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Review Article Opioid Use Disorder

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Digital Therapeutics in the Management of Opioid Use Disorder: A Systematic Review of Emerging Technologies and Clinical Outcomes

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Opioid use disorder (OUD) poses a major global health burden, with limited access to traditional treatment in many regions. Digital therapeutics (DTx) offer, noble tools to support treatment, enhance accessibility, and improve outcomes. The objective of the study is to systematically review the current evidence on the effectiveness, implementation, and outcomes of DTx in the management of OUD. A thorough literature was searched using PubMed, Science direct, Google scholar et cetera, from 2015 to 2024, which included randomize control trials (RCTs), systemic reviews, and real-world studies evaluating digital interventions (apps, telehealth, AI based tools, wearable for OUD. A total of 28 studies met the inclusion criteria. digital interventions demonstrated improved treatment, retention, opioid abstinence, and patient engagement. Tools such as reset-O, telehealth, CBT based mobile apps and AI driven systems showed efficacy in both clinical and real-world settings. Barriers included digital, illiteracy, privacy, concerns, and Limited regulatory frameworks, especially in low- and middle-income countries. Digital therapeutics present a promising adjunct or alternative to conventional OUD treatment. Tailored implementation, Cultural Adaptation, and regulatory support are essential for maximizing their impact.

Keywords: Digital health, Opioid Addiction, Telepsychiatry, CBT Apps, reSET-O, AI in addiction, mHealth, Digital Interventions

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Introduction

OUD continues to escalate as global health issue, with over 60 million people estimated to have used opioids in 2022, & millions other suffering from opioid dependence & its complications.[1] Traditional treatment modalities such as methadone, like psychosocial interventions & Cognitive Behavioral therapy (CBT), are well established but face challenges in accessibility, stigma & patient retention.[2] Digital therapeutics (DTx) have developed as innovative, scalable, & evidence-based healthcare delivery options, including treatment for OUD. These include smartphone applications, prescription digital treatments (e.g., reSET-O), AIpowered behavioral tracking tools, wearable biosensors, & telehealth systems. These tools are designed to promote abstinence, increase treatment adherence, & provide real-time behavioral interventions.[3],[4] Promise of DTx resides in its ability to fill treatment gaps, particularly in rural & underprivileged populations. For example, reSET-O, first FDA-approved digital therapy for OUD, demonstrated enhanced abstinence & retention rates in both randomized controlled trials & realworld appl. (Maricich et al., 2021).[5] Similarly, telemedicine & mHealth initiatives increased access to MOUD during COVID-19 epidemic,

With results equivalent to in-person therapy.[6] Despite the good results, challenges to widespread adoption persist. These include worries regarding digital literacy, data privacy, governmental backing, and a lack of culturally appropriate solutions in low-and middle-income nations like India.[7]

The objective of this systematic review is to consolidate and evaluate the current data on digital therapy tools in OUD treatment, focusing on their clinical efficacy, real-world applicability, and the problems associated with implementation across various settings.

Materials and Methods

Search strategy

This systematic review was carried out in accordance with the PRISMA guidelines. [Figure 1]. A comprehensive literature search was conducted using the PubMed, ScienceDirect, and Google Scholar databases for publications published between January 2015 and March 2024. The following search terms and Boolean operators were used: ("Opioid Use Disorder" OR "OUD") AND ("digital therapeutics" OR "mobile health" OR "telemedicine" OR "CBT apps" OR "reSET-O" OR "AI in addiction" OR "wearables" OR "digital health").



Figure 1: PRISMA 2020 flowchart.

Inclusion and Exclusion Criteria

Inclusion criteria: Research published in peerreviewed journals between 2015 and 2024. Randomized controlled trials (RCTs), systematic reviews, meta-analyses, and real-world implementation research. Digital treatments for managing OUD and human participants.

Exclusion criteria for non-English language publications. Case studies, editorials, and conference abstracts without complete text. Studies that are not focused on OUD or do not include digital therapy approaches.

Data Extraction and synthesis

To identify appropriate studies, two independent authors examined the titles and abstracts. Full texts were retrieved for potentially relevant articles. Disagreements were resolved by discussion and consensus. The following key data were extracted: study design and location, sample size and demographic characteristics, kind of digital intervention, Outcome measures (e.g., abstinence rates, treatment retention, engagement, satisfaction), key findings, and conclusions.

Quality Assessment

The Cochrane Risk of Bias Tool was used to assess the quality and risk of bias for all included RCTs. Systematic reviews and observational studies were assessed using the AMSTAR 2 checklist and the Newcastle-Ottawa Scale, respectively.

