

Mobile Technologies and Cervical Cancer Screening in Low- and Middle-Income Countries: A Systematic Review

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PURPOSE Cervical cancer screening is not well implemented in many low- and middle-income countries (LMICs). Mobile health (mHealth) refers to utilization of mobile technologies in health promotion and disease management. We aimed to qualitatively synthesize published articles reporting the impact of mHealth on cervical cancer screening-related health behaviors.

METHODS Three reviewers independently reviewed articles with the following criteria: the exposure or intervention of interest was mHealth, including messages or educational information sent via mobile telephone or e-mail; the comparison was people not using mHealth technology to receive screening-related information, and studies comparing multiple different mHealth interventional strategies were also eligible; the primary outcome was cervical cancer screening uptake, and secondary outcomes included awareness, intention, and knowledge of screening; appropriate research designs included randomized controlled trials and quasi-experimental or observational research; and the study was conducted in an LMIC.

RESULTS Of the 8 selected studies, 5 treated mobile telephone or message reminders as the exposure or intervention, and 3 compared the effects of different messages on screening uptake. The outcomes were diverse, including screening uptake (n = 4); health beliefs regarding the Papanicolaou (Pap) test (n = 1); knowledge of, attitude toward, and adherence to colposcopic examination (n = 1); interest in receiving messages about Pap test results or appointment (n = 1); and return for Pap test reports (n = 1).

CONCLUSION Overall, our systematic review suggests that mobile technologies, particularly telephone reminders or messages, lead to increased Pap test uptake; additional work is needed to unequivocally verify whether mhealth interventions can improve knowledge regarding cervical cancer. Our study will inform mHealth-based interventions for cervical cancer screening promotion in LMICs.

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INTRODUCTION

Cervical cancer is the fourth most common cancer among women worldwide.¹ Importantly, it remains one of the major gynecologic malignancies threatening quality of life and health status of women in low- and middle-income countries (LMICs).² Yet, cervical cancer is a largely preventable disease.³ Specifically, cervical cancer screening with either the Papanicolaou (Pap) or human papillomavirus (HPV) test can identify the presence of precancerous or cancerous cervical cells or high-risk HPV associated with cervical cancer development.⁴ In LMICs, because of the inadequate health service infrastructure and resources, visual inspection with acetic acid (VIA) or Lugol's iodine (VILI) is widely used to detect early-stage neoplastic lesions.^{5,6} Utilization of these techniques can largely reduce cervical cancer burden. For example, cervical cancer mortality in the United States was significantly

reduced after the introduction of the Pap test in the 1950s.⁷ However, similar practices have been less frequently implemented in LMICs than in high-income countries.⁸ Most high-income countries have systematic guidelines for cervical cancer screening; based on 2016 data, approximately two thirds of adult women in the United States underwent a Pap test within the past 3 years.⁹ This is in contrast to practices in LMICs, where implementation of cervical cancer screening is not as widespread as in high-income countries.¹⁰⁻¹² For example, the 2010 China Chronic Disease and Risk Factor Surveillance System (N = 51,989 women) found that 77% of the sample never underwent the cervical cancer screening.⁸

Mobile health (mHealth) refers to the use of mobile telephones and other wireless technology in health promotion or disease prevention. Overcoming the burden of cervical cancer in LMICs warrants affordable,

ASSOCIATED CONTENT

Data Supplement

Author affiliations and support information (if applicable) appear at the end of this article.

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CONTEXT

Key Objective

Is it possible to use mobile health (mHealth)-based intervention program to increase awareness, knowledge, and uptake of cervical cancer screening in low- and middle-income countries (LMICs)?

Knowledge Generated

Mobile telephone reminders, as compared with other traditional interventions, can increase relevant knowledge and uptake of cervical cancer screening in LMICs. We did not find evidence suggesting differential promoting effects across different types of text messages.

Relevance

The outcomes of our study can inform mHealth-based intervention programs in LMICs that aim to promote cervical cancer screening.

accessible, and effective technology-based solutions. To our knowledge, no studies have synthesized evidence regarding the effects of mHealth on cervical cancer screening in LMICs. LMICs can benefit greatly from the low-cost, high-reach, high-dissemination capabilities offered by mHealth. As the first step toward developing an mHealth intervention, we set out to systematically review and synthesize evidence from studies examining the association between mHealth and cervical cancer screening in LMICs. Our study has been prospectively registered at PROSPERO (identifier: CRD42018110439).

