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Editorial board member of *World Journal of Psychiatry*, Dr. Lenin Pavón is an immunologist, having received his PhD at Instituto Politécnico Nacional (Mexico City, Mexico). Since 2000, he has served as Head of the Department of Psychoimmunology at Instituto Nacional de Psiquiatría "Ramón de la Fuente Muñiz" in Mexico City. His career-long work has focused on studying the neuroendocrine-immune interactions in stress, major depression, fibromyalgia, schizophrenia, multiple sclerosis, and Alzheimer's disease. Dr. Pavón's research efforts have yielded a variety of professional awards as well as two books, 88 articles in indexed journals, and 19 chapters in specialized books. At present, he is a member of the editorial board of several international journals and scientific societies. His ongoing research interests involve inflammatory mediators' participation in psychiatric disorders and immunomodulation by neurotransmitters and hormones. (L-Editor: Filipodia)

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Effects of smartphone-based interventions and monitoring on bipolar disorder: A systematic review and meta-analysis

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Abstract

BACKGROUND

Recently, there has been a range of studies about smartphone-based interventions and monitoring for reducing symptoms of bipolar disorder (BD). However, their efficacy for BD remains unclear.

AIM

To compare the effect of smartphone-based interventions and monitoring with control methods in treating patients with BD.

METHODS

A systematic literature search was performed on PubMed, Embase, Clinical trials, psycINFO, Web of Science, and Cochrane Library. Randomized clinical trials (RCTs) or single-group trials in which smartphone-based interventions and monitoring were compared with control methods or baseline in patients with symptoms of BD were included. Data were synthesized using a random-effects or a fixed-effects model to analyze the effects of psychological interventions and monitoring delivered *via* smartphone on psychiatric symptoms in patients with BD. The primary outcome measures were set for mania and depression symptoms. Subgroups were created to explore which aspects of smartphone interventions are relevant to the greater or lesser efficacy of treating symptoms.

RESULTS

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We identified ten articles, including seven RCTs (985 participants) and three single-group trials (169 participants). Analysis of the between-group study showed that smartphone-based interventions were effective in reducing manic [$g = -0.19$, 95% confidence interval (CI): -0.33 to -0.04 , $P = 0.01$] and depressive ($g = -0.28$, 95% CI: -0.55 to -0.01 , $P < 0.05$) symptoms. In within-group analysis, smartphone-based interventions significantly reduced manic ($g = 0.17$, 95% CI: 0.04 to 0.30 , $P < 0.01$) and depressive ($g = 0.48$, 95% CI: 0.18 to 0.78) symptoms compared to the baseline. Nevertheless, smartphone-based monitoring systems significantly reduced manic ($g = 0.27$, 95% CI: 0.02 to 0.51 , $P < 0.05$) but not depressive symptoms. Subgroup analysis indicated that the interventions with psychoeducation had positive effects on depressive ($g = -0.62$, 95% CI: -0.81 to -0.43 , $P < 0.01$) and manic ($g = -0.24$, 95% CI: -0.43 to -0.06 , $P = 0.01$) symptoms compared to the controlled conditions, while the interventions without psychoeducation did not ($P > 0.05$). The contacts between therapists and patients that contributed to the implementation of psychological therapy reduced depression symptoms ($g = -0.47$, 95% CI: -0.75 to -0.18 , $P = 0.01$).

CONCLUSION

Smartphone-based interventions and monitoring have a significant positive impact on depressive and manic symptoms of BD patients in between-group and within-group analysis.

Key Words: Smartphone; Bipolar disorder; Monitoring; Interventions; Meta-analysis; Systematic review

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Core Tip: To date no study has used meta-analysis to pool the existing evidence to examine the efficacy of smartphone-based interventions and monitoring for bipolar disorder. This is the first systematic review and meta-analysis investigating the effects of interventions and monitoring delivered *via* smartphone on bipolar disorder.

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INTRODUCTION

Bipolar disorder (BD) is a serious mood disorder that is characterized by depressive, manic, and mixed episodes^[1]. It is estimated that the lifetime prevalence of BD is 1.0% for bipolar disorder I and 1.1% for bipolar disorder II^[2]. Patients who do not receive adequate and appropriate treatment are then at risk for experiencing plentiful social and occupational impairments, even suicidal behavior^[3].

Fortunately, many available treatments aim to cure manic or depressive episodes and stabilize mood^[4]. However, there is a range of obstacles towards professional BD treatment. The transportation of mental health services and high-quality medications make treatment unaffordable^[5]. Additionally, medication and psychotherapy are not acceptable to some patients who are ashamed of receiving treatment for mood disorder, so the therapeutic effectiveness is limited^[6,7]. Therefore, innovative methods of treatment are urgently needed.

Digital technology may represent a feasible and novel solution. The majority of adults own smartphones. In recent years, the number of smartphone users has continually increased^[8]. Additionally, using smartphones as a tool for psychological treatment is well-accepted for most people because it is cost-effective^[9]. Use of smartphone also breaks the limitation of distance between patients and therapists, therefore it is a viable method to apply smartphones to treat someone who has difficulty in accessing health care^[10]. Meanwhile, privacy and individuation are

guaranteed in the use of smartphones so that treatment acceptance and compliance are enhanced^[11]. Therefore, people can accept smartphone-based interventions at any time they want.

This promising therapeutic potential of smartphones has aroused the interest of different organizations such as governments, advocacy groups, technology companies, and research groups internationally^[12,13]. Recently, there has been a range of articles^[14-23] about smartphone mental health interventions used for reducing symptoms of BD. Although the idea of smartphone use in health care is becoming more popular, it is difficult to separate actual efficacy from overstated aspirational claims. With thousands of mental disorder treatments readily available, it is urgent to find strong evidence to prove it useful, making sure that people have an understanding of smartphone-based interventions for BD.