Results

Overview of Included Studies

After screening and full-text evaluation, 28 of the 357 records found during the database search met the criteria for eligibility requirements. They included: Ten randomized controlled trials (RCTs), six systematic reviews or meta-analyses, and 12 real-world implementation or observational studies. These studies included both high-income countries like the United States and Canada, as well as middle-income situations like India.

| Study (Year) | Sample & | Digital Intervention | Comparator | Primary/Secondary | Key Findings (Effect Size) |
|------------------|-------------------|-------------------------------------|----------------------|-------------------------|---|
| | Population | | | Outcomes | |
| Maricich et | N=170 adults | reSET-0 (12-week prescription | Treatment-as-usual | Primary: Opioid | Significantly higher opioid abstinence in digital |
| al., 2021 | with DSM-IV | digital therapeutic; 67 interactive | (TAU): | abstinence (urine- | group (77.3% vs 62.1% abstinent in weeks 9-12, |
| (reSET-O trial- | OUD on | CRA modules + contingency | buprenorphine + | negative) during | p = 0.02; OR \sim 2.08). Retention improved (hazard |
| USA) | buprenorphine | management via mobile app) | biweekly clinician | weeks 9–12; | of dropout 0.49 vs TAU). No difference in adverse |
| | in outpatient | | visits + urine | Treatment retention. | events. |
| | treatment. | | monitoring + CM | Secondary: Adverse | |
| | | | vouchers. | events. | |
| Shi et al., | N=20 adults | CBT4CBT-Buprenorphine | Standard | Primary: Treatment | Digital CBT group had higher opioid-negative urine |
| 2019 | with OUD (DSM- | (web-based CBT program tailored | buprenorphine | retention; | rates (91% vs 64% at end of 12 weeks, p \approx 0.05) |
| (CBT4CBT- | 5) on office- | for OUD, self-guided modules) | treatment alone | Opioid/cocaine | and higher overall drug abstinence (82% vs 30%, |
| Bup- USA) [8] | based | plus standard buprenorphine | (office-based, | abstinence (urine | p = 0.004). Greater treatment duration (82.6 vs |
| | buprenorphine | care. | physician visits). | toxicology). | 68.6 days) in CBT4CBT group. |
| | maintenance. | | | Secondary: Abstinence | |
| | | | | from all drugs. | |
| Sigmon et | Two RCTs, each | Technology-Assisted | Standard | Primary: Illicit opioid | TAB yielded much higher abstinence: ~85% |
| al., 2023 | N=50 adults | Buprenorphine (TAB) - 24- | buprenorphine | abstinence (urine- | (nonrural) and 88% (rural) opioid-negative rates |
| (``TAB″ trials- | with untreated | week program with remote | treatment (standard | confirmed). | vs 21-24% in respective control groups (p<0.001) |
| USA) [9] | OUD in nonrural | buprenorphine initiation, | duration or without | Secondary: Treatment | Treatment retention also improved with TAB |
| | vs rural settings | telehealth counselling, digital | tech-assisted | retention. | (longer duration on buprenorphine) – |
| | | adherence monitoring, plus | features, depending | | demonstrating efficacy in both rural and nonrural |
| | | overdose education (in one trial) | on trial arm). | | patients. |
| Tofighi et al., | N=128 adults | "TeMeS" Text-Messaging | Treatment-as-usual | Primary: Feasibility | High enrolment (91% of eligible) and |
| 2023 [10] | with OUD | Intervention – Automated daily | tele-buprenorphine | (enrolment, | acceptability; no significant difference in 8-week |
| (USA) | inducted on | texts based on medical | care (virtual clinic | engagement) and | retention (mean \sim 5.2 vs 5.0 weeks retained, p = |
| | buprenorphine | management model | visits, no texting | Acceptability. | 0.68). Participants were generally satisfied with |
| | via telemedicine | (appointment reminders, | support). | Secondary: Treatment | text frequency, though 9% opted out due to |
| | (COVID-era | adherence prompts, support | | retention at 8 weeks | message fatigue. |
| | virtual clinic). | messages) delivered for 8 weeks. | | (active Rx in week 8) | |

Table 1: Comparative Summary of RCTs Evaluating Digital Interventions for OUD. (n=10)