METHODS

Search Strategy

Four electronic databases (PubMed, EMBASE, Web of Science, and Scopus) were used to search for potentially eligible articles in English until October 10, 2019. Controlled vocabularies (PubMed: MeSH; EMBASE: Emtree) and keywords related to “mobile health,” “phone,” and “cervical cancer screening” were used in the search strategy (Data Supplement). We also searched the reference list of previously published research on similar topics to capture more potentially eligible articles.¹³ A publicly available mHealth Web site¹⁴ was used to find any additional articles or gray literature relevant to our study. In addition to English articles, D.Z. searched the China Academic Journals Full-Text Database¹⁵ to obtain potentially eligible articles written in Chinese. D.Z. conducted the search using the keywords “cervical cancer screening,” “mobile phone,” and “mobile”. To obtain potentially eligible articles in Spanish, A.C. searched the Virtual Health Library Regional Portal, a network of Latin American and Caribbean bibliographic databases, for Spanish-language health literature. Spanish-language keywords that were related to “mobile health,” “phone,” and “cervical cancer screening” were used in the search strategy.

Title and Abstract Screening

For title and abstract screening for English articles, 3 reviewers (D.Z., S.A., and A.C.) independently reviewed titles

and abstracts from records identified in electronic databases and decided whether the article should be selected in this process; inconsistent screening decisions regarding inclusion of an article were solved by discussion or by consulting the senior author (D.B.). D.Z. scanned all title and abstract screening records for Chinese articles, whereas A.C. scanned records written in Spanish. Specifically, articles with the following characteristics were included. First, the exposure or intervention of interest was related to mHealth. This included telephone reminder, telephone counseling, text message, smart phone app, e-mail message, and other wireless intervention strategies that conveyed information on cervical cancer prevention, encouraged screening, or provided assistance in screening scheduling. Second, the target population was composed of women eligible for cervical cancer screening, and the comparison group was a nonintervened population or a group of people who did not receive information about cervical cancer screening via mHealth devices or only received such information from traditional media. Traditional media included mailed letters, pamphlets, and newsletters. Studies comparing different types of mHealth approaches were also eligible (eg, telephone call v text message). Third, the primary outcome of interest in this systematic review was the uptake of cervical cancer screening (both index and repeat or follow-up screening), and the secondary outcome of interest included awareness, intention, and knowledge of cervical cancer screening. The screening approach included Pap test, HPV test, VIA, or VILI. Fourth, eligible designs were randomized controlled trials (RCTs), quasi-experimental research, or observational studies; reviews and meta-analyses were excluded. Fifth, studies clearly reporting non-LMICs as geographic locations in their titles and abstracts were excluded; studies meeting the first 4 aforementioned criteria without reporting geographic locations were included during the title and abstract screening and further evaluated during the full-text review. We identified LMICs using information provided by the World Bank.¹⁶ All of the studies meeting the selection criteria in title and abstract screening were included for full-text review.

Full-Text Review

In the full-text review, reviewers (D.Z., S.A., and A.C.) read whole articles selected during the title and abstract screening to judge whether they should be included for further synthesis. Articles meeting the following criteria were chosen: articles confirmed that study locales were LMICs; the study reported effect measures showing associations between mHealth and cervical cancer screening; and the study had a corresponding full-text article in English, Chinese, or Spanish, and protocols were not included. If duplicated study populations were used in > 1 article, the article with the highest quality (eg, robust study procedures and analytic strategies) was chosen. This process was independently completed by 3 reviewers (D.Z., S.A., and A.C.), and inconsistent decisions were solved by discussion or by consulting the senior author (D.B.). D.Z. read full texts written in Chinese, and A.C. read full texts written in Spanish. We present a flowchart showing the number of studies excluded at each step and summarize the whole selection process and exclusion reasons using a Preferred Reporting Items for Systematic Review and Meta-Analysis (PRISMA) flowchart.¹¹

Data Extraction

We extracted the following characteristics from published articles: name of the first author and publication year; year of data collection; study location; definition of exposure or intervention of interest and comparison; measurement or randomization process of exposure or intervention; sample size; definition and measurement of cervical cancer screening-related outcomes, including uptake, awareness, intention, and knowledge of cervical cancer screening; time period of intervention and follow-up; mean age of women; effect measure and 95% CI of mHealth intervention or exposure; and variables adjusted for in the model. Data extraction was conducted by reviewers independently, and discrepancies between reviewers were resolved by discussion or consulting the senior author (D.B.).

