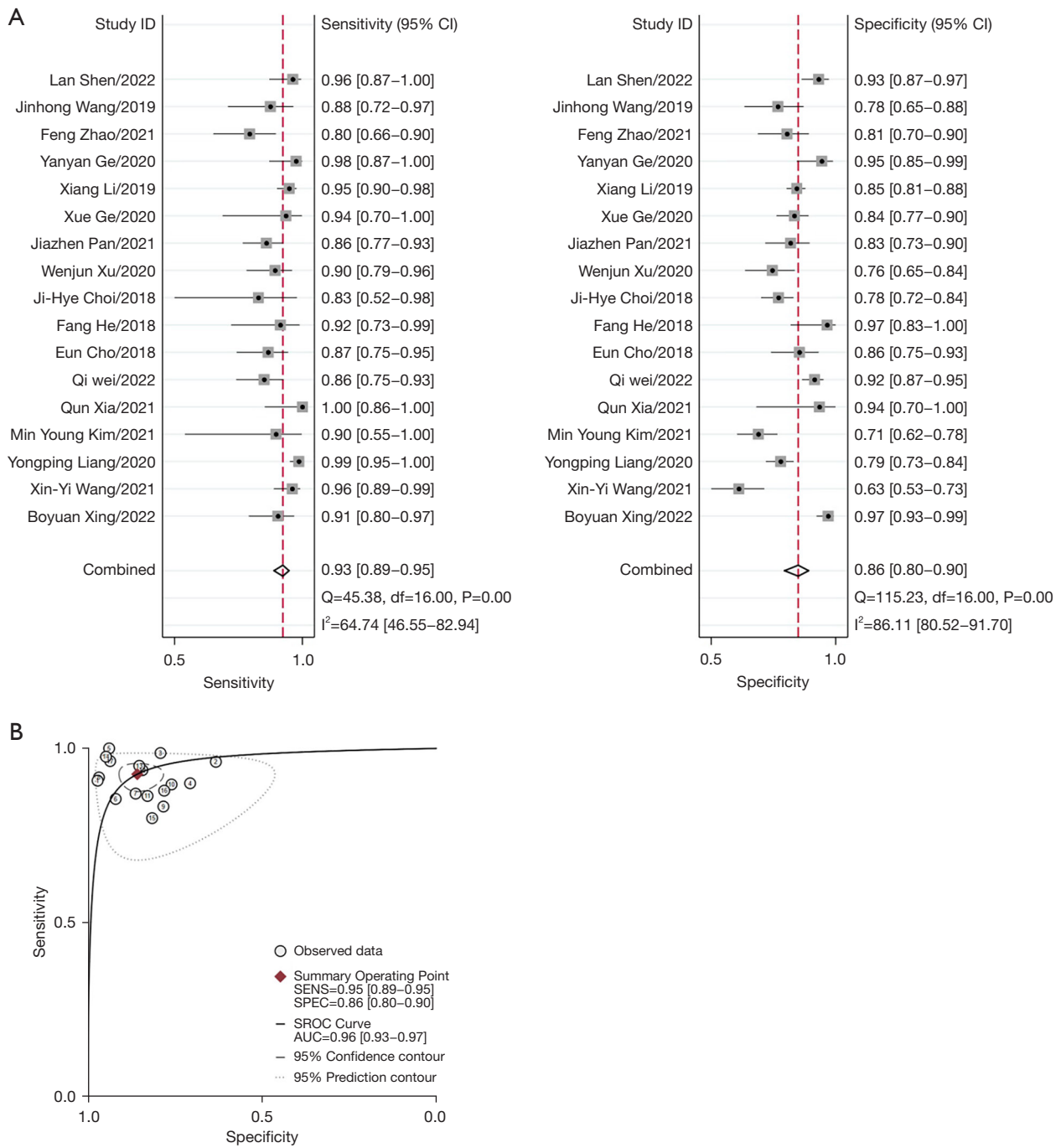
Figure 2 Methodological quality graphs (risk of bias and applicability concerns) of the included studies as percentages.