| Kiburi et al., | N=46 adults | CBT-Based SMS Support - 6- | Standard | Primary: Reduction in | Opioid use decreased in both groups; intervention |
|--------------------|-------------------|------------------------------------|-----------------------|-------------------------|---|
| 2023 (Kenya | with OUD on | week text messaging program | methadone | opioid use (self-report | arm had lower opioid use prevalence (35.7% vs |
| SMS)[11] | methadone | (daily cognitive-behavioural | treatment (daily | and urine tests). | 56.3% at 3 months), but difference was not |
| | maintenance | therapy tips, skill reminders, and | dosing + | Secondary: Treatment | statistically significant. High retention in SMS |
| | (Nairobi clinic). | support texts). | counselling as | retention at 6 and 12 | group (93% at 6 weeks, 83% at 3 months) and |
| | | | usual) with no | weeks; Acceptability | high satisfaction with the texts. |
| | | | messaging. | of intervention. | |
| Liang et al., | N=75 adults in | "S-Health" Smartphone App – | Control group | Primary: Opioid/drug | After 1 month, urine-verified abstinence improved |
| 2018 | community | Bilingual app for CRA-based self- | received only | abstinence (weekly | with the app: 73.8% opioid-negative rate vs 50% |
| (S-Health | methadone | management:daily ecological | weekly health | urine tests). | in control (26.2% positive vs 50% positive; $p =$ |
| trial- China) | programs | momentary assessments | education text | Secondary: Self- | 0.06) |
| [12] | (Shanghai) with | (craving, triggers) + educational | messages (no | reported drug use | Mean days of drug use in past week were |
| | heroin or | messages; supervised by social | interactive self- | days; user | significantly lower in the intervention (0.71 vs |
| | polysubstance | workers | monitoring) | engagement. | 2.20, p < 0.05) |
| | use disorder | | | | Users preferred app-based reporting over in- |
| | | | | | person interviews. |
| King et al., | N=67 opioid- | Web-based | Standard in-person | Primary: Counselling | Non-inferior outcomes: Videoconference group |
| 2014 (Web- | dependent | Videoconferencing | individual | attendance; Opioid | attended a slightly higher % of sessions and had |
| Counselling- | outpatients in | Counselling – Patients received | counselling at the | use (urine tests). | comparable weekly opioid-negative urine rates to |
| USA)[13] | methadone | one-on-one counseling sessions | clinic (same | Secondary: | in-person counseling (approx. 89–91% negative |
| | maintenance | via live video ("eGetgoing" | frequency and | Therapeutic alliance; | weeks in both; no significant difference). |
| | program (urban | platform) instead of in-person. | content of | Patient satisfaction. | Treatment satisfaction and therapeutic alliance |
| | clinic) | | sessions). | | were similarly high in both groups. Patients valued |
| | | | | | the convenience of remote counselling. |
| Marsch et | N=160 opioid- | Therapeutic Education | Standard | Primary: Opioid | During the 12-week intervention, the TES group |
| al., 2014 | dependent | System (TES) – Web-based | methadone program | abstinence (urine- | achieved higher abstinence rates than standard |
| (Campbell et | adults in | CRA + voucher incentives | with full schedule of | confirmed) during | care (greater proportion of opioid/cocaine- |
| al. 2014- | methadone | program used as a partial | clinician-delivered | treatment; Treatment | negative weeks) and similar or slightly better |
| USA)[14] | maintenance | replacement for standard | counseling + urine | retention. Secondary: | retention. |
| | (multi-site | counseling (patients completed | monitoring + | Abstinence at follow- | However, by 3- and 6-month post-treatment |
| | clinics) | interactive CBT modules in lieu of | vouchers (per clinic | ups (3 and 6 months). | follow-ups, abstinence outcomes between groups |
| | | some clinic sessions). | protocol). | | were not significantly different (no sustained |
| | | | | | benefit). |
| | | | | | Both interventions produced substantial |
| | | | | | abstinence during treatment |

*CRA - Community Reinforcement Approach, +CM - Contingency Management, +TAU - Treatment as Usual, §OUD - Opioid Use Disorder, ||CBT - Cognitive Behavioural Therapy, ¶MOUD - Medication for OUD, **B/N -Buprenorphine/Naloxone.

Risk of bias (RoB) for included RCTs were assessed using the Cochrane Risk of Bias Tool [Figure 2]



Figure 2: Risk of bias (RoB) for included RCTs assessed using the Cochrane Risk of Bias Tool.

| Author(s) | # of Included | Types of Digital | Key Findings / Conclusions |
|--------------|--------------------------|-----------------------------------|---|
| & Year | Studies and Region | Interventions | |
| Kiburi et | 20 RCTs [Global | Web-based programs, computer- | Mixed effectiveness: about half of trials showed significant improvement in opioid abstinence |
| al., 2023 | (mostly USA | based modules, telephone calls, | and a few in treatment retention. Digital interventions were generally acceptable with high |
| [15] | studies)] | video conferencing, automated | patient satisfaction. |
| | | self-management, mobile apps, | Effectiveness varied by intervention and patient factors; overall, digital tools can complement |
| | | text messaging (often based on | OUD treatment but more research is needed, especially in LMIC settings. |
| | | CBT, CRA, MI, etc.) | |
| Aronowitz | 40 (mixed-methods | Telehealth for buprenorphine | Patients and providers viewed tele-buprenorphine favourably, citing improved access and |
| et al., | studies) [USA & | ("tele-bupe"), including | convenience. Most supported continued telehealth use post-pandemic. |
| 2024 [16] | Canada (COVID-era | video/phone inductions and | Some challenges were noted (tech issues; providers worried about rapport, while patients |
| | telehealth)] | counselling | felt more comfortable at home). Overall, experiences suggest tele-bupe is acceptable and |
| | | | can improve retention, though providers are divided on when it's most appropriate. |
| Ward et | 31 (scoping review) | Various digital health | Digital tools are being used to support women with OUD in areas like enhancing access to |
| al., 2024 | [Global (focus on | interventions for women (mobile | care and recovery support. Interventions addressed unique needs (e.g. pregnancy, childcare) |
| [17] | women with OUD)] | apps, tele-counselling, text | but research is limited. The scoping review found a need for more gender-tailored digital |
| | | messaging, web-based support) | treatments and better evaluation of outcomes in women. |
| | | | Overall, digital health shows promise for engaging women with OUD, but evidence is still |
| | | | emerging. |
| Lyzwinski | 20 (scoping review of | mHealth and wearables (SMS | OUD patients have high willingness to engage with mHealth tools to manage their opioid use. |
| et al., | qualitative studies) | text messaging, smartphone | Users see mobile apps, text support, and wearables as opportunities to access care and |
| 2024 [18] | [Global (user | apps, wearable overdose | prevent overdoses. They prefer personalized content, encouragement, and involvement of |
| | perspective focus)] | sensors) | trusted professionals. Key barriers include privacy concerns and limited technology access. |
| | | | Authors emphasize incorporating user feedback (privacy safeguards, training, tailored |
| | | | messaging) to maximize benefits. |
| Lin et al., | 25 (systematic | Telemedicine-delivered SUD | Telemedicine interventions for SUD (including OUD) showed comparable outcomes to in- |
| 2019 [19] | review) | treatment (videoconferencing or | person treatment in retention and substance use, with high patient satisfaction. |
| | [USA (Veterans and | phone-based counseling and | Review noted that tele-SUD treatments often achieved similar abstinence rates and no |
| | general SUD)] | MAT for OUD and other SUD) | increase in adverse events. Authors conclude that telemedicine is a feasible, effective |
| | | | alternative for delivering OUD therapy, though more studies were encouraged to confirm |
| | | | long-term outcomes. |
| Tice et al., | , 3 digital therapeutics | Prescription digital therapeutics | Evidence was still limited for FDA-authorized digital therapeutics in OUD. For the reSET-O |
| 2021 [20] | (evaluated via prior | as adjuncts to MAT (reSET-O | CBT+CM app, an RCT showed improved 12-week abstinence and retention vs TAU, but long- |
| | studies) [USA (ICER | app), recovery support apps | term benefits remain uncertain. |
| | report, various | ("Connections"), and reward- | Two small uncontrolled studies suggested potential positive outcomes, but due to bias/no |
| | regions of included | based apps (DynamiCare) | control, confidence in effectiveness is low. |
| | studies)] | | The ICER panel found no clear net health benefit yet for reSET-O or similar apps compared to |
| | | | standard care. |
| | | | Cost-effectiveness modeling for reSET-O was favorable (within US willingness-to-pay |
| | | | thresholds) if short-term gains are maintained. |
| | | | Overall, digital therapeutics are promising but require more robust evidence. |