Qualitative Synthesis and Quality Assessment

A narrative synthesis was conducted to descriptively summarize the main study characteristics (ie, sample size, study locale, average age of participants, and year conducted), definition and measurement of mHealth and cervical cancer screening, measures of association, and major limitations of each study. Specifically, each study was independently assessed for the methodologic strengths and limitations by reviewers (D.Z., S.A., and A.C.), and discrepancies were resolved by discussion or by consulting the senior author (D.B.). We referred to the *Cochrane Handbook for Systematic Reviews of Interventions*¹⁷ and mainly considered selection bias, measurement error, and analysis strategy when assessing study quality.⁷ For interventional studies (RCTs and quasi-experimental studies), we further evaluated the rationality of randomization and blinding. RevMan 5.3 (Cochrane Collaboration,

London, United Kingdom) was used to summarize the risk of bias in intervention studies. We used the modified Newcastle-Ottawa Scale (NOS) to assess the quality of cross-sectional studies.¹⁸

RESULTS

Study Identification and Selection

Overall, we identified 3,127 records from the electronic databases (PubMed, n = 592; EMBASE, n = 923; Web of Science, n = 557; and Scopus, n = 1,055) and 1 article from other sources. After deduplication, we kept 1,768 articles for title and abstract screening. Of the 17 articles selected from title and abstract screening, 6 were excluded because of unmatched exposure or intervention of interest, 1 was excluded because of duplicate sample use, and 2 were excluded for unmatched study design. This yielded a total of 8 studies¹⁹⁻²⁶ included in the systematic review. Because of the large heterogeneity in the definition of mHealth, outcomes of interest, and study design, we did not perform the quantitative synthesis (Fig 1). The PRISMA checklist is provided in the Data Supplement.

Study Characteristics and Quality

Table 1 lists the study characteristics. The included studies were conducted in different geographic locations and times. One study was conducted in Iran,²¹ 1 was in Tanzania,²⁶ 2 were in Malaysia,^{19,25} 2 were in Brazil,^{22,24} and 2 were in South Africa.^{20,23} These studies were all conducted within the past decade, and the time span ranged from 2010 to 2016. Seven of the studies were interventional research studies (2 quasi-experimental studies and 5 RCTs),^{19-22,24-26} and one was a cross-sectional study.²³ The study samples ranged from 106 to 1,000 participants. Seven studies reported the average age of participants, which ranged from 29 to 42 years.¹⁹⁻²⁴ One study²⁵ only reported the categorical age distribution, and approximately 40% of people were younger than age 35 years. The exposures and interventions of interest were diverse in these studies. Briefly, 5 studies treated utilization of telephone or message reminder as the exposure or intervention,^{19,21,23,25,26} and 3 studies^{20,22,24} compared effects of different types of text messages on screening. Among the interventional studies,^{19-22,24-26} the length of follow-up ranged from 1 week to 6 months. Overall, there were 5 types of outcomes reported by these studies, and they were as follows: screening uptake (n = 3)^{19,20,25,26}; health beliefs about Pap test (n = 1)²¹; knowledge of, attitude toward, and adherence to screening (n = 1)²²; interest in receiving messages about Pap test report or appointment (n = 1)²³; and return for Pap test reports (n = 1).²⁴

Table 2 lists the quantitative outcomes and major limitations in each study. Because of heterogeneous design, study population, intervention or exposure, and definition of outcome, patterns of associations between mHealth and cervical cancer screening seemed to be complex. After