Recent meta-analyses have suggested that smartphone interventions can have a positive impact on physical diseases, such as diabetes^[24]. Also, the efficacy of smartphone interventions for a series of mood disorders is explained with recent meta-analyses, such as depression, anxiety, and post-traumatic stress disorder^[25-27]. To date, no study has used meta-analysis to pool the existing evidence to examine the efficacy of smartphone interventions for BD. Moreover, there are three reviews related to the smartphone monitoring systems collecting data to predict severity of symptoms^[28-30]. However, the clinical effect of the smartphone-based monitoring system on symptoms of BD patients has yet to be established. More recently, several single-group trials are relevant to the effectiveness of the smartphone-based intervention for BD. However, no systematic review and meta-analysis has examined the efficacy of the smartphone-based intervention and monitoring for symptoms in single-group trials. Nevertheless, there is evidence that the estimates from high-quality single-group trials can overcome the paucity of prospective randomized evidence. The estimated results may be similar to those of randomized controlled trials (RCTs). Pooling of high-quality single-group trials may be as accurate as pooling of RCTs^[31-34].

We conducted the present meta-analysis to provide the first overall estimated effects of smartphone-based interventions and monitoring for reducing symptoms in BD patients. We conducted between-group analyses using the data extracted from the RCTs. We also performed within-group analyses of RCTs and single-group trials. Subgroups were created to explore which aspects of smartphone interventions are relevant to the greater or lesser efficacy of treating symptoms.

MATERIALS AND METHODS

This review and meta-analysis followed the PRISMA statement for the transparency and comprehension of the methodology and results reporting^[35]. The PROSPERO ID of this meta-analysis is CRD42018092539.

Literature search

We searched PubMed, Embase, Clinical trials, psycINFO, Web of Science, and Cochrane Library from 1993 to August 1, 2019. In the case of any other eligible studies, we looked up reference lists of related reviews and articles. Besides, key researchers in this field were asked whether they knew about unpublished trials.

Study selection

The included studies fulfilled the following criteria: (1) All articles in English language; (2) Patients with a diagnosis of BD. Nevertheless, participants with BD and other types of mood disorders that had no influence on the cure of BD at the same time were also included; (3) Intervention: Website, smartphone-based apps, instruction of therapists *via* smartphone to facilitate psychotherapy; (4) RCTs and single-group studies; and (5) Outcomes: Reported either mania or depression symptom severity scores before and after intervention. The exclusion criteria were: (1) The intervention was not clearly defined; (2) Specific outcomes were not reported; (3) Unavailability of full text; and (4) Studies investigating the feasibility and satisfaction of smartphone-based intervention.

Two authors (Jia-Yuan Liu and Kang-Kang Xu) selected the articles according to the inclusion criteria after retrieval and screening of the relevant citations in full text. To identify the applicable articles, they read the abstracts and titles. A third reviewer resolved any disagreement that emerged in the process of searching and selection. The procedure of specific inclusion and extraction is shown in [Figure 1](#).

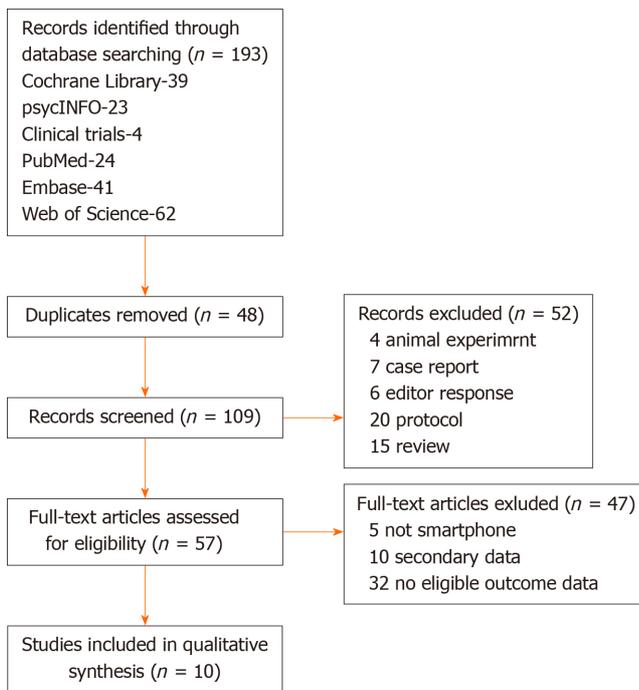


Figure 1 Study selection.

Data extraction

An extraction form was used for each article to collect the following data: (1) Participant information: Sample size, mean age, inclusion criteria, and diagnosis; (2) Study design: Trial quality, whether controlled or not; (3) Smartphone intervention: Study duration, details of intervention, and frequency of intervention; and (4) Effects on mania and depression: Changes in total depressive and manic symptoms scored before and after interventions using any clinically validated rating scale.

Risk of bias

The Cochrane risk of bias assessment tool was used for assessment of the RCT methodological quality. Bias risk in each domain of selection, performance, detection, attrition, and reporting ranking as high, low, or unknown was assessed independently by two reviewers. And inconsistencies were resolved through discussion. For publication bias, it was inappropriate to make a funnel plot to determine it because the number of included trials was no more than 10^[36]. We did not have enough studies to make a meaningful funnel plot that was proposed by Egger *et al*^[37].

