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# Leveraging mHealth for the Treatment and Management of PLHIV

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**Objective:** The objective of this systematic review was to analyze published literature from the last five years to assess facilitators and barriers to the adoption of mHealth as interventions to treat and manage HIV for PLHIV (people living with HIV). The primary outcomes were physical and mental conditions. The secondary outcomes were behavior based (substance use, care engagement, and healthy habits).

**Methods:** Four databases (PubMed, CINAHL, Web of Science, and ScienceDirect) were queried on 9/2/2022 for peer-reviewed studies on the treatment and management of PLHIV with mHealth as the intervention. The review was conducted in accordance with the Kruse Protocol and reported in accordance with PRISMA 2020.

**Results:** Five mHealth interventions were identified across 32 studies that resulted in improvements in physical health, mental health, care engagement, and behavior change. mHealth interventions offer both convenience and privacy, meet a digital preference, increase health knowledge, decrease healthcare utilization, and increase quality of life. Barriers are cost of technology and incentives, training of staff, security concerns, digital literacy gap, distribution of technology, technical issues, usability, and visual cues are not available over the phone.

**Conclusion:** mHealth offers interventions to improve physical health, mental health, care engagement, and behavior for PLHIV. There are many advantages to this intervention and very few barriers to its adoption. The barriers are strong, however, and should be addressed through policy. Further research should focus on specific apps for younger versus older PLHIV, based on preferences and the digital literacy gap.

Keywords: mHealth, eHealth, telehealth, telemedicine, human immunodeficiency virus, HIV

# Plain Language Summary

mHealth is being used to help manage the symptoms and spread of HIV. This modality is being used in at least seven countries. Thirtytwo studies from seven countries were analyzed for the effectiveness of leveraging mHealth. Five interventions were identified in the literature. These resulted in improvements in physical and mental health, care management, and behavior change. This intervention offers both convenience and privacy. It meets a digital preference of many users, increases the health knowledge of users, decreases healthcare utilization, and increases quality of life. While several barriers to adoption were noted in the literature, the facilitators to adoption outweighed the barriers. mHealth should be seen as a viable option for the management of symptoms and behavior change to improve care and reduce the spread of HIV.

#### Introduction

#### Rationale

Human immunodeficiency virus (HIV) affects the white-blood cells, targeting the immune system, which in turn, weakens the body's defense against sickness, infection, and some cancers.<sup>1</sup> The prevalence of PLHIV in the world was 38.4 million, at the end of 2021 (486 per 100,000), two-thirds of whom (25.6 million) reside in the African region. Also, in 2021, 650,000 died of HIV and approximately 1.5 million more acquired the condition (incidence is difficult to

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identify because testing is neither ubiquitous nor universal). There is no cure for HIV, but it is no longer a death sentence. The statistics show that transmission of the disease continues to outpace deaths, so the prevalence of people living with HIV (PLHIV) continues to climb.

HIV is transmitted through bodily fluids such as blood, breast milk, semen, and vaginal secretions. PLHIV taking antiretroviral therapy (ART) are virally suppressed and do not transmit HIV to their sexual partners.<sup>1</sup> Acquired immunode-ficiency syndrome (AIDS) is an advanced stage of HIV, which is defined by the development of cancers, infections, and other long-term conditions. To limit transmission of HIV, it is vital to educate the population about healthy sexual habits, getting tested, and if positive, to develop new healthy habits for both sexual activity, treatment and management of HIV symptoms. mHealth can help in this regard. HIV is disproportionately found in low-to-middle-income countries due to funding for and access to prevention and treatment services.<sup>2</sup> Additionally, sexualized drug use, or *chemsex*, is a behavioral factor common in men who have sex with men (MSM) community, which increases the risk of HIV transmission.<sup>3</sup>

mHealth is a subset of telemedicine. Telemedicine and telehealth are defined by the World Health Organization (WHO) as healing at a distance through the use of information communication technologies to improve health outcomes.<sup>4</sup> The WHO does not distinguish between telemedicine and telehealth, so these terms may be used interchangeably in this study. mHealth, specifically, is a component of eHealth that enables the practice of medicine and public or population health through mobile devices, such as phones, tablets, or patient monitoring devices.<sup>5</sup> Mobile devices have blurred the lines between computers and tablets because the processing power of the two have become similar. Many applications work the same on these two modalities.

mHealth has been used for the management of many conditions such as HIV, and it is associated with high satisfaction.<sup>6–8</sup> It is a convenient modality of care and education due to the prevalence of smartphones across the spectrum of country wealth.<sup>7</sup> It can deliver education through text messages, or simple message system (SMS), track risky behaviors and drug cravings, and remind patients to take medication.<sup>9</sup>

A systematic literature review was published in 2022 that analyzed six studies from three databases over five years to assess the tele-education capabilities of mHealth.<sup>10</sup> It focused on patients with HIV/AIDS and their families during treatment. It is concluded that the development of mHealth applications for the treatment and management of HIV can provide rigorous monitoring, research, and evaluation. The group of articles for analysis was very small, so its external validity is questionable.

A systematic literature review was published in 2021 that analyzed 20 studies from five databases over five years to assess the ability of mHealth for HIV prevention. It focused on emerging adults in sub-Saharan Africa.<sup>11</sup> It identified themes of social and structural drivers for transmission and current gaps in understanding HIV prevention. It did not provide an assessment of effectiveness.

#### Objectives

The purpose of this review is to analyze the facilitators and barriers to the adoption of mHealth interventions for the treatment and management of PLHIV through examination of published, peer-reviewed literature over the last five years. The primary outcome is treatment (reminders for ART, etc.) and management (primary and secondary symptoms of physical health and mental health) of PLHIV. The secondary outcome is the management of personal behavior to develop healthy habits for both the PLHIV and their partner (condom use, sexual agreements, substance use, etc.).

