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The emergence of digital mental health in low-income and middle-income countries: A review of recent advances and implications for the treatment and prevention of mental disorders

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Abstract

In low-income and middle-income countries (LMICs), emerging digital mental health interventions should be accompanied by regular and comprehensive assessment of available scientific evidence. This review aims to support efforts to monitor progress in digital mental health research, ensuring new evidence can guide researchers, clinicians, policymakers and program managers positioned to adopt and implement these digitally-enabled treatments. In accordance with PRISMA guidelines, an electronic database search from 2016 to 2020 yielded 37 digital intervention studies for detection, diagnosis, prevention, treatment, and/or management of a broad range of mental disorders in 13 LMICs. This date range was selected to update previous reviews. Most studies involved online interventions and many reported feasibility and acceptability, reflected by participant satisfaction or program adherence. About half the studies

Conflict of Interest

The authors report no conflict of interest.

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Author Statement

HC designed the study and wrote the search strategy and protocol. RA and JAN reviewed the search strategy and study protocol. HC and KA completed the literature search and screened records for relevant articles for inclusion. HC compiled the final list of included studies. KA and JAN reviewed the list of included studies to confirm eligibility. HC and KA extracted the data from the included studies. RA, JAN, and DD reviewed the data tables to confirm accuracy of data extraction. DD conducted the methodological quality assessment of the included studies. HC wrote the first draft of the manuscript. RA and JAN assisted with the interpretation of the findings and contributed to multiple revisions of the manuscript. KA and DD made revisions to subsequent versions of the manuscript. All authors reviewed the final list of included studies, and contributed to and approved the final manuscript for submission.

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(N=23) reported clinical benefits based on changes in mental health. For depression and mood disorders, some digital interventions showed improvements in depressive symptoms, quality of life, treatment adherence, and recovery. However, sample sizes were small and studies focused primarily on adults. Further limiting generalizability was the lack of consistency in clinical assessment and measurement tools between studies. No studies reported worsening symptoms, negative acceptability or dissatisfaction with digital interventions, suggesting possible publication bias. While digital interventions show promise, it remains difficult to conclude that digital interventions are effective from these studies, as it is prudent to exercise caution before drawing conclusions about clinical effectiveness. This review reflects continued growth in digital mental health research in LMICs and further highlights the need for rigorous evaluation of effectiveness and cost-effectiveness.

Keywords

mental health; low- and middle-income country; digital technology; digital psychiatry; global health

1. Introduction

Mental disorders are a leading contributor to the global burden of disease, with mental illness accounting for 32.4% of years lived with disability and 13% of disability-adjusted life-years (Vigo et al., 2016). In low-income and middle-income countries (LMICs), there is a significant care gap (Pathare et al., 2018), with up to 90% of people with mental disorders not receiving adequate treatment and follow up (Becker and Kleinman, 2012; Patel et al., 2010; Patel and Prince, 2010). Globally, technology continues to emerge as a promising tool for offering direct support to individuals living with mental disorders, improving quality of services, and making evidence-based treatments more widely available (Bhugra et al., 2017). Digital health technologies typically encompass any form of remote or online platform or mobile device that can be used or designed to deliver a mental health intervention. This includes smartphone apps, online programs, text messaging, telepsychiatry, and wearable devices such as smart watches (Lipschitz et al., 2019). Effective digital health innovations present an opportunity to address the global discrepancy in mental healthcare provision.

In LMICs, a lack of awareness and adequate mental health services, social welfare and legislation to protect those with mental illness from stigma and discrimination are barriers to accessing services and promoting recovery from mental illness (Knaak et al., 2017; Koschorke et al., 2014; Semrau et al., 2015). Moving towards digital and online services may present an opportunity to enable individuals to access adequate mental healthcare in communities where mental health services may not be otherwise available. In cases where stigma and discrimination are particularly severe, such that an individual may fear social isolation or other inhumane responses to their mental illness, such as restraints (Semrau et al., 2015), digital technologies could present safe and confidential ways for people to access mental health interventions.

In high-income countries, the application of technology in healthcare is increasing rapidly (Price et al., 2014), with emphasis on new digital approaches for supporting mental and

behavioural health care (Bhugra et al., 2017). Digital applications have demonstrated promise in reducing symptoms of mental disorders such as depression (Firth et al., 2017), psychotic disorders (Gire et al., 2017) and other severe mental illnesses (Naslund et al., 2015), and improving medication adherence (Rootes-Murdy et al., 2018). These include smartphone delivery of cognitive behavioural therapy (Baumel et al., 2016), and apps for the early identification of relapse in psychotic disorders (Ben-Zeev et al., 2018). However, depending on digital mental health interventions that have been developed and studied in higher income countries, namely 'WEIRD' countries (i.e., Western, Educated, Industrialized, Rich, and Democratic), may bias conclusions and limit generalizability (Henrich et al., 2010). Therefore, to address this limitation, it is necessary to explore studies that have employed digital mental health interventions in LMICs.

Mounting research from lower-income countries suggests the potential benefit of digital mental health technologies in these settings (Fairburn and Patel, 2017; Naslund et al., 2019), as reflected in recent reviews. However, these reviews have also highlighted gaps in the evidence. For instance, a 2017 review found 49 studies that supported the feasibility, acceptability, and potential clinical effectiveness of online, mobile, and text-messaging services aimed at the treatment or prevention of mental disorders in over 20 LMICs (Naslund et al., 2017). It was found that online, text-messaging, and telephone support interventions showed promise in decreasing mental health symptoms. However, many included studies also did not include adequate comparison conditions, were not rigorously designed trials, or reported limited information on average effect sizes, the relative effectiveness of mental health interventions for different populations, and the variability in these effects across studies. For instance, fewer than half of the 49 studies summarized in the 2017 review reported significant improvements in mental health outcomes or symptoms. In general, insufficient details were reported in many studies to enable replication, with the review concluding that more research is required to determine clinical effectiveness of technological interventions and how to scale up and integrate use of technology into existing health systems (Naslund et al., 2017).

Furthermore, Kaonga and Morgan (2019) reviewed studies of digital tools including mobile and online modalities for mental health care, focused on improving psychological support and treatment adherence in resource-limited settings. This review did not provide comprehensive summaries of the included studies, which employed a wide range of designs, and were mostly pilot studies, without consistent reporting of outcomes. Importantly, of the 67 studies included, only 31 were from LMICs, making it difficult to determine whether the acceptability and potential clinical benefits of digital mental health programs from low-resource settings in higher income countries can generalize to LMICs (Kaonga and Morgan, 2019). Of the 31 studies from LMICs, only 4 reported significant improvements in mental health outcomes, highlighting a gap with reporting on effectiveness (Kaonga and Morgan, 2019).

Lastly, a review by Jimenez-Molina et al. (2019) identified 22 studies of Internet-based interventions focused on prevention and treatment of mental disorders including depression, anxiety, and substance use in Latin America. This review suggests that there were favorable short-term results in less than half of the interventions including reduction in stress, anxiety,

and depression. However, the studies mostly had short term follow-up and reported difficulty in participant retention and adherence to the online treatment protocols. Nine of the included studies reported improvements on mental health outcomes, with many studies unable to report effectiveness due to high attrition rates and methodological limitations including unclear sample groups (Jiménez-Molina et al., 2019). The limited reporting of clinical efficacy among these Internet interventions, combined with methodological challenges, makes it difficult to inform efforts to scale up these programs in real world settings.

These recent reviews of published studies highlight a mix of promising examples as well as important future directions for the field. Another important consideration is that there continues to be an unprecedented increase in the number of available commercial mental health technologies, which raises concerns that the emergence of new digital applications and devices has not been accompanied by similar progress in scientific evidence to support their use for addressing mental disorders (Naslund et al., 2019). For instance, over 10,000 mental health apps are available for download via commercial app stores (Torous et al., 2019); though, most of these commercially available apps have no reliable or high-quality scientific evidence as support (Larsen et al., 2019; Lau et al., 2020). In LMICs, this concern is especially relevant as growth of technology is even less likely to be accompanied by robust research evidence. This stems from general neglect and under-prioritisation of mental health on research agendas in many LMICs (Becker and Kleinman, 2013). Furthermore, governance and regulation of mental healthcare systems and policies in LMICs are low (Petersen et al., 2017), suggesting that regulatory bodies in place in many countries may be insufficient to match the unprecedented increase in the development of mental health technologies. Additionally, public access to commercial digital mental health applications is becoming increasingly easy given that wireless connectivity is becoming more widely available even in rural areas, and that smartphone prices continue to decrease (We Are Social, 2020).

To respond to this rapid evolution of the field of digital health technologies in global mental health, regular and comprehensive assessment of studies as they become available is required. This will also contribute further to the development of mental healthcare technologies that are safe, evidence-based and offer potential for the greatest benefit to those living with mental disorders in LMICs. In the current review, we aimed to address the need for regular monitoring of digital mental health research in low income settings by specifically expanding on these recent reviews that summarise use of digital mental health interventions in LMICs (Jiménez-Molina et al., 2019; Kaonga and Morgan, 2019; Naslund et al., 2017). We build on these studies not only by updating the review of published studies in this field, but also by providing a comprehensive synthesis and assessment of methodological quality of current research findings from LMICs.

A large number of new interventions studies have been published since the review by Naslund et al. (2017), so the current review accounts for the increasing use of technology and smartphone access in many LMICs. The current review also expands upon the review by Kaonga and Morgan (2019) by including studies of interventions applied specifically in LMICs (rather than low-resource settings in higher-income countries), and expands on the review by Jiménez-Molina et al. (2019) by including studies beyond Latin America.

Furthermore, to address a key gap in the prior reviews, we aim to provide a comprehensive summary of the potential clinical benefits of digital mental health interventions in LMICs. Our goal is to contribute to ensuring that new studies reporting clinical effectiveness are summarized in a way that can guide both researchers and clinicians, as well as policymakers and program managers responsible for the adoption and implementation of safe and effective new services into practice.

2. Methods

2.1 Search Strategy

This review adhered to the Preferred Reporting Items for Systematic and Meta-Analyses (PRISMA) reporting guidelines (Moher et al., 2009). A search was undertaken using Medline, with time limits placed between 2016/08/01 and 2019/07/16. The search was subsequently updated to include relevant studies from 2019/07/16 to 2020/08/28. Studies prior to 2016/08/01 were not considered in this search as these were previously summarised in the review by Naslund et al. (2017), the earliest of the three prior reviews. The search strategy was adapted from the search strategies described in prior reviews (Jiménez-Molina et al., 2019; Kaonga and Morgan, 2019; Naslund et al., 2017). The complete list of search terms are in Table 1.

2.2 Inclusion & Exclusion Criteria

Studies included in the review evaluated *technological interventions* for the *prevention* or *treatment* of *mental disorders*. '*Technological interventions*' involved digital, online or mobile technologies, including telephones (e.g., telepsychiatry), mobile phones (e.g., SMS texting services) and smartphones (e.g., smartphone applications), programs delivered via the Internet (e.g., websites for internet-delivered CBT), and/or wearable technologies. The main component of the interventions must be technology-based for studies to be included. '*Mental disorder'* encompassed any specifically defined (ICD or DSM) common or serious psychological disorders, as well as other more general psychological health terms such as 'stress', 'psychological distress', '(improving) mental health/ill-health' or 'suicide', as long as a clear definition of the concept and a measurement instrument were given. Interventions could target multiple disorders or symptom clusters in the same study, as long as they were all clearly defined. Mental disorders had to be the primary disorder for a paper to be included.

Outcomes included clinical effectiveness, feasibility, acceptability, efficacy, costs and/or cost-effectiveness, patient, carer or mental health-worker satisfaction, as well as other clinical or mental health related outcomes, such as quality of life, self-esteem, coping and functioning. While outcomes could be varied in this search, the primary outcome must be directly related to the patients themselves. For example, a study investigating a digital intervention targeting caregiver well-being as the primary outcome was excluded. However, if caregiver well-being was an outcome secondary to patient well-being, the study was included. Study designs were also varied, including randomised controlled trials, pilot trials, case control studies and naturalistic studies; however, trial protocols, descriptions of

intervention development, discussion or opinion pieces, qualitative content analysis and/or review articles were excluded.