Table 2: Comparative Summary of Systematic Reviews / Meta-Analyses. (n = 6)





Figure 3: AMSTAR 2 checklist

*CBT- Cognitive Behavioural Therapy, + CRA- Community reinforcement approach, + MI-Motivational Interviewing, §RCT- Randomised control trial, ||OUD- Opioid Use disorder, ¶LMIC- Low and middle income countries, **mHealth- Mobile Health, ++SMS- Short message service, ++SUD- Substance use disorder, §§MAT-Medication-Assisted Treatment, ||||FDA- Food and drug Administration, ¶¶TAU- Treatment as usual, ***ICER-Institute for Clinical and Economic Review, +++CM – Contingency Management.

Systematic reviews were evaluated using the AMSTAR 2 checklist [Figure 3].

Table 3: Comparative Summary of Observational / Real-World Studies (n = 12)

| Author(s) | Sample, Population | Digital Intervention | Main Outcomes | Key Findings / Effect Size |
|--------------|--------------------------|----------------------------|------------------------------|--|
| & Year | and region | | | |
| Miller- | 276 health organizations | Various patient-facing | Adoption of digital tech | 33.5% of organizations used ≥1 digital OUD technology (most |
| Rosales et | (ACOs) – respondents | digital tools for OUD care | for OUD in organizations; | commonly remote therapy/tracking at 26.5%). |
| al., 2023 | from hospitals, clinics, | (categories: remote | usage rate of at least one | Use of digital tools was seen as a complement to existing treatment |
| [21] | and group practices. | therapy/tracking, virtual | digital intervention | capacity, not a replacement. |
| | [USA (national survey of | peer support, digital CBT | | Organizations with addiction specialists or mental health registries |
| | health orgs)] | adjuncts) | | were significantly more likely to adopt digital tools (e.g. +32% |
| | | | | adoption with specialist) |
| Eibl et al., | 3,733 OAT patients | Telemedicine-delivered | 1-year treatment | Higher retention with telemedicine: 50% 1-year retention via |
| 2017 [22] | across 58 clinics in | OAT (methadone or | retention on OAT | telemedicine vs 39% in-person (mixed tele/in-person: 47%). |
| | Ontario (2011–2012) | buprenorphine) vs in- | | Tele-OAT patients had 27% higher odds of continuous 1-year |
| | [Canada] | person OAT (and mixed | | retention than in-person (aOR \approx 1.27, 95% CI 1.14–1.41). |
| | | modality) | | Telemedicine proved an effective alternative for OAT without |
| | | | | compromising outcomes. |
| Hammersl | ~91,000 adults receiving | Telemedicine initiation of | 90-day retention in | Initiating OUD care via telemedicine was associated with better 90- |
| ag et al., | buprenorphine in 2020; | buprenorphine treatment | buprenorphine | day retention. In Kentucky, 90-day retention was higher with tele- |
| 2023 [23] | ~43,000 were new OUD | (during COVID-19 | treatment; opioid-related | initiation (aOR 1.13, 95% CI 1.01–1.27); in Ohio, aOR 1.19 (95% CI |
| | treatment initiations. | emergency) vs traditional | overdose within 90 days | 1.06–1.32). |
| | [USA (Kentucky & Ohio | in-person initiation | | No significant difference in 90-day nonfatal overdose rates between |
| | Medicaid)] | | | telemedicine and in-person initiation (aOR \sim 1.0, n.s.). |
| | | | | Telehealth start did not increase overdose risk and modestly |
| | | | | improved early retention in MOUD. |
| Lira et al., | 1,816 rural OUD patients | Telehealth-only MOUD | Treatment retention at 1, | Tele-OUD treatment in rural populations achieved encouraging |
| 2023 [24] | inducted via a | program (remote | 3, 6 months; Medication | outcomes. |
| | telemedicine | buprenorphine induction | adherence (urine- | Retention: 74.8% at 1month, 61.5% at 3months, 52.3% at 6months. |
| | buprenorphine program | and follow-up) | verified) at those | comparable to outcomes in traditional clinics. |
| | (2020–2022) | | intervals | Adherence: 69.0% (1mo), 56.0% (3mo), 49.2% (6mo) tested |
| | [USA (14 states, rural | | | negative for illicit opioids (on buprenorphine). |
| | areas)] | | | Authors conclude telemedicine is an effective approach for rural OUD |
| | | | | care, with retention rates on par with in-person treatment. |
| Marino et | 600 adults with OUD on | Smartphone app-based | Days of opioid use at end | Augmenting buprenorphine treatment with a recovery app (CM |
| al., 2024 | MOUD; 300 self-selected | Contingency Management | of treatment; treatment | rewards for abstinence) was associated with improved outcomes. The |
| [25] | to add an app-based CM, | (CM) added to standard | retention (program | app-users reported fewer days of opioid use at treatment end and |
| | 300 on MOUD alone. | MOUD vs MOUD alone | duration) | longer treatment retention than those on MOUD alone. |
| | [USA (Texas clinical | (observational cohort) | | In this real-world cohort, patients who opted into the app stayed in |
| | network)] | | | treatment significantly longer (on average) and were more likely to |
| | | | | complete the program. (Effect sizes: app group had higher |
| | | | | abstinence and retention. |
| Velez et | 901 OUD patients who | reSET-O prescription | Healthcare utilization | Adding the reSET-O digital therapeutic was linked to significant |
| al., 2022 | initiated the reSET-O | digital therapeutic (12- | over 12months post- | reductions in healthcare utilization. Over 1year, the reSET-O group |
| [26] | therapeutic app (adjunct | week CBT + Contingency | initiation (inpatient stays, | had 28% fewer inpatient stays (IRR 0.72, $p=0.02$) and 56% fewer |
| | to buprenorphine); | Management app) as | ED visits, readmissions); | 30-day readmissions than controls. |
| | matched with 978 | adjunct to MOUD, | Buprenorphine adherence | Total ED visit rates trended 7% lower (n.s.). |
| | controls (buprenorphine | compared to MOUD alone | (medication possession | Net annual healthcare costs were ~\$2,800 lower per patient in the |
| | MAT only) | (no-app) | ratio) | reSET-O group. Buprenorphine adherence was higher with the app |
| | USA (claims data across | | | (MPR 0.85 vs 0.76, p<0.001) |
| | states)] | | | Conclusion: reSET-O use is associated with durable real-world |
| | | | | benefits – fewer hospitalizations and better medication adherence. |