# **Materials and Methods**

#### Eligibility Criteria

To be eligible for this review, studies must have been published in peer-reviewed academic journals over the last five years, used adult PLHIV for participants, and used mHealth in either a treatment or management role. Five years was chosen due to the rapid growth of mHealth technology. Articles were eliminated if they did not address these elements of the study objective. As established by published precedent, other reviews were eliminated from the group for analysis to prevent confounding the results.<sup>12,13</sup>

# Information Sources

We queried four research databases: PubMed (MEDLINE), the Cumulative Index to Nursing and Allied Health Literature (CINAHL), Web of Science, and Science Direct on 9/2/2022. MEDLINE was excluded from all databases except PubMed to avoid duplicates. These databases were chosen due to their common availability, exhaustive ability to query existing literature, and the ability for other scientists to duplicate this work. We chose only published literature to ensure the work was peer-reviewed.

#### Search Strategy

We created a Boolean search string to combine key terms listed in the Medical Subject Headings (MeSH) of the US Library of Medicine. We used the same search strategy in all databases: (mHealth OR telemedicine) AND (hiv OR aids) AND (prevention OR management). We used similar filter strategies because not all databases have the same filter tools.

### **Selection Process**

In accordance with the Kruse Protocol, we searched using key terms in all databases, filtered the results, and screened the abstracts for applicability.<sup>14</sup> The Kruse Protocol was chosen because it is a published protocol from which 47 other systematic literature reviews were published. At least two, but no more than three reviewers screened all abstracts. Studies that did not address the research objective were omitted.

# Data Collection Process

We used a standardized Excel spreadsheet as a data extraction tool collecting additional data at each step of the process. This spreadsheet was standardized in the Kruse Protocol and provides fields that are valuable to both clinicians and administrators. Three consensus meetings were held to identify articles for full analysis, data extraction, and theme identification.

#### Data Items

In accordance with the Kruse Protocol, we collected the following fields at each process step: Google Scholar search (date of publication, authors, study title, journal, impact factor from Journal Citations Reports, study design, key terms, experimental intervention, results, and comments from each reviewer); filter article step (the number of results before and after each filter applied in all four databases); abstract screening step (database source, date of publication, authors, study title, journal, screening decision for each reviewer, notes about rejections, consensus meeting one, determination of screening decision, and a set of rejection criteria); analysis step (database source, date of publication, authors, study title, participants, experimental intervention, results compared with a control group, medical outcomes, study design, sample size, bias effect size, country of origin, statistics used, patient satisfaction, facilitators to adoption, barriers to adoption, and the strength and quality of evidence).

# Study Risk and Reporting of Bias Assessment

We observed individual cases of bias and combined these observations with the quality assessment of each study using the Johns Hopkins Nursing Evidence-Based Practice tool (JHNEBP).<sup>15</sup> Strength of Evidence is defined by the JHNEBP as follows: Level I studies are RCTs or experiments with controls and randomization, Level II studies are quasi-experimental (control group but no randomization), Level III studies are observational, qualitative, or other non-experimental methods, and Levels IV and V are opinions. Levels IV and V were not accepted for this study. We considered the instances of bias in how to interpret the results because bias can limit external validity.<sup>16</sup>

#### Effect Measures

Because we accepted three qualitative, two non-experimental, and one observational studies, we were unable to standardize summary measures, as would be performed in a meta-analysis. Measures of effect are summarized in tables for those studies in which it was reported. Measures of effect were reported as Cohen's *d*, Odds Ratios, *beta*, and Wald's *w*. For those studies that reported an effect size, a weighted average effect size was calculated.

# Synthesis Methods

Reviewers also performed a thematic analysis to make sense of the data extracted.<sup>17</sup> The same or similar observations were consolidated into themes. These themes, and the individual observations that did not fit into themes, were tabulated into affinity matrices for further analysis.

# Additional Analyses and Certainty Assessment

Effect sizes were tabulated and included in the data extraction step. Certainty assessment was performed by combining the narrative analysis with the effect sizes. The frequency of observations was tabulated. Frequency of themes is not intended to imply importance: It only provides the probability of encountering the theme in the group of articles for analysis.

# Results

# Study Selection

Figure 1 illustrates the study selection process. The query of four databases resulted in 6151 results; however, 5715 of these results were duplicates. After filtering and screening, reviewers were left with 32 articles eligible for review. Many



Figure I Study selection process.

records were eliminated as "not germane" because they did not address the objective statement. The kappa statistic was calculated to report agreement among reviewers (k=0.88, strong agreement).<sup>18,19</sup>

#### **Study Characteristics**

Following the PRISMA checklist and the Kruse Protocol, our group extracted data and created tables to summarize findings. As established in the literature, a summary table is provided in Table 1: PICOS (participants, intervention, comparison (to control or other group), observation, study design). Of the 32 studies analyzed over the 5-year period, two were from 2017,<sup>20,21</sup> five were from 2018,<sup>22–26</sup> ten were from 2019,<sup>27–36</sup> seven were from 2020,<sup>37–43</sup> five were from 2021,<sup>44–48</sup> and three were from 2022.<sup>49–51</sup> All the studies used adults as participants (>18). About 59% (19/32) used an mHealth app, 16% (5/32) used eHealth over any platform, 13% (4/32) used telephone (counseling, educating, psychotherapy, video), 6% (2/32) used mHealth SMS and the same number used telemedicine over mobile platform. Of the group for analysis, 26 of the studies used strong methodologies such as RCT or robust experiments. No quasi-experimental studies were analyzed, but 6 of the studies were either qualitative, non-experimental, or observational.

### Risk of Bias in and Across Studies

The JHNEBP quality assessment tool identified the following both strength and quality of evidence. This tool qualifies strength through methodologies. Our group for the analysis consisted of 81% (26/32) Level I (RCTs and other robust experiments) and 18% (6/32) Level III (qualitative, non-experimental, and observational). The JHNEBP tool assesses the quality of evidence by sample size, consistency of results, control groups, consistency of conclusions, and adequate literature reviews. Our group of articles for analysis consisted of 88% (28/32) Level A and 13% (4/32) Level B. There were no Level C studies in the group for analysis.

Our group of reviewers also noted instances of study bias. There were 31 cases of selection bias and 29 cases of sample bias, which affect the internal and external validity, respectively. Selection bias occurred when participants were collected from one location in one country (convenience sample), and sample bias occurred when participants consisted of a high percentage of one gender or race.