Low- and middle-income countries (LMICs) were defined according to the 2015 World Bank classification. Additionally, countries were included in the search that had recently transitioned from middle- to high-income classification (e.g., Chile, Uruguay), to allow for more inclusive eligibility criteria.

2.3 Study selection & Data extraction

The lead researcher (HFC) screened titles and abstracts to identify articles to undergo full-text review. No language restrictions were applied, and English abstracts were obtained for non-English articles. Studies written in Spanish were directly translated by HFC, and any other non-English language articles were translated using Google Translate. Potentially eligible studies were shared with coauthors (RA and JAN). HFC extracted data including country of origin, sample description, method design and duration, outcome measures and main results, and summarised these details in a table. KA and DD updated the literature search and conducted data extraction for new eligible studies before adding these to the tables. JAN reviewed the table of included studies to confirm eligibility and accuracy in data extraction. HFC, KA and DD finalized the summary table. Due to heterogeneity of the included studies and reported outcomes, and given that this review aimed to summarise recent evidence, a quantitative synthesis of the findings was not possible. Fourteen studies assessing technological interventions for smoking (tobacco) cessation were identified in this search. Due to widespread prevalence of tobacco smoking, along with its relative social acceptance and lack of sanctioning in LMICs, tobacco cessation is often assessed separately from other mental health conditions and addiction in research. Therefore, these 14 studies will be summarised separately.

2.4 Quality Assessment

To assess the quality of included studies, we used the MQRS (Methodological Quality Rating Scale) along with the scoring instructions provided to assess each of the included studies (Miller & Wilbourne, 2002). The scale measures 12 dimensions of methodological quality with cumulative ratings ranging from 0 (lowest quality) to 16 (highest quality) and has been widely used to evaluate study quality in mental health services research, including in prior reviews (Cabassa et al., 2017; Fortuna et al., 2020; Whiteman et al., 2016). Studies with a quality rating score of at least 14 are considered to be high-quality studies (Miller & Wilbourne, 2002).

3. Results

3.1 Characteristics of Included Studies

We identified 37 studies of digital technology for the detection, diagnosis, prevention, treatment, and/or management of mental disorders in 13 LMICs across 7 geographical regions (East Asia, n = 10; South Asia, n = 5; Central or Latin America, n = 11; Middle East, n = 3; Eastern Europe or Western/Central Asia, n = 3; South-East Asia, n = 2; Africa, n = 3). We arranged studies by disorder type, into the following 5 categories: depression and

mood disorders; anxiety disorders and post-traumatic stress disorder; alcohol or substance use disorders; community mental health services including common and severe mental disorders; and suicide prevention. In the sections below, we summarize key findings for each category of included studies. Studies targeting more than one type of disorders were summarised in both tables and are mentioned in subsequent relevant sections in the text.

3.2 Digital interventions for depression and mood disorders

Nearly half of included studies (48%; n = 18 out of 37) targeted depression or mood disorders. As summarized in Table 2, these studies primarily focused on depression in adults, and used online interventions. Arjadi et al. (2018) described an Internetbased behavioural activation programme for major depression in Indonesia, which led to improvements in depressive symptoms and remission rates at follow-up. An online mindfulness intervention from China led to improvements in depression compared to control (Ma et al., 2018). Osorio et al. (2017) described an online intervention for symptom monitoring to prevent relapse and encourage treatment retention, and found that depressive symptoms did not increase during the trial. Tulbure et al. (2018a) implemented an internetsupported intervention for depression in Romania and observed improvement in depressive symptoms. Furthermore, Tulbure et al. (2018b) tested a web-based transdiagnostic intervention in Romania and found improvements in mood disorder symptoms and quality of life measures. One study by Anttila et al. (2019) targeting adolescent depression trialled an internet-delivered program for high school psychosocial wellbeing in Thailand, though no significant differences in depression emerged post-intervention. Yang et al. (2019) tested a mindfulness intervention delivered over WeChat, a highly popular social media platform in China for perinatal depression, which improved depressive symptomatology. Yeung et al. (2018) describe a Chinese-adapted version of MoodGYM, an online CBT-based intervention, which resulted in improved depressive symptoms compared to the care-asusual control (Yeung et al., 2018).

In another study from China, Guo et al. (2020) tested a 6-week internet-based program for mindfulness and self-compassion. At 3 months and 1 year following the intervention, participants showed decreases in depressive symptoms. A study in Iran demonstrated that an online course addressing common causes of depression significantly improved depressive symptoms in a 12-week follow up for female high school students (Moeini et al., 2019). Furthermore, a study on students implementing a 10-week online intervention in Nigeria demonstrated significant reduction in depressive symptoms compared to baseline (Ofoegbu et al., 2020). A multicenter intervention using a digital kiosk to support depression screening, diagnosis, and care in primary care facilities in Colombia resulted in an increase in the percentage of patients diagnosed with depression (Torrey et al., 2020).

Four studies used mobile phones for depression. In Nigeria, one study piloted mobiletelephone supported collaborative stepped care for depression, with the mobile-phone intervention group showing significantly lower depressive symptoms and quality of life compared to the control (Adewuya et al., 2019). Menezes et al. (2019) piloted a mobilephone delivered emotional control intervention (CONEMO) in Peru and Brazil, and found that a high proportion of participants in each country achieved recovery from depressive

symptoms at follow up (Menezes et al., 2019). In India, a guided self-help smartphone app based on principles of cognitive behavioural and interpersonal therapies was piloted, with improvements in depression and well-being emerging post-intervention (Mehrotra et al., 2018).

Two studies tested telepsychiatry through videoconferencing, including a Remote Collaborative Care framework for depression in Chile using shared health records, remote supervision and telephone support, with one study targeting adolescent depression (Martínez et al., 2018b) and the other targeting adult depression in rural areas (Rojas et al., 2018). Both reported positive treatment adherence and satisfaction, though depressive symptoms did not differ compared to the control groups. Finally, Maulik et al. (2017) trialled a digital decision support system (*SMART*) designed to help healthcare workers recognise, diagnose and manage common mental disorders including depression. The authors report a significant reduction in depression symptoms in patients screened using the *SMART* system.

3.3 Digital interventions for anxiety disorders and post-traumatic stress disorder

Table 3 lists studies evaluating Internet-based interventions for anxiety disorders including panic disorder, child and adolescent anxiety disorder, social anxiety, and post-traumatic stress disorder (PTSD). Ma et al. (2016) reported significant improvements in anxiety scores in China following 8 weeks of an online mindfulness intervention. In a study of an electronic decision support system in India to screen, diagnose, and manage common mental disorders, findings after three months of using the intervention demonstrated a significant reduction in anxiety scores (Maulik et al., 2017). In Romania, Ciuca et al. (2018) found that 12 weeks of online CBT appeared to reduce symptoms of panic disorder compared to a waitlist control (Ciuca et al., 2018). In a study of online CBT for child and adolescent anxiety disorders in Iran, anxiety symptoms at 12 weeks were lower compared to the control group (Karbasi and Haratian, 2018). In China, Kishimoto et al. (2016) tested an 8-week online CBT program for social anxiety disorder in adults (Kishimoto et al., 2016). At 8 weeks, symptoms of anxiety reduced in both experimental groups compared to the waitlist control (Kishimoto et al., 2016). In Brazil, Coelhoso et al. (2019) found that an 8-week intervention administered through a well-being mobile app significantly decreased overall and work-related stress in addition to significantly increasing general well-being in adult women participants.

Of the studies targeting PTSD, one from Mexico used virtual reality, where victims or witnesses of violent crimes were exposed to virtual reality scenarios and asked to relay the traumatic event in as many details as possible (Cárdenas-López et al., 2016). At 12-weeks, participants showed improvement in measures of stress, anxiety and depression. In another study, Wang et al. (2016) tested an adapted version of the online My Trauma Recovery self-help intervention for traumas of diverse nature in China, including natural disaster (i.e., earthquake) (Wang et al., 2016). The triggers and self-talk modules of the programme were associated with improvements in PTSD and depression, coping self-efficacy and social functioning, and the relaxation module was associated with improvements in PTSD. In a study from Iran, war veterans with PTSD received daily texts with information about signs and symptoms of PTSD, relaxation techniques, diet, suicidal thoughts and behaviour

(Darvish et al., 2019). At 6 months, improvements in PTSD symptoms and quality of life were observed, as well as lower frequency of recurrence of post-traumatic symptoms.

3.4 Digital intervetions for alcohol use and substance use disorders

As summarized in Table 4, four studies targeted alcohol use disorders, and three studies targeted substance use, including heroin, marijuana and methamphetamine. In Brazil, Baldin et al. (2018) tested a web intervention for binge-drinking in nightclub attendees (Baldin et al., 2018). Intervention participants were exposed to normative feedback relating to alcohol consumption, risk of dependence, financial consequences, driving while under the influence, and tips to reduce damage. After 6 months, there appeared to be reductions in problem drinking among intervention participants. Bedendo et al. (2019) found that a web-based personalised normative feedback intervention for college students could reduce the number of typical drinks consumed compared to the control group (Bedendo et al., 2019). In India, Nandyal et al. (2019) evaluated the feasibility and acceptability of using mobile telephones to follow up with patients with Alcohol Dependence Syndrome. It was found that after 6 months, the collection of outcomes was improved by 35% and that almost 30% of patients were able to remain completely abstinent from alcohol during this follow-up period. A multicenter study in Colombia, also mentioned in section 3.2 above, implemented a digital kiosk to identify and treat alcohol use disorder and found that this intervention increased the percentage of patients diagnosed with the disorder across different primary care sites (Torrey et al., 2020).

Among the studies targeting substance use, one piloted S-Health, a smartphone app based on principles of CBT to support recovery from addiction, with a sample consisting principally of people with heroin addiction in China (Liang et al., 2018). Participants received daily messages and surveys through the app to measure cravings, mood, triggers and other aspects relating to cognitive components of addiction. The intervention group had fewer positive urine tests (indicating relapse) and fewer self-reported days using drugs compared to the control group. Another study from China compared an app-delivered cognitive addiction therapy (CCAT) for methamphetamine use disorder focused on ameliorating patients' cognitive functioning and reducing drug-related cognitive biases with regular detoxification treatment (Zhu et al., 2018). Participants in the CCAT group showed higher performance on measures of cognitive function than the control, though no substance use-related outcomes were reported. In a 3-arm study from Mexico, individuals with substance use disorders including marijuana, inhalants and cocaine dependence, were allocated to a web-based program for drug abuse and concurrent depression (PAADD) based on the transtheoretical model of change along with elements of CBT, a self-help guide based on ASSIST (Alcohol, Smoking and Substance Involvement Screening Test), or care-as-usual (Tiburcio et al., 2018). There were decreases in the average days of drug use and severity of dependence across all conditions, though no significant differences between groups.