Statistical analysis

Statistical analyses were performed with Review Manager version 5.3 (free software downloaded from <http://www.cochrane.org>). Between-group analysis was conducted for RCTs. The differences in changes in manic and depressive symptoms between smartphone-based intervention and control groups were pooled for calculation of the overall effect size using Hedges' g ^[38] with 95% confidence intervals (CIs). We conducted heterogeneity tests among these studies. In case of $I^2 > 50\%$ and $P \leq 0.05$, which indicated the presence of high heterogeneity, we chose a random-effects model; otherwise, we used a fixed-effects model. We next implemented within-group effect size of smartphone-based interventions and monitoring on manic and depressive symptoms. We computed Hedges' g statistic as the estimate of within-group effect size for changes from pre- to post-treatment with RCTs and the single-group trials. Lastly, we conducted subgroup analyses to investigate the different effects of components of smartphone-based interventions.

RESULTS

Study selection

A total of 193 articles were retrieved. Eighty-four duplicate articles were excluded. After reading the abstract and title of the remaining 109, we removed 52 case reports, editor responses, reviews, and studies that included animal experiments. Full versions were retrieved for 57 papers, of which ten met eligibility criteria, including seven RCTs and three single-group trials.

Characteristics of included studies

Table 1 displays the full details of the included studies. Available data were extracted from seven RCTs with two arms^[15-19,21] or three arms^[22] and three trials with single arms^[14,20,23]. All but five articles had graphically reported data, which were extracted with graphical digitizer (Get Data Graph Digitizer) from the reported figures^[14,18,19-21]. Two of the ten eligible studies included patients with BD and patients with other mood disorders^[15,21]. Eight studies included only participants with BD diagnosed according to DSM or ICD-10^[14,16-20,22,23]. The mean age of the sample ranged from 16 to 59 years (median, 38 years). Three studies were related to monitoring systems^[14,18,19]. Three articles highlighted the heterogeneity in the analysis of the depressive symptoms^[14,18,19]. The smartphone interventions and monitoring lasted from 4 wk to 12 mo. Manic and depressive symptoms were measured as an outcome in ten articles.

Risk of bias assessment

The results from the Cochrane Risk of Bias assessment are displayed in Figure 2. The most frequent risk factor for bias was inadequate blinding of participants and personnel with only five of seven studies using the blinding method for which the participants would not be aware of their treatment or control status, as well as the hypothesized outcomes of the trial.

Between-group effect sizes of smartphone interventions on manic and depressive symptoms

The pooled effect size of smartphone-based interventions on manic symptom changes in comparison to control conditions is shown in Figure 3A. Meta-analysis indicated a positive effect size of smartphone-based interventions for reducing manic symptoms compared to control groups ($k = 6$, $n = 785$, $g = -0.19$, 95%CI: -0.33 to -0.04 , $P = 0.01$, $I^2 = 0$). There was no heterogeneity across the studies, therefore, we chose a fixed-effects model. Smartphone-based interventions were significantly more efficacious than control conditions in improving depressive symptoms ($k = 8$, $n = 985$, $g = -0.28$, 95%CI: -0.55 to -0.01 , $P < 0.05$, $I^2 = 75\%$) (Figure 3B). We found high heterogeneity among the studies. Therefore, sensitivity analyses were conducted to explore the reasons for this high heterogeneity. After excluding one article^[18], the heterogeneity was lower, while the pooled effect size was still significant ($k = 6$, $n = 863$, $g = -0.38$, 95%CI: -0.61 to -0.14 , $P < 0.01$, $I^2 = 63\%$).

Within-group effect sizes of smartphone-based interventions on manic and depressive symptoms

In within-group analysis, comparing changes from baseline to post-treatment scores, smartphone-based interventions resulted in significant improvements in manic symptoms ($k = 8$, $n = 914$, $g = 0.17$, 95%CI: 0.04 to 0.30 , $P < 0.01$, $I^2 = 0$) (Figure 4A). Heterogeneity did not exist, and the fixed-effects model was chosen. In within-group analysis, smartphone-based interventions significantly reduced depressive symptoms ($k = 11$, $n = 1154$, $g = 0.48$, 95%CI: 0.18 to 0.78 , $I^2 = 81\%$) after the interventions (Figure 4B). High heterogeneity across article data was found. Therefore, sensitivity analyses were conducted to examine the reasons for the high heterogeneity. After removing two articles^[14,21], the heterogeneity was lowered, whereas the effect size was still significant ($k = 9$, $n = 932$, $g = 0.25$, 95%CI: 0.10 to 0.39 , $I^2 = 15\%$).

Within-group effect sizes of smartphone-based monitoring on manic and depressive symptoms

In order to investigate whether monitoring *via* smartphone had positive effects on manic and depressive symptoms, we performed within-group analyses. Smartphone-based monitoring appeared effective for manic symptoms ($k = 3$, $n = 257$, $g = 0.27$, 95%CI: 0.02 to 0.51 , $P < 0.05$, $I^2 = 0$) compared with changes from baseline to post-treatment scores (Figure 5). The pooled effect size from smartphone-based monitoring