#### **Results of Individual Studies**

Table 2 summarizes the results of individual studies through the themes identified in the thematic analysis. An observation-to-theme match is provided in <u>Appendices A</u> and <u>B</u>. Additional observations and data collected (sample size, bias, effect size, country of origin, statistics used, and JHNEBP strength and quality of evidence) are provided in <u>Appendix C</u>. The average sample size for all studies was 270, and for Level I studies it was 280. The weighted average effect size was 1,84 (large). Studies originated in seven countries.

# Results of Syntheses, Additional Analysis, and Certainty of Evidence

Our team conducted a thematic analysis to make sense of the data extracted. Although thematic analysis is often associated with qualitative research, other systematic reviews in the literature used this technique to make sense of the observations collected, regardless of the methodology used in the studies analyzed.<sup>52,53</sup> Themes and observations are tabulated in affinity matrices for interpretation.

# Patient Satisfaction

Although participant satisfaction was not always reported, there were zero reports of dissatisfaction with the technologyrelated interventions. In our group for analysis, 30/32 (94%) reported some level of satisfaction, while only two studies did not report user satisfaction. Users were not bothered by SMS messages or reminders. They worked well with the technology. The only report that was not entirely positive was that apps need to be updated to work better with a younger population.

Table I PICOS (Participants, Intervention, Comparison t	o the Control, Medical Outcome, Study Design), Sorted Chronologically by Author
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Authors	Participants	Experimental Intervention	Results (Compared to Control Group)	Medical Outcomes Reported	Study Design
Heckman et al <sup>20</sup>	eckman et al <sup>20</sup> Older adults average age 51.9, 75% Caucasian, all with HIV and 81% with depressive disorder		Intervention reported significantly lower depressive symptoms (BDI, $p = 0.012$ ) and fewer interpersonal problems (IIP, $p = 0.002$ ) than the control	Decreased depression and interpersonal problems	RCT
Himelhoch, et al <sup>21</sup>	Older adults average age 47.3 with HIV + drug use, 66% female (as assigned at birth)	mHealth app (Heart2HAART)	No statistical difference in medicine adherence	Decreased drug cravings, equally as effective with medication adherence	Robust experiment
Kalichman et al <sup>22</sup>	Adults with HIV, 76% male (as assigned at birth)	mHealth app (B-TasP)	Lower HIV RNA ( $p = 0.01$ ), greater cART adherence ( $p = 0.01$ ), fewer indicators of GTI ( $p = 0.05$ ), decreased substance abuse than control	Reduced HIV RNA, increase in medication adherence, decreased genital tract inflammation, and decreased substance abuse	RCT
Sayegh et al <sup>23</sup>	Young adults average age 20.4, 62% male (as assigned at birth), 70% African American, with HIV	mHealth app	Intervention group reported significant decreases in perceived stress ( $p = 0.02$ ), illicit substance use ( $p = 0.05$ ), a short-term decrease in depression ( $p = 0.02$ ), and a short-term increases in self-efficacy ( $p = 0.04$ ) in comparison to control.	Decreased stress, decreased depression, decreased substance use, increase in self- efficacy, and fewer physician visits	Non- experimental (no randomization and no control)
Schnall et al <sup>24</sup>	Older adults average age 50.4, 52.5% female (as assigned at birth), 68.8% Caucasian, with HIV	mHealth app (mVIP)	Improvement in anxiety ( $p = 0.001$ ), depression ( $p = 0.001$ ), neuropathy ( $p = 0.002$ ), fever/chills/sweat ( $p = 0.037$ ), weight loss/wasting ( $p = 0.020$ ), and medication adherence over the control group	Decrease in anxiety, decrease in depression, improvement in neuropathy, decrease in fever/chills/sweat, improvement in weight loss, and improvement in medication adherence	Robust experiment
Stonbraker et al <sup>25</sup> Older adults with HIV, average age 54.4, 60% female (as assigned at birth), 85% African American		mHealth app (VIP- HANA)	No control group. Good usability, but needs improvement	Not reported	Non- experimental (no randomization, no control)
van der Kop et al <sup>26</sup>	Adults with HIV, 60% female (as assigned at birth)	mHealth SMS	No effect on retention or treatment outcomes	No effect on treatment outcomes	RCT
Basaran et al <sup>27</sup>			Decreased risk-taking behavior compared with the control group	Decreased risk-taking behavior	RCT
Brantley et al <sup>28</sup>	Adults with HIV, 69.3% Black, 69.3% male (as assigned at birth)	videoconferencing	No significant improvement in the linkage to care	None reported	Robust experiment

ho et al <sup>29</sup>	Older adults with HIV, average	mHealth app to pill	No control group. Increased medication adherence,	Increased medication adherence and	Robust
	age 47.6, 79% African	bottle (Wise App)	and increased self-efficacy	increased self-efficacy	experiment
	American, 60% female (as				
	assigned at birth)				
lirshfield et al <sup>30</sup>	Adult men (as assigned at	eHealth video-based	Reduced risky behavior more than the control	Reduced risky sexual behavior	Robust
	birth) with HIV, average age	intervention (Sex	group		experiment
	39, 50% Caucasian	Positive)			
uo et al <sup>31</sup>	Adults with HIV, average age	mHealth app (CARE +	Increased odds of viral suppression (not statistically	Increased viral suppression and increased care	Robust
	41.5, 58% male (as assigned at	Corrections)	significant) and increase in care engagement	engagement	experiment
	birth), 19% transgender (male-		compared with control group		
	to-female), 85% Black				
Curth et al <sup>32</sup>	Adults with HIV, 80% female	eHealth	Reduced viral load ( $p = 0.0007$ ) compared with the	Reduced viral load	RCT
	(as assigned at birth), 100%		control group		
	Black, average age 37.5				
i et al <sup>33</sup>	Adults with HIV, 100%	mHealth WeChat	Reduced suicide rate ( $p = 0.02$ ), reduced stress and	Reduced suicide, reduced stress, and reduced	RCT
	Chinese, 92.3% male (as	(Run4Love)	depressive symptoms ( $p = 0.001$ ) compared to	depression	
	assigned at birth), average age		control		
	27.5				
arna et al <sup>34</sup>	Adult females (as assigned at	mHealth app	Retention in counseling was higher in the	Increased retention, increased HIV testing of	RCT
	birth) with HIV, 100% Black,		intervention, HIV testing of infants was higher in the	infants, increased medication adherence, and	
	average age 25		intervention group, and positive infant HIV tests	decreased HIV transmission to infants	
			were lower in intervention because of medication		
			adherence		
oni et al <sup>35</sup>	Adult males (as assigned at	mHealth app	32% lower condomless anal intercourse (CAI) in	Decrease in sexual risk behaviors	RCT
	birth) with HIV, 100% Black,	(HealthMpowerment,	the intervention group (short-term effect only).		
	average age 24.3	HMP)	Among HIV-positive participants, it was 82% lower		
			CAI than the control.		
hu et al <sup>36</sup>	Adults with HIV, 100%	mHealth WeChat app	Decreased depressive symptoms, positive coping,	Decreased depression, increased coping, and	RCT
	Chinese, average age 27.5,	(Run4Love) + phone	and decreased HIV stigma compared to the control	decreased HIV stigma	
	92.3% male (as assigned at	calls			
	birth)				
arroso et al <sup>37</sup>	Older adults with HIV, average	mHealth CBT stress	Decreased fatigue compared to the control	Decreased fatigue	RCT
	age 51.2, 66.5% Black, 63%	management			
	male (as assigned at birth)				