3.5 Digital interventions for community mental health services including common and severe mental disorders

Four studies aimed to improve community based mental health services for a range of disorders including schizophrenia and psychotic disorders using SMS reminders and

digital support systems for frontline healthcare workers (Table 5). In India, Singh et al. (2017) conducted a randomized controlled trial of a text-messaging intervention to improve follow-up for patients at a community mental health clinic (Singh et al., 2017). Results showed a higher proportion of participants receiving text-message reminders attended their appointment compared to the control group. A pilot study from South Africa tested a multicomponent intervention for adults with psychotic disorders which integrated care-as-usual, a psychoeducation session, and text message reminders for doctor appointments (Sibeko et al., 2017). At follow up, intervention participants appeared to show better understanding of their diagnosis, and were less likely to miss their first clinic visit and to experience relapse in the 9 months following hospital discharge, though differences compared to usual care were not significant. In a second study from India, the feasibility and initial clinical effectiveness of an Internet-based computerized Clinical Decision Support System for healthcare workers, based on the DSM-IV and ICD-10 criteria, was piloted in rural areas for people with a various psychiatric diagnoses (e.g., mood disorders, substance and alcohol use disorders, stress disorders and psychotic disorders) (Malhotra et al., 2019). The online intervention included 3 separate modules for diagnosis, management, and follow-up, and emerged as feasible and acceptable to patients and health care workers, and achieved high sensitivity in screening for all disorders, but low positive predictive values for most disorders except substance and alcohol use. Lastly, a randomized controlled trial from China tested LEAN (Lay health supporters, *E*-platform, Award and iNtegration) for adults with schizophrenia in rural, resource-poor communities (Xu et al., 2019). This programme used text messages to remind participants of medication, provide health education, and promote self-monitoring of early signs of relapse. At 6 months, the intervention group was significantly more likely to have adhered to medication than the control group, and showed a significant reduction in risk of relapse and hospital readmission.

3.6 Digital interventions for suicide prevention

Two studies focused on suicide prevention using mobile and web-based interventions (Table 6). In the first study, college students in China and Australia completed two mini-modules covering risk factors, warning signs, stigma surrounding suicide, and perceived barriers to help-seeking (Han et al., 2018). Compared to an attentional control group, the experimental group showed improvement in suicide literacy, though there were no differences in help-seeking beliefs and intentions. In Mexico, Martinez-Miranda et al. (2019) conducted a pilot study to determine feasibility and acceptability of an intervention using a virtual conversational agent accessible by mobile app for the prevention and detection of suicidal behaviours (Martínez-Miranda et al., 2019). At 8 weeks, participants favourably rated the emotional competence of the virtual conversational agent and showed high programme adherence (though no outcomes related to change in suicidal behaviors were reported).

3.7 Quality Assessment of Included Studies

As outlined in Table 7, the quality scores of the included studies varied widely, from 2 to 13, with an average of 8.6 (SD= 2.6) and a median of 9. A total of 17 studies scored at least 10 on the MQRS, suggesting moderate quality. Of these, only two studies had scores of 13. No studies were considered high quality, meeting the criteria of 14 or greater. Among the moderate quality studies, most used randomized controlled designs and reported

well-defined outcome measures. The remaining studies scored below 10, with the main drivers of lower quality being the short follow up length, low follow up rate, and lack of independent outcome assessment.

4. Discussion

We identified 37 studies published between 2016 and 2020 evaluating digital interventions for the treatment and prevention of mental disorders in LMICs. Our review expands on recent reviews (Jiménez-Molina et al., 2019; Kaonga and Morgan, 2019; Naslund et al., 2017), reflecting continued growth in digital mental health research in LMICs. Included studies were primarily from East Asia (i.e., China) (Guo et al., 2020; Han et al., 2018; Kishimoto et al., 2016; Liang et al., 2018; Ma et al., 2018; Wang et al., 2016; Xu et al., 2019; Yang et al., 2019; Yeung et al., 2018; Zhu et al., 2018) and Central or Latin America (Baldin et al., 2018; Bedendo et al., 2019; Cárdenas-López et al., 2016; Coelhoso et al., 2019; Martínez et al., 2018b; Martínez-Miranda et al., 2019; Menezes et al., 2019; Osorio et al., 2017; Rojas et al., 2018; Tiburcio et al., 2018; Torrey et al., 2020), followed by South Asia (i.e. India) (Malhotra et al., 2019; Maulik et al., 2017; Mehrotra et al., 2018; Nandyal et al., 2019; Singh et al., 2017). Fewer studies were from Africa (Adewuya et al., 2019; Ofoegbu et al., 2020; Sibeko et al., 2017), Southeast Asia (Anttila et al, 2019; Arjadi et al., 2018), Eastern Europe (Ciuca et al., 2018; Tulbure et al., 2018a; Tulbure et al., 2018b), or the Middle East (Darvish et al., 2019; Karbasi & Haratian, 2018; Moeini et al., 2019). No studies were identified from Western or Central Asia. Our review has the strength of adding new evidence to support potential benefits of digital interventions across various cultural contexts and a broad range of mental health problems. However, our findings suggest that, while lower-income countries in some geographical regions are producing a relatively high amount of research, other areas such as Africa and Western, Central and Southeast Asia are less well-served by research.

4.1 Which mental disorders are being targeted by digital intervention studies in LMICs?

Nearly half the studies evaluated digital interventions targeting depression and mood disorders, which is expected as depression accounts for the highest mental health-related global burden of disease (Rehm and Shield, 2019), and is consistent with prior reviews of digital mental health interventions in LMICs and other low-resource settings (Kaonga and Morgan, 2019; Naslund et al., 2017) as well as high income countries (Lal and Adair, 2014). Remaining studies targeted anxiety disorders and post-traumatic stress disorder, alcohol use and substance use disorders, and community mental health services including care for psychotic disorders. Only two studies addressed suicide prevention, which is perhaps unsurprising considering the social and religious stigma, and legal implications, associated with suicide in many countries (Khan, 2005; Uddin et al., 2019).

4.2 Are digital mental health interventions acceptable in LMICs?

Several studies reported outcomes relating to feasibility and acceptability, including program usage, adherence to interventions, satisfaction or favorability. Consistent with prior research, digital interventions appeared to be well received by participants, with favourable reporting of satisfaction and acceptance (Adewuya et al., 2019; Anttila et al., 2019; Cárdenas-López

et al., 2016; Han et al., 2018; Malhotra et al., 2019; Martínez-Miranda et al., 2019; Nandyal et al., 2019; Osorio et al., 2017). These favourable satisfaction ratings suggest that digital and online delivery of mental health interventions may be suitable and acceptable for diverse patient populations, given the variability in geographical locations in the included studies. Participant-reported acceptability is also important as this may increase intervention engagement and sustained use, which in turn could contribute to greater improvements in desired mental health outcomes. Future studies of digital technology for mental health should explore specific intervention features associated with participant satisfaction and acceptability, and if these features vary by different contexts in LMICs, and whether modifying these features can result in greater intervention uptake and improvements in clinical outcomes.

4.3 Are digital interventions effective in treating mental health conditions in LMICs?

This updated review found a greater proportion of current studies reporting potential effectiveness and overall change in mental health outcomes, suggesting that there have been improvements in design and implementation of digital mental health interventions in LMICs as well as application of more rigorous research methods as the field continues to advance. This is also reflected by the large number of randomized controlled trials (n = 27; 73%); yet, despite moderate quality ratings observed in close to half of the included studies, there were no studies considered high quality on the MQRS (Miller & Wilbourne, 2002). Importantly, nearly two thirds of included studies (n = 23) reported improvements in mental health measures at follow-up (Adewuya et al., 2019; Arjadi et al., 2018; Bedendo et al., 2019; Ciuca et al., 2018; Coelhoso et al., 2019; Guo et al., 2020; Karbasi and Haratian, 2018; Darvish et al., 2019; Kishimoto et al., 2016; Liang et al., 2018; Ma et al., 2018; Maulik et al., 2017; Mehrotra et al., 2018; Menezes et al., 2019; Moeini et al., 2019; Ofoegbu et al., 2020; Tulbure et al., 2018a; Tulbure et al., 2018b; Wang et al., 2016; Xu et al., 2019; Yang et al., 2019; Yeung et al., 2018; Zhu et al., 2018). Of these studies, 15 involved online (n = 14) or computer software (n = 1) interventions, 7 were mobile apps, and one was an SMS text-messaging service. Most studies targeted depression or anxiety disorders (n = 20), with a further two studies addressing alcohol or substance abuse, and one study targeted schizophrenia.

Where effect sizes were given, these were generally promising. Studies of online interventions reported effect sizes, including Tulbure et al. (2018a), who reported Cohen's d values between 0.45 and 1.89, indicating a reduction in depressive symptoms after using online CBT in Romania, that was maintained at 6 months. An online mindfulness intervention in China reported reductions in anxiety and depression with medium to large effect sizes (d = 0.44 - 0.75) (Ma et al., 2018). Kishimoto et al. (2016) noted reductions in anxiety with their internet CBT intervention, with effect sizes between d = 0.29 and 1.03. These studies suggest that online interventions could be effective in reducing symptoms of anxiety and depression in some settings. However, only Tulbure et al. (2018a) had a follow-up period of 6 months, while Ma et al. (2018) had no follow-up period. Without a meaningful follow-up period, it is not possible to determine whether effects are sustained over time, a notable limitation reflected in the methodological quality assessment for many of the included studies, where 23 studies (62%) had less than 6-months follow up.

Considering the high recurrence rates and chronicity of common mental disorders, it is crucial to understand whether mental health interventions are effective in the long term (Koopmans et al., 2011).

It is difficult to draw conclusions about whether digital interventions are more effective for treating some mental disorders than others, due to the disproportionate number of studies targeting each disorder type. The finding that most studies evaluating digital interventions for common mental disorders reported significant findings (n = 20 out of 23) is promising. These findings also point to the potential applicability of digital technologies in treating mental disorders across diverse social and cultural contexts. However, many of these studies were limited by small sample sizes, short follow-up periods, and only 8 of the studies reporting positive findings included standardised effect sizes.

Additionally, where control groups were present, it is important to note that several studies reported no differences between intervention and control groups (Anttila et al., 2019; Baldin et al., 2018; Cárdenas-López et al., 2016; Martínez et al., 2018b; Rojas et al., 2018; Sibeko et al., 2017; Tiburcio et al., 2018). The nature of the interventions trialled in these studies was varied, with two telepsychiatry interventions, one SMS-based intervention, three web-based and one virtual reality intervention reporting no statistically significant differences in clinical outcomes. However, several of these studies reported other positive findings which could be of potential clinical interest. For example, although there was no significant improvement in symptoms, participants with psychotic disorders receiving Sibeko et al.'s (2017) SMS intervention reported better understanding their diagnosis than control participants. This may present clinical utility, as cognitive insight and awareness in psychosis may be associated with more favourable therapy outcomes (Perivoliotis et al., 2009). In two telepsychiatry studies, Rojas et al. (2018) and Martinez et al. (2018) reported better medication adherence in the intervention groups compared with the control. Although these findings were not paired with statistically significant improvements in depressive symptoms, they could suggest clinical utility of digital services in enhancing treatment adherence among patients with depression. Furthermore, the heterogeneity in the types of outcome measures, and study durations limits the possibility of conducting a meta-analysis on the included studies, which would be necessary to determine the efficacy of digital mental health interventions in LMICs. This demonstrates that despite increasing volumes of research on the topic and frequent reporting of clinical outcomes, there remain limitations for drawing conclusions about the relative effectiveness of these digital mental health interventions and the variability across diverse settings and population groups.

4.4 Which technologies are most suitable to deliver interventions?

More than half of the studies used online interventions accessible from a mobile device or computer, involving either new content developed specifically for the study or adaptations to existing evidence-based psychosocial treatments such as CBT or mindfulness therapy (Anttila et al., 2019; Arjadi et al., 2018; Baldin et al., 2018; Bedendo et al., 2019; Ciuca et al., 2018; Guo et al., 2020; Han et al., 2018; Karbasi and Haratian, 2018; Kishimoto et al., 2016; Ma et al., 2018; Moeini et al., 2019; Ofoegbu et al., 2020; Osorio et al., 2017; Tiburcio et al., 2018; Tulbure et al., 2018a; Tulbure et al., 2018b; Wang et al., 2016; Yang

et al., 2019; Yeung et al., 2018). For example, one study described cultural adaptations to an existing program, such as adapting a Chinese version of My Trauma Recovery (Wang et al., 2016), while another study used a well-established and popular social media platform called WeChat to deliver a mindfulness intervention to pregnant women in China (Yang et al., 2019). Interestingly, this study by Yang et al. (2019) was the only study identified in this review using a popular social media platform, which highlights a gap in the current literature. With most of the world's social media users located in LMICs, there may be opportunities to leverage the widespread reach of these platforms to support mental health promotion efforts and service delivery (Naslund et al., 2020). Our findings illustrate continued growth in the use of online interventions for mental disorders, and further confirm the feasibility and acceptability of these programs in low-resource settings, as well as offer new insights about the potential clinical effectiveness for common mental disorders such as depression or anxiety (Jiménez-Molina et al., 2019; Martínez et al., 2018a). Our findings also align with global efforts to increase accessibility to affordable broadband (e.g., the United Nations' Sustainable Development Goal (SDG) 9.c: "strive to provide universal and affordable access to the Internet in least developed countries by 2020' (United Nations Development Programme, 2016)) and highlight the promise of leveraging Internet to deliver mental health care to individuals in lower resource areas of the world.