**Sensitivity analysis**

Sensitivity analyses were conducted separately for the senior and junior radiologist groups. After included literature were

removed successively, the remaining literature was combined to calculate the value of  $I^2$  for sensitivity analysis. and no significant effect of the literature on the results was found, indicating that the results of this meta-analysis are stable.



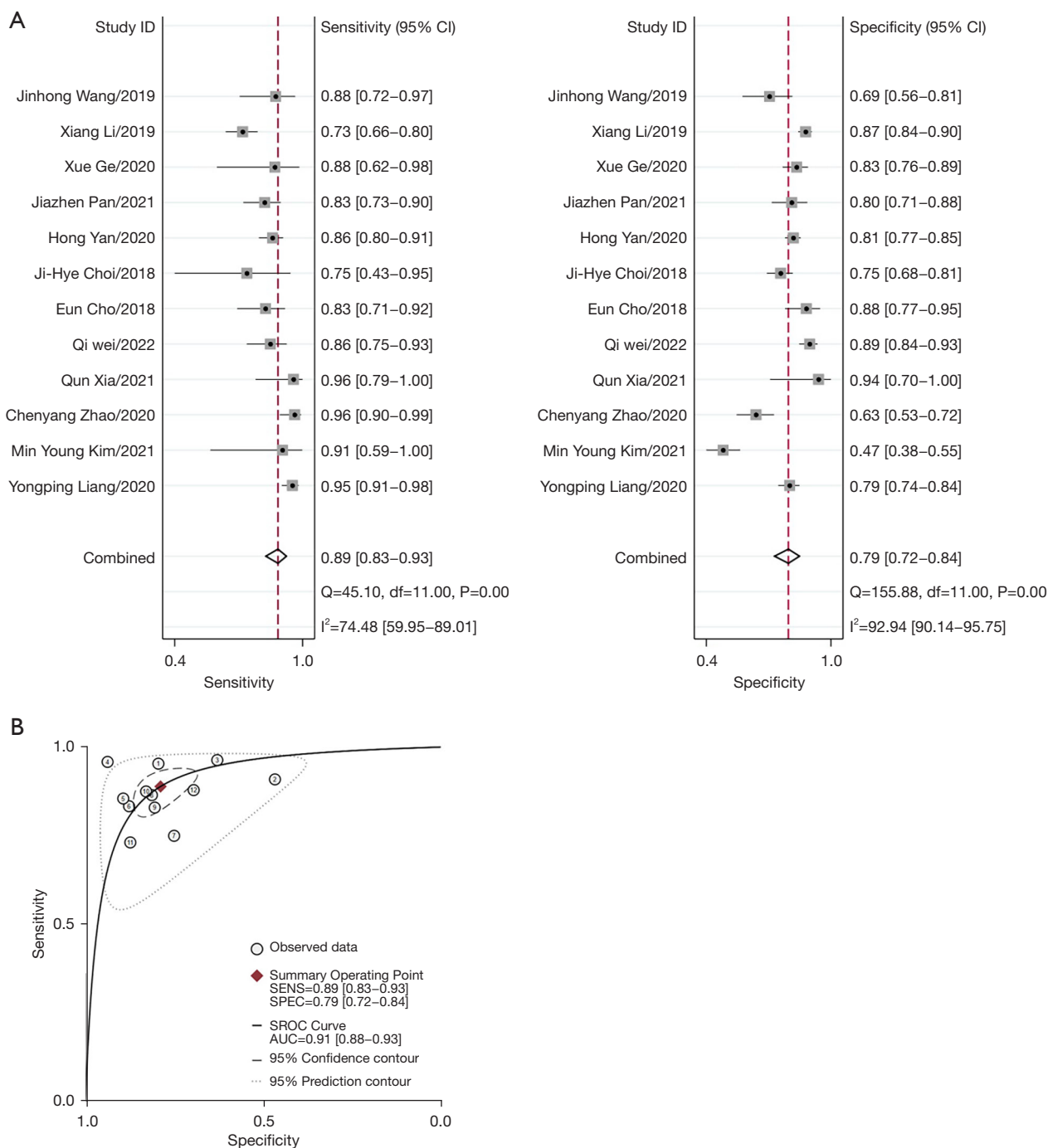


**Figure 3** (A) Forest plot of the sensitivity and specificity of US + S-detect for identifying breast cancer in the senior radiologists group. (B) SROC curve of US + S-detect for identifying breast cancer in the senior radiologist’s group. SENS, sensitivity; SPEC, specificity; SROC, summary receiver operating characteristic; AUC, the area under the curve; US, ultrasound.

**Clinical value analysis**

Fagan diagrams were constructed (Figure 5), assuming a pre-test probability of 50%, and post-test probabilities of

87% and 81% were obtained after calculation for the senior and junior radiologist groups, respectively. This indicated that the application of US + S-detect has a high clinical



**Figure 4** (A) Forest plot of the sensitivity and specificity of US + S-detect for identifying breast cancer in the junior radiologists group. (B) SROC of US + S-detect for identifying breast cancer in the junior group. SENS, sensitivity; SPEC, specificity; SROC, summary receiver operating characteristic; AUC, the area under the curve; US, ultrasound.

**Table 2** Subgroup analysis of the diagnostic performance of US + S-detect in the senior radiologist group

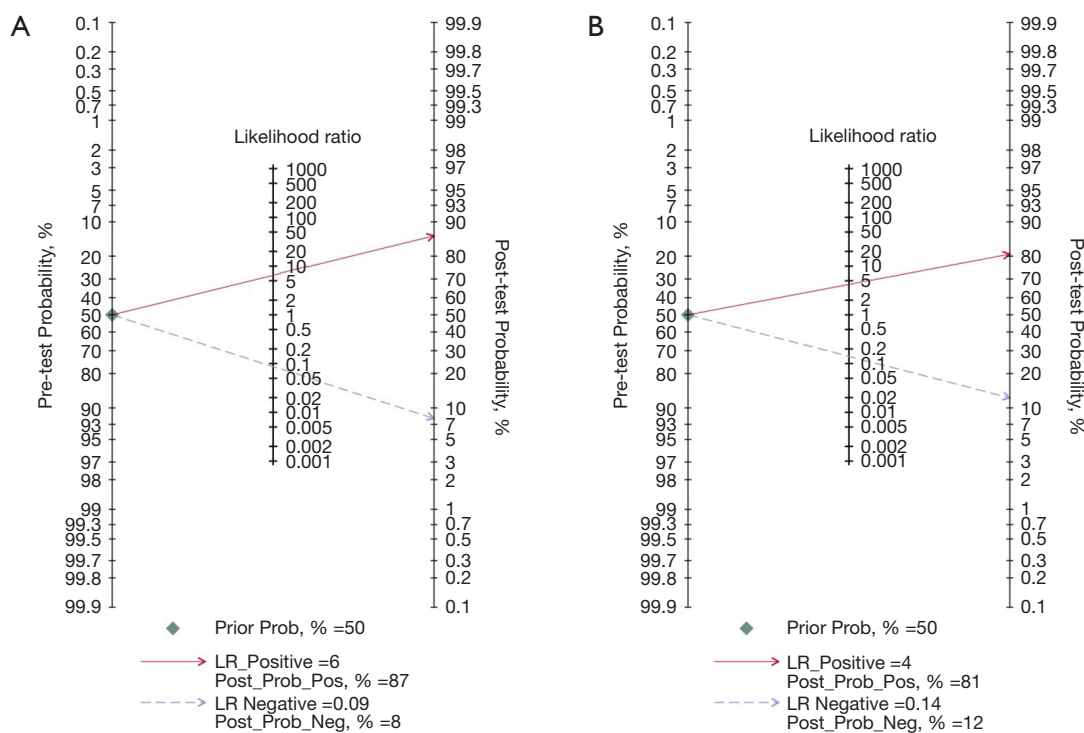
Covariate/subgroup	Studies, n	Sensitivity (95% CI)	I <sup>2</sup>	Specificity (95% CI)	I <sup>2</sup>
Reference test					
One method	7	0.95 (0.89–0.98)	73.21	0.85 (0.76–0.91)	84.51
Two methods	10	0.91 (0.87–0.94)	58.69	0.86 (0.79–0.91)	88.00
No. of lesion					
≥200	4	0.95 (0.89–0.98)	83.24	0.84 (0.78–0.89)	83.63
<200	13	0.92 (0.88–0.94)	52.94	0.87 (0.80–0.91)	88.93
BI-RADS diagnostic cut-offs					
BI-RADS 4a and 4b	11	0.93 (0.89–0.96)	73.46	0.87 (0.81–0.91)	76.90
BI-RADS 3 and others	6	0.92 (0.86–0.95)	33.37	0.83 (0.72–0.90)	90.35
US combined with the S-detect method					
Objective	12	0.94 (0.91–0.97)	66.63	0.87 (0.80–0.92)	87.34
Subjective and others	5	0.84 (0.78–0.89)	0	1.83 (0.75–0.89)	86.40
The methodological quality of the study					
High-quality	6	0.96 (0.90–0.98)	74.08	0.85 (0.77–0.90)	88.11
Acceptable-quality	11	0.91 (0.87–0.94)	45.62	0.85 (0.78–0.90)	85.09

US, ultrasound; BI-RADS, breast imaging reporting and data system.

**Table 3** Subgroup analysis of the diagnostic performance of US + S-detect in the junior radiologist group

Covariate/subgroup	Studies, n	Sensitivity (95% CI)	I <sup>2</sup>	Specificity (95% CI)	I <sup>2</sup>
Reference test					
One method	6	0.90 (0.84–0.94)	57.31	0.76 (0.63–0.85)	93.86
Two methods	6	0.86 (0.77–0.92)	76.41	0.82 (0.74–0.88)	90.42
No. of lesion					
≥200	5	0.87 (0.78–0.92)	86.80	0.83 (0.78–0.87)	82.43
<200	7	0.90 (0.83–0.94)	44.67	0.75 (0.63–0.84)	92.87
BI-RADS diagnostic cut-offs					
BI-RADS 4a and 4b	7	0.88 (0.81–0.92)	82.17	0.69 (0.56–0.82)	65.06
BI-RADS 3 and others	5	0.91 (0.83–0.95)	54.43	0.75 (0.58–0.87)	95.74
US combined with S-detect method					
Objective	7	0.90 (0.82–0.95)	86.31	0.78 (0.67–0.86)	95.99
Subjective and others	5	0.86 (0.80–0.90)	0	1.78 (0.64–0.88)	96.14
The methodological quality of the study					
High-quality	7	0.91 (0.83–0.96)	85.80	0.90 (0.82–0.95)	95.99
Acceptable-quality	5	0.84 (0.78–0.88)	0	0.79 (0.74–0.84)	58.12

US, ultrasound; BI-RADS, breast imaging reporting and data system.



**Figure 5** Fagan diagrams of US + S-detect for the diagnosis of breast cancer. (A) Senior radiologists group. (B) Junior radiologists group. LR, likelihood ratio; US, ultrasound.

application for the diagnosis of breast cancer.

**Discussion**

US is widely used for the diagnosis of breast lesions, but it is influenced by the subjective factors of the operator. Furthermore, it has been reported that when using the BI-RADS vocabulary for the assessment of breast masses, the specificity of less experienced radiologists is significantly lower than that of experienced radiologists (37). With the development of medical imaging technology, the feasibility of artificial intelligence technology applied to the diagnosis of breast diseases has been reported by several articles. S-detect reduces the influence of the subjective factors of radiologists to a certain extent, making the results more objective (38). In this meta-analysis, we collected domestic and international articles on the combined diagnosis of US + S-detect by senior and junior radiologists to investigate whether the combination of both could improve the discrimination of benign and malignant breast masses.

A previous meta-analysis (39) reported that the pooled sensitivity and specificity of S-detect in diagnosing breast

cancer were 0.82 and 0.86, respectively, and could be a complement to conventional US. In our meta-analysis, we selected 19 articles in which US + S-detect were combined to diagnose breast masses; seventeen of these articles evaluated the diagnostic performance of US + S-detect for breast cancer by senior radiologists and twelve articles evaluated the diagnostic performance of US + S-detect for breast cancer by junior radiologists. The diagnostic indexes of US + S-detect by senior and junior radiologists were respectively calculated and quantitatively pooled by meta-analysis. The literature included in both groups had no threshold effect, but the heterogeneity was significant, so a random-effects model was used for the combined analysis of data. The results showed that in the senior radiologist group, the pooled sensitivity, specificity, and AUC of US + S-detect were 0.93 (95% CI: 0.89–0.95), 0.86 (95% CI: 0.80–0.90), and 0.96 (95% CI: 0.93–0.97). In the junior radiologist group, pooled sensitivity, pooled specificity, and AUC of US + S-detect were 0.89 (95% CI: 0.83–0.93), 0.79 (95% CI: 0.72–0.84) and 0.91 (95% CI: 0.88–0.93).

It was suggested that the diagnostic performance was high in both groups; pooled sensitivity, the pooled

specificity and AUC were all higher in the senior radiologist group than in the junior radiologist group, and the AUC value was significantly different between the two groups ( $P < 0.01$ ). Some articles showed that the improvement was more significant in the junior radiologist group compared to US (19-21,23). Meanwhile, we analyzed the clinical application value of US + S-detect by plotting Fagan diagrams, setting the pre-test probability at 50% and the post-test probability at 87% and 81% for the senior and junior groups, respectively. The pre-test probability refers to the prevalence rate estimated by the clinician based on the patient's medical history, signs, and personal experience before the patient undergoes this test. The more the post-test probability increases compared to the pre-test probability, the more valuable the test is in terms of clinical application. Therefore, the US + S-detect is of high clinical value for the diagnosis of breast cancer by both senior and junior radiologists.

The malignancy rate of breast masses classified as BI-RADS 4a is approximately 2–10%, and most category 4a breast masses, although benign, may be subject to unnecessary biopsy (40,41). A previous meta-analysis by Park *et al.* (42) reported that downgrading of BI-RADS category 4a breast masses to BI-RADS category 3 when US elastography was combined with US for diagnosis reduced unnecessary biopsies in 41.1% of breast category 4a nodules. Combining S-detect can also reduce the biopsy rate by downgrading BI-RADS 4a masses. Combining S-detect to diagnose breast masses reduced the rate of unnecessary biopsies (22,23,25), but downgrading can also result in false-negative results for malignant nodes, so further training in S-detect and the empirical judgment of radiologists is also important.

Since the results of the heterogeneity test showed that the  $I^2$  values of sensitivity and specificity were  $>50\%$  for both the senior and junior radiologist groups, indicating a pronounced heterogeneity between the original studies, the causes of heterogeneity need to be explored. The results of the subgroup analysis to determine the factors that may lead to heterogeneity. It showed that the heterogeneity for sensitivity of the BI-RADS diagnostic cut-offs, US combined with the S-detect method, and the methodological quality of the study may lead to the sources of heterogeneity of sensitivity in the senior radiologist groups. In the junior radiologist group, The No. of lesion, US combined with the S-detect method, and the methodological quality of the study indicated that these three factors may be a source of heterogeneity of sensitivity

in the original study. The sources of heterogeneity were further explored by sensitivity analysis, and no significant effect of the literature on the results was observed, indicating good stability of the meta-analysis.

Limitations of this meta-analysis: (I) the study populations of most of the original studies were Chinese patients, leading to the possibility of geographical bias, and thus, more high-quality studies from other countries should be included in future studies to further validate the reliability of the diagnostic results. (II) There was a high degree of heterogeneity in the included literature, and some of the original studies included cases from a retrospective study method; therefore, more high-quality prospective multicenter studies are needed to further explore the diagnostic efficacy of US + S-detect for breast cancer in-depth.

## Conclusions

In this study, we collected diagnostic studies from domestic and foreign countries for meta-analysis to evaluate the diagnostic performance of US + S-detect for breast cancer by radiologists of different seniority levels. The combined results showed that the sensitivity and the AUC were all high in the senior and junior radiologist groups, which have a high clinical application value. However, the results of this study are somewhat heterogeneous, and thus, it is still necessary to collect more articles and accumulate a larger sample size for validation.

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## Footnote

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*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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