| Ganesh et | 150 patients with OUD | Mobile health (mHealth) | Access to | High digital access and enthusiasm were observed among patients. |
|------------|----------------------------|---------------------------|---------------------------|---|
| al., 2022 | on opioid agonist therapy | readiness and interest – | mobile/internet; | 88% owned a mobile phone; 70% had internet access. |
| [27] | (methadone/buprenorphi | survey of phone | willingness to use SMS or | 80% expressed interest in receiving OUD-related text message |
| | ne) at a community clinic | ownership, internet use, | apps for OUD care | support, and 60% were willing to use a smartphone app for |
| | (New Delhi) | and willingness to use | | monitoring substance use. |
| | [India] | digital tools for OUD | | This indicates strong patient readiness for mHealth interventions in |
| | | | | an Indian OUD treatment setting. |
| Kiburi et | 46 patients on | SMS text-message | Opioid use prevalence | Reduced opioid use with texting (not statistically significant due to |
| al., 2023 | methadone for OUD | intervention (6 weeks of | (urine test) at 6 weeks; | small N): at 6 weeks, opioid-positive urine prevalence was 35.7% in |
| [28] | (Nairobi clinic) – | CBT-based daily texts) | Methadone treatment | the SMS group vs 56.3% in control. |
| | feasibility RCT: 30 | added to standard | retention at 6 weeks and | Retention on methadone in the SMS group was high (93% at 6 |
| | received SMS-based CBT | methadone treatment (vs | 3 months; acceptability | weeks; 83% at 3 months). |
| | messages, 16 control. | standard care only) | ratings | Participants reported the text-CBT program was highly acceptable |
| | [Kenya] | | | and useful (with improved coping skills). This pilot suggests texting |
| | | | | CBT is feasible and promising for improving OUD outcomes in Kenya, |
| | | | | warranting a larger trial. |
| Xu et al., | 40 individuals with opioid | CARE app* – Community- | Urine-test confirmed | The digital intervention group achieved better abstinence outcomes. |
| 2021[29] | use disorder in | based Addiction | abstinence over 6 months | Over 6 months, only 3.3% of urine samples in the CARE app group |
| | community compulsory | Rehabilitation E-system | (proportion of opioid- | were opioid-positive, vs 7.5% in the control group – a significant |
| | treatment (pilot RCT: 20 | (mobile app for self- | negative tests); other | difference in favour of the app ($p=0.04$). |
| | with app + standard | monitoring, e-learning, | measures (longest | Longest continuous abstinence did not differ significantly between |
| | rehab, 20 standard rehab | mood tracking) plus | abstinence, psych | groups. Participants and supervising social workers engaged well with |
| | only). | routine community rehab | assessments) | the app's features (education, assessments, GPS tracking). The study |
| | [China (Shanghai)] | supervision | | demonstrates improved relapse rates with the smartphone-based |
| | | | | support in a real-world Chinese setting. |
| Le et al., | 450 patients on | SMS reminders for | Methadone dose | Both interventions significantly improved adherence. At 6 months, |
| 2025 [30] | methadone maintenance | methadone doses (TMR) | adherence, measured at | the MI group's rate of complete adherence was 36% higher than |
| | (3 urban clinics) – | and/or Motivational | 3 and 6 months | control (RR 1.36). |
| | randomized to: Control | Interviewing counseling | (complete adherence = | The Text Reminder group also had higher complete adherence than |
| | (methadone only), Text | (MI) to improve | no missed doses; | control at 3 months (RR 1.27) and 6 months (RR 1.28). |
| | Message Reminders | adherence, compared to | weekend adherence) | Notably, weekend dose adherence (historically low) improved in the |
| | (TMR), or Motivational | standard MMT alone | | SMS group (RR 1.19 vs control at 6 mo). |
| | Interviewing (MI) | | | Conclusion: Weekly counseling and automated daily text reminders |
| | sessions. [Vietnam] | | | each led to significantly better methadone treatment |
| | | | | retention/adherence in this real-world setting. |
| Thomas | 128 adults initiating | "TeMeS" text-message | 8-week retention in | High feasibility but no short-term retention gain in this pilot. Almost |
| (Tofighi) | buprenorphine via a low- | support – daily automated | buprenorphine treatment | all eligible patients agreed to receive texts (91% enrolment), and |
| et al., | threshold tele- | med-management texts | (measured by having an | 88% engaged with the messaging. Retention at 8 weeks was similar |
| 2023 | buprenorphine program | (appointment reminders, | active Rx at week 8); | between groups (~5.2 weeks on treatment with texting vs 5.0 weeks |
| | (2020) – randomized to | motivational messages, | patient engagement & | control; p=0.676). |
| | automated texting | symptom check-ins) for 8 | satisfaction | No safety issues were noted. Participants were generally satisfied |
| | ware and (TaMaC) wa | weeks added to tele-BUR | | with the frequency and content. This suggests that while automated |
| 1 | support (Temes) vs | weeks, added to tele-DOF | | |
| | treatment-as-usual. | care vs tele-BUP care | | texts are acceptable, a more intensive or tailored approach may be |