Table 1 Characteristics of included studies

Ref.	Sample type	n	Age (yr)	Female	Study design	Design	Measure	Smartphone intervention details
Depp <i>et al</i> ^[17]	Outpatient diagnosed with BD	41, 41	47.5	58.50%	RCT	10 wk of PRISM (self-management and monitoring app) <i>vs</i> paper and pencil monitoring	YMRS, MADRS	Mood monitoring and self-management APP
Faurholt-Jepsen <i>et al</i> ^[18]	BD patients previously treated for affective disorder	85, 44	43.1	88.40%	RCT	9 mo of MONARCA II (monitoring app) <i>vs</i> controlled conditions	YMRS, HDRS	Self-monitoring APP
Ben-Zeev, <i>et al</i> ^[15]	People with mental illness	75, 74	49	41%	RCT	3 mo of FOCUS (self-management app) <i>vs</i> clinical treat	PSYRATS, BDI-II	App and phone call from specialist to facilitate APP use
Faurholt-Jepsen <i>et al</i> ^[19]	Outpatients diagnosed with BD	33, 34	29.3	67.10%	RCT	6 mo of MONARCA (self-monitoring app) <i>vs</i> Controlled conditions	YMRS, HAMD-17	Self-monitoring APP
Gliddon <i>et al</i> ^[22]	Participants with bipolar disorder	96, 95, 87	39.5	82%	RCT	9 mo of Mood Swings 2.0 <i>vs</i> control conditions	YMRS, MADRS	Mood Swing is a web site
Celano, <i>et al</i> ^[16]	BD patients with an acute depressive episode	14, 11	45	68%	RCT	4 wk of telephone-based positive psychology <i>vs</i> the control group	None, QIDS-SR	Phone call from specialist to instruct the self-management
Kilbourne, <i>et al</i> ^[21]	People with unipolar and bipolar disorder	115, 123	41.3	66%	RCT	12 mo of CCM (psychosocial intervention) <i>vs</i> usual care from provider	None, PHQ-9	Phone call from care manager to facilitate the psychosocial intervention
Depp <i>et al</i> ^[23]	Outpatients with bipolar disorder	10	41	/	Single-group trials	2 wk of PRISM (monitoring and self-management app)	YMRS, MADRS	Mood monitoring and self-management APP
Beiwinkel, <i>et al</i> ^[14]	Outpatients with bipolar disorder	14	47.2	38%	Single-group trials	12 mo of a smartphone-based monitoring system	YMRS, HAMD	Monitoring APP
Miklowitz <i>et al</i> ^[20]	Patients with bipolar disorder	19	37.2	68.40%	Single-group trials	4 mo of FIMM (psychoeducation and monitoring)	YMRS, QIDS-SR	Monitoring and self-management APP

BD: Bipolar disorder; RCT: Randomized clinical trial; ASRM: Altman self-rating mania; APP: Application; BDI: Beck depression inventory; BHS: Beck hopelessness scale; HAMD-17: 17-item hamilton depression rating scale; HRSD: Hamilton rating scale for depression; MADRS: Montgomery-Asberg depression rating scale; PHQ-9: Nine-item patient health questionnaire; PSYRATS: Psychotic symptom rating scales; QIDS-SR: Quick inventory of depressive symptoms-self report; YMRS: Young mania rating scale.

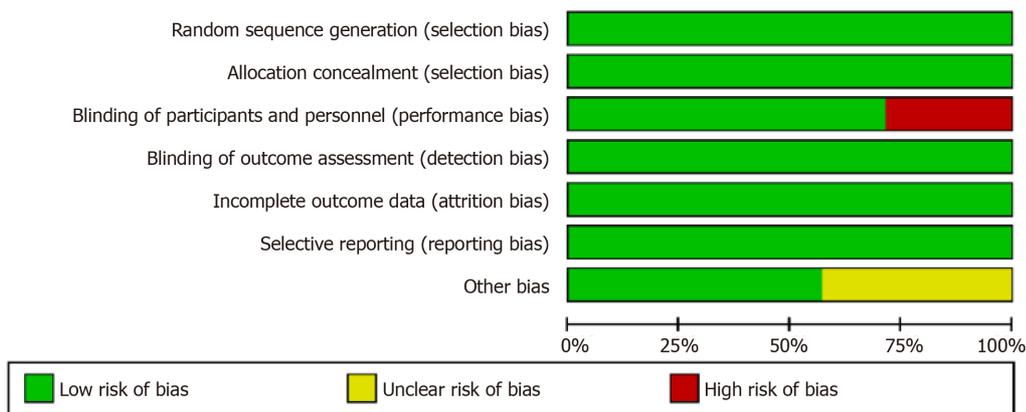


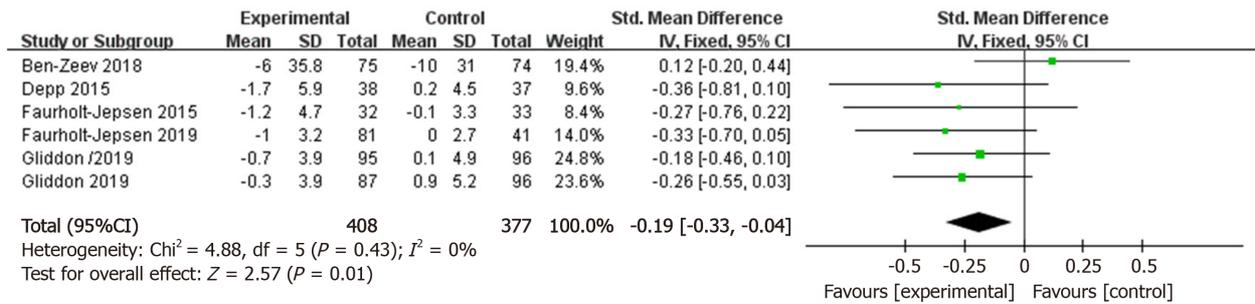
Figure 2 Quality assessments of randomized clinical trials.

on depressive symptoms was near to significant ($k = 3, n = 257, g = 1.26, 95\%CI: -0.14$ to $2.67, P = 0.08, I^2 = 94\%$).