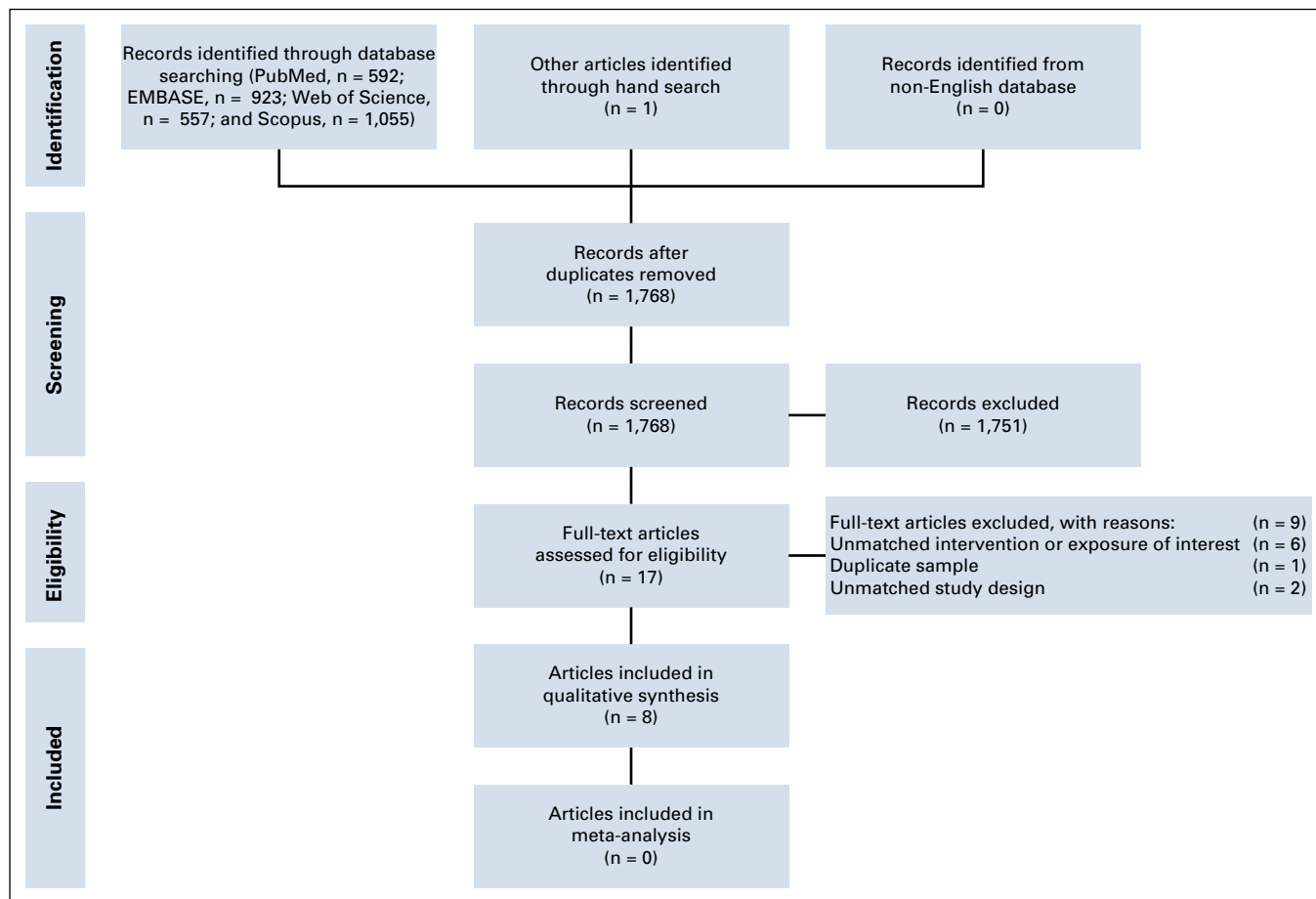


FIG 1. Flowchart of study identification and selection.

a 24-week observation, Abdullah and Su¹⁹ reported that women receiving invitation letter and telephone reminder were more likely (odds ratio [OR], 2.44; 95% CI, 1.29 to 4.62) to undergo a Pap test compared with women only receiving an invitation letter. Adonis et al²⁰ found there was no difference in likelihood of Pap test utilization across people receiving different types of text messages. In particular, as compared with a text message focusing on screening recommendation, messages focusing on the importance of regular screening (OR, 0.57; 95% CI, 0.26 to 1.19) and messages focusing on the risk of cervical cancer (OR, 0.87; 95% CI, 0.47 to 1.66) did not seem to affect Pap test uptake. In a 60-day RCT, Erwin et al²⁶ found that sending multiple short message service (SMS) messages with eVoucher (v single SMS) significantly increased screening attendance (OR, 4.7; 95% CI, 2.9 to 7.4); the results also showed that participants receiving 15 SMS messages were more likely to attend screening (OR, 3.0; 95% CI, 1.5 to 6.2). A study conducted in Iran applied a comprehensive set of intervention that incorporated text messages, electronic posters, infographics, podcasts, and video tutorials in a quasi-experimental study for 4 weeks.²¹ Consequently, such interventions have been found to positively affect health beliefs and cervical cancer screening

knowledge among married women. The intervention versus control group score difference in knowledge was 8.18 points, whereas the corresponding score difference in perceived susceptibility was 4.07. The score differences were 7.78, 2.99, -19.17, and 3.18 for perceived severity, perceived benefits, perceived barriers, and health motivation, respectively. Lima et al²² found there was no difference ($P > .05$) in the rates of knowledge of (-3.7%) and attitude toward (-2.0%) colposcopic examination between women receiving educational versus reminder messages; in addition, although the study found there was an increase in uptake of colposcopic examination in both groups, the increase was more substantial among women receiving reminder messages (reminder text message, 66.8%; educational text message, 57.5%; rate difference, 9.3%; $P = .03$). A cross-sectional survey reported that women using mobile telephone text messages had a higher interest in receiving appointment reminders via SMS (OR, 14.19; 95% CI, 1.72 to 117.13).²³ Nicolau et al²⁴ reported that an educational message introducing knowledge about cervical cancer (OR, 1.37; 95% CI, 1.22 to 1.54) and a reminder message (OR, 1.40; 95% CI, 1.25 to 1.57) could increase the likelihood of returning to the clinic to receive Pap test results; because their effect measures