Table I (Continued).
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Authors	Participants         Experimental         Results (Compared to Control Group)           Intervention         Intervention         Intervention		Results (Compared to Control Group)	Medical Outcomes Reported	Study Design	
Carey et al <sup>38</sup>	y et al <sup>38</sup> Older adults with HIV, 50% Telephone male (as assigned at birth), mindfu 52% male (gender identity), average age 47.5, 43% Caucasian		Improved medication adherence, increased mindfulness, reduced sexual risk behavior, reduced anxiety, reduced depressive symptoms, decreased stress, decreased impulsivity – no statistical difference with control	Improved medication adherence, increased mindfulness, reduced sexual risk behavior, reduced anxiety, reduced depressive symptoms, decreased stress, and decreased impulsivity	RCT	
Fahey et al <sup>39</sup>	Adults (18–35) with HIV, 100% Black, 62.3% female (as assigned at birth)	mHealth app	Intervention increased retention in treatment and achieved viral suppression, incentive size and viral suppression were positively correlated ( $P$ -trend = 0.0032)	Increase in viral suppression and program retention	RCT	
Guo et al <sup>40</sup>	Adults with HIV, 100% Chinese, 92.3% male (as assigned at birth), average age 27.5	mHealth WeChat app (Run4Love)	Significant reduction in CES-D score in the intervention group ( $p$ <0.001)	Reduction in depression symptoms	RCT	
Marhefka et al <sup>41</sup>			No control group.     Not reported       Older participants less likely to use technology,     Hispanics less likely to use technology for HIV       issues     Image: Control of the second secon		Observational	
Uhrig et al <sup>42</sup>	Adults with HIV, average age 42, 86% Black, 58% male (as assigned at birth), and 18% transgender		Participant preferred the customized messages	Not reported	RCT	
Zeng et al <sup>43</sup>	Adults with HIV, 94.7% male (as assigned at birth), 100% Chinese, average age 28	with HIV, 94.7% male mHealth app Decreased depressive symptom gned at birth), 100%		Decreased depression	RCT	
Li et al <sup>44</sup>			Increased quality of life ( $p = 0.001$ ), reduced HIV- related stigma ( $p = 0.003$ ), and reduced depressive symptoms ( $p = 0.001$ )		RCT	
Plomer et al <sup>45</sup>	Adults males (as assigned at birth) with HIV, median age 40	mHealth app (PrEP)	No control group. Users appreciated the confidence the app gave them to enjoy their sex lives	Increased confidence, increased patient-to- provider relationship	Qualitative	
Policarpo et al <sup>46</sup>	icarpo et al <sup>46</sup> Older adults average age 54, Video and phone 74.2% male (as assigned at counseling on diet birth), non-alcohol fatty liver (standard vs disease (NAFLD) patients with HIV		74.2% male (as assigned at birth), non-alcohol fatty liver disease (NAFLD) patients withcounseling on diet (standard vs0.01), but both groups gained weight, but the intervention gained less (p < 0.002)gaineddisease (NAFLD) patients withMediterranean)MediterraneanMediterraneanMediterranean		RCT	

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Twimukye et al <sup>47</sup>	Young adults (18–24) with HIV,	mHealth app (CFLU)	No control group.	Increased medication adherence,	Qualitative
	76% female (as assigned at		Increased medication adherence, strengthened	strengthened relationships with provider, and	
	birth)		relationships with provider, increased health	increased health knowledge	
			knowledge		
Yelverton et al <sup>48</sup>	Adult carers for PLWH	Telehealth	No control group.	Not reported	Qualitative
			Security concerns, digital literacy gap		
Guo et al <sup>49</sup>	Adults with HIV, 92.3% male	mHealth WeChat app	Decreased depressive symptoms ( $p = 0.002$ )	Decreased depression	RCT
	(as assigned at birth), 100%	(Run4Love)			
	Chinese, average age 28.3				
Stephenson et al <sup>50</sup>	Adult males with HIV, average	Telehealth couples	Couples in the intervention group reported safer	Decreased interpersonal problems	RCT
	age 30.4, 75% Caucasian, 100%	counseling and testing	sexual agreements ( $p = 0.007$ ), lower odds of		
	male (as assigned at birth)	(CHTC)	discordant relationships ( $p = 0.048$ ), lower odds of		
			breaking their sexual agreement ( $p = 0.000$ )		
Zeng et al <sup>51</sup>	Adults with HIV, 92.3% male	mHealth WeChat app	Increased quality of life through positive coping ( $p$ =	Increased quality of life	RCT
	(as assigned at birth), 100%	(Run4Love)	0.006) over control		
	Chinese, average age 27.5				