Subgroup analysis

In order to explore which details of smartphone interventions make them effective for

A



B

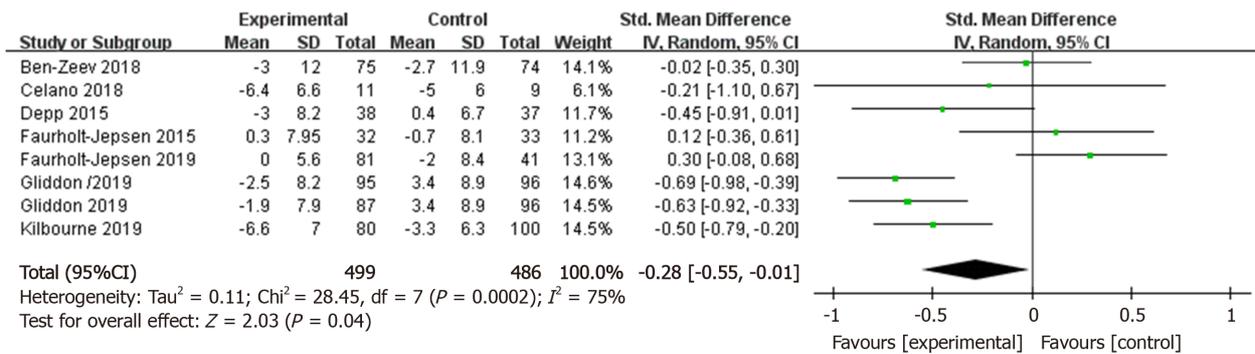


Figure 3 Between-group effect size of smartphone-based interventions on bipolar disorder. A: Meta-analysis of effects of smartphone-based interventions on manic symptoms; B: Meta-analysis of effects of smartphone-based interventions on depressive symptoms.

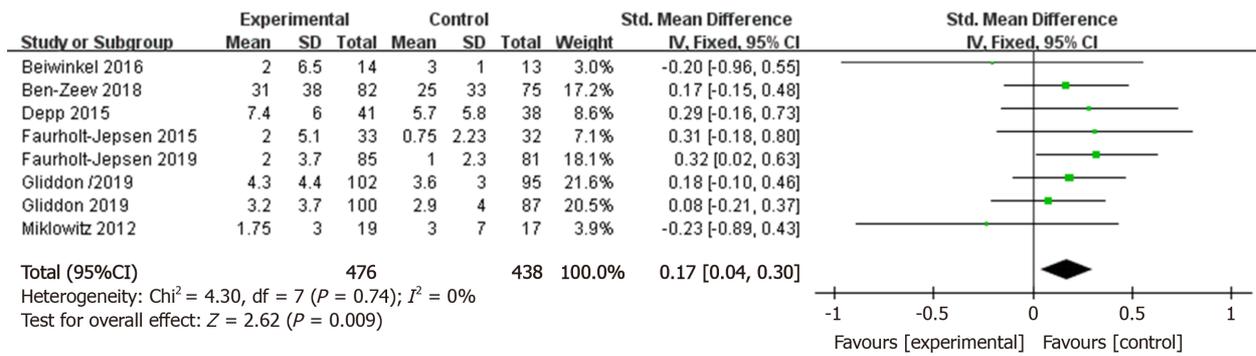
manic and depressive symptoms, we conducted further subgroup analyses. Smartphone interventions that involved psychoeducation had positive effects on manic symptoms ($k = 3$, $n = 449$, $g = -0.24$, 95%CI: -0.43 to -0.06, $P = 0.01$, $I^2 = 0$) compared to control groups, while those without psychoeducation did not ($k = 3$, $n = 336$, $g = -0.13$, 95%CI: -0.43 to 0.17, $P = 0.39$, $I^2 = 45\%$) (Figure 6A). There was no significant subgroup difference ($c^2 = 0.36$, $\text{df} = 1$, $P = 0.55$) in the effect size obtained for smartphone interventions that involved psychoeducation compared to those without. Our meta-analysis revealed a positive effect size of smartphone mental health interventions for reducing depressive symptoms ($k = 3$, $n = 449$, $g = 0.62$, 95%CI: -0.81 to -0.43, $P < 0.01$, $I^2 = 0$) compared to control groups, while those without psychoeducation were not significantly more efficacious than control conditions in improving depressive symptoms ($k = 5$, $n = 536$, $g = -0.06$, 95%CI: -0.39 to 0.26, $P = 0.71$, $I^2 = 67\%$) (Figure 6B). There was a significant subgroup difference ($c^2 = 8.52$, $\text{df} = 1$, $P = 0.004$) in the overall effect size gained for smartphone interventions that involved psychoeducation compared to that did not.

Interventions that involved instruction *via* smartphone had a significantly great effect size on depression outcomes compared to control conditions ($k = 2$, $n = 200$, $g = -0.47$, 95%CI: -0.75 to -0.18, $P = 0.01$, $I^2 = 0$), whereas those without instruction through smartphone did not reduce manic and depressive symptoms compared to control groups ($k = 6$, $n = 785$, $g = -0.24$, 95%CI: -0.58 to 0.10, $P = 0.17$, $I^2 = 82\%$) (Figure 7). There was no significant subgroup difference ($c^2 = 1$, $\text{df} = 1$, $P = 0.32$) in the pooled effect size obtained for interventions that involved instruction *via* smartphone compared to those that did not.

DISCUSSION

As far as we know, this is the first meta-analysis to examine the effects of smartphone-based interventions and monitoring on BD. Ten studies with a total sample of 1028 participants who received smartphone-based treatment or monitoring were used for meta-analysis. The studies were comprehensive in this field. The number of included articles for this meta-analysis was similar to that of a previous meta-analysis that found positive effects from smartphone apps across nine studies with 1837 participants with anxiety^[25]. The literature base for BD appears larger than that for other affective disorders, such as post-traumatic stress disorder, according to a recent meta-analysis in this field that included only five relevant articles^[27]. Furthermore, six