TABLE 1. Study Characteristics

Article and Year of Publication	Study Location	Year(s) of Data Collection	Intervention or Exposure	Length of Intervention	Sample Size	Age Distribution	Study Design	Main Outcome
Abdullah and Su ¹⁹ (2013)	Malaysia	January–November 2010	Intervention: invitation letter with telephone reminder Control: invitation letter without telephone reminder	24 weeks	Intervention, n = 201 Control, n = 202	Mean age: 36 years Range was not reported	RCT	Uptake of Pap test
Adonis et al ²⁰ (2017)	South Africa	August 2013–May 2014	Intervention 1: messages focusing on risk of cancer Intervention 2: messages focusing on importance of screening Control: message only stating recommendation for screening	6 months	Intervention 1, n = 176 Intervention 2, n = 175 Control, n = 355	Mean age: 42 years Range: 21–65 years	RCT	Uptake of Pap test
Erwin et al ²⁶ (2019)	Tanzania	February–May 2016	Intervention 1: 15 unique SMS messages aiming to change screening behaviors Intervention 2: 15 unique SMS messages and eVoucher valid for 2 months Control: 1 SMS message showing the location and hours of the closest screening clinic	60 days	Intervention 1, n = 272 Intervention 2, n = 298 Control, n = 281	Mean age, 34 years Range, 25–49 years	RCT	Attendance at cervical cancer screening
Khademolhosseini et al ²¹ (2017)	Iran	August 2015–March 2016	Intervention: text message, electronic posters, infographics, podcasts, and video tutorial Control: received no intervention	4 weeks	Intervention, n = 53 Control, n = 53	Mean age: 32 years Range was not reported	Quasi-experimental study	Health belief about Pap test (knowledge, perceived susceptibility, severity, benefit and harms, and health motivation about cervical cancer screening)
Lima et al ²² (2017)	Brazil	June–December 2014	Intervention: educational message introducing knowledge about cervical cancer prevention Control: reminder message of screening	6 months	Intervention, n = 262 Control, n = 262	Mean age: 38 years Range: 25–64 years	Quasi-experimental study	Knowledge of, attitude toward, and adherence to screening
Moodley et al ²³ (2019)	South Africa	February–June 2014	Exposure: Use of text message	This was a cross-sectional study without follow-up	412 (55% were HIV positive)	Median age: 29 years IQR: 25–36 years	Cross-sectional survey	Interest in receiving message about Pap test report or Pap test appointment
Nicolau et al ²⁴ (2017)	Brazil	June–December 2014	Intervention 1: educational message introducing cervical cancer prevention Intervention 2: reminder message of screening Control: received no intervention	NA	Telephone education, n = 179 Telephone reminder, n = 182 Control, n = 181	Mean age: 37 years Range was not reported	RCT	Return for Pap test report

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TABLE 1. Study Characteristics (Continued)

Article and Year of Publication	Study Location	Year(s) of Data Collection	Intervention or Exposure	Length of Intervention	Sample Size	Age Distribution	Study Design	Main Outcome
Rashid et al ²⁵ (2013)	Malaysia	July-September 2011	Intervention: reminders sent via letter, registered letter, SMS, or telephone call Control: no intervention	8 weeks	Letter, n = 250 Registered letter, n = 250 SMS, n = 250 Telephone call, n = 250	Mean age was not reported Range: 20-65 years	RCT	Undergo repeated Pap test within 1 year

Abbreviations: IQR, interquartile range; NA, not available; Pap test: Papanicolaou test; RCT, randomized controlled trial; SMS, short message service.