Table 2 Summary of Analysis,	Sorted	Chronologically	by Au	uthor
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Authors	Intervention Theme	Results Themes	Medical Outcomes Themes	Patient Satisfaction Themes	Facilitator Themes	Barrier Themes
Heckman et al <sup>20</sup>	Telephone	Improved mental health	Improved mental health	Satisfied	Convenience	Cost of technology
		conditions	conditions			
		Decreased interpersonal	Decreased interpersonal		No HIV stigma associated	Training of staff
		problems	problems		with clinic	
					Improvement in medical	Visual cues not available
					condition(s)	over phone
Himelhoch et al <sup>21</sup>	mHealth app	Decreased substance use	Decreased substance use	Satisfied	Convenience	Cost of technology
		Increased care engagement	Increased care engagement		Meets a digital preference of	Training of staff
					patient	
IX 11: 1		No statistical difference			Increases care engagement	
Kalichman et al <sup>22</sup>	mHealth app	Reduced HIV RNA	Reduced HIV RNA	Satisfied	Convenience	Cost of technology
		Increased care engagement	Increased care engagement		Meets a digital preference of patient	Training of staff
		Increased physical health	Increased physical health		Improvement in medical	
		conditions	conditions		condition(s)	
		Decreased substance use	Decreased substance use		Increases care engagement	
Sayegh, et al <sup>23</sup>	mHealth app	Improved mental health	Improved mental health	Satisfied	Convenience	Cost of technology
		conditions	conditions			
		Decreased substance use	Decreased substance use		Meets a digital preference of	Training of staff
					patient	
		Improved mental health	Improved mental health		No HIV stigma associated	
		conditions	conditions		with clinic	
		Increased care engagement	Increased care engagement		Improvement in medical condition(s)	
		Decreased physician visits	Decreased physician visits		Increases care engagement	
Schnall et al <sup>24</sup>	mHealth app	Improved mental health	Improved mental health	Satisfied	Convenience	Cost of technology
		conditions	conditions			
		Improved mental health	Improved mental health		Meets a digital preference of	Training of staff
		conditions	conditions		patient	
		Increased physical health	Increased physical health		Improvement in medical	
		conditions	conditions		condition(s)	
		Increased physical health	Increased physical health		Low healthcare utilization	
		conditions	conditions			
		Increased care engagement	Increased care engagement		Increases care engagement	

tonbraker et al <sup>25</sup>	mHealth app	Good usability of app	Not reported	Satisfied	Convenience	Cost of technology
					Meets a digital preference of	Training of staff
					patient	
					Ease of use	Usability
					Ease of use	
van der Kop et al <sup>26</sup>	mHealth SMS	No effect on treatment	No effect on treatment	Satisfied	Convenience	Cost of technology
		outcomes	outcomes		Meets a digital preference of	Training of staff
					patient	
Basaran et al <sup>27</sup>	eHealth	Changed behavior	Changed behavior	Satisfied	Convenience	Cost of technology
					Meets a digital preference of	Training of staff
					patient	
Brantley et al <sup>28</sup>	eHealth	No statistical difference	Not reported	Satisfied	Convenience	Cost of technology
					Meets a digital preference of	Training of staff
					patient	
					No HIV stigma associated	
					with clinic	
Cho et al <sup>29</sup>	mHealth app	Increased care engagement	Increased care engagement	Satisfied	Convenience	Cost of technology
					Meets a digital preference of	Training of staff
					patient	
					No HIV stigma associated	
					with clinic	
					Increases care engagement	
Hirshfield et al <sup>30</sup>	eHealth	Changed behavior	Changed behavior	Satisfied	Convenience	Cost of technology
					Meets a digital preference of	Training of staff
					patient	
					No HIV stigma associated	
					with clinic	
Kuo et al <sup>31</sup>	mHealth app	Reduced HIV RNA	Reduced HIV RNA	Satisfied	Convenience	Cost of technology
		No statistical difference	Increased care engagement		Meets a digital preference of	Training of staff
					patient	
		Increased care engagement			No HIV stigma associated	
					with clinic	
					Increases care engagement	
					Improvement in medical	
					condition(s)	

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Authors	Intervention Theme	Results Themes	Medical Outcomes Themes	Patient Satisfaction Themes	Facilitator Themes	Barrier Themes
Kurth et al <sup>32</sup>	eHealth	Reduced HIV RNA	Reduced HIV RNA	Satisfied	Convenience Meets a digital preference of patient No HIV stigma associated with clinic Improvement in medical condition(s)	Cost of technology Training of staff
Li et al <sup>33</sup>	mHealth app	Improved mental health conditions Improved mental health conditions Improved mental health conditions	Improved mental health conditions Improved mental health conditions Improved mental health conditions	Satisfied	Convenience Meets a digital preference of patient No HIV stigma associated with clinic Improvement in medical condition(s)	Cost of technology Training of staff
Sarna et al <sup>34</sup>	mHealth app	Increased care engagement Increased HIV testing of infants Decreased HIV transmission to infants Increased care engagement	Increased care engagement Increased HIV testing of infants Decreased HIV transmission to infants Increased care engagement	Satisfied	Convenience Meets a digital preference of patient No HIV stigma associated with clinic Increases care engagement	Cost of technology Training of staff
Soni et al <sup>35</sup>	mHealth app	Changed behavior	Changed behavior	Satisfied	Convenience Meets a digital preference of patient No HIV stigma associated with clinic	Cost of technology Training of staff
Zhu et al <sup>36</sup>	mHealth app	Improved mental health conditions Improved mental health conditions Decreased HIV stigma	Improved mental health conditions Improved mental health conditions Decreased HIV stigma	Satisfied	Convenience Meets a digital preference of patient No HIV stigma associated with clinic Improvement in medical condition(s)	Cost of technology Training of staff