A



B

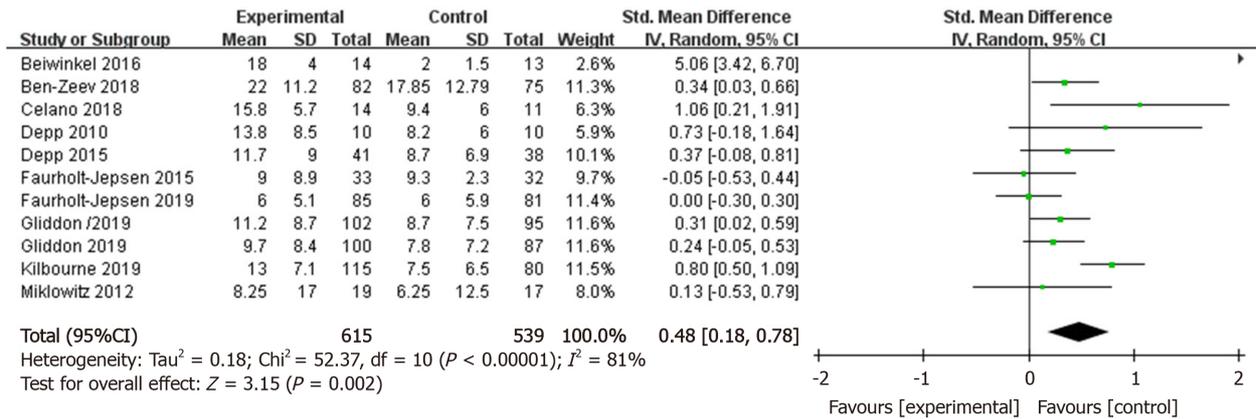


Figure 4 Within-group effect size of smartphone-based interventions on bipolar disorder. A: Meta-analysis of effects of smartphone-based interventions on manic symptoms; B: Meta-analysis of effects of smartphone-based interventions on depressive symptoms.

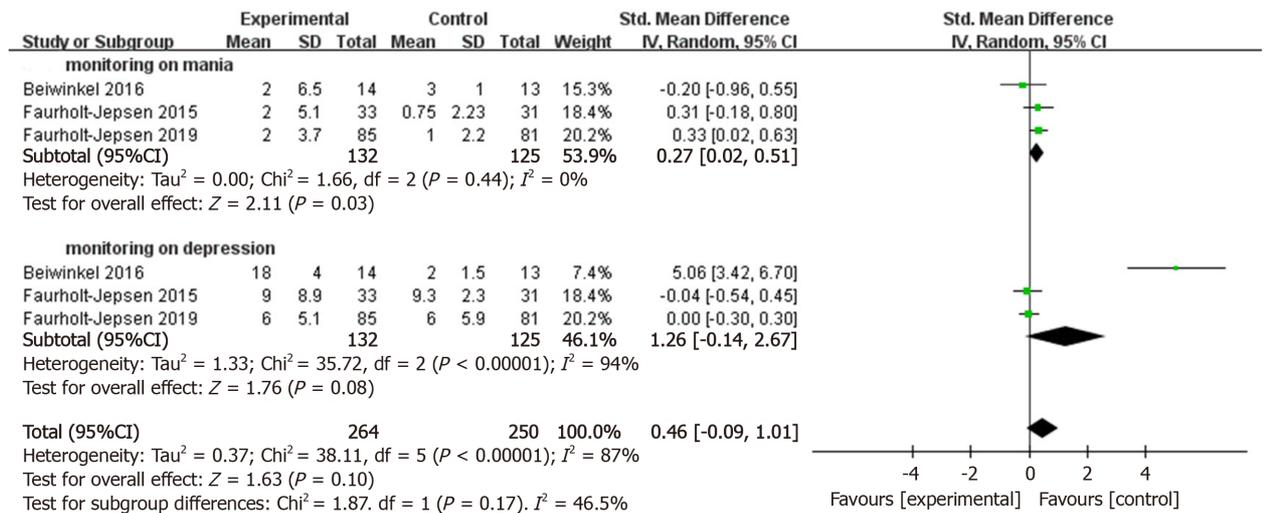
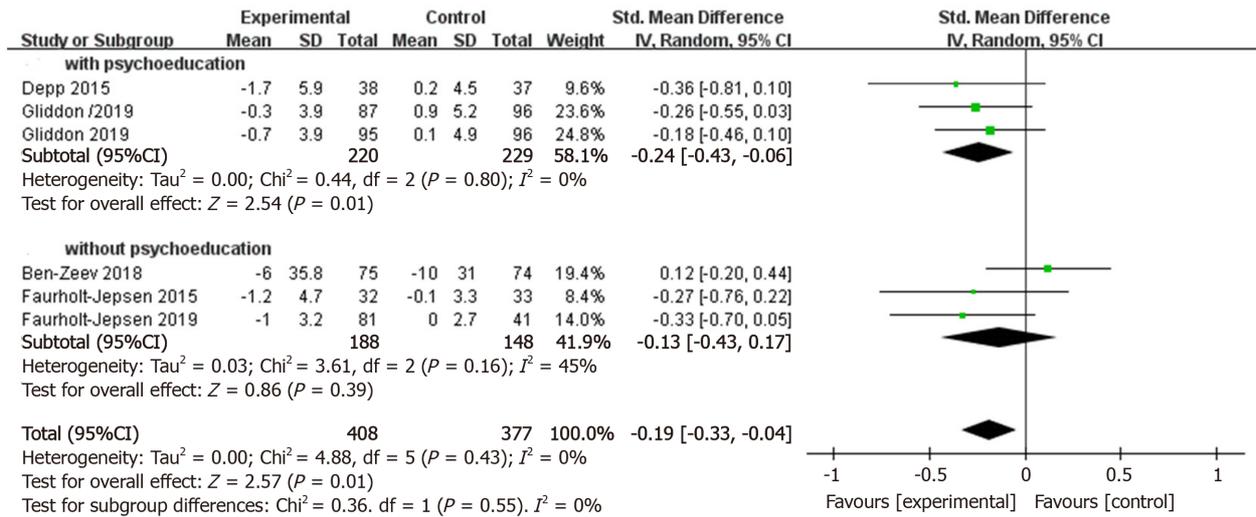


Figure 5 Within-group effect size of smartphone-based monitoring on manic and depressive symptoms.

of our ten articles were published within the past 5 years^[14-16,18,21,22], which means that our analysis reflected the effectiveness of the most updated smartphone-based intervention tools.