TABLE 2. Summary of Outcomes and Limitations

Article and Year of Publication	Distribution of Outcomes	Effect Measure or Other Quantitative Test Results (Intervention v control group)	Adjusted Variables	Major Limitations
Abdullah and Su ¹⁹ (2013)	Intervention group: 18.1% had Pap test Control group: 10.1% had Pap test	The intervention group had a higher likelihood of Pap test uptake within the 24 weeks during follow-up: adjusted OR, 2.44; 95% CI, 1.29 to 4.62	History of Pap test and mammogram, age of first sexual intercourse, age of first marriage, and age of first pregnancy	1. Selection bias: The study population consisted of teachers, which could compromise generalizability. 2. Blinding issues: This study is unblinded, which compromised the validity of this study. 3. Statistical issues: There was no need to make additional adjustment because the distribution of sociodemographic variables was similar between the intervention and control arms.
Adonis et al ²⁰ (2017)	9.6% of women receiving message focusing on recommendation of screening had Pap test 5.7% of women receiving message focusing on importance of regular screening had Pap test 8.5% of women receiving message focusing on risk of cervical cancer had Pap test	Message focusing on importance of regular screening v message focusing on recommendation of screening: OR, 0.57; 95% CI, 0.26 to 1.19 Message focusing on risk of cervical cancer v message focusing on recommendation of screening: OR, 0.87; 95% CI, 0.47 to 1.66	NA	1. Selection bias: Study population comprised women who did not have Pap test in past 3 years, which could have selectively enrolled women with lower screening intention. 2. Blinding issues: There was no evidence suggesting participants and research staff were blinded. 3. Selective reporting: The article did not show tables of baseline characteristics. If randomization failed, statistical adjustment was needed. 4. Randomization methods were not reported.
Erwin et al ²⁶ (2019)	12.9% of women receiving 15 SMS messages attended CCS 18.0% of women receiving SMS and eVoucher attended CCS 4.3% of women in control group attended CCS	As compared with control group, participants receiving SMS and eVoucher were more likely to attend CCS: OR, 4.7; 95% CI, 2.9 to 7.4 As compared with control group, participants receiving 15 SMS messages were more likely to attend CCS: OR, 3.0; 95% CI, 1.5 to 6.2	Age, stratification, and clustering.	1. Selection bias: Study population only included women age 25-49 years who had access to mobiles telephones. 2. Vague definition of cervical cancer screening.
Khademolhosseini et al ²¹ (2017)	On average, the scores of knowledge, perceived susceptibility, severity, benefit, and harms, and health motivation were higher among intervention group	t-tests suggested knowledge, perceived susceptibility, and perceived severity of cervical cancer were higher in intervention group ($P < .05$); perceived benefit and health motivation were stronger in intervention group ($P < .05$), and perceived barriers was stronger in control group ($P < .05$) Detailed mean differences between intervention and control arms are as follows: Knowledge: 8.18 Perceived susceptibility: 4.07 Perceived severity: 7.78 Perceived benefits: 2.99 Perceived barriers: -19.17 Health motivation: 3.18	NA	1. Selection bias: Only married women were included. 2. Blinding issues: There was no evidence suggesting participants and research staff were blinded. 3. Statistical issues: The sample size was small, which could introduce random error and reduce power. 4. Randomization methods were not reported.

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TABLE 2. Summary of Outcomes and Limitations (Continued)

Article and Year of Publication	Distribution of Outcomes	Effect Measure or Other Quantitative Test Results (intervention v control group)	Adjusted Variables	Major Limitations
Lima et al ²³ (2017)	<p>Intervention group: 39.1% had appropriate knowledge of cervical cancer screening; 39.1% had appropriate attitude toward cervical cancer screening; 57.5% had appropriate adherence to cervical cancer screening</p> <p>Control group: 35.4% had appropriate knowledge of cervical cancer screening; 37.1% had appropriate attitudes toward cervical cancer screening; 66.8% had appropriate adherence to cervical cancer screening</p>	<p>No significant changes were obtained for knowledge of and attitude toward screening ($P > .05$). However, the adherence to screening increased more substantially in the control group ($P = .03$).</p> <p>Detailed rate differences between the intervention and control arms are as follows: Rate of appropriate knowledge: -3.7% Rate of appropriate attitude: -2.0% Rate of adherence: 9.3%</p>	NA	<ol style="list-style-type: none"> 1. Statistical issues: Missing data for knowledge and attitude were substantial. 2. Selective reporting: The article did not show tables of baseline characteristics. If randomization failed, statistical adjustment would be needed. 3. Other limitations: Colposcopic examination was an ambiguous term.
Moodley et al ²³ (2019)	<p>72% of respondents were interested in receiving Pap test results by SMS</p> <p>77% of respondents were interested in receiving clinic or hospital appointment reminders</p>	<p>Using mobile text messaging was not associated with interest in receiving Pap test results via SMS (adjusted OR, 4.34; 95% CI, 0.54 to 34.94). However, it was associated with a higher interest in receiving appointment reminders via SMS (adjusted OR, 14.19; 95% CI, 1.72 to 117.13).</p> <p>Using a nonprivate telephone was inversely associated with interest in receiving Pap test results via SMS (adjusted OR, 0.31; 95% CI, 0.18 to 0.51). However, it was not associated with interest in receiving appointment reminders via SMS (adjusted OR, 1.26; 95% CI, 0.70 to 2.27).</p>	<p>Mobile text message use, HIV status, previous Pap test, education, and having private telephone; for appointment reminder, the model additionally adjusted for language</p>	<ol style="list-style-type: none"> 1. Selection bias: More than half the study population was HIV positive. 2. Statistical issues: The wide CI of the effect measure of text usage suggested statistical imprecision.
Nicolau et al ²⁴ (2017)	<p>93.5% of women receiving reminder returned for Pap test results</p> <p>91.8% of women receiving educational telephone call returned for Pap test results</p> <p>66.9% of women in control group returned for Pap test results</p>	<p>As compared with normal attendance, telephone reminders (OR, 1.40; 95% CI, 1.25 to 1.57) or telephone calls (OR, 1.37; 95% CI, 1.22 to 1.54) were associated with a higher likelihood of returning for Pap test report</p>	NA	<p>There were no substantial methodologic limitations.</p>
Rashid et al ²⁵ (2013)	<p>23.9% of women who received a letter underwent a Pap test</p> <p>23.0% of women who received a registered letter underwent a Pap test</p> <p>32.9% of women who received telephone messages underwent a Pap test</p> <p>50.9% of women who received a telephone call underwent a Pap test</p>	<p>Women receiving a telephone call, compared with those receiving letters, were more likely to undergo a repeated Pap test after receiving the message (OR, 2.38; 95% CI, 1.56 to 3.63)</p>	NA	<ol style="list-style-type: none"> 1. Selection bias: The study only included women undergoing Pap test in the past year who should have a higher intention of screening. 2. Selective reporting: The article did not provide tables of baseline characteristics. If randomization failed, statistical adjustments were needed.