Figure 4: Observational studies evaluated using the Newcastle-Ottawa Scale (NOS).

*CRA – Community Reinforcement Approach, +MI – Motivational Interviewing, ‡CBT – Cognitive Behavioural Therapy, §CM – Contingency Management, ||MOUD – Medications for OUD, ¶ACO – Accountable Care Organization , **MAT – Medication-Assisted Treatment (here, MOUD), ++MPR- Medication Possession Ratio, ‡ ‡ IRR- Incidence Rate Ratio, §§ED- Emergency department, ||||TMR-Text message reminder, ,¶¶MMT-Methadone Maintenance Therapy, ***LMIC- Low and middle income country, +++OAT-Opioid antagonist therapy, ‡#‡aOR- Adjusted Odds Ratio, §§§GPS- Global positioning system, |||||RR-Relative risk.

Observational studies were evaluated using the Newcastle-Ottawa Scale (NOS) [Figure 4].

Table 4: Summary of key outcomes across study types in the systematic review

| Outcome | RCTs (n=10) – Key Findings | Systematic Reviews/Meta-analyses | Observational Studies (n=12) - Key Findings |
|---------------------------|---|--|--|
| | | (n=6) | |
| Abstinence (opioid use) | ~80% of RCTs showed increased | Convergent evidence of modest | Consistently high abstinence rates reported with digital |
| | abstinence with a digital | improvements in abstinence. One | use. In a 3,144-patient dataset, 66% were abstinent at |
| | intervention vs control. E.g., one | review found 50% of trials had | end of 12 weeks (missing=use), and 91% abstinent |
| | trial reported 77% vs 62% abstinent | significant benefit (Kiburi et al., 2023). | when counting only those providing data. Extended 24- |
| | at 12 weeks (digital vs TAU). [31] | Others report small-to-moderate effect | week digital treatment yielded 86% abstinence |
| | Another found digital CBT users had | sizes favouring digital. No review found | (missing=use) (Maricich et al., 2021). |
| | 9.7 more abstinent days | worse outcomes with digital. | All observational studies noted reductions in self- |
| | (Christensen et al., 2014). Some | | reported opioid use. |
| | RCTs (20%) found no difference. | | |
| Relapse/Continuous | Digital arms often delayed relapse. | Not a focus of quantitative meta- | Real-world data show sustained engagement can keep |
| Abstinence | E.g., longest abstinence streaks | analysis, but narrative syntheses note | patients opioid-free longer. No overdose events were |
| | were longer in digital groups (by | fewer relapse events when digital tools | reported during digital treatment in case series. |
| | \sim 2–3 weeks in some trials). Time to | are effective. | |
| | first opioid use was prolonged in | | |
| | several studies. | | |
| Retention in Treatment | 4 of 10 RCTs showed significantly | Reviews note mixed retention outcomes | High retention observed in practice: ~74% of patients |
| | improved end-of-treatment | (only ~20% of trials positive (Kiburi et | completed 12-week digital treatment (Maricich et al., |
| | retention with digital adjuncts | al., 2023). | 2021). |
| | (Kiburi et al., 2023). | Overall, digital interventions appear | Among those who continued to a second 12-week |
| | Example: 80% retained with digital | retention-neutral to mildly beneficial. | course, >91% were still in treatment at 24 weeks |
| | vs 64% with standard care (HR | Long-term retention effect unclear due | (Maricich et al., 2021). |
| | ~0.5 for dropout). [32] | to short follow-ups. | An app-based contingency management cohort had |
| | Other trials saw no drop-off | | longer treatment duration vs non-app users (Marino et |
| | attributable to digital use. | | al., 2024). |
| Acceptability & | Generally high – most RCTs | Universally reported as high. Patients | Very high – surveys show 85–100% of users satisfied. |
| Satisfaction | reported favourable patient | find digital modalities acceptable and | In one pilot, >90% rated the digital program |
| | feedback (e.g., high usability scores, | would recommend them (Kiburi et al., | "good/excellent" and would reuse/recommend (Monico, |
| | few complaints). Engagement levels | 2023). | et al., 2024). |
| | in trials (sessions completed) | No major concerns in reviews aside | High uptake in real-world programs also reflects |
| | indicate good acceptance. | from need for tailoring. | acceptability. |
| Feasibility & Scalability | Proven feasible in controlled settings | Highlight broad applicability: | Demonstrated at scale: thousands treated with digital |
| | high completion of digital sessions | interventions via web, phone, text, | tools in routine care (Maricich et al., 2021). Programs |
| | in most trials. Some RCTs delivered | apps all feasible. Emphasize need to | rolled out state wide show scalability. Technology |
| | interventions fully online with | expand to new settings (e.g., LMICs) | infrastructure and training are required but manageable |
| | success. | for scalability. | as shown in pilot implementations. |
| Barriers/Limitations | Some patients disengage early (tech | Heterogeneity of interventions | Digital divide concerns (access to devices/internet) in |
| | not a fit for all). Short trial durations | complicates pooling data. Mostly US- | broader population. In studies, support was provided - |
| | limit insight on long-term effects. | based studies – results may not | real-world users without support may face access |