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Barroso et al <sup>37</sup>	mHealth app	Increased physical health	Increased physical health	Satisfied	Convenience	Cost of technology
		conditions	conditions		Meets a digital preference of	Training of staff
					patient	
					No HIV stigma associated	
					with clinic	
					Improvement in medical	
					condition(s)	
Carey et al <sup>38</sup>	Telephone	Increased care engagement	Increased care engagement	Satisfied	Convenience	Cost of technology
		Changed behavior	Changed behavior		Meets a digital preference of	Training of staff
					patient	
		Changed behavior	Changed behavior		No HIV stigma associated	
					with clinic	
		Improved mental health	Improved mental health		Improvement in medical	
		conditions	conditions		condition(s)	
		Improved mental health	Improved mental health		Increases care engagement	
		conditions	conditions			
		Improved mental health	Improved mental health			
		conditions	conditions			
		Changed behavior	Changed behavior			
		No statistical difference				
Fahey et al <sup>39</sup>	mHealth app	Increased care engagement	Increased care engagement	Satisfied	Convenience	Cost of technology
		Reduced HIV RNA	Reduced HIV RNA		Meets a digital preference of	Cost of incentives
					patient	
		Changed behavior	Changed behavior		No HIV stigma associated	Training of staff
					with clinic	
					Improvement in medical	
					condition(s)	
Guo et al <sup>40</sup>	mHealth app	Improved mental health	Improved mental health	Satisfied	Convenience	Cost of technology
		conditions	conditions		Meets a digital preference of	Training of staff
					patient	
					No HIV stigma associated	
					with clinic	
					Improvement in medical	
					condition(s)	

(Continued)

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Table 2	(Continued).
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Authors	Intervention Theme	Results Themes	Medical Outcomes Themes	Patient Satisfaction Themes	Facilitator Themes	Barrier Themes
Marhefka et al <sup>41</sup>	eHealth	Digital literacy gap	Not reported	Not reported	Convenience	Digital literacy gap
		Security concerns			Meets a digital preference of	Security concerns
					patient	
					No HIV stigma associated	
Uhrig et al <sup>42</sup>	mHealth SMS		Net we ented	Satisfied	with clinic Convenience	Cost of tools also
Onrig et al	mHealth SMS	Users preferred customizable	Not reported	Satisfied		Cost of technology Training of staff
		messages			Meets a digital preference of patient	Training of staff
					No HIV stigma associated	
					with clinic	
					Customizable messages	
Zeng et al <sup>43</sup>	mHealth app	Improved mental health	Improved mental health	Satisfied	Convenience	Cost of technology
	in lealer upp	conditions	conditions	batisfied	Meets a digital preference of	Training of staff
		conditions	conditions		patient	
					No HIV stigma associated	
					with clinic	
					Improvement in medical	
					condition(s)	
Li et al <sup>44</sup>	mHealth app	Increased quality of life	Increased quality of life	Satisfied	Convenience	Cost of technology
		Decreased HIV stigma	Decreased HIV stigma		Meets a digital preference of	Training of staff
		_	-		patient	-
		Improved mental health	Improved mental health		No HIV stigma associated	
		conditions	conditions		with clinic	
					Improvement in medical	
					condition(s)	
Plomer et al <sup>45</sup>	Telephone	Increased quality of life	Increased quality of life	Satisfied	Convenience	Cost of technology
					Meets a digital preference of	Training of staff
					patient	
					No HIV stigma associated	Security concerns
					with clinic	
					Increased quality of life	
Policarpo et al <sup>46</sup>	Telephone	Changed behavior	Changed behavior	Satisfied	Convenience	Cost of technology
					Meets a digital preference of	Training of staff
					patient	

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Twimukye et al <sup>47</sup>	mHealth app	Increased care engagement	Increased care engagement	Satisfied	Convenience	Cost of technology
,		Increased care engagement	Increased care engagement		Meets a digital preference of	Training of staff
					patient	U U
		Increased health knowledge	Increased health knowledge		Increased health knowledge	Needs to be youth friendly
					Increases care engagement	Technical issues
					Increases care engagement	
Yelverton, et al <sup>48</sup>	Telemedicine	Security concerns	Not reported	Not reported	Convenience	Cost of technology
		Digital literacy gap			Meets a digital preference of	Training of staff
					patient	Distribution of technology
						Digital literacy gap
						Low reimbursement
						Security concerns
						Socio-economic status of
						patient
Guo et al <sup>49</sup>	mHealth app	Improved mental health	Improved mental health	Satisfied	Convenience	Cost of technology
		conditions	conditions		Meets a digital preference of	Training of staff
					patient	
					No HIV stigma associated	
					with clinic	
					Improvement in medical	
					condition(s)	
Stephenson et al <sup>50</sup>	Telemedicine	Changed behavior	Decreased interpersonal	Satisfied	Convenience	Cost of technology
		Decreased interpersonal	problems		Meets a digital preference of	Training of staff
		problems			patient	
		Changed behavior			No HIV stigma associated	
					with clinic	
					Improved behaviors	
Zeng et al <sup>51</sup>	mHealth app	Increased quality of life	Increased quality of life	Satisfied	Convenience	Cost of technology
		Improved mental health	Improved mental health		Meets a digital preference of	Training of staff
		conditions	conditions		patient	
					No HIV stigma associated	
					with clinic	
					Increased quality of life	

## Results of Interventions, Compared with Control Groups

Table 3 summarizes the results of studies, compared with control groups (where appropriate). Six studies were qualitative, non-experimental, or observational studies. Although these studies did not have control groups, their results were still reported. Twelve themes and six individual observations were identified by the reviewers for a total of 74 observations. Because these are themes, there were multiple instances of one theme in the same article, which will be explained. There were 18/74 (24%) instances of "improved mental health conditions" which included the following observations: Decreased anxiety, decreased depression, decreased stress, reduced suicidal ideations, and increased coping. 20,23,24,33,36,38,40,43,44,49,51 There were 13/74 (18%) instances of "increased care engagement" which included the following: increased patient-toprovider relationship, increased retention in treatment program, improved medication adherence, and increased selfefficacy.<sup>21-24,29,31,34,38,39,47</sup> There were 10/74 (14%) instances of changed behavior, which included the following: decreased risk-taking behavior, decreased risky sex behavior, increased mindfulness, decreased impulsivity, decreased condomless anal intercourse, changed diet, and safer sex agreements with partner.<sup>27,30,35,38,39,46,50</sup> There were 4/74 (5%) instances of three themes: increased physical health outcomes, which included fever, chills, sweats, neuropathy, weight loss, fatigue, and genital tract inflammation,<sup>22,24,37</sup> no statistical difference in improvement with the control group, which means it is equally as effective as traditional care, <sup>21,28,31,38</sup> and reduced HIV RNA or increased viral suppression. <sup>22,31,32,39</sup> There were 3/74 (4%) instances of two themes: decreased substance use, which included a decrease in drug cravings.<sup>21–23</sup> and an increased quality of life, which included peace of mind or increased confidence to enjoy a sex life.<sup>44,45,51</sup> There were 2/74 (3%) instances of four themes: decreased PLHIV stigma<sup>36,44</sup> and decreased interpersonal problems, which included fewer discordant relationships,<sup>20,50</sup> highlighting a digital literacy gap<sup>41,48</sup> and security or privacy concerns.<sup>41,48</sup> The last two results are also listed as barriers to adoption. The following observations only occurred once in the literature: decreased HIV transmission to infants, decreased physician visits, good usability of app, increased health knowledge, increased HIV testing of infants, no effect on treatment outcomes, and users preferred customizable messages.<sup>23,25,26,34,42,47</sup>