Our analysis found that smartphone interventions had a positive effect on BD between and within groups. Meta-analysis indicated a positive effect of smartphone-based interventions for reducing manic ($g = -0.19$) and depressive ($g = -0.28$) symptoms compared to control groups. In within-group analyses, the effects of smartphone-based interventions appeared effective for manic ($g = 0.17$) and depressive ($g = 0.48$) symptoms. Similar positive effects were observed in the meta-analysis of smartphone interventions for other mood disorders, such as anxiety,

A



B

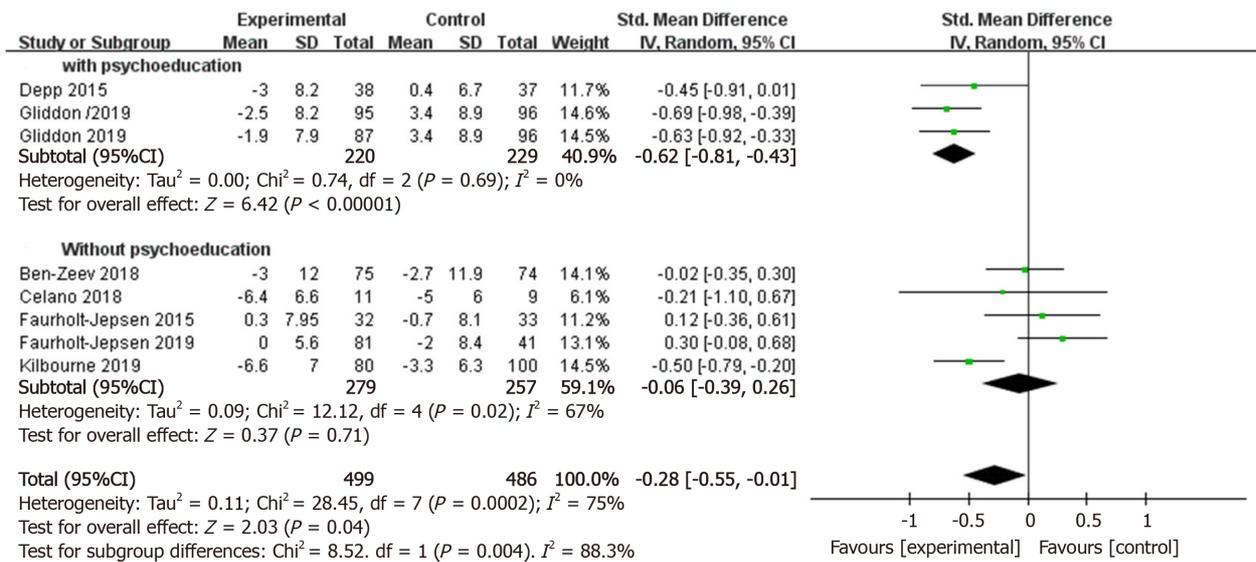


Figure 6 Subgroup analysis of interventions with psychoeducation on bipolar disorder. A: Meta-analysis of effects of smartphone-based interventions on manic symptoms; B: Meta-analysis of effects of smartphone-based interventions on depressive symptoms.

depression, post-traumatic stress disorder, and obsessive compulsive disorder^[25-27,37]. These findings indicate that delivering psychological treatments *via* smartphone devices is an efficacious and promising method to treat BD. The potential mechanism by which smartphone interventions reduce symptoms of BD might be explained by using these devices to support and enhance the delivery of existing face-to-face therapy^[25]. The use of smartphone-based interventions may provide creative and innovative interventions, filling the gap between feasibility and the demand for treatment^[39]. Additionally, interventions *via* smartphone devices may remedy some limitations of existing traditional therapy in the future, with the advantages of convenience and lower treatment cost.

Our analysis showed that BD participants with mania who used the smartphone monitoring systems experienced benefits (g = 0.27). Meanwhile, a recent review of the smartphone application for mood disorders had similar finding that smartphone application with mood monitoring features reduces depressive symptoms^[26]. Additionally, a previous study proposed that using a digital device in itself may produce some psychological benefits^[40]. The smartphone-based monitoring system was used to collect data on phone usage, social communication, social activity, and mobility, which were irrelevant to psychotherapy. To some extent, using the smartphone-based monitoring system was equivalent to only using the device itself. Therefore, our results that monitoring systems have positive effect on manic symptoms may contribute to the hypothesis that smartphones may offer some psychological benefits. There were three reviews related to the smartphone monitoring