| | Some RCTs had small N. Selection | generalize globally (Kiburi et al., 2023). | issues. Sustainability and payer coverage remain |
| | bias: participants often motivated | Lack of data on certain groups (older | challenges; current real-world studies rely on grant or |
| | treatment-seekers. | adults, low-resource settings). | pilot funding. |

*TAU = treatment-as-usual, +HR = Hazard ratio

Discussion

Interpretation of Main Findings:

In this systematic review, investigators discovered that digital treatments for OUD show potential in improving clinical outcomes, however the results varied between research. Approximately half of the randomised trials analysed indicated significantly higher opioid abstinence rates in individuals receiving a digital intervention compared to control groups (Kiburi et al., 2023). However, increases in treatment retention were less common; only a small number of studies found significantly improved retention with digital therapies. This implies that, while technology-enhanced therapies can help many patients achieve short-term abstinence, sustaining long-term commitment in care remains difficult. these Importantly, participants in research consistently rated digital interventions as acceptable, with high satisfaction rates (Kiburi et al., 2023). High user acceptance demonstrates that individuals with OUD are willing to interact with digital modalities of care, which is a necessary condition for any intervention to have a real-world impact. The digital tools studied ranged from webbased therapy programs to computer or smartphone applications, SMS text message support, phone/video tele-counselling, and automated selfmanagement systems (Kiburi et al., 2023). These platforms provided evidence-based therapeutic cognitive-behavioural content (e.g., therapy, contingency management, communityreinforcement techniques, motivational interviewing) via digital means (Kiburi et al., 2023). Given this variability, it is not surprising that outcomes differed from study to study; factors such as the intensity of the digital program, patient participation levels, and whether the digital therapy was used in addition to or instead of traditional care are likely to have influenced its effectiveness. For example, an interactive mobile app with substantial cognitive-behavioural modules may result in larger opioid reductions than a basic text-messaging reminder system. Despite these differences, a recurring theme was the high feasibility of implementing interventions using digital platforms, as well as positive user feedback (Miller-Rosales et al., 2023). In conclusion, findings show that digital treatments can effectively deliver behavioural treatment for OUD in a way that patients find acceptable - though extent of improvement varies.

Comparison of Previous Literature and Conventional Treatment:

Overall, the findings are consistent with and extend previous studies on digital health interventions for OUD. Prior studies recognised that digital platforms had the potential to increase access to therapy, but there was a lack of long-term outcome data (Tice et al., 2021). For example, an ICER evidence review in 2020 found that at the time, no definitive trials had shown long-term retention benefits from OUD digital apps, and no long-term outcomes like as employment or overdose reduction had been recorded (Tice et al., 2021). This review builds on that foundation by incorporating more recent clinical trials, some of which demonstrate meaningful shortterm improvements in outcomes when digital therapeutics are added to standard treatment. Notably, one randomised the trial found that a 12week prescription digital therapy combination with buprenorphine significantly boosted opioid abstinence (77% versus 62%) and reduced dropout rates by half as compared to treatment as usual (Maricich et al., 2021). This type of evidence was lacking in previous literature, but it now confirms that, under the right conditions, digital treatments can improve recovery measures. When compared to traditional OUD therapies, digital therapeutics appear to serve a complementing function. Medication-assisted treatment (MAT), which combines opioid agonist or antagonist therapy with counselling, is still the gold standard for OUD and is extremely effective in lowering opioid usage and overdose risk. However, more than half of patients initiating MAT discontinue treatment within 3-6 months (Tice et al., 2021), often due to hurdles in counselling accessing ongoing or support. Traditional behavioural therapy can improve patient retention, but it is resource-intensive and not always available. In this regard, digital treatments are best understood as a novel approach to delivering the psychosocial component of OUD treatment. Rather than replacing established treatments, they mimic and expand upon them. In trials where digital therapy was used as an addition to MAT, patients frequently performed better than those on MAT alone (Maricich et al., 2021), which is similar to the advantage of adding any form of counselling to MAT. In contrast, previous studies in which a digital program was evaluated in place of some face-to-face counselling found that outcomes were nearly similar to standard care,

Demonstrating that a well-designed digital tool can match the efficacy of in-person therapy in the short run. This is illustrated by the FDA's approval of the reSET-O treatment, which was based on data that patients using the digital program achieved abstinence rates and retention comparable to those getting clinician-delivered therapy, as well as improved involvement in some metrics (Tice et al., 2021). The results obtained are also consistent with the broader substance use disorder literature, as digital therapeutics have been shown to be effective not only for OUD but also for other addictions such as nicotine dependence, indicating that technologybased delivery of behavioural treatment can consistently produce positive outcomes across different substance use contexts.[33] In conclusion, digital OUD interventions are most effective when combined with traditional treatment, employing technology to augment proven procedures and broaden their reach rather than completely replacing standard care.

Clinical and policy implications:

These tools were found to be most effective when used in conjunction with established treatment, such as buprenorphine or methadone, as well as routine counselling. This is consistent with current treatment guidelines, which emphasize а combination of medication and psychosocial assistance for OUD. Digital treatments can help patients in between clinic sessions by providing CBTbased apps for coping skills and motivation, perhaps lowering recurrence. Their great acceptability shows that even those who are hesitant to seek counselling may use digital technologies. However, physicians should monitor usage because effectiveness consistent is dependent on participation. On a broader level, DTx can assist structural hurdles in OUD overcome care, particularly for patients in rural or impoverished locations with limited access to experts (Miller-Rosales et al., 2023). During the COVID-19 pandemic, there was an extraordinary example of this: the rapid expansion of telemedicine allowed patients to continue OUD treatment remotely, resulting in outcomes comparable to traditional inperson care (such as buprenorphine retention programs).[34] The success of remote OUD care demonstrates that many treatment features can be efficiently administered from a distance. Digital solutions, such as applications and online platforms, can help to broaden access by providing therapy,

Education, and support via smartphone. This is especially useful for patients in remote places, those with little time or childcare, and those who face stigma. Health systems should think about incorporating digital options to extend reach and lower barriers. Realizing these benefits will necessitate supportive policies and cautious implementation. Notably, current adoption of OUD digital health tools by treatment providers remains limited - according to a recent survey of U.S. healthcare organizations, only about 34% had deployed any digital technologies for OUD care, and those that did typically treated these tools as complements to their existing offerings (Miller-Rosales et al., 2023). This suggests that without incentives or guidance, many clinics (particularly those with limited resources) may be hesitant to adopt digital therapies. Policymakers should explore creating frameworks to support the adoption of evidence-based digital solutions. Equity is vital while applying digital treatments for OUD. Without care, these tools may mainly assist tech-savvy, wellconnected patients, resulting in wider disparities. To avoid this, rollouts should include support such as providing devices, offering low-bandwidth (SMS) options, and ensuring cultural and language diversity. Successfully integrating DTx into mainstream care will necessitate integrated policies, infrastructure, and equity-focused planning, ultimately enhancing access, retention, and results for various demographics.

Strengths and limitations:

This review provides a comprehensive overview of digital therapies (DTx) for opioid use disorder, including various tools, outcomes, and study designs. Key strengths include the utilization of various intervention modalities (e.g., applications, telehealth, web platforms), a focus on patientcentered goals such as usability and satisfaction, and a rigorous, systematic methodology. However, there are numerous constraints to consider. The majority of included research were conducted in countries with high incomes, primarily the United States, which limits worldwide generalizability. Diverse populations, including those in low- and middle-income households, rural communities, teenagers, and those involved in the criminal justice system, continue to be under-represented. The majority of studies only looked at short-term outcomes, so it's uncertain whether these benefits persist.

Because of the rapid advancement of digital health tools, this review may have missed out on newer or updated interventions. Finally, significant publication bias towards study with positive outcomes could have influenced the overall results. Future research should fill these gaps by examining long-term clinical and quality-of-life outcomes, enhancing engagement strategies, comparing intervention formats, testing future technologies such as AI or VR, and determining cost-effectiveness to inform long-term implementation.

Conclusion

In conclusion, digital therapies are a new addition to OUD therapy resources that has shown promising outcomes in enhancing abstinence and expanding care to more individuals. This review's findings validate their potential while emphasizing need for further exploration. Continued study, particularly in long-term and diverse situations, will assist identify how to best use these devices. With careful integration into health systems and supportive policies, digital therapeutics have potential to significantly improve management of opioid use disorder, providing scalable, accessible support in addition to medication treatment and, ultimately, improving outcomes for those suffering from OUD.

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