Table 3 Summary of Results Compared to the Contro	Table 3 Su	ummary of	Results	Compared	to	the	Control
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Results Themes and Observations	Frequency
Improved mental health conditions <sup>20,23,24,33,34,38,40,43,44,49,51</sup> *	18
Increased care engagement <sup>21-24,29,31,34,38,39,47</sup> *	13
Changed behavior <sup>27,30,35,38,39,46,50</sup> *	10
Increased physical health conditions <sup>22,24,37</sup> *	4
No statistical difference <sup>21,28,31,38</sup>	4
Reduced HIV RNA <sup>22,31,32,39</sup>	4
Decreased substance use <sup>21-23</sup>	3
Increased quality of life <sup>44,45,51</sup>	3
Decreased HIV stigma <sup>36,44</sup>	2
Decreased interpersonal problems <sup>20,50</sup>	2
Digital literacy gap <sup>41,48</sup>	2
Security concerns <sup>40,47</sup>	2
Decreased HIV transmission to infants <sup>34</sup>	I
Decreased physician visits <sup>23</sup>	I
Good usability of app <sup>25</sup>	I.
Increased health knowledge <sup>47</sup>	I.
Increased HIV testing of infants <sup>34</sup>	I
No effect on treatment outcomes <sup>26</sup>	I
Users preferred customizable messages <sup>42</sup>	I
Total	74

Notes: \*Multiple occurrences observed in one study.

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#### Medical Outcomes Commensurate with the Intervention

Table 4 summarizes the medical outcomes commensurate with the interventions. Eight themes and six individual observations were identified by the reviewers for a total of 67 occurrences in the literature. The Results compared to the control group and medical outcomes were highly similar, but they are focused on themes and observations for the provider. For instance, observations about security, usability, and literacy gap are not included.

## Facilitators to the Intervention of mHealth to Manage HIV

Table 5 summarizes the facilitators observed. Seven themes and four individual observations were identified by the reviewers for a total of 119 occurrences in the literature. Convenience was identified in 32/119 (27%) instances, which included no travel and managing HIV from the comfort of the home.<sup>20–51</sup> The intervention meets a digital preference of the patient and was identified 31/119 (26%) times.<sup>21–51</sup> The intervention frees the patient from a typical HIV stigma usually felt when visiting the clinic occurred 23/119 (19%) times.<sup>20,23,28–45,49–51</sup> There were 15/119 (13%) instances of improvements in medical conditions, which included the following: weight loss, chills, sweats, fever, neuropathy, fatigue, genital tract inflammation, anxiety, depression,

Table	4 Medical	Outcomes	Commensurate	with	the	Interventions
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Medical Outcomes Themes and Observations	Frequency
Improved mental health conditions <sup>20,23,24,33,36,38,40,43,44,49,51</sup> *	18
Increased care engagement <sup>21–24,29,31,34,38,39,47</sup> *	13
Changed behavior <sup>27,30,35,38,39,46,50</sup> *	8
Increased physical health conditions <sup>22,24,37</sup> *	4
Reduced HIV RNA <sup>22,31,32,39</sup>	4
Decreased substance use <sup>21–23</sup>	3
Increased quality of life <sup>44,45,51</sup>	3
Decreased HIV stigma <sup>36,44</sup>	2
Decreased interpersonal problems <sup>20,50</sup>	2
Decreased HIV transmission to infants <sup>34</sup>	I
Decreased physician visits <sup>23</sup>	I
Increased health knowledge <sup>47</sup>	I
Increased HIV testing of infants <sup>34</sup>	I
No effect on treatment outcomes <sup>26</sup>	I
Not reported <sup>25,28,41,42,48</sup>	5
Total	67

Notes: \*Multiple occurrences observed in one study.

Facilitator Themes and Observations	Frequency
Convenience <sup>20–51</sup>	32
Meets a digital preference of patient <sup>21–51</sup>	31
No HIV stigma usually associated with clinic visit <sup>20,23,28–45,49–51</sup>	23
Improvement in medical condition(s) <sup>20,22–24,31–33,36–40,43,44,49</sup>	15
Increases care engagement <sup>21–24,29,31,34,38,47</sup>	10
Ease of use <sup>25</sup> *	2
Increased quality of life <sup>45,51</sup>	2
Customizable messages <sup>42</sup>	I
Improved behaviors <sup>50</sup>	I
Increased health knowledge <sup>47</sup>	I
Low healthcare utilization <sup>24</sup>	I

#### Table 5 Facilitators to the Intervention of mHealth to Manage HIV

Notes: \*Multiple occurrences observed in one study.

stress, suicide ideation, coping, substance use, and substance cravings.<sup>20,22–24,31–33,36–40,43,44,49</sup> There were 10/119 (8%) instances of increased care engagement which included building relationships with provider, treatment adherence, medication adherence, self-efficacy, and viral load.<sup>21–24,29,31,34,38,47</sup> There were 2/119 (2%) instances of two themes: ease of use, which included look and feel,<sup>25</sup> and increased quality of life, which included increased confidence to enjoy a sex life.<sup>45,51</sup> The following observations only occurred once in the literature: customizable messages, improved behaviors, increased health knowledge, and low healthcare utilization.<sup>24,42,47,50</sup>