Mobile phones represented another widely used approach for intervention delivery in the included studies, comprising smartphone applications, text-messaging service (SMS), or phone calls (Adewuya et al., 2019; Coelhoso et al., 2019; Darvish et al., 2019; Liang et al., 2018; Martínez-Miranda et al., 2019; Mehrotra et al., 2018; Menezes et al., 2019; Nandyal et al., 2019; Sibeko et al., 2017; Singh et al., 2017; Zhu et al., 2018). The finding that smartphone delivered mental health interventions are increasingly being tested and reported is promising, especially as mental health apps are continuously being released to the public with little to no empirical evidence behind them (Larsen et al., 2019). Text message and mobile applications appeared to show promise in supporting care for a multitude of disorders, from mild depression to PTSD and psychotic disorders. This evidence suggests that mobile phone applications, if designed and tested rigorously, may provide a means for treating and managing a range of mental illnesses across a variety of contexts and clinical settings. Mounting research on the use of mobile health applications could inform the development and implementation of stronger regulations to guide the safe use of these technologies in lower resource settings. For example, if relevant governing bodies in different countries can draw from lessons learned in empirical research, it may help guide new regulations to ensure that mental health apps being released for easy download to the general public are subject to stringent privacy protections and methodological criteria, including effectiveness, safety and feasibility (Jacobson et al., 2020).

4.5 How do findings from digital mental health studies translate into practice in LMICs?

Despite accumulating evidence presented in this review, uptake by clinics and integration of digital technology into mental health care practice remains disappointingly low, even in higher-income settings (Torous et al., 2018). Efforts to increase global connectivity and accessibility to the Internet and mobile devices are underway (United Nations Development Programme, 2016), though remain behind international targets set under the Connect 2020

Agenda (ITU, 2020). Significant gaps in access to Internet and mobile technologies persist across many regions and are often neglected (Lutz, 2019). The low access to mobile phones or mobile Internet among women when compared to men is a persistent challenge that is especially apparent in impoverished settings globally (Naslund et al., 2019; Rowntree, 2018; Välimäki et al., 2016). Interestingly, with digital technology emerging as a promising approach for promoting mental health and treating some mental disorders, as reflected in this review and other recent reviews, there may be opportunities to align mental health care priorities with existing sustainable development priorities related to increasing digital technology access. For instance, goals to increase access to technology (outlined in SDG 9) could complement efforts to increase access to evidence-based mental health services (SDG 3) (United Nations Development Programme, 2016), thereby potentially allowing individuals at elevated risk of mental health problems, such as suicide, access to online mental health resources (Scherr et al., 2019). As many barriers persist to the implementation and delivery of in-person mental health services and integration of mental health care into existing health systems, by highlighting this important alignment of technological development and mental health care priorities to policymakers and other key decision makers, it may be possible to advocate for the roll out and implementation of promising digital mental health services in LMICs. This also highlights the potential importance of involving frontline health workers in co-creating and designing digital mental health interventions to ensure that new technologies are contextually relevant and to support successful adoption and seemless integration into existing health systems and community mental health services (Khan et al., 2020; Muke et al., 2019; Naslund et al., 2019; Raviola et al., 2019). Such efforts would still require careful consideration of the best evidence available for digital mental health, the need to address privacy and ethical challenges, recognition of stigma, and specific needs and preferences of population groups in greatest need of quality mental health care (Jacobson et al., 2020; Merchant et al., 2020).

4.6 Limitations

Our review has limitations to consider. Our goal was to offer an update over recent reviews on the emerging use of digital technology for treating mental disorders in LMICs. Although this allowed for a broad scope, as reflected by the wide variation in study design and outcomes reported, this made a quantitative synthesis of findings and meta-analysis not possible. Given the heterogeneity of included studies, an assessment of risk of bias was not conducted, further increasing risk of selection bias, though we conducted an assessment of methodological quality of the included studies. Furthermore, we limited our search to MEDLINE, as the prior reviews identified a large number of entries that were not focused on use of digital technology for mental health. Although use of MEDLINE may provide a sufficient and accurate search on its own (Rice et al., 2016) and allows for the use of MeSH headings to improve search efficiency (Chang, Heskett & Davidson, 2006), the lack of complementary searches of other databases has the potential for leaving relevant studies undetected. We found that there were no studies that reported a worsening of symptoms in any intervention group, and no studies reporting negative acceptability or dissatisfaction with digital interventions among participants. The lack of negative findings may reflect the hype surrounding digital mental health and suggests publication bias, whereby only positive findings are published. Additionally, we found that the included studies generally

had low methodological quality as indicated by our MQRS scores. Even though the MQRS rating scale threshold is arbitrarily set at 14 or greater, this quality scale was developed based on the independent assessment of over 360 clinical trials (Miller & Wilbourne, 2002). Therefore, this illustrates the low methodological quality of the current studies relative to existing clinical research, thereby highlighting continued challenges with reliably interpreting research findings in digital mental health research.

The studies in this review largely targeted adults (ages 18–65), making it difficult to draw conclusions about the acceptability of digital interventions for children and adolescents, or the elderly, which are demographic groups at increased risk for mental disorders. Interestingly, we found only one study reporting cost-effectiveness of the digital intervention compared to the control condition (Adewuya et al., 2019). The low number of studies assessing costs indicates an important area for future research and highlights the need to investigate such characteristics as validity, reliability, feasibility, and effectiveness in the context of costs. Determining cost-effectiveness of digital mental health interventions is crucial towards gaining support from investors and stakeholders, and thereby supporting the successful delivery and implementation of digital mental health interventions. Finally, despite not excluding non-English language studies in our search, we recognise that papers in languages other than English may have been missed.

5. Conclusions & Directions for Future Research

Our review builds on previous reviews by tracking the growth in digital mental health research in LMICs over the last four years. Our findings further confirm the feasibility and acceptability of digital mental health interventions, and provide new insights on effectiveness of digital approaches in low-resource settings. Importantly, our findings are also consistent with mounting evidence of digital mental health technologies from higher income countries (Firth et al., 2017; Henson et al., 2019; Lal and Adair, 2014; Naslund et al., 2015). Therefore, our review combined with recent emerging evidence highlights the broader opportunities for digital interventions to target the burden of mental illness and global shortages in the provision of evidence-based mental healthcare services. In recognizing that digital technology holds potential to reduce demand for in-person clinic attention, improve treatment adherence, reduce symptoms of mental illness, lower rates of relapse and readmission, provide on-demand psychological support in the absence of in-person care, and provide access for even the most hard-to-reach groups (such as rural areas, areas of conflict or natural disasters, and people affected by stigma), there is need for continued rigorous research on effectiveness and cost-effectiveness.

To date, digital mental health research has largely been centered in specific regions or health systems, with limited collaboration across diverse contexts, cultures, or countries. The need for collaboration between researchers, clinicians, engineers, persons with lived experience, policymakers, and other stakeholders is clear; however, new initiatives should engage stakeholders across both higher and lower income countries. Future endeavours should step beyond a focus on common mental disorders, to include studies of varied groups and disorders to reflect the mental health needs and care disparities across low-income settings (Merchant et al., 2020). There is a need for rigorous randomized controlled trials

with larger sample sizes to allow for greater confidence in the clinical significance and power of findings and allow for synthesis of high quality evidence. Studies should report standardised effect sizes as well as statistical significance, to ensure that findings can be compared across studies and to allow for meaningful conclusions to be drawn about effectiveness. Furthermore, barriers to accessing care, such as cultural interpretations of mental illness, stigma, discrimination and financial difficulties are relevant and should be taken into consideration by researchers embarking on intervention trials. Such efforts could generate novel findings about mechanisms of action for achieving clinically meaningful outcomes, improve acceptability of interventions, ensure generalizability of findings, and accelerate efforts to sustain digital mental health interventions and integrate high quality mental health care into underresourced health systems.

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- Review builds on recent reviews and rapid growth of digital mental health in LMICs.
- Review identified 37 studies from 13 LMICs published from 2016 to 2020.
- Studies of digital programs to detect, diagnose, prevent and treat mental disorders.
- Findings confirm feasibility and acceptability of digital mental health in LMICs.
- Continued need for rigorous effectiveness and cost-effectiveness studies.

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Table 1.