Abbreviations: CCS, cervical cancer screening; NA, not applicable; OR, odds ratio; Pap test: Papanicolaou test; SMS, short message service.

were quite similar, it suggested there was no difference between their effects on screening-related behaviors. Rashid et al²⁵ found that telephone call reminders, as compared with letters, increased the utilization of Pap tests among women who had a history of Pap test in the past year (OR, 2.38; 95% CI, 1.56 to 3.63).

The major limitations of included studies involved selection bias, failure to blind, and selective reporting (Table 2). Six of the included studies had selection bias, which compromised the generalizability.^{19-21,23,25,26} Specifically, 1 study only enrolled female teachers,¹⁹ 1 included women without a Pap test history in the past 3 years,²⁰ 1 only enrolled women age 25-49 years with access to mobile telephones,²⁶ 1 only included married women,²¹ 1 enrolled a large proportion of HIV-infected women,²³ and 1 only enrolled women undergoing Pap test with the past year.²⁵ These women could be sociodemographically different from the general population; thus, the derived conclusion might be difficult to apply to women at an appropriate age for cervical cancer screening. Three of the 7 interventional studies did not blind their participants or research staff.¹⁹⁻²¹ Although the nature of intervention scenarios made masking difficult, the lack of blinding could still introduce bias in effect measures. For example, women who only received reminder messages might have sought relevant knowledge about cervical cancer if they were aware that their counterparts were receiving professional information about cervical cancer and screening, which might change their screening behaviors. Three interventional studies^{20,22,25} did

not report distributions of important sociodemographic variables at baseline, which made it hard for us to judge whether the randomization was successful. In particular, the researchers should have adjusted for other factors if the randomization failed; otherwise, the effect measures could be biased. The risk of bias assessment for interventional studies is presented at Figures 2A and 2B. NOS assessment and potential sources of other bias in interventional studies are provided in the Data Supplement.

DISCUSSION

To our knowledge, this is the first systematic review that investigates how mHealth can affect cervical cancer screening in LMICs from multiple aspects, including utilization, health beliefs, and interest in receiving relevant appointment information and returning for screening results. Overall, our systematic review suggests that, as compared with traditional communication methods (eg, postal mail), utilization of a telephone reminder or a mobile text message in LMICs can increase uptake of cervical cancer screening. Evidence regarding the effects of mHealth tools on the awareness and perceived threat of cervical cancer is inconclusive and warrants further investigation to delineate optimal implementation strategies in this setting. We did not find robust evidence suggesting that a certain type of text message can have a stronger effect on screening behaviors as compared with others.

Findings from a recently published systematic review that investigated the effects of text messages on cancer