#### Barriers to the Intervention of mHealth for the Management of HIV

Very few barriers were identified in the literature that could create a theme. The burden of cost, for the equipment, the app, and the incentives, appeared in 32/75 (42%) of occurrences.<sup>20–51</sup> The requirement to train staff appeared in 31/75 (41%) of occurrences.<sup>20–40,42–51</sup> Security concerns appeared in 3/75 (4%) of occurrences.<sup>41,45,48</sup> The existence of a digital literacy gap appeared in 2/75 (3%) of occurrences.<sup>41,48</sup> Seven individual observations appeared in 1/75 (1%) of occurrences: distribution of technology, low reimbursement, needs to be youth friendly, socio-economic status of patients, technical issues, usability, and visual cues not available over the phone.<sup>20,25,47,48</sup>

#### Interactions Between Observations

The telephone interventions were predominantly used with older adults as the participants.<sup>20,21,24,29,37,38,46</sup> These interventions were effective with this older population, and they resulted in improved mental health conditions, decreased interpersonal problems, increased physical health, decreased substance use, increased care engagement, and changed behavior. This intervention is a good solution in the face of the digital literacy gap.<sup>41,48</sup> One study even highlighted the preference of older participants away from mHealth apps.<sup>41</sup> Young adults and mid-range adults were comfortable with mHealth apps and eHealth apps used on mobile devices.

# Discussion

#### Summary of Evidence

This systematic literature review analyzed 32 studies from seven countries published over the last five years to analyze the facilitators and barriers to the use of mHealth for the treatment and management of PLHIV. Five interventions were studied (mHealth app, mHealth SMS, eHealth, telephone, and telemedicine). The lines between mHealth and eHealth are blurred due to the robust capabilities of mobile phones and other devices. mHealth apps, eHealth apps, and mHealth SMS comprised 71% (26/32) of the modalities analyzed.<sup>21–37,39,41,42,44,47</sup> Also, 26/32 (71%) used either RCT or other robust experiments as the modality.<sup>20–22,24,26–40,42–44,46,49–51</sup> Although effectiveness was not one of the objectives of this review, the results of physical and mental health (primary outcomes),<sup>20,22–24,31–33,36–40,43,44,49</sup> changes in behavior (secondary outcomes),<sup>27,30,35,38,39,46,50</sup> and increases in care engagement,<sup>21–24,29,31,34,38,47</sup> are all excellent cases for the effectiveness of this modality.

Significantly more facilitators were identified than barriers. The largest barrier is one of the convenience and preference. Managing PLHIV symptoms through an mHealth app reduces the number of times a patient must visit the HIV clinic, which reduces the stigma of coming into the clinic.<sup>20,23,28–45,49–51</sup> The lack of stigma may also play a role in why mHealth and eHealth apps increase engagement in care.<sup>21–24,29,31,34,38,47</sup> Users were very pleased with the modality, interfaced with it well, and commented on ease of use.<sup>25</sup> There were zero reports of negative satisfaction with the interventions. The only negative comments were about making the apps more friendly to a younger audience,<sup>47</sup> the digital literacy gap,<sup>41,48</sup> fixing small usability challenges,<sup>25,47</sup> and helping users feel more comfortable about the security of their data.<sup>41,45,48</sup> There are larger institutional barriers to the adoption of this modality in terms of the cost of technology,<sup>20–51</sup> training of staff,<sup>20–40,42–51</sup> and low reimbursement.<sup>48</sup>

Future research should focus on distinct apps for age groups. Some apps could be developed to be more friendly to younger audiences, and other apps could be simplified for an older audience. Because ART is so vital to limit the spread of HIV, and mHealth is effective in the improvement of medication adherence,<sup>21,22,24,29,31,32,34,39</sup> future research should integrate ART reminders into other apps. The customization of SMS messages was popular with one study,<sup>42</sup> so future

research on SMS interventions should allow customization. Because risky behavior and substance abuse are socialized into many PLHIV circles,<sup>27,30,35,38,46,50</sup> it is vital that additional research seeks behavior changing interventions.

The results of this review should enable providers to adopt effective mHealth interventions for their PLHIV population with confidence. The results should help PLHIV develop healthy habits and limit transmission of HIV. Administrators should feel confident that the infrastructure necessary to adopt telemedicine practices like mHealth interventions is effective at improving physical health, mental health, care engagement, and changing behavior.

#### Limitations

We queried four research databases in order to limit sample bias. However, had we used other databases, we may have identified other studies with additional mHealth interventions for the treatment and management of PLHIV. We used only peer-reviewed, published literature in order to control for validity, but this poses a risk of publication bias. Opening the search up to the grey literature could have controlled for publication bias, but it may have introduced additional validity issues. We identified several instances of both selection and sample bias. We did not determine these instances were significant enough to discount the studies from analysis; however, these forms of bias threaten internal and external validity. To control for design bias, we used a published protocol for the conduct of this systematic literature review.

# Conclusions

mHealth offers several interventions that are effective in the treatment and management of PLHIV. These interventions create improvements in physical health, mental health, care engagement and behavior management. While a few barriers stand in the way of universal adoption, there are many facilitators to the adoption of mHealth interventions that far outweigh the barriers. mHealth interventions are necessary to develop an atmosphere of ART adherence and non-risky sexual behavior to treat and manage HIV and limit the spread of the condition.

# **Protocol and Registration**

This review is conducted in accordance with the Kruse Protocol for writing a systematic review. This protocol was published in 2019, and from it, 47 systematic literature reviews have been published. This systematic literature review is reported in accordance with the Preferred Reporting Items for Systematic Reviews and Meta-Analysis (PRISMA 2020). It is registered with PROSPERO: CRD42021266719.

# **Data Sharing Statement**

Data from this study can be obtained by contacting the lead author.

# **Author Contributions**

All authors made a significant contribution to the work reported, whether that is in the conception, study design, execution, acquisition of data, analysis and interpretation, or in all these areas; took part in drafting, revising or critically reviewing the article; gave final approval of the version to be published; have agreed on the journal to which the article has been submitted; and agree to be accountable for all aspects of the work.

# Disclosure

The authors declare that they have no competing interests.

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