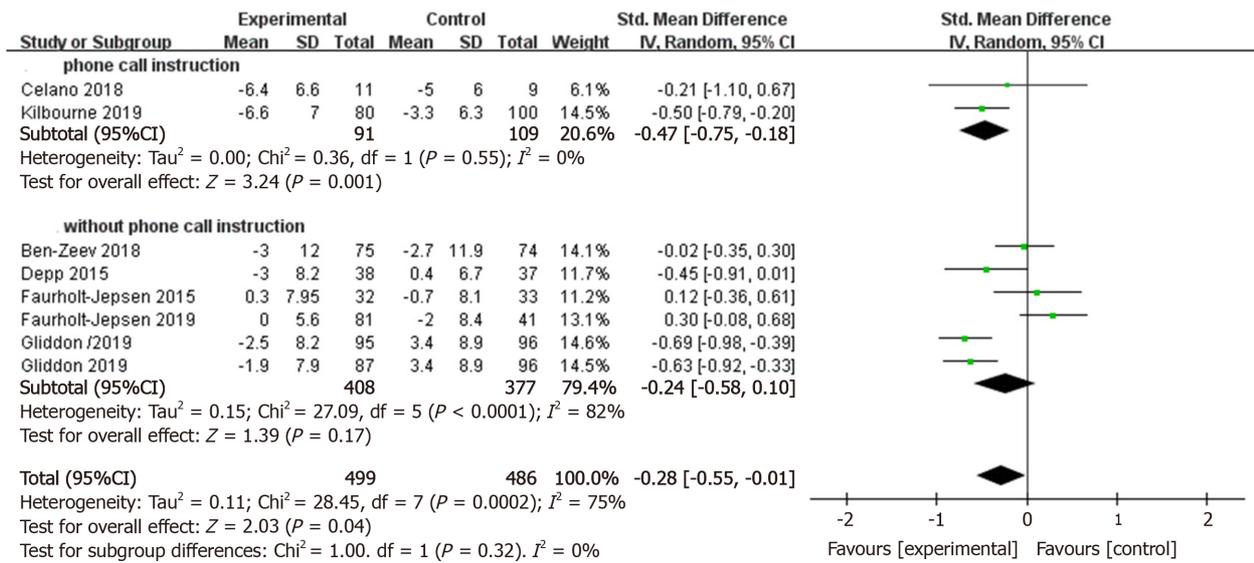


Figure 7 Subgroup analysis of interventions with instruction on depressive symptoms.

system collecting data to predict severity of symptoms^[28-30]. Therefore, we conclude that the monitoring system may contribute to treatment of BD and predictions of clinically assessed depressive and manic symptoms. In the future, using smartphone-based monitoring and treatment simultaneously may form a complete treatment system that is convenient and efficient.

With regards to intervention features, in our subgroup analysis, smartphone interventions that involved psychoeducation had great effects on manic ($g = -0.24$) and depressive ($g = -0.62$) symptoms of BD compared to control conditions. A similar finding was reported in a recent meta-analysis about psychoeducation for the management of BD patients^[41]. The effects of smartphone interventions that involved instruction from the therapist ($g = -0.47$) appeared significant, while the effect size of those without instruction was not significant. A previous study found that therapist guidance bolstered the effectiveness of smartphone interventions^[42]. Instruction from therapists may be conducive to improvement of patient engagement and completion rate of therapy. Although smartphone-based interventions reduced symptoms, there are numerous factors that may affect outcomes of smartphone interventions. Therefore, to design these interventions to be effective in the future, more analyses are needed.

In general, smartphone-based interventions and monitoring reduced symptoms in BD patients. BD is always associated with poor quality of life in addition to personal suffering from depression^[43]. Although there are many treatments for BD, there is a limitation in the availability of therapy. The use of smartphones is accessible and affordable^[44], which ensures the implementation of mental health interventions and monitoring for BD. It enhances and facilitates the delivery of existing therapy programs and makes them more acceptable. However, there are some complex issues generated by the use of smartphones and the technology itself, including low resource settings somewhere, privacy policy, clinical utility, commercialization, and evolving technology^[45-48]. Therefore, in order to achieve satisfactory therapeutic effects in the use of smartphone interventions for BD patients, the influential factors in the whole treatment process should be studied and resolved in the future. Additionally, communication between the users and designers of smartphone-based interventions is not adequate. In the future, the treatment methods will need to be optimized according to the users' feedback and the researchers' thoughts.

Our study had a few limitations. First, two studies in this meta-analysis were considered to have a high risk of bias because of inadequate blinding of participants and personnel^[15,18]. Second, the absolute number of available applications is unknown and the applications are frequently updated. It is impossible to examine total smartphone interventions. Third, there were only three eligible articles regarding smartphone-based monitoring patterns^[14,18,19], which was not sufficient to provide definitive results. Future research that directly tests the monitoring effectiveness against BD would add value to our results. Lastly, to explore potential factors relevant to smartphone interventions, details about demographics, severity of BD, and

engagement need to be collected. Nevertheless, with limited data that can be extracted, we could not conduct subgroup analysis on these variables that might impact statistical results. In the future, a more standard data reporting format needs to be implemented with research data, thus improving the validity and reliability of meta-analysis.

CONCLUSION

In summary, smartphone-based interventions are effective in reducing manic and depressive symptoms. Nevertheless, the smartphone-based monitoring systems only worked for participants with manic but not depressive symptoms. Our results contribute to the literature on smartphone-based interventions and monitoring for manic and depressive symptoms in BD patients, although much work remains to be done. This meta-analysis shows that smartphone-based intervention and monitoring have the potential to enhance the methods of treatment with its characteristics of low expenditure and highly-convenience in addition to available medications and psychological interventions.

ARTICLE HIGHLIGHTS

Research background

Recently, a range of studies about smartphone-based interventions and monitoring for reducing symptoms of bipolar disorder (BD) have been published. However, their efficacy for BD remains unclear.

Research motivation

The present study aimed to assess randomized controlled trials and single-group trials of smartphone-based interventions and monitoring for reducing the symptoms of BD.

Research objectives

The main objective was to update and evaluate innovative treatment suggestions for BD.

Research methods

We performed a systematic literature search on PubMed, Embase, Clinical trials, psycINFO, Web of Science, and Cochrane Library. Randomized clinical trials or single-group trials in which smartphone-based interventions and monitoring were compared with control methods or baseline in patients with symptoms of BD were included. We synthesized data using a random-effects or a fixed-effects model by Review Manager version 5.3 to analyze the effects of psychological interventions and monitoring delivered *via* smartphone on psychiatric symptoms in patients with BD. The primary outcome measures were set for mania and depression symptoms. The subgroups were created to explