

Sensitivity and Specificity for Skin Cancer Diagnosis in Primary Care Providers: a Systematic Literature Review and Meta-analysis of Educational Interventions and Diagnostic Algorithms

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Accepted: 20 June 2022 / Published online: 14 July 2022 © The Author(s) under exclusive licence to American Association for Cancer Education 2022

Abstract

Background In areas without convenient access to dermatology care, primary care providers (PCPs) serve as an important patient resource for early skin cancer detection. To determine the most effective strategy for skin cancer detection training in PCPs, we conducted a systematic review of educational interventions and performed a meta-analysis on sensitivity and specificity outcomes in PCPs.

Objectives To summarize data on skin cancer sensitivity and specificity outcomes for PCP-targeted training programs and diagnostic algorithms. Our PCP cohort included practicing physicians, trainee physicians, and advanced practice practitioners. **Methods** A literature search was performed in MEDLINE, Embase, Web of Science, and the Cochrane Library for relevant English-language articles published worldwide from 2000 onward. Results were screened for eligibility, and overlapping datasets were reconciled. Data extracted included the educational intervention, diagnostic algorithm, and outcomes of interest (sensitivity and specificity). Outcomes were pooled across interventions that taught the same diagnostic algorithm. A bivariate model was fit to compare different interventions/algorithms. This review followed the PRISMA guidelines.

Results In total, 21 articles were included in this review, encompassing over 58,610 assessments of skin lesions by about 1529 participants worldwide. Training programs that implemented the triage-amalgamated dermoscopic algorithm (TADA) demonstrated high pooled sensitivity (91.7%) and high pooled specificity (81.4%) among PCPs.

Conclusions and Relevance Overall, this systematic review and meta-analysis showed that dermoscopy training in PCPs was generally associated with gains in skin cancer sensitivity without loss of specificity. Clinically, this correlates with fewer skin cancers overlooked by PCPs and fewer excisions of benign lesions.

Keywords Systematic literature review \cdot Meta-analysis \cdot Sensitivity \cdot Specificity \cdot Skin cancer \cdot Melanoma \cdot Early detection \cdot Primary care provider \cdot Advanced practice practitioner \cdot Nurse practitioner \cdot Physician assistant \cdot Skin cancer education \cdot Skin cancer detection \cdot Skin cancer diagnosis \cdot Secondary prevention

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Background

Skin cancer represents a major public health burden in the U.S. with an estimated 9000 Americans being diagnosed each day [1, 2]. Keratinocyte carcinoma (KC, i.e., basal cell carcinoma, squamous cell carcinoma) comprises a vast majority of skin cancer diagnoses with over 5.4 million KC cases diagnosed in over 3.3 million Americans in 2012 [2]. While the incidence of KC is much greater than that of melanoma, the latter has a significantly higher mortality rate [3]. For the year 2022, the American Cancer Society estimated 197,700 new cases of melanoma (99,780 invasive and 97,920 in situ) and 7650 deaths from melanoma [3]. In addition to its risks of morbidity and mortality for

As the global incidence of skin cancer continues to increase [2, 5], the early detection, or secondary prevention, of skin cancer becomes important to efforts seeking to decrease patient morbidity and mortality, especially for those with melanoma. If melanoma is diagnosed early and treated in a timely manner, the 5-year survival rate is highly favorable, nearing 99% [6]. However, once a melanoma metastasizes beyond its site of origin and spreads to regional lymph nodes (stage III disease) or distant organs (stage IV disease), the rates decrease to 65% and 25%, respectively. [6]

While dermatologists are specially trained in skin cancer detection, many Americans lack timely access to a dermatologist for diagnostic skin cancer examinations and procedures. Known barriers to dermatology care include patient socioeconomic status, race/ethnicity, residence in a rural county, and insurance type [7, 8]. On the county level, dermatologists tend to be concentrated in urban counties with higher median incomes [7]. Only 1 in 10 dermatologists practice in rural counties, and 88% of rural counties have 0 dermatologists [7]. With respect to race/ethnicity, most counties with populations reflecting African American, Hispanic, and/or Native American ethnic majorities were also found to have 0 dermatologists. [7]

Many Americans are thus frequently left without convenient access to specialized dermatology care for important skin cancer-related services. Patients without access may suffer delays in diagnosis and treatment, increasing the risk of disease progression [9]. While virtual solutions such as teledermatology help improve access [10], patients still require in-person visits for diagnostic procedures [7]. In these areas, primary care providers (PCPs), who work on the frontline of healthcare delivery, engage in skin cancer behavioral diagnosis and management.

However, many PCPs do not receive sufficient training in skin cancer detection during their post-graduate medical education [11]. Insufficient training may lead to numerous skin cancers being inadvertently overlooked or high volumes of benign skin lesions being needlessly excised [9]. While diagnosis of melanoma in its earlier stages is associated with decreased mortality, excision of benign lesions performed in the course of identifying melanoma not only bears financial consequences but also contributes to patient morbidity [9]. Undergoing a biopsy procedure and waiting for the pathology results may induce a sense of anxiety in patients, and the procedure itself will invariably result in a physical scar. [9]

Objectives

For the above reasons, numerous efforts have been devoted to the development of educational interventions and diagnostic aids (e.g., algorithms, mnemonics) that support sensitive yet specific skin cancer diagnosis by PCPs. To determine the effectiveness of these initiatives, we conducted a systematic review and quantitative synthesis of skin cancer educational interventions and diagnostic algorithms evaluated in PCP populations.

Our review comprised articles published between January 2000 and June 2021, and our PCP educational cohort included practicing physicians, trainee physicians, and advanced practice practitioners (APPs). We analyzed their skin cancer sensitivity and specificity outcomes through a meta-analysis. By comparing these key learning outcomes across multiple studies, this review will inform the future development of PCP-targeted programs seeking to adopt evidence-based approaches for skin cancer detection training.

Methods

This review was conducted in accordance with the PRISMA (preferred reporting items for systematic reviews and meta-analyses) guidelines. [12]

Eligibility Criteria

The inclusion and exclusion criteria for this review are listed in Online Resource 1. Studies deemed eligible for inclusion evaluated skin cancer detection training programs or diagnostic algorithms in PCPs. Eligible studies measured the effectiveness of the particular program/algorithm in terms of sensitivity and specificity for skin cancer diagnosis, including melanoma diagnosis. Studies that did not explicitly report these outcomes but provided sufficient data for calculations by the research team were included. All studies that involved technology deemed inaccessible to most PCPs or assessed computer-aided diagnosis were excluded.

For our population of interest, primary care physicians were defined as MDs or DOs practicing in family medicine, internal medicine, medicine/pediatrics, or obstetrics/gynecology. Studies that involved PCP trainees and/or APPs, such as nurse practitioners (NPs) or physician assistants (PAs), in the educational cohort were included. Studies that involved majority (> 50%) non-PCPs (e.g., dermatology physicians/ trainees, medical students, laypeople) without segregation of data between PCP and non-PCP participants were excluded.

Data Sources

A medical research librarian (D.P.F.) searched MEDLINE (Ovid), Embase (Ovid), Web of Science (Clarivate), and the Cochrane Library (Wiley) for relevant articles published from January 1, 2000, to June 22, 2021. For each database, the librarian developed and tailored a search strategy in consultation with the research team and selected controlled vocabulary (MeSH and Emtree) and natural language terms for the concepts of *melanoma*, *dermoscopy*, and *diagnostic algorithm*. The search strategy implemented for each database is shown in Online Resource 2.

Searches were limited to the English language, but no other limiters or published search filters were used. Grey literature (e.g., conference proceedings, dissertations, reports, unpublished data) were included in addition to peer-reviewed articles. Previous review articles related to PCP-targeted training programs on skin cancer detection were excluded from data analysis but were closely examined by the team to identify relevant manuscripts not found during the search process. EndNote X9 (Clarivate) was used to deduplicate search results, and all unique records were identified and uploaded to Rayyan, a web-based software developed to help filter and manage search results. [13]

Study Selection

Two authors (T. T. and N. G.) independently reviewed all results generated from the search process for study eligibility, as depicted in the PRISMA flow diagram in Fig. 1. Titles and abstracts were screened using Rayyan [13]. For studies that passed the initial screening, full-text manuscripts were

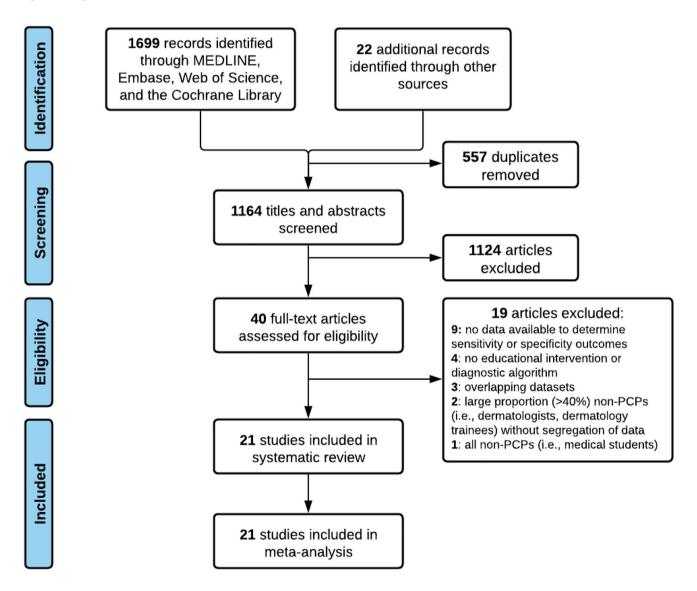


Fig. 1 PRISMA flow diagram of the study selection process. Abbreviations: *PRISMA*, preferred reporting items for systematic reviews and meta-analyses; *MEDLINE* Medical Literature Analysis and Retrieval System Online; *PCP*, primary care provider

retrieved and independently assessed for eligibility. A third author (K. C. N.) provided the final decision in the event of disagreement.

Data Extraction

For articles deemed appropriate for inclusion, two authors (T. T. and N. G.) independently reviewed the full-text manuscripts plus any supplemental materials and independently extracted data. Data extracted included characterization of the educational cohort, educational intervention, and diagnostic algorithm (if applicable) as well as reported outcome measures. Studies that randomized participants to different educational exposures were treated as separate educational cohorts per their allocation.

The primary outcomes of interest were sensitivity (proportion of malignant skin lesions correctly diagnosed) and specificity (proportion of benign skin lesions correctly diagnosed). Related outcomes of interest included the total number of true positives (TP, number of malignant lesions correctly classified as malignant by the participant), false negatives (FN, number of malignant lesions incorrectly classified as benign), true negatives (TN, number of benign lesions correctly classified as benign), and false positives (FP, number of benign lesions incorrectly classified as malignant). If these values were not reported, the research team made reasonable efforts to calculate them using available published data. All calculations performed by the research team are shown in Online Resource 3. In some cases, values were extrapolated from graphical displays or obtained from correspondence with original investigators.

Data Analysis

To compare outcomes across multiple educational interventions and diagnostic algorithms, a meta-analysis was performed by a biostatistician (R. L.B.). For the purposes of this study, dermoscopic and clinical interventions were considered separately. Dermoscopic interventions trained participants in the use of dermoscopy for skin cancer diagnosis, whereas clinical interventions taught skin cancer diagnosis using the naked eye.

Data was aggregated across interventions by diagnostic algorithm. Pooled sensitivity and specificity outcomes were estimated using a bivariate linear mixed model with known variances of random effects, and variance components were estimated using restricted maximum likelihood [14]. For studies that provided both pre- and post-intervention data, odds ratios (ORs) were calculated and visualized using forest plots. The post-intervention datasets used for the metaanalyses have been made available in Online Resource 4. Statistical analysis and figure production were performed using the statistical software R (Version 4.1.1).

Results

The literature search retrieved a total of 1699 records (MED-LINE, n = 403; Embase, n = 632; Web of Science, n = 631; and the Cochrane Library, n = 33). The team also identified 22 additional records for screening from reference lists [15, 16]. Following de-duplication, 1164 unique records were identified and screened for eligibility (Fig. 1). In the initial round of screening, 1124 records were excluded based on their titles/abstracts, leaving 40 manuscripts for full-text review. Another 19 articles were then excluded for reasons listed in the PRISMA flow diagram (Fig. 1), the most common being insufficient reporting of the outcomes of interest without means for calculations by the team. For articles with overlapping datasets, the most recently published dataset was favored for inclusion. Ultimately, 21 articles were selected for inclusion in the quantitative synthesis and meta-analysis [17-37]. All the included articles were peer-reviewed publications. While grey literature were included in the search and selection process, none was eventually included in the quantitative synthesis or meta-analysis.

Study Designs

An overview of the 21 included articles is provided in Table 1 (dermoscopic interventions) and Table 2 (clinical interventions). The following study designs are represented in this review: 3 randomized controlled trials (RCTs) [18–20], 1 prospective cohort study [29], and 17 prospective cross-sectional studies. [17, 21–28, 30–37]

Study Populations

This review encompassed studies from 12 different countries: Australia [17, 20], Belgium [28, 34], Canada [24], Colombia [25], Ireland [31], Italy [18, 33], the Netherlands [19, 27], Serbia [26], Spain [18], Switzerland [32], the UK [35], and the USA [21–23, 29, 30, 36, 37]. All articles in this review recruited practicing PCPs. Five included APPs, such as NPs [17, 22, 29, 30, 37] and PAs [22], with one of these comprising only NPs [37]. Three included PCP trainees [27, 28, 31] with two of these comprising only PCP trainees [27, 31]. Three also included minority (<50%) non-PCPs (e.g., general surgeons, plastic surgeons) in the educational cohort. [26, 30, 36]

Educational Interventions

To improve diagnostic accuracy by PCPs, many interventions taught skin cancer detection with a dermoscopy training component [17–27]. In terms of delivery method, most

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|---|--------------|-----------------------|---|------------------------------------|-------------------|--|------------------------------------|--------------------------------------|--------------------------------------|
| Author, year, loca- tion | Group no | Group no Study design | Educational inter- vention | Diagnostic algo- rithm | Delivery format | No. observations per participant | No. participants in group | Post- intervention sensitivity | Post- intervention specificity |
| Bourne et al., 2012, Australia [17] | la | DA: CS (P) A | Dermoscopy train- ing | 3-point checklist | Live | 45 lesions with der- moscopic images | 4 (3 PCPs, 1 NP) | 36.2% | 65.3% |
| Argenziano et al., 2006, Italy and Spain [18] | 2a | RCT | Dermoscopy train- ing + clinical training | 3-point checklist | Live | 33.4 mean no. lesions evaluated in person | 36 PCPs | 79.2% | 71.8% |
| Koelink et al., 2014, The Netherlands [19] | 3a | RCT | Dermoscopy train- ing+clinical training | 7-point checklist | Live | 7.2 mean no. lesions evaluated in person | 27 PCPs | 65.9% | 46.0% |
| Bourne et al., 2012, Australia [17] | lb | DA: CS (P) | Dermoscopy train- ing | BLINCK algorithm | Live | 45 lesions with clinical and der- moscopic images | 4 (3 PCPs, 1 NP) | 52.7% | 89.9% |
| Bourne et al., 2012, Australia [17] | 1c | DA: CS (P) | Dermoscopy train- ing | Menzies method | Live | 50 lesions with der- moscopic images | 4 (3 PCPs, 1 NP) | 49.3% | 73.4% |
| Westerhoff et al., 2000, Australia [20] | 4 | RCT | Dermoscopy train- ing (melanoma detection) | Menzies method | Live | 100 atypical PSL with dermoscopic and clinical images | 37 PCPs | 75.9% | 57.8% |
| Seiverling et al., 2021, USA [21] | 5 | DA: CS (P) B/A | Dermoscopy train- ing | TADA | Live | 30 lesions with der- moscopic images | 96 PCPs | 87.0% | 76.4% |
| Susong et al., 2020, USA [22] | 6 | DA: CS (P) B/A | Dermoscopy train- ing (live delivery) | TADA | Live or web-based | 30 lesions with der- moscopic images | 102 PCPs, NPs and PAs ^a | 89.5% | 87.7% |
| Rogers et al., 2016, USA [23] | ٢ | DA: CS (P) A | Dermoscopy train- ing | TADA | Live | 50 lesions with der- moscopic images | 49 (41 PCPs, 8 other) | 93.7% | 72.1% |
| Sawyers et al., 2020, Canada [24] | × | DA: CS (P) B/A | Dermoscopy train- ing | TADA (step I) | Live | 50 lesions with dermoscopic and clinical images | 31 PCPs | 94.7% | 86.1% |
| De Bedout et al., 2021, Colombia [25] | 6 | DA: CS (P) B/A | Dermoscopy train- ing + clinical training | Not specified | Live | 50 lesions with clinical and der- moscopic images | 21 PCPs (rural) | 83.8% | 77.9% |
| Bandic et al., 2020, Serbia [26] | 10 | DA: CS (P) A | Teledermoscopy training + clinical training | Not specified | Live | 24.2 mean no. PSL evaluated in person | 5 (3 PCPs, 2 sur- geons) | 89.1% | 92.0% |
| Secker et al., 2017, The Netherlands [27] | 11 | DA: CS (P) B/A | Dermoscopy train- ing + clinical training | Not specified | Live | 20 lesions with clinical ± dermo- scopic images ^b | 293 PCPs (trainees included) | 59.2% | 64.3% |

^a59 participants in live version, 43 participants in web-based version; ^bdermoscopic images: – in pre-test, + in post-test. Abbreviations: *study designs—RCT*, randomized controlled trial; *DA*, diagnostic accuracy; *CS (P)*, cross-sectional (prospective); *B/A*, before/after; *A*, after only. *Interventions/algorithms—BLINCK*, benign, lonely, irregular, nervous/change, known clues; *TADA*, triage amalgamated diagnostic algorithm; *ABCDE*, asymmetry, border, color, diameter, evolving. Lesions—*PSL*, pigmented skin lesions. Participants—*PCP*, primary care provider; *NP*, nurse practitioner; *PA*, physician assistant

Description Springer

| Author, year, loca- tion | Group no | Group no Study design | Educational inter- vention | Diagnostic algo- rithm | Delivery format | No. observations per participant | No. participants in group | Post- intervention sensitivity | Post- intervention specificity |
|---|----------|-------------------------|--|--|-----------------|--|---|--------------------------------------|--------------------------------------|
| Harkemanne et al., 2020, Belgium [28] | 12 | DA: CS (P) B/A | Clinical training (melanoma detec- tion) | ABCD, ugly duck- ling sign, pattern analysis | Live | 15 PSL with clinical images | 56 (29 PCPs and 27 PCP trainees) | 78.9% | 76.4% |
| Eide et al., 2013, USA [29] | 13 | DA: 1-cohort (P) B/A | Clinical training (INFORMED) | ABCDE. ugly duck- ling sign | Web-based | 25 lesions with clinical images | 54 (53 PCPs, 1 NP) | 48.8% | 56.7% |
| Argenziano et al., 2006, Italy and Spain [18] | 2b | RCT | Clinical training | ABCD | Live | 36.0 mean no. lesions evaluated in person | 36 PCPs | 54.1% | 71.3% |
| Orfaly et al., 2021, USA [30] | 14 | DA: CS (P) B/A | Clinical training (INFORMED) | Not specified | Web-based | 48 PSL with clinical 40 (? PCPs, ≤14 images NPs, ≤5 derma tologists, ? oth | 40 (? PCPs, ≤14 NPs, ≤5 derma- tologists, ? other) | 93.3% | 84.6% |
| Beecher et al., 2018, Ireland [31] | 15 | DA: CS (P) B/A | Clinical training | Not specified | Live | 40 lesions with clinical images | 23 PCP trainees | 70.0% | %0.69 |
| Koelink et al., 2014, The Netherlands [19] | 3b | RCT | Clinical training | Not specified | Live | 8.5 mean no. lesions evaluated in person | 27 PCPs | 60.9% | 35.2% |
| Badertscher et al., 2013, Switzerland [32] | 16 | DA: CS (P) B/A | Clinical training | Not specified | Live | 36 lesions with clinical images | 78 PCPs | 82.7% | 76.6% |
| Carli et al., 2005, Italy [33] | 17 | DA: CS (P) B/A | Clinical training (melanoma detec- tion) | Not specified | Live | 15 PSL with clinical 41 PCPs images | 41 PCPs | 74.4% | 84.4% |
| Brochez et al., 2001, Belgium [34] | 18 | DA: CS (P) B/A | Clinical training (melanoma detec- tion) | Not specified | Live | 13 PSL with clinical 160 PCPs images | 160 PCPs | 84.0% | 70.0% |
| Bedlow et al., 2001, UK [35] | 19 | DA: CS (P) B/A | Clinical training | Not specified | Live | 30 lesions with clinical images | 17 PCPs | 75.9% | 61.8% |
| Harris et al., 2001, USA [36] | 20 | DA: CS (P) B/A | Clinical training | Not specified | Web-based | 15 lesions with clinical images | 354 (232 PCPs, 8 dermatologists, 114 other ^a) | 91.0% | 89.0% |
| Oliveria et al., 2001 USA [37] | 21 | DA: CS (P) B/A | Clinical training | Not specified | Live | 34 lesions with clinical images | 5 NPs | 100.0% | 73.7% |

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interventions evaluated this review provided live/synchronous instruction with the exception of three that provided web-based/asynchronous instruction only [29, 30, 36]. Of these three, two utilized an online curriculum called INFORMED (INternet curriculum FOR Melanoma Early Detection) that was rigorously designed to improve the skin cancer diagnostic skills of PCPs but is no longer available. [29, 30]

Diagnostic Algorithms

For skin cancer diagnosis, diagnostic algorithms refer to mnemonic aids that may be used during the evaluation of concerning skin lesions. These algorithms tend to emphasize the most salient diagnostic features. Clinical algorithms identified in this review included the ABCD(E) algorithm [18, 26, 28, 29] and the ugly duckling sign [28, 29]. Since these algorithms are commonly taught in medical education, it is likely that most training programs included some mention of ABCD(E) and the ugly duckling sign even if not explicitly stated. Dermoscopic algorithms identified in this review included the 3-point checklist [17, 18], 7-point checklist [19], BLINCK algorithm [17], Menzies method [17, 20], and triage amalgamated dermoscopic algorithm (TADA) [21–24]. Each algorithm/method is described in further detail in Online Resource 5. In addition to dermoscopic patterns and features, BLINCK also considered findings from the patient's history and clinical examination (e.g., whether the spot is different from others, the patient's degree of concern). [17]

Sensitivity and Specificity

In sum, this review encompassed over 58,610 assessments of skin lesions by about 1529 PCPs worldwide. Pooled sensitivity analyses relied on the total number of TP and FN across all participants for a given diagnostic algorithm, and pooled specificity on the total number of TN and FP. Pooled analyses results for both the dermoscopic and clinical interventions can be found in Online Resource 6.

Among the dermoscopic interventions, TADA was found to exhibit a high pooled sensitivity (91.7%) and high pooled specificity (81.5%) on over 10,800 total skin lesion assessments by 278 participants including NPs and PAs. In addition, the dermoscopy training programs developed by Bandic et al. [26] (121 skin lesion assessments by five participants including two general surgeons) and De Bedout et al. [25] (952 skin lesion assessments by 21 rural PCPs) had relatively high post-intervention sensitivity (89.1% and 83.8%, respectively) and specificity (92.0% and 77.9%, respectively).

Among the clinical interventions, those developed by Oliveria et al. [37], Orfaly et al. [30], and Harris et al. [36] all

demonstrated sensitivity outcomes exceeding 90% (100%, 93.3%, and 91.0%, respectively). Of these three, the Orfaly et al. [30] and Harris et al. [36] interventions also exhibited high specificity (84.6% and 89.0%, respectively), while the Oliveria et al. [37] intervention exhibited a lower specificity (73.7%). Of note, the Orfaly et al. [30] intervention utilized the INFORMED curriculum.

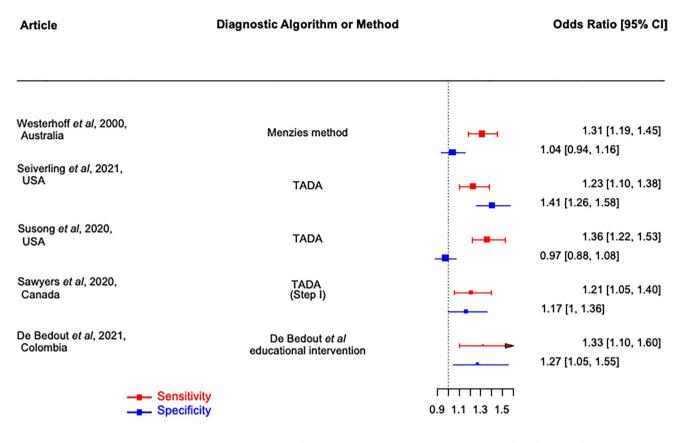
Relevant outcomes of studies reporting both pre- and post-intervention data were visualized using forest plots. Figure 2 displays pre- vs. post-intervention data for dermoscopic interventions in terms of ORs. A positive OR suggests that outcomes favor the post-intervention condition, whereas a negative OR suggests the contrary. Educational cohorts that used TADA demonstrated improvement in sensitivity for skin cancer on post-intervention assessments compared to baseline [21, 22, 24]. Among the TADA interventions, specificity for skin cancer either remained the same [24] or increased [21, 24]. The clinical interventions evaluated in this review also generally resulted in improvements in sensitivity for skin cancer without significant loss of specificity. Forest plots for the clinical interventions are displayed in Online Resource 7.

Discussion

In regions with limited access to specialized dermatology care, PCPs engage in skin cancer detection, diagnosis, and management [38], but there is currently no standardized curriculum available to PCPs that teach early skin cancer detection [16]. The potential negative consequences of insufficient training-whether overlooking skin cancers or needlessly excising benign lesions-highlight the need for evidencebased programs designed to improve skin cancer diagnosis by PCPs [9]. To determine the most effective strategy for skin cancer education in PCPs, we conducted a systematic review of educational interventions and diagnostic aids evaluated in cohorts with over a majority ($\geq 60\%$) PCPs. We determined the effectiveness of a particular curriculum/ algorithm using participants' post-intervention sensitivity and specificity outcomes. To our knowledge, this is the first meta-analysis of sensitivity and specific outcomes for PCPtargeted training programs on skin cancer detection.

In this review, we identified several interventions that resulted in relatively high (>70%) sensitivity and specificity for skin cancer in PCPs [18, 21–26, 28, 30, 33, 34, 36, 37]. However, our analyses were complicated by heterogeneity in some educational cohorts. PCPs overall differed in terms of their career stage (e.g., trainee vs. attending physician), specialty (e.g., family medicine vs. internal medicine), years of experience evaluating skin lesions, and previous skin cancer detection or dermoscopy training [21]. Some educational cohorts also included non-PCPs





Favors Pre-Intervention $\leftarrow \rightarrow$ Favors Post-Intervention

Fig. 2 Forest plot of pre- vs. post-intervention sensitivity and specificity outcomes for <u>dermoscopic</u> educational interventions. Abbreviations: *TADA*, triage amalgamated dermoscopic algorithm

[23, 26, 30, 36]. Two interventions, in particular, included a number of dermatologists in their cohorts without segregation of data: the Orfaly et al. intervention [30] with ≤ 5 ($\leq 12.5\%$) dermatologists and the Harris et al. intervention [36] with 8 (2.3%) dermatologists. These two interventions were included in data analysis owing to the relatively small number of dermatologists in each cohort, but the presence of the dermatologists' data may have positively skewed educational outcomes.

Of the 21 interventions included in this review, 52.4% (11/21) provided dermoscopy training, or instruction in the use of a dermatoscope (a non-invasive visualization tool) for skin examinations. With appropriate training, the use of dermoscopy in evaluating suspicious skin lesions improves the ability of PCPs to accurately diagnose and appropriately triage patients [39, 40]. Ongoing efforts are seeking to develop consensus-based proficiency standards for dermoscopy that are specific to the practice needs of PCPs.

In evaluating the effectiveness of a particular curriculum/algorithm, it is important to consider its sensitivity or specificity for skin cancer given the clinical relevance of these measures. While auditing participants' real-world clinical assessments of skin lesions would constitute best practice, educational outcomes are often evaluated using assessments containing clinical and/or dermoscopic images of skin lesions. In this review, only three (14.3%) studies used participants' in-person evaluations of suspicious skin lesions in real-life clinical practice [18, 19, 26], and the remainder used sets of clinical and/or dermoscopic images. Ideally, image sets used in these assessments would undergo formal validation by a panel of experts who determine whether images are of appropriate quality for PCP learners and whether image sets are of similar difficulty. Otherwise, lesions later deemed "problematic" may complicate analyses, as was the case for one article [36]. Classification of benign and malignant diagnoses on these assessments should also be consistent, especially for suspicious or borderline lesions (e.g., squamous cell carcinoma in situ, keratoacanthoma, actinic keratosis, dysplastic nevus).

Study Limitations

Our meta-analysis was limited by the number of articles providing enough data to calculate TP, FN, TN, and FP across all participants. In this review, many of these values were estimated based on the reported number of participants, number of test items, and percentage of malignant and benign lesions diagnosed correctly. However, some participants may have submitted incomplete assessments with missing answers, so TP, FN, TN, and FP may be slightly overestimated for some studies. In addition, this review only included articles published in the English language and may have missed reports of educational interventions and/ or diagnostic algorithms published in other languages. Relevant articles of interest may have also been inadvertently overlooked during the screening process.

Conclusions

While PCPs play an important role in skin cancer detection in underserved areas, they may require additional training to accurately diagnose and appropriately manage skin cancer. To determine the effectiveness of different training programs, it is important to evaluate participants' diagnostic performance in terms of sensitivity and specificity. However, this review identified relatively few PCP-targeted skin cancer educational interventions reporting these clinically relevant outcomes. To support further rigorous investigations of cancer detection education for PCPs, future studies should utilize validated instruments with a sufficient number of test items and segregate outcomes data between PCPs and non-PCPs. Among the training programs evaluated in this review, those that implemented TADA were found to demonstrate high sensitivity and specificity for skin cancer among PCP participants.

Supplementary Information The online version contains supplementary material available at https://doi.org/10.1007/s13187-022-02194-4.

Author Contribution KCN, NG, and TT contributed to the study conceptualization and design. Literature search was performed by KCN, DPF, NG, and TT. Study selection and data collection were performed by TT, NG, and KCN. Data analysis was performed by RLB, TT, and KCN. The first draft of the manuscript was written by TT and NG, and all authors commented on previous versions of the manuscript. All authors reviewed and approved the final manuscript.

Funding This work is supported in part by the generous philanthropic contributions of the Lyda Hill Foundation to The University of Texas MD Anderson Cancer Center Moon Shots ProgramTM. This work is

also supported in part by a National Cancer Institute (NCI) Cancer Center Support Grant (Award No. P30 CA016672). The funding sources were not involved in the design or conduct of the study; the collection, management, analysis, or interpretation of the data; the preparation, review, or approval of the manuscript; or the decision to submit the manuscript for publication.

Data Availability All data compiled and analyzed in this review has been made available to the reader.

Declarations

Ethics Approval This is a secondary review of literature which has been published. Ethics approval was not required.

Consent to Participate This review of published literature did not directly involve any human subjects. In this study, informed consent was not required.

Conflict of Interest The authors declare no competing interests.

References

- US Department of Health and Human Services (2014) The surgeon general's call to action to prevent skin cancer. Office of the Surgeon General (US), Washington (DC). Skin cancer as a major public health problem. Available from: https://www.ncbi.nlm.nih. gov/books/NBK247164/
- Rogers HW, Weinstock MA, Feldman SR, Coldiron BM (2015) Incidence estimate of nonmelanoma skin cancer (keratinocyte carcinomas) in the U.S. population, 2012. JAMA Dermatol 151(10):1081–1086. https://doi.org/10.1001/jamadermatol.2015. 1187
- Siegel RL, Miller KD, Fuchs HE, Jemal A (2022) Cancer statistics, 2022. CA Cancer J Clin 72(1):7–33. https://doi.org/10.3322/ caac.21708
- Guy GP Jr, Machlin SR, Ekwueme DU, Yabroff KR (2015) Prevalence and costs of skin cancer treatment in the US, 2002–2006 and 2007–2011. Am J Prev Med 48(2):183–187. https://doi.org/ 10.1016/j.amepre.2014.08.036
- Apalla Z, Lallas A, Sotiriou E, Lazaridou E, Ioannides D (2017) Epidemiological trends in skin cancer. Dermatol pract conceptual 7(2):1–6. https://doi.org/10.5826/dpc.0702a01
- Howlader NNAM, Noone AM, Krapcho ME, Miller D, Brest A, Yu ME, Ruhl J, Tatalovich Z, Mariotto A, Lewis DR, Chen HS (2019) SEER cancer statistics review, 1975–2016. National Cancer Institute, p 1
- Vaidya T, Zubritsky L, Alikhan A, Housholder A (2018) Socioeconomic and geographic barriers to dermatology care in urban and rural US populations. J Am Acad Dermatol 78(2):406–408. https://doi.org/10.1016/j.jaad.2017.07.050
- Alghothani L, Jacks SK, Vander Horst A, Zirwas MJ (2012) Disparities in access to dermatologic care according to insurance type. Arch Dermatol 148(8):956–957. https://doi.org/10.1001/ archdermatol.2012.804
- Nelson KC, Swetter SM, Saboda K, Chen SC, Curiel-Lewandrowski C (2019) Evaluation of the number-needed-to-biopsy metric for the diagnosis of cutaneous melanoma: a systematic review and meta-analysis. JAMA Dermatol 155(10):1167–1174. https:// doi.org/10.1001/jamadermatol.2019.1514
- 10. Perera E, Xu C, Manoharan S (2012) Real-life teledermatology cases. In: Soyer H, Binder M, Smith A, Wurm E (eds) Telemedicine

in Dermatology. Springer, Berlin, Heidelberg. https://doi.org/10. 1007/978-3-642-20801-0_15

- Moore MM, Geller AC, Zhang Z et al (2006) Skin cancer examination teaching in US medical education. Arch Dermatol 142(4):439– 444. https://doi.org/10.1001/archderm.142.4.439
- Moher D, Liberati A, Tetzlaff J, Altman DG (2009) Preferred reporting items for systematic reviews and meta-analyses: The PRISMA statement. BMJ 339:b2535. https://doi.org/10.1136/bmj.b2535
- Ouzzani M, Hammady H, Fedorowicz Z, Elmagarmid A (2016) Rayyan—a web and mobile app for systematic reviews. Syst Rev 5(210):10. https://doi.org/10.1186/s13643-016-0384-4
- Reitsma JB, Glas AS, Rutjes AW, Scholten RJ, Bossuyt PM, Zwinderman AH (2005) Bivariate analysis of sensitivity and specificity produces informative summary measures in diagnostic reviews. J Clin Epidemiol 58(10):982–990. https://doi.org/10.1016/j.jclinepi. 2005.02.022
- Harkemanne E, Baeck M, Tromme I (2021) Training general practitioners in melanoma diagnosis: A scoping review of the literature. BMJ Open 11(3):e043926. https://doi.org/10.1136/bmjop en-2020-043926
- 16 Najmi M, Brown AE, Harrington SR, Farris D, Sepulveda S, Nelson KC (2021) A systematic review and synthesis of qualitative and quantitative studies evaluating provider, patient, and health care system-related barriers to diagnostic skin cancer examinations. Arch Dermatol Res. https://doi.org/10.1007/s00403-021-02224-z
- Bourne P, Rosendahl C, Keir J, Cameron A (2012) BLINCK-A diagnostic algorithm for skin cancer diagnosis combining clinical features with dermatoscopy findings. Dermatol Pract Conceptual 2(2):202a12. https://doi.org/10.5826/dpc.0202a12
- Argenziano G, Puig S, Zalaudek I et al (2006) Dermoscopy improves accuracy of primary care physicians to triage lesions suggestive of skin cancer. J Clin Oncol 24(12):1877–1882. https://doi.org/10. 1200/jco.2005.05.0864
- Koelink CJ, Vermeulen KM, Kollen BJ et al (2014) Diagnostic accuracy and cost-effectiveness of dermoscopy in primary care: A cluster randomized clinical trial. J Eur Acad Dermatol Venereol 28(11):1442–1449. https://doi.org/10.1111/jdv.12306
- Westerhoff K, McCarthy WH, Menzies SW (2000) Increase in the sensitivity for melanoma diagnosis by primary care physicians using skin surface microscopy. Br J Dermatol 143(5):1016–1020. https:// doi.org/10.1046/j.1365-2133.2000.03836.x
- 21 Seiverling E, Ahrns H, Stevens K et al (2021) Dermoscopic lotus of learning: implementation and dissemination of a multimodal dermoscopy curriculum for primary care. Short Report. J Med Educ Curric Dev 8:1–4. https://doi.org/10.1177/2382120521989983
- Susong JR, Ahrns HT, Daugherty A, Marghoob AA, Seiverling EV (2020) Evaluation of a virtual basic dermatology curriculum for dermoscopy by using the triage amalgamated dermoscopic algorithm for novice dermoscopists. J Am Acad Dermatol 83(2):590–592. https://doi.org/10.1016/j.jaad.2019.05.097
- Rogers T, Marino ML, Dusza SW et al (2016) A clinical aid for detecting skin cancer: the triage amalgamated dermoscopic algorithm (TADA). J Am Board Fam Med 29(6):694–701. https://doi. org/10.3122/jabfm.2016.06.160079
- Sawyers EA, Wigle DT, Marghoob AA, Blum A (2020) Dermoscopy training effect on diagnostic accuracy of skin lesions in Canadian family medicine physicians using the triage amalgamated dermoscopic algorithm. Dermatol Pract Conceptual 10(2):e2020035. https://doi.org/10.5826/dpc.1002a35
- De Bedout V, Williams NM, Muñoz AM et al (2021) Skin cancer and dermoscopy training for primary care physicians: A pilot study. Dermatol Pract Conceptual 11(1):e2021145. https://doi.org/10.5826/ dpc.1101a145
- Bandic J, Kovacevic S, Karabeg R, Lazarov A, Opric D (2020) Teledermoscopy for skin cancer prevention: a comparative study of clinical and teledermoscopic diagnosis. Acta Informatica Medica 28(1):37–41. https://doi.org/10.5455/aim.2020.28.37-41

- Secker LJ, Buis PA, Bergman W, Kukutsch NA (2017) Effect of a dermoscopy training course on the accuracy of primary care physicians in diagnosing pigmented lesions. Acta Derm Venereol 97(2):263–265. https://doi.org/10.2340/00015555-2526
- 28 Harkemanne E, Goublomme N, Sawadogo K, Tromme I (2020) Early melanoma detection in primary care: clinical recognition of melanoma is not enough, one must also learn the basics [online ahead of print]. J Cancer Educ. https://doi.org/10.1007/ s13187-020-01897-w
- Eide MJ, Asgari MM, Fletcher SW et al (2013) Effects on skills and practice from a web-based skin cancer course for primary care providers. J Am Board Fam Med 26(6):648–657. https://doi.org/10. 3122/jabfm.2013.06.130108
- Orfaly VE, Berry EG, Stoos ER et al (2021) Melanoma toolkit for early detection for primary care providers: a pilot study. Pigment Cell Melanoma Res 34(5):984–986. https://doi.org/10.1111/pcmr. 12968
- Beecher SM, Keogh C, Healy C (2018) Dedicated general practitioner education sessions can improve diagnostic capabilities and may have a positive effect on referral patterns for common skin lesions. Ir J Med Sci 187(4):959–963. https://doi.org/10.1007/ s11845-018-1788-z
- Badertscher N, Braun RP, Held U et al (2013) Diagnostic competence of Swiss general practitioners in skin cancer. Swiss Med Wkly 143:w13834. https://doi.org/10.4414/smw.2013.13834
- Carli P, De Giorgi V, Crocetti E, Caldini L, Ressel C, Giannotti B (2005) Diagnostic and referral accuracy of family doctors in melanoma screening: effect of a short formal training. Eur J Cancer Prev 14(1):51–55. https://doi.org/10.1097/00008469-200502000-00007
- Brochez L, Verhaeghe E, Bleyen L, Naeyaert JM (2001) Diagnostic ability of general practitioners and dermatologists in discriminating pigmented skin lesions. J Am Acad Dermatol 44(6):979–986. https://doi.org/10.1067/mjd.2001.113442
- Bedlow AJ, Cliff S, Melia J, Moss SM, Seyan R, Harland CC (2000) Impact of skin cancer education on general practitioners' diagnostic skills. Clin Exp Dermatol 25(2):115–118. https://doi.org/10.1046/j. 1365-2230.2000.00590.x
- Harris JM, Salasche SJ, Harris RB (2001) Can Internet-based continuing medical education improve physicians' skin cancer knowledge and skills? J Gen Intern Med 16(1):50–56. https://doi.org/10. 1111/j.1525-1497.2001.00615.x
- Oliveria SA, Nehal KS, Christos PJ, Sharma N, Tromberg JS, Halpern AC (2001) Using nurse practitioners for skin cancer screening: a pilot study. Am J Prev Med 21(3):214–217. https://doi.org/10.1016/ s0749-3797(01)00354-3
- Prasad S, Black S, Chen HW et al (2022) Beliefs and barriers to care of primary care practitioners toward melanoma screening and education in rural Texas. J Am Acad Dermatol. https://doi.org/10. 1016/j.jaad.2021.12.059
- Pagnanelli G, Soyer HP, Argenziano G et al (2003) Diagnosis of pigmented skin lesions by dermoscopy: web-based training improves diagnostic performance of non-experts. Br J Dermatol 148(4):698– 702. https://doi.org/10.1046/j.1365-2133.2003.05168.x
- Herschorn A (2012) Dermoscopy for melanoma detection in family practice. Can Fam Physician 58(7):740-5-e372-8
- Argenziano G, Puig S, Zalaudek I et al (2006) Dermoscopy improves accuracy of primary care physicians to triage lesions suggestive of skin cancer. J Clin Oncol 24(12):1877–1882. https://doi.org/10. 1200/jco.2005.05.0864

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