Terms used in Medline search

Search	Search Terms [*]
"Mental illness"	(((("serious mental illness" OR "serious and persistent mental illness" OR "severe mental illness" OR "mental illness" OR "mental health" OR "mental disorder" OR "schizophrenia" OR "bipolar disorder" OR "schizoaffective disorder" OR "major depressive disorder" OR "major depressive disorder" OR "depression" OR "anxiety" OR "affective disorder" OR "psychotic disorders" OR "psychosis" OR "posttraumatic stress disorder" OR "ptsd" OR "stress disorder" OR "antipsychotic disorders" OR "psychosis" OR "posttraumatic stress disorder" OR "substance related disorder" OR "alcohol" OR "alcohol "OR "substance use" OR "substance use disorder" OR "creack cocaine" OR "marijuana" OR "cannabis" OR "opioid" OR "heroin" OR "morphine" OR "tobacco" OR "street drugs" OR "drug abuse" OR "illicit drugs" OR "recreational drugs" OR "smoking" OR "cigarette")) OR ("mental disorders" [MeSH] OR "Creack or "MeSH] OR "Schizophrenia and Disorders with Psychotic Features" [MeSH] OR "Psychotic Disorders" [MeSH] OR "Schizophrenia and Disorders with Psychotic Features" [MeSH] OR "Psychotic Disorders" [MeSH] OR "CocaineRelated Disorders" [MeSH] OR "Marijuana Abuse" [MeSH] OR "OpioidRelated Disorders" [MeSH] OR "CocaineRelated Disorders" [MeSH] OR "Marijuana Abuse" [MeSH] OR "OpioidRelated Disorders" [MeSH] OR "Tobacco Use Disorder" [MeSH] OR "Street Drugs" [MeSH] OR "Crack Cocaine" [MeSH] OR "Smoking" [MeSH] OR "Tobacco Use" [MeSH])
"Technology"	(("Telemetry" OR "Telemedicine" OR "Telepsychiatry" OR "Telehealth" OR "Telecare" OR "Telemental health" OR "eHealth" OR "uHealth" OR "Connected Health" OR "online" OR "internet" OR "Internet health" OR "Web Browser" OR "Website" OR "Webbased" OR "Social Media" OR "Facebook" OR "Mobile Health" OR "Mobile Technology" OR "Mobile phone" OR "Cellular Phone" OR "Cellphone" OR "Bartphone" OR "text message" OR "SMS" OR "text messaging" OR "Artificial Intelligence" OR "Game Theory" OR "UserComputer Interface" OR "Computer Simulation" OR "Speech Recognition Software" OR "Telemedicine" [MeSH] OR "Internet [MeSH] OR "Meb Browser" [MeSH] OR "Social Media" [MeSH] OR "Cellular Phone" [MeSH] OR "Text Messaging" [MeSH] OR "Artificial Intelligence" [MeSH] OR "Social Media" [MeSH] OR "UserComputer Interface" [MeSH] OR "Artificial Intelligence" [MeSH] OR "Game Theory" [MeSH] OR "UserComputer Interface" [MeSH] OR "Computer Simulation" [MeSH] OR "Speech Recognition Software" [MeSH] OR "Therapy, ComputerAssisted" [MeSH] OR "Computer Simulation" [MeSH] OR "Speech Recognition Software" [MeSH] OR "Telemedicine" [MeSH] OR "Computer Simulation" [MeSH] OR "Speech Recognition Software" [MeSH] OR "Telemedicine" [MeSH] OR "Computer Simulation" [MeSH] OR "Speech Recognition Software" [MeSH] OR "Therapy, ComputerAssisted" [MeSH] OR "Remote Sensing Technology" [MeSH])
"Low- and middle-income countries"	(((("Afghanistan" OR "Benin" OR "Burkina Faso" OR "Burundi" OR "Cambodia" OR "Central African Republic" OR "Chad" OR "Comoros" OR "Congo" OR "Democratic Republic of Congo" OR "Eritrea" OR "Ethiopia" OR "The Gambia" OR "Guinea" OR "GuineaBissau" OR "Haiti" OR "Democratic Republic of Korea" OR "Liberia" OR "Madagascar" OR "Malawi" OR "Mali" OR "Mozambique" OR "Nepal" OR "Niger" OR "Rwanda" OR "Sierra Leone" OR "Somalia" OR "South Sudan" OR "Tanzania" OR "Togo" OR "Uganda" OR "Zimbabwe")) OR ("Armenia" OR "Bangladesh" OR "Bhutan" OR "Bolivia" OR "Cabo Verde" OR "Cape Verde" OR "Cameroon" OR "Republic of Congo" OR "Cote d'Ivoire" OR "Djibouti" OR "Egypt" OR "EI Salvador" OR "Georgia" OR "Ghana" OR "Guatemala" OR"Guyana" OR "Honduras" OR "India" OR "Indonesia" OR "Kenya" OR "Kiribati" OR "Kosovo" OR "Kyrgyz Republic" OR "Kyrgyzstan" OR "Lao PDR" OR "Laos" OR "Lesotho" OR "Mauritania" OR "Micronesia" OR "Moldova" OR "Morocco" OR "Myanmar" OR "Nicaragua" OR "Solomon Islands" OR "Sri Lanka" OR "Sudan" OR "Sanoa" OR "Syrian" OR "Syrian Arab Republic" OR "Tajikistan" OR "TimorLeste" OR "American Samoa" OR "Sanoa" OR "Vietnam" OR "Belarus" OR "Genada" OR "Gosnia and Herzegovina" OR "American Samoa" OR "Bangladesh" OR "Gabon" OR "Genada" OR "Grava OR "Depointe" OR "Dilpines" OR "Sanoa" OR "China" OR "Gabon" OR "Genada" OR "Gusan and Herzegovina" OR "American Samoa" OR "Angola" OR "China" OR "Gabon" OR "Grenada" OR "Iran" OR "Tana" OR "Dominica" OR "Dominican Republic" OR "Ecuador" OR "Fiji" OR "Gabon" OR "Malayia" OR "Maranan" OR "Panamai" OR "Mauritan" OR "Maranai" OR "Malagais" OR "Mongolia" OR "Montenegro" "Gabon" OR "Grenada" OR "Iran" OR "Iraq" OR "Jamaica" OR "Jordan" OR "Kazakhstan" OR "Libya" OR "Macedonia" OR "Malaysia" OR "Maldives" OR "Marshall Islands" OR "Mauritans" OR "Mexico" OR "Mongolia" OR "Montenegro" OR "Namibia" OR "Halawa" OR "Panama" OR "Paraguay" OR "Parau" OR "Tonga" OR "Turkey" OR "St. Lucia" OR "St. Vincent and the Grenadines" OR "Mauritans" OR "Tonga" OR "Tunisia" OR "Turkey" OR "Turkmenistan" O

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Authors & Country	Disorder	Trial type	Sample	Methods	Instruments	Outcome measures	Time points	Findings
Adewuya et al. (2019); Nigeria	Depression	Pilot RCT	895 patients (average age 34.9 years; age range 18–60; 55% female) at 10 primary care centres in Lagos, diagnosed with depression	Mobile telephone adherence support for management of depression (McHPriC Project). Clinics were randomised either to mobile telephone supported Collaborative Stepped Care (mCSC) or control group, consisting of ordinary CSC (oCSC)	PHQ-9, WHODAS, Quality of Life (QoL) Brief version, attendance logs	Rate of adherence to intervention, recovery rate, old, cost- effectivenes, and acceptance	Baseline, 6 and 12 months	mCSC group had significantly better intervention adherence rate than oCSC at 6 months (90% vs 67.8%; 95% CI 1.22–1.40) and at 12 months (78.1% vs 59.2%; 96% CI 1.20–1.43), mCSC had higher depression recovery rate, QoL scores, retention in treatment, cost-effectiveness and acceptance of intervention
Anttila et al. (2019); Thailand	Adolescent psychologica I well-being	Quasi- experimental (feasibility)	167 adolescent high- school students (aged 15.9 years; average 15.9 years; 68.3% female) at three high schools in Southem Thailand	7-week web-based programme (<i>DepisNet-Thai</i>) for supporting the mental wellbeing (depression and stress) of students in a high school setting. Includes 5 weekly modules each lasting 50 minutes (Psychological stress, physical wellbeing, Me and my family, Me and my fricanships, Me and my society). Intervention group completed the programme in groups of 11–14 with a tutor per group. Active control programme due programme independently; passive control had no access to programme	PHQ-9, PSS, CSQ-8	Depression, psychological stress, satisfaction, acceptance and usage of service	Baseline, 11 weeks from baseline	No statistically significant changes emerged at follow-up for satisfaction between the groups. Difference between programme users (group and independent) and non-users (passive control group) were non- significant. Acceptance was higher among those who used the programme who used the programme independently (resolback did not differ significantly between groups
Arjadi et al. (2018); Indonesia	Major depressive disorder (MDD) or persistent depressive disorder (PDD)	RCT	313 participants (over 16 years, average 24.5 years, 81% female) recruited in community settings, scoring 10+ on the PHQ-9, and with internet access	Internet-based behavioural activation and lay counsellor support by email or chat (Guided Act and Feel Indonesia; GAF- ID). Sessions lasted 30–60 minutes per week for 10 weeks for each participant. Control group received online psychoeducation without counselling	РНQ-9; SCID	Self-reported PHQ-9 scores (depressive symptoms)	Baseline, 10- weeks (post- test), 3- and 6- months follow-up	At 10 weeks (post-test, PHQ-9 scores were significantly lower in the GAF-ID group than the control group (mean difference -1.26 , $p = .017$). GAF-ID group had 50% higher chance of remission at 10 weeks (p <.0001). These effects were sustained at 3- and 6- month follow-up

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Authors & Country	Disorder	Trial type	Sample	Methods	Instruments	Outcome measures	Time points	Findings
Guo et al. (2020); China	Postpartum Depression	RCT	284 participants (pregnant women aged 18–401, 2nd or 3rd trimester after 34 weeks, 9 or above on EPDS, have internet and fluent in Chinese)	Internet-based program (6- week MBSP). Contained 6 sequential steps with various guiding exercises. 10 hours of training with 36 episodes, 6 episodes a week each lasting about 15 minutes.	EPDS, BDI, MAAS, SCS, CPBQ, PSI, WHO-5	Mindfulness, self- compassion, depressive symptoms	Baseline,3 months from baseline, 1 year from baseline	Intervention group showed significant ($p < 0.05$) improvement at 3 months and further improved at 1 year compared to control. Intervention group dropped EPDS scores below 9, seeing most depressive symptoms alleviated and falling within reference range after one year.
Ma et al. (2018); China*	Psychological distress (depression and anxiety)	RCT	192 participants (aged 18-47; average 27.8 years; 42% female) recruited by web-based advertisements	8-week online mindfulness intervention, with participants divided into 3 conditions: group mindfulness (GMBI), self- directed (SDMBI) and a discussion group (DG), as well as a blank control group	FFMQ, DERS, SRAS, SRDS	Mindfulness, self-reported symptoms of anxiety and depression, regulation	Baseline, 8 weeks (post- test)	Mindfulness, emotion regulation and depression scores improved significantly in the GMB1 and SDMB1 groups pre- to post-test, with medium to large effect sizes (Cohen's d values between 0.44 - 0.75). Between-group analysis showed that these analysis showed that these outcome changes were highest in GMB1 groups than all others
Martinez et al. (2018); Chile	Adolescent depression	RCT	143 adolescents aged between 13– 19 years (average 15.4 years; 81.1% female), recruited from primary care centres in the Araucanía region	Remote Collaborative Depression Care (RCDC) for adolescents, consisting of 3 months of continuous remote supervision by psychiatrists, shared electronic health records, and telephone support. Control received EUC	BDI. KIDSCREEN	Depressive symptoms, health-related QoL, adherence to medications	Baseline, 12- weeks (post- test)	Patients showed higher adherence to pharmacological treatment. RCDC participants were more satisfied with psychological assistance than the EUC control group. No significant diffreences emerged in depressive or QoL scores between the groups
Maulik et al. (2017); India *	Common Mental Disorders (CMDs)	Pre-post evaluation	238 participants (aged 18–92; average 39.7 years; 58.6% female) with CMDs, recruited from clinics in 30 rural villages	SMART: Systematic Medical Appraisal Referral and Treatment. This project trained lay villege health workers and primary care physicians to screen, diagnose and manage individuals with CMDs using an electronic decision support system	PHQ-9, GAD-7, Knowledge, Attitude and Behaviour. Barriers to Access Care Fvaluation; Treatment Stigma Subscale	Change in mental health utilisation and depression/ adviety scores, stigma, mental health awareness	Baseline, 3- months (post- test)	Significant reduction in depression scores between start and end of intervention among those who screened positive at the beginning of trial. Service utilisation increased, as well as awareness of mental health and stigma

Findings	Depression scores decreased significantly from baseline to post- assessment, with large effect sizes ($r = .68 -$.82, $p < .01$). Well-being also increased within this timeframe.	At post-test, 65% of participants in Sao Paulo and 81% in Lima had achieved 50% reduction in depressive symptoms	Intervention group showed significant improvement in depressive symptoms in first follow-up (from 22.6 to 18.5 mean score on CES-D). These attenuated by 24 week (to 19.5 mean).	First follow-up showed significant reduction in depressive symptoms of intervention group. Further follow-up indicated significant reduction in the intervention group, indicating GIAI was significantly efficacious in reducing depression.	About half the sample made partial use of the programme; though only one participant completed all scheduled online monitoring. There was no reported increase in depressive
Time points	Baseline, mid- way through intervention, post-test, and 2-month follow-up	Baseline, 6 weeks (post- test)	Baseline, 12 week, 24 week	Baseline, 10- weeks, 14 weeks follow- up	Baseline, every 2 weeks, post-test (8 weeks).
Outcome measures	Depression, functioning, self-esteern, well-being	Depressive symptoms, functioning	Depressive symptoms, psychosocial measures	Depressive symptoms	Depressive symptoms, usage of programme.
Instruments	BDI-II, PHQ-9, WSAS, RS- ES, WBI	PHQ-9, WHODAS	CES-D, PSSS- R	BDI-II	PHQ-9, semi- structured interview
Methods	Practise and Use Self-Help for Depression (PUSH-D): a guided self-help app with 10 modules based on CBT, IPT, supportive psychology, Participants completed the intervention for between 6–8 weeks for 2 hours a week.	Emotional Control intervention (CONEMO): mobile-phone delivered, with support of a nurse/ nursing assistant. Consists of 18 behavioural activation sessions over 6 weeks (3 sessions per week)	2-phased DAD online course, first phase was qualitative study consisting of 2 interviews and group discussion. Second phase was preliminary website developed on data collected from phase 1, addressing common depressive themes using multimedia modules	10-week online intervention with GIAI, regular guidance from therapists via phone and internet who were staff of federal universities, and followed ICBT principles. Online modules included psychoeducation, peer support, cognitive disputation, role play, behavioral homework, and depression management.	ASCENSO: online programme to support depression treatment and relapse prevention. The program includes reminder emails and web-based modules for symptom motules for symptom
Sample	78 participants (aged over 17, average 32.3 years; 48.7% female) with major depression or dysthymia and significant impairment in functioning.	66 patients (21 in Sao Paulo, 42 in Lima) at Family Health Care clinics in São Paulo and Lima , aged over 21 and with hypertension or diabetes and PHQ-9 > 10	128 participants, high school students in Iran Hamadan City (all female, mild to moderate depressive symptoms, 15–17 years)	192 EdTech students identified as having depression in Nigeria (62% female, age 19-29 years)	15 patients discharged from treatment for MDD (18–65 years, n=10 female)
Trial type	Pilot pre-post evaluation	Pilot pre-post evaluation	RCT	Group randomized trial	Pilot feasibility/ acceptability study
Disorder	Depression	Depression	Depression	Depression	Depression
Authors & Country	Mehrotra et al. (2018); India	Menezes et al. (2019); Brazil & Peru	Moeini et al. (2019); Iran	Ofoegbu et al. (2020); Nigeria	Osorio et al. (2017); Colombia

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Findings	symptoms during the trial. Participants reported favourable opinions	At 6 months follow-up, RCDC group showed higher satisfaction (OR 1.94; 95% CI 1.25–3.00) and better treatment adherence (OR 1.81; 95% CI 1.02–3.19) than CAU. No statistically significant differences emerged in depressive symptoms between the two groups.	Primary care doctors trained with DIADA increased the percentage of patients diagnosed as having depression from next to 0% to 17% at different locations	Significant differences emerged in outcomes among the 3 conditions (Cohen's d. 45–1.89) in favour of the interventions. No significant differences emerged between C-CBT and R-CBT	Relative to waidlist control, the intervention group appeared to show improvements in the primary outcome measures, anxiety sensitivity, symptom interference and QoL
Time points		Baseline, 3- and 6-months follow-up	Baseline, 6 months for second location, 1 year for first location	Baseline, post- test and 6- months follow-up	Baseline, 10 weeks (post- test), 6-month follow-up
Outcome measures		Depressive symptoms, treatment adherence, health-related QoL	Depression symptoms, unhealthy substance use	Depressive and anxious symptoms, QoL	Symptoms of anxiety and mood disorder, QoL, perfectionism, anxiety, sensitivity, social and work adjustment
Instruments		BDI-I, SF-36, SMAQ	6-ОНА	BDI-II, ATQ, QIDS-SR, BAI, QOLI, GQ, DURI	BDI, PSWQ, SPI, QOLI, WSAS, ODSIS, APS- R, PCL-5, YBOCS
Methods		Remote Collaborative Care for Depression (RCDC) for people living in rural areas in Chile. Programme used shared electronic used shared electronic supervision and telephone monitoring. Once a week a specialist mental health team reviewed patient data and provided remote assistance. The intervention lasted 3 months. Control group received CAU	5-year multicenter intervention, DIADA, aiming to evaluate implementation of technology-supported screening, diagnosis, and care of depression among adults in primary care adults in primary care at six sites in Colombia. New primary care site implements the intervention every 6 months.	Two internet-supported interventions for depression: conventional CBT (C-CBT) or religious CBT (R-CBT). Control was waitlist	Web-based transdiagnostic intervention based on Unified Protocol for Transdiagnostic Treatment of Emotional Disorders that addresses common anxiety and depression. Programme lasted 10 weeks. Control was waitlist
Sample		250 patients from community hospitals in rural locations, (between 18–70 years, 86.4% female) with major depressive episode	2656 patients screened (1943 urban, 713 small town, 62% females, 37% 18–44 years and 63% older than 45)	79 participants (over 18 years, average 32.1 years, 82.3% female), recruited from local and national media, with BDI score between 14–50 and a diagnosis of current MDD or dysthymia	105 participants recruited from local and national media and diagnosed with anxiety and/or affective disorder
Trial type		Open-label (nonrandomised) trial	RCT	RCT	RCT
Disorder		Depression (major depressive episode)	Depression and Alcohol Use Disorder	Depression	Depression and anxiety (transdiagnostic intervention)
Authors & Country		Rojas et al. (2018); Chile	Torrey et al. (2020); Colombia **	Tulbure et al. (2018a); Romania	Tulbure et al. (2018b); Romania

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Findings	Participants in the intervention group showed greater declines in depressive and anxious symptoms (p<.001) compared with those in the control group, as well as a significant improvement in mindfulness skills	Intervention group showed a significantly higher improvement in depressive symptoms than control group (11 vs 5.9 points post-test, $p=.02$), with a medium effect size ($d=.6$)	
Time points	Baseline, 8 weeks (post- test)	Baseline, 5 weeks (post- test)	
Outcome measures	Anxiety, depression and mindfulness	Depressive symptoms	
Instruments	gads, phq, ffmq	CES-D	
Methods	Online mindfulness intervention, delivered over WeChat. 4 sessions, each lasting ~40 minutes, over 8 weeks. Control group received routine prenatal care	Chinese translated version of MoodGYM along with CAU. MoodGYM is a computerised version of CBT for depression, consisting of 5 interactive modules. Participants completed 1 per week, and received weekly reminder emails or phone calls. CAU CAU	
Sample	123 pregnant women over the age of 18 (average 31.3 years), recruited from outpatient clinics of women's hospitals, and screening positive for depressive or anxious symptoms	75 adults from outpatients clinics, with significant depressive symptoms (as judged by a clinician), aged over 18 (average 33 years; 77% female)	
Trial type	RCT	Pilot RCT	
Disorder	Maternal (perinatal) depression and anxiety	Depression	
Authors & Country	Yang et al. (2019); China	Yeung et al. (2018); China	*

* Indicates the presence of this study in Table 2 for depression and mood disorders in addition to Table 3 for anxiety disorders

J Psychiatr Res. Author manuscript; available in PMC 2022 January 31.

** Indicates the presence of this study in Table 2 for depression and mood disorders in addition to Table 4 for substance use and alcohol use disorders APS-R: Almost Perfect Scale-Revised

ATQ: Automatic Thoughts Questionnaire

BAI: Beck's Anxiety Inventory

BDI/BDI-II: Beck's Depression Inventory (Version I or II)

CES-D: Centre for Epidemiologic Studies Depression Scale

CPBQ: Child Prosocial Behavior Questionnaire

DURI: Duke University Religion Index

EPDS: Edinburgh Postnatal Depression Scale

FFMQ: Five Facet Mindfulness Questionnaire

GADS: Generalised Anxiety Disorder Scale

GQ: Gratitude Questionnaire

KIDSCREEN: quality of life questionnaire for children and adolescents

MAAS: Mindful Attention Awareness Scale

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WHO-5: World Health Organization- Five Well-Being Index

WSAS: Work and Social Adjustment Scale YBOCS: Yale-Brown Obsessive Compulsive Scale

WHODAS: WHO Disability Assessment Schedule

WBI: Well-Being Index

Findings	Post-treatment evaluation revealed a 30% improvement in measures of stress, anxiety and depression in both PTSD and ASD groups. A significant effect of time emerged pre- to post-test (p<.001), but there were no significant differences across groups (p>.05)	Both active treatment conditions (guided and unguided iCBT) showed superior symptom outcomes (guided iCBT vs control: d=1.04-1.36; non-guided vs control: d=.07-1.06). Treatment gains were maintained at follow-up and guided iCBT became superior to unguided iCBT at 6 month follow-up (p<.05)	The intervention group and control group showed significant increase in general well-being as a function of time (p=0.006). However, only the intervention group showed significant increase in work-related well-being (P-0.001) and significant reduction in work-related stress (p=0.004) and overall stress (p<0.001)	Intervention group post- test scores were significantly lower than
Time points	Baseline, 12 weeks (post-test)	Baseline, 12 weeks (post-test), 6-month follow-up	Baseline, 4 weeks (midinterven tion), and 8 weeks after baseline (postintervention)	Baseline, 3 months (post-test)
Outcome measures	PTSD and anxiety symptoms, depressive symptoms, satisfaction with intervention	Self-report panic symptoms	Symptoms of Stress, 10-Item Perceived Stress Scale, 5- Item World Health Health Organization Well-Being Index	Symptoms of anxiety
Instruments	CAPS-I, PSS- SR, BDI, Treatment satisfaction questionnaire	PDSS-SR, PDSQ	PSS-10, WHO-5	SCARED
Methods	Participants were separated into PTSD and ASD. Intervention consisted of 610 individual sessions once a week, lasting 90 mins aech, with a clinical psychologist. Sessions 4–10 psychologist. Sessions 4–10 consisted of virtual reality scenario exposure. During sessions, participants were asked to talk about the traumatic event in first person in as much detail as possible	Internet CBT (iCBT): 12 weeks of guided (via real- time video sessions) or unguided internet CBT. Control group was waitlist	Well-being mobile app had an 8-week program with 4 classes per week with a brief theoretical portion and a 15-min guided practice. The active control app also provided 4 assessments per week to encourage self- observation in participants and also conducted web- based questionnaires to assess subjective levels of stress and well-being at baseline, midintervention, and postintervention.	iCBT for anxiety disorders, consisting of 7 stages in a 3-month period. Paricinants did 1 hour each
Sample	9 individuals (aged 18–65 years) who were victims or witnesses of assault, kidnapping, or other criminal violence, recruited from a psychological assistance centre	111 participants (aged 18-65, average 35.2 years, 68% female) recruited through media and GP recommendation s, diagnosed with panic disorder	148 women aged 20-60 years who completed high school, owned a mobile device, recruited from private tertiary care hospital in Sao Paulo, Brazil	30 girls aged 10–18 (average 14.2 years) with anxiety disorders, recrinited from clinics
Trial type	Nonrandomised open-label trial	RCT	2-arm RCT	RCT
Disorder	PTSD and Acute Stress Disorder (ASD) associated with trauma	Panic disorder	Psychological distress	Child and adolescent anxiety
Authors & Country	Cárdenas- López et al. (2016); Mexico	Ciuca et al. (2018); Romania	Coelhoso et al. (2019); Brazil	Karbasi & Haratian (2018); Iran

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Table 3.

Digital interventions for anxiety disorders and post-traumatic stress disorder

Authors & Country	Disorder	Trial type	Sample	Methods	Instruments	Outcome measures	Time points	Findings
			of child and adolescent psychiatry in Isfahan	day of assignments for each stage. Control was CAU				control group scores (p<.001)
Kishimoto et al. (2016); China	Social anxiety	Controlled trial	197 participants (average 24.6 years; 73% female), 75 of which had social anxiety disorder	iCBT for 8 weeks with and without guidance (weekly email by supervising therapist). Control was waitlist	SIAS, SPS, BDI	Symptoms of social anxiety	Baseline, 8 weeks (post-test)	For SIAS scores, both iCBT conditions were superior to watelist control (guided vs waitlist: $p<001$; unguided vs waitlist: p<001), with small to large effect sizes (d = 0.29 - 1.03). No difference arose between the two iCBT conditions were superior to waitlist control ($p<001$), while the two iCBT conditions were superior to waitlist control ($p<001$), while the two iCBT conditions
Darvish et al. (2019); Iran	War-related PTSD	RCT	66 male war veterans with PTSD (aged 42– 61 years; average 48.4 years), recruited from the outpatient clinic of a hospital	Text messaging-based psychiatric nursing programme. Texts were sent daily to participants for 6 months. Content of messages covered: welcome message, description of PTSD symptoms, methods of symptoms, methods of accepting/dealing with PTSD, self-control, relaxation, deep breathing, insomnia, diet, suicidal thoughts. Control group received CAU	SRS-PTSD, SF-36	PTSD symptom severity and QoL	Baseline, 6 months (post-test)	In the intervention group, a significant improvement in PTSD symptoms was observed pre- to post- intervention (p=.001). Frequency of recurrence of PTSD symptoms was significantly lower in the intervention group at 6 months (p=.03)
Ma et al. (2018); China*	Psychological distress (depression and anxiety)	RCT	192 participants (aged 18-47; average 27.8 years: 42% female) recruited by web- based advertisements	8-week online mindfulness intervention, with participants divided into 3 conditions: group mindfulness (GMBI), self- directed (SDMBI) and a discussion group (DG), as well as a blank control group	FFMQ, DERS, SRAS, SRDS	Mindfulness, self-reported symptoms of anxiety and depression, regulation	Baseline, 8 weeks (post-test)	Mindfulness, emotion regulation and anxiety scores improved significantly in the GMB1 and SDMB1 groups pre- to post-test, with medium to large effect sizes (Cohen's effect sizes (Cohen's analysis showed that these outcome changes were highest in GMB1 groups than all others
Maulik et al. (2017); India*	Common Mental	Pre-post evaluation	238 participants (aged 18–92; average 39.7 years; 58.6% female)	SMART: Systematic Medical Appraisal Referral and Treatment. This project	PHQ-9, GAD-7, Knowledge,	Change in mental health utilisation and	Baseline, 3-months (post-test)	Significant reduction in anxiety scores between start and end of

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Findings	intervention among those who screened positive at the beginning of trial. Service utilisation increased, as well as awareness of mental health and stigma	2 of the intervention modules showed consistent associated with improvement in PTSD and depressive symptoms, coping self-efficacy and social functioning. Other modules, such as relaxation, were associated with improvement in PTSD but a negative change in two social variables	
Time points		Baseline, 1 month (post-test), 3-month follow-up	
Outcome measures	depression/ anxiety scores, stigma, mental health awareness	PTSD symptoms, depressive symptoms, functioning	
Instruments	Attitude and Behaviour, Barriers to Access Care Evaluation; Treatment Stigma Subscale	PDS, SCL-D, CSS, SFI	
Methods	trained lay village health workers and primary care physicians to screen, diagnose and manage individuals with CMDs using an electronic decision support system	Chinese version of My Trauma Recovery (CMTR): a web-based self-help intervention for trauma. Participants accessed the programme for 1 month on their own or guided by volunteers in a counselling centre	
Sample	with CMDs, recruited from clinics in 30 rural villages	56 urban survivors of different trauma types and 90 rural survivors of 2008 Sichuan earthquake (aged 16–70; 24.5% female)	
Trial type		PTSD associated with different categories of trauma	
Disorder	Disorders (CMDs)	Pre-post clinical evaluation	
Authors & Country		Wang et al. (2016); China	*

"Indicates the presence of this study in Table 2 for depression and mood disorders in addition to Table 3 for anxiety disorders

CAPS-I: Clinician-Administered PTSD Scale

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CSS: Crisis Support Scale

PDS: Posttraumatic Diagnostic Scale

PDSS-SR: Panic Disorder Severity Scale - Self-Report

PDSQ: Psychiatric and Diagnostic Screening Questionnaire

PSS-SR: PTSD Symptom Scale - Self-Report

PSS-10: Perceived Stress Scale-10 Item

SCARED: Screen for Child Anxiety Related Emotional Disorders

SCL-D: Symptom Checklist for Depression

SFI: Social Functioning and Impairment Questionnaire

SIAS:Social Interaction Anxiety Scale

SPS: Social Phobia Scale

SRS-PTSD: Self-Rating Scale for Post-Traumatic Stress Disorder

STAI: State-Trait Anxiety Inventory for Adults

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Table 4.

Digital interventions for substance use and alcohol use disorders

Findings	A significant reduction in binge-drinking during the week was observed in 38% of participants in the experimental group at 6 months; this reduction was not seen in the control group. However, upon observation of group- time interaction, the results are not significant and the reduction in 'lack of control' when drinking was marginally significant only	PNF intervention reduced the number of typical drinks consumed at all follow-up points compared to control (all p values <0.5). Low motivation students receiving PNF experienced an increase in AUDIT score at T3 (p<.001)	At post-test, 26.2% of intervention group and 50% of control group had positive urine tests (p=.06). The number of self-reported days using drugs in the intervention group was lower than controls (p<.05)	After 6 months, 27,5% of participants had complete abstinence. 72% of participants had lapses/ relapses, and 62% were relapses, and 62% were
Time points	Baseline, 6 months	Baseline, 1–3- and 6months.	Baseline, 4 weeks (post-test)	Baseline, 6 months
Outcome measures	Various measures of drinking behaviour	AUDIT score, typical number of drinks consumed	Cravings, affect, trigger thoughts, place and situations, responses to triggers, social context, urine test-confirmed abstinence	ADS symptoms, consumption of alcohol
Instruments	AUDIT	AUDIT	I-PANAS-SF, urine tests, Addiction Severity Index	Short alcohol dependence data questionnaire, rotter's scale for locus of
Methods	Web-based intervention consisting of exposure to normative feedback regarding alcohol consumption, risk, money spent, drink-driving, risk classification (AUD) and tips to reduce alcohol-related damage. Control group had no intervention and just completed a questionnaire about drinking behaviour (AUDIT)	Web-based Personalised Normative Feedback (PNF) intervention, consisting of several components: 1) drinking profile (AUDIT risk level, binge-drinking frequency); 2) gender-specific normative comparisons; 3) practical costs (e.g. money spent on alcohol); 4) alcohol-related consequences; 5) low-risk drinking limits and protective strategies. Students took 5– 10 minutes to complete the intervention at each time point. Control group received the basic assessment only (no PNF)	S-Health: a bilingual smartphone app based on principles of CBT to support addiction recovery. Participants received text messages and daily surveys on cravings, affect, triggers, responses to triggers, and social contexts	After initial assessment of patients in psychiatry department of tertiary care hospital, a follow-up was conducted after 6 months through a combination of in-
Sample	465 nightclub patrons from Sao Paulo, classified as problem drinkers (aged 18–55; average 24.7 years, 35.5% female)	4460 college students (between 18–30 years; average 22 years; 51% female), divided into high and low motivation, nigh and low motivation, and Facebook and Facebook	75 participants (aged 18– 65; average 41,6 years; 29.3% female) who had used heroin in the last 30 days. Recruited from methadone maintenance treatment clinics and a social worker consortium	51 participants (diagnosed with ADS, age 22–58)
Trial type	RCT	RCT	Pilot RCT	Feasibility/ ac ceptability study
Disorder	Binge drinking	Alcohol use disorder	Substance use (principally heroin but also other)	Alcohol Dependence Syndrome (ADS)
Authors & Country	Baldin et al. (2018); Brazil	Bedendo et al. (2019); Brazil	Liang et al. (2018); China	Nandyal et al. (2019); India

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Findings	the time of follow-up. The collection of outcomes was improved by 35.2%. The lower rate of abstinence is lower rate of abstinence is hypothesized to be caused by the ability to contact patients via mobile phone for patients who dropped out of hospital visits.	No significant difference arose in average number of days of substance use or severity of substance abuse between the three conditions. Both outcomes decreased independent of group, with moderate to strong effect size (between $d = -0.7$ and -1.1). A reduction in depressive symptoms was also observed, again independent of group	Primary care doctors trained with DIADA increased the percentage of patients diagnosed as having Alcohol Use Disorder from next to 0% to 2% at different locations	Compared to control, CCAT significantly improved cognitive function (p=.01) and impulsive risk-decision making (p<.001) in experimental group
Time points		Baseline, 8 weeks (post-test), 1-month follow-up	Baseline, 6 months for second location, 1 year for first location	Baseline, 4 weeks (post-test)
Outcome measures		Number of days of substance use, severity of substance use, depressive symptoms, readiness to change	Depression symptoms, unhealthy substance use	Cognitive function, impulsive risk- decision making attention bias
Instruments	control, clinical institute of withdrawal assessment of alcohol scalerevised	ASSIST, Timeline Followback, DAST-20, RCQ	6-ОНА	CogState test, ISL, CPAL, DDT, IGT, BART
Methods	person and phone interviews with patients and caregivers.	Web-based help programme for drug abuse and concurrent depression (PAADD), based on the transtheoretical model of change, and incorporating elements of CBT. Completion of programme required approx. 8 weeks if used for 1 hour per week. A second group received ASSIST self-help guide. Control group received CAU	5-year multicenter intervention, DIADA, aiming to evaluate implementation of technology- supported screening, diagnosis, and care of depression among adults presenting or primary care visits at aix sites in Colombia. New primary care site implements the intervention every 6 months.	Mobile-based cognitive addiction therapy app (CCAT) (and detox) based on cognitive rehabilitation therapy, with the aim of improving cognitive function and eliminating 33 years). Patients had moderate to severe addiction drug-related attention bias. Intervention was delivered for 4 weeks: 20 sessions total, 5 times a week, each session approx. 60 mins. Control group received CAU
Sample		74 participants (over 17 years; average age not given, but most aged 18–25; 12.2% female) recruited from two addiction treatment centres	2656 patients screened (1943 urban, 713 small town, 62% females, 37% 18-44 years and 63% older than 45)	40 male patients at a compulsory methamphetamine renabilitation centre in Shanghai (aged 18–49; avenge 33 years). Patients had moderate to severe addiction
Trial type		RCT	RCT	RCT
Disorder		Substance use (marijuana, inhalants, cocaine)	Alcohol Use Disorder and Depression	Substance use (methampheta mine)
Authors & Country		Tiburcio et al. (2018); Mexico	Torrey et al. (2020); Colombia **	Zhu et al. (2018); China

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** Indicates the presence of this study in Table 2 for depression and mood disorders in addition to Table 4 for substance use and alcohol use disorders

ASSIST: Alcohol, Smoking and Substance Involvement Screening Test

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Table 5.

Digital interventions for community mental health services including common and severe mental disorders

Authors & Country	Disorder	Trial type	Sample	Methods	Instruments	Outcome measures	Time points	Findings
Malhotra et al. (2019); India	Mental health problems (general) in remote areas of India	Feasibility/ initial clinical effectiveness study	2594 psychiatric patients in rural areas. Most common diagnoses were mood disorders, substance or alcohol use disorders, anxiety and stress-related disorders and psychotic disorders	Internet-based, computerized Clinical Decision Support System (CDSS): a digital diagnostic and management application to improve delivery of mental healthcare in rural areas. Included 3 modules for diagnosis, management and follow-up respectively, based on DSM- IV and ICD-10 criteria. The aim was to replicate a routine clinical situation. Psychologist or social worker used the app to interview patients, choose medication or psychotherapy as per CDSS guidelines. Patient was assessed at follow-up using the relevant module	MINI/MINI- KID, measures of validity, reliability, feasibility and satisfaction	Validity, reliability and feasibility of CDSS; satisfaction with intervention	Unclear. Follow-up timepoints varied depending on the nature of a participant's illness and treatment	Sensitivity of screening tool was high for all disorders (ranging from 48–99%). Positive predictive values were low for most disorders except alcohol and substance use disorders. Online diagnostic module was acceptable/ satisfactory to patients and interviewers
Sibeko et al. (2017); South Africa	Psychotic disorder (various)	Pilot RCT	77 participants (average 35.5 years; 28.6% female) with schizophrenia spectrum disorder, bipolar mood disorder, or substance use induced psychotic disorder, recruited from Valkenburg hospital	A treatment- partner and text message intervention. Incorporated CAU with addition of 1) a treatment partner contracting and psychoeducation session and 2) text message reminders of clinic appointments. The aim of the intervention was to improve adherence to mental health services for people with psychotic disorder.	PANSS, CGI, GAF, CAN,, EUROQUOL, VAS, MARS	Clinic attendance, adherence, symptoms, readmission rate at 9 months	Baseline, post- discharge, 3-month follow-up	At 3-month follow-up, intervention participants understood their diagnosis better than CAU group. CAU group CAU group were more likely to miss their first clinic visit and relapse in the 9 months following discharge, and to show worsening PANSS scores. Efficacy outcomes tended to

Authors & Country	Disorder	Trial type	Sample	Methods	Instruments	Outcome measures	Time points	Findings
				Control group received CAU				favour the intervention but did not reach statistical significance
Singh et al. (2017); India	Unspecified diagnoses (various mental illnesses)	RCT	214 patients (40.6% female)	Intervention to improve follow- up rates at a community mental health clinic. SMS group received SMS 1 day before appointment at clinic. Control group received no SMS. At second-level intervention patients from both groups who missed first appointment were given a voice call reminder	Measure of attendance	Attendance at clinic	No fixed time points	At first SMS intervention level, 62.3% of SMS group reached their appointment, compared to 45.4% of the CAU group. At second- level intervention, 66 of 88 came for follow-up. Diagnosis of alcohol dependence was significantly associated with missing appointments
Xu et al. (2019); China	Schizophrenia	RCT	278 community- dwelling villagers with schizophrenia (average 46 years; 55.4% female)	LEAN intervention: Lay health supporters, Eplatform, Award and iNtegration). This featured recruitment of lay health support and text messages for medication reminders, psycho- and health education, monitoring of early signs of relapse, and linkage to primary healthcare for people with schizophrenia in rural resource- poor China. Control group received CAU	Score of adherence to antipsychotic medication, BARS, DAI, CGI for schizophrenia, WHODAS, GASS	Medication adherence, patient symptoms, functioning, relapse, re- hospitalisation, death for any reason, violence	Baseline, 6 months	Medication adherence was significantly improved at 6 months (0.48 in control group, 0.61 in intervention; representing a 27% improvement in the LEAN group), with moderate effect size of d = 0.38 reported, as well as a significantly reduced risk of relapse and re- hospitalisation with the intervention. The two groups did not differ on other outcomes

BARS: Brief Adherence Rating Scale

CAN: Camberwell Assessment of Unmet Needs

CGI: Clinical Global Impressions

DAI: Drug Attitude Inventory

EUROQUOL: Quality of Life measurement

GAF: Global Assessment of Function

GASS: Glasgow Antipsychotic Side-Effect Scale

MARS: Medication Adherence Rating Scale

VAS: Visual Analogue Scale

Findings	No significant difference was found between control and experimental group on professional help- seeking beliefs and intentions. Both groups' help-seeking attiudes increased during attiudes increased during attudes increased during attudes increased at follow-up). A significant improvement in sucide literacy at post-test was found for the experimental group (p=.015)	ECA emotional competence and adherence rate were rated positively
Time points	Baseline, post-test, 1- month follow-up	Baseline, follow-up each 2 weeks, 8 weeks (post- test)
Outcome measures	Professional help- seeking beliefs, intentions and attitudes; suicide literacy, stigma, social support, self- reliance, preparedness to assist a suicidal fried, evaluation of program	Acceptability of program, emotional competence of ECA, adherence
Instruments	GHSQ, Attitudes Towards Seeking Professional Psychological Help Scale: LSS, SSS; BHSS, PHQ-9	Unstructured brief interviews; Likert-based questionnaire to assess ECA features
Methods	Brief, two-module online psychoeducational program (<i>ProHelp</i>) to encourage help-seeking behaviour for university studens with suicidal ideation and behaviour. The suicidal ideation and behaviour. Thutes, and addressed risk factors, warning signs, stigna and perceived barriers to help- seeking, with the aim of preventing suicide in university students. Control group completed an attentional control programme	Embodied virtual conversational agent used as the main interface of HelPath: a mobile application addressed to individuals detected with suicidal behaviours (ideation, planning or attempt). Participants were invited to use HelPath daily for 8 weeks, as well as a fortnightly interview face-to-face with the specialist
Sample	156 Chinese students and 101 Australian students (18–30 years, average 19.3 years, 79.5% female) from universities in China and Australia	18 participants (20–53 years; average 31.5 years; 11 female) recruited by mental health specialists
Trial type	RCT	Exploratory pilot acceptability study
Disorder	Suicidal thoughts, tendencies and behaviour	Suicidal behaviour
Authors & Country	Han et al. (2018); China (and Australia)	Martinez- Miranda et al. (2019); Mexico

BHSS: Barriers to Help-Seeking Scale

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CSQ-8: Client Satisfaction Questionnaire

DERS: Difficulties in Emotion Regulation Scale

HHS: Herth Hope Scale

GAD-7: Generalised Anxiety Disorder Questionnaire

GHSQ: General Help-Seeking Questionnaire

LSS: Literacy of Suicide Scale

MINI/MINIKID: Mini International Neuropsychiatric Interview

MLQ: Meaning in Life Questionnaire

PSS: Perceived Stress Scale

SRAS: Self-Rating Anxiety Scale

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Table 6.

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Digital interventions for suicide prevention

Author Manuscript SRDS: Self-Rating Depression Scale	STS: Self-Transcendence Scale	SSS: Stigma of Suicide Scale	ZSAS: Zung Self-Rating Anxiety Scale
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Table 7.

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Assessment of methodological quality of included studies.

Total Quality Score (0	Low to 16 High)		13	8	6	13	5	10	7	2	10	Π	8	5
12	L. Collateral (0-1)		0	0	0	0	0	0	0	0	0	0	0	0
11	K. Study Site (0-1)		0	1	0	0	0	0	0	0	1	0	0	0
10	J. Analyses (0-1)		1	1	1	1	1	1	1	0	1	1	1	1
6	I. Independent (0–1)		0	0	0	0	0	0	0	0	0	0	0	0
×	H. Dropouts (0-1)		1	1	0	1	0	1	0	0	1	1	0	0
7	G. Objective Measurement of Outcomes (0-1)		_	0	1	-	1	1		0	I	-	1	1
9	F. Follow- Up Rate (0–2)		2	2	2	5	0	2	0	0	7	2	0	0
w	E. Follow- Up Length (0-2)		2	0	1	2	0	0	0	0	0	1	0	1
4	D. Quality Control (0-1)		1	0	0	1	0	0	1	0	1	1	1	1
3	C. Baseline (0-1)		1	1	1	1	0	1	1	1	1	1	1	0
7	B. Replicability (0-1)	sra	1	1	0	1	0	1	0	1	1	0	1	1
1	A. Study Design (0-3)	ood Disorde	n	1	ŝ	б	3	3	ŝ	0	1	3	3	0
Quality Criteria*	Author (year)	Depression & Mo	Adewuya et al. (2019); Nigeria	Anttila et al. (2019); Thailand	Arjadi et al. (2018); Indonesia	Guo et al. (2020); China	Ma et al. (2018); China [*]	Martinez et al. (2018); Chile	Maulik et al. (2017); India [*]	Mehrotra et al. (2018); India	Menezes et al. (2019); Brazil & Peru	Moeini et al. (2019); Iran	Ofoegbu et al. (2020); Nigeria	Osorio et al. (2017);

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Total Quality Score (0	Low to 16 High)	12	9	6	10	10	10		5	6	6	4	5	L	5
12	L. Collateral (0-1)	0	0	0	0	0	0		0	0	0	0	0	0	0
11	K. Study Site (0-1)	0	1	0	0	0	0		0	0	0	0	0	0	0
10	J. Analyses (0-1)	1	1	1	1	1	1		1	1	1	1	1	1	1
6	I. Independent (0-1)	1	0	0	0	0	0		0	0	0	0	0	0	0
×	H. Dropouts (0-1)	1	0	1	1	0	1		0	1	1	0	0	0	0
7	G. Objective Measurement of Outcomes (0-1)	1	1	1	1	1	1		1	1	1	1	1	1	1
9	F. Follow- Up Rate (0–2)	2	0	0	0	2	1		0	1	1	0	0	1	0
S	E. Follow- Up Length (0-2)	1	1	1	1	0	0		0	1	0	0	0	0	0
4	D. Quality Control (0-1)	1	1	1	1	1	1		0	0	0	1	1	0	0
e	C. Baseline (0-1)	1	0	0	1	1	1		0	1	1	0	1	0	0
6	B. Replicability (0-1)	1	0	1	1	1	1	tress disorders	1	0	1	0	0	1	0
1	A. Study Design (0-3)	2	1	3	3	3	3	raumatic S	2	3	3	1	1	3	3
Quality Criteria*	Author (year)	Rojas et al. (2018); Chile	Torrey et al. (2020); Colombia**	Tulbure et al. (2018a); Romania	Tulbure et al. (2018b); Romania	Yang et al. (2019); China	Yeung et al. (2018); China	Anxiety & Post-t	Cárdenas- López et al. (2016); Mexico	Ciuca et al. (2018); Romania	Coelhoso et al. (2019); Brazil	Karbasi & Haratian (2018); Iran	Kishimoto et al. (2016); China	Darvish et al. (2019); Iran	Ma et al. (2018); China*

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Quality Criteria*	1	2	3	4	2	9	7	8	6	10	11	12	Total Quality Score (0
Author (year)	A. Study Design (0–3)	B. Replicability (0-1)	C. Baseline (0–1)	D. Quality Control (0-1)	E. Follow- Up Length (0-2)	F. Follow- Up Rate (0-2)	G. Objective Measurement of Outcomes (0-1)	H. Dropouts (0-1)	I. Independent (0-1)	J. Analyses (0-1)	K. Study Site (0-1)	L. Collateral (0-1)	Low to 16 High)
Maulik et al. (2017); India *	e,	0	1	1	0	0	-	0	0	1	0	0	7
Wang et al. (2016); China	-	-	0	0	0	2	-	0	0	1	0	0	9
Substance Use a	nd Alcohol	Use Disorders											
Baldin et al. (2018); Brazil	m	-	1	1	1	0	-	1	0	1	0	0	10
Bedendo et al. (2019); Brazil	ŝ	1	1	1	1	0	-	1	0	1	0	0	10
Liang et al. (2018); China	r,	-	1	0	0	2	-	1	0	1	0	0	10
Nandyal et al. (2019); India	2	1	0	0	1	2	1	1	0	1	0	0	6
Tiburcio et al. (2018); Mexico	ŝ	1	1	1	0	1	-	1	0	1	1	0	11
Torrey et al. (2020); Colombia**	1	0	0	1	1	0	1	0	0	1	1	0	6
Zhu et al. (2018); China	3	1	0	1	0	2	1	1	0	1	0	0	10
Severe Mental L	Disorders												
Malhotra et al. (2019); India	3	0	0	1	1	1	-	1	0	1	0	0	6
Sibeko et al. (2017); South Africa	3	1	1	0	1	2	1	1	0	1	0	0	11
Singh et al. (2017); India	3	0	0	0	0	0	1	0	0	1	0	0	5
Xu et al. (2019); China	3	1	1	1	1	2	1	1	0	1	0	0	12
Suicide Preventi	ion												

Total Quality Score (0	Low to 16 High)	10	8	
12	L. Collateral (0-1)	0	0	
11	K. Study Site (0–1)	0	0	
10	J. Analyses (0-1)	1	1	
6	I. Independent (0–1)	0	0	
8	H. Dropouts (0-1)	1	1	
7	G. Objective Measurement of Outcomes (0-1)	1	1	
9	F. Follow- Up Rate (0–2)	2	2	
5	E. Follow- Up Length (0–2)	0	0	
4	D. Quality Control (0–1)	1	1	
3	C. Baseline (0-1)	1	0	
2	B. Replicability (0-1)	0	1	
1	A. Study Design (0–3)	3	1	
Quality Criteria [*]	Author (year)	Han et al. (2018); China (and Australia)	Martinez- Miranda et al. (2019); Mexico	-

* MQRS (Methodological Quality Rating Scale) measures 12 dimensions of methodological quality with cumulative ratings ranging from 0 (poorest quality) to 16 (highest quality), with scores of 14 considered high quality (Miller & Wilbourne, 2002).

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