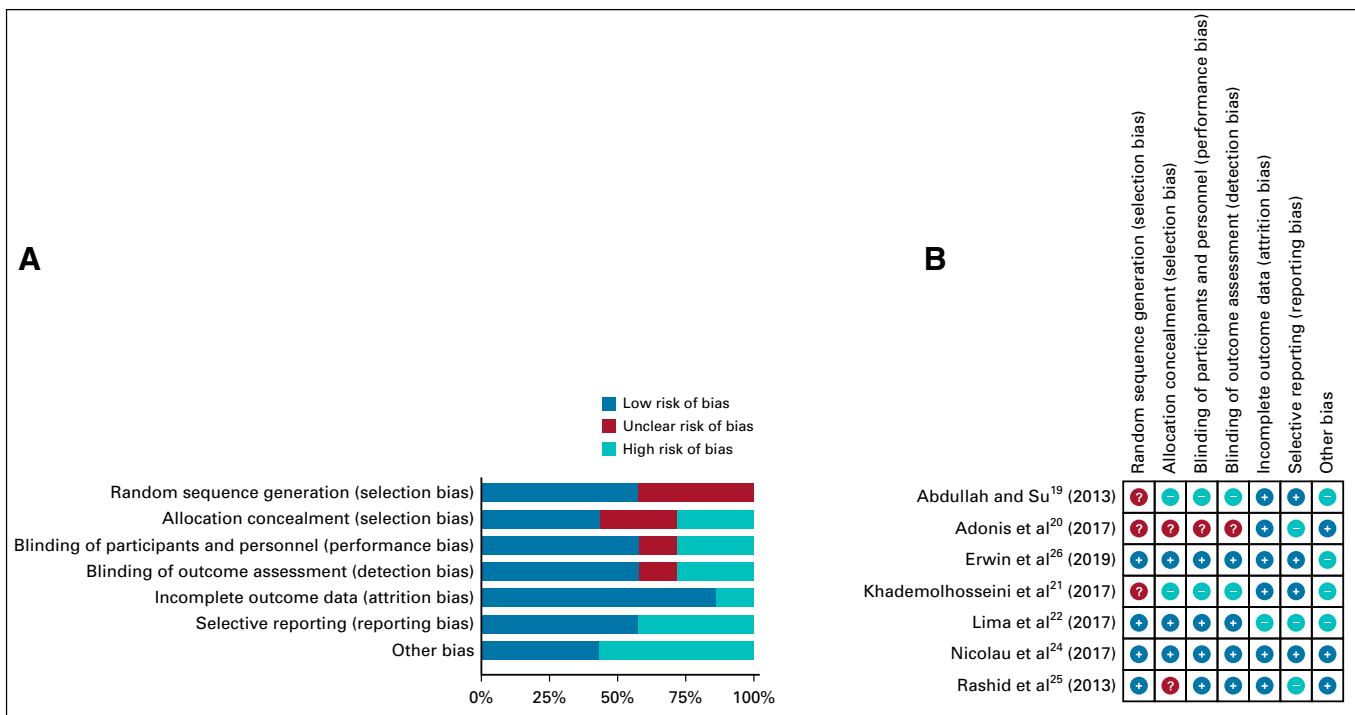


FIG 2. (A) Overall summary of bias of interventional studies. (B) Summary of bias in each interventional study. ?, unclear risk of bias; +, low risk of bias; -, high risk of bias.

screening rates are consistent with our results. Uy et al²⁷ synthesized 9 published articles (8 in developed countries and 1 in an LMIC) and concluded that text messaging interventions could increase screening rate for breast (5 studies), cervical (1 study), and colorectal (3 studies) cancer.

This systematic review has some limitations. First, because of the heterogeneity across primary studies and different types of mHealth interventions, we cannot quantitatively synthesize the effect measures, making the intervention effectiveness ambiguous. Second, 3 of the included studies^{21,23,24} did not measure uptake of cervical cancer screening directly, but treated health beliefs about the Pap test, interest in receiving screening appointment, and return for Pap test results as the outcomes of interest. These outcomes can only reflect the awareness of cervical cancer and potential intention of screening and may not guarantee the screening utilization. Third, LMICs consist of countries with differential economic and developmental status. According to the classification of World Bank,¹⁶ the eight included studies were all conducted in LMICs with better economic situations, which makes our evidence less generalizable to other LMICs with lower economic situations. Fourth, some LMICs may have an organized screening program; however, studies included in this systematic review have insufficient information on how the presence of organized screening programs affected the effectiveness of mHealth interventions targeting cervical cancer screening. It will be important to explore this issue in future research. Furthermore, five of the interventional studies^{19-21,24,25} treated Pap test-related behaviors as the

outcomes of interest, and 1 study²² used colposcopic examination as the outcome of interest. However, the cytologic test requires advanced medical equipment and laboratory training, and some low-resource areas do not have such infrastructure, which can reduce accuracy of the cytologic test.¹² This suggests that future research should examine whether mHealth technology or other wireless devices can affect utilization of HPV testing, which has better screening accuracy in LMICs.²⁸

The burden of cervical cancer is higher in LMICs and the corresponding screening rate is lower compared with developed countries,^{8,9} suggesting that an effective and convenient intervention approach is needed in these areas to promote cervical cancer screening. Mobile telephones are much cheaper and portable when compared with laptops, making them easier to use during daily communication. These characteristics demonstrate the potential of such devices to spread knowledge of cervical cancer prevention and promote screening utilization in LMICs. Our results can be informative by providing health practitioners in LMICs with the evidence necessary to establish cost-effective cervical cancer screening promotion programs using mHealth technology. Future studies should explore how mHealth can modify women's screening behaviors in LMICs with worse economic situations and examine the effectiveness of other mobile devices or technologies, such as telephone apps. Moreover, because numerous cultural and spiritual factors across LMICs influence the uptake of mHealth interventions related to cervical cancer screening, further research is paramount to evaluate their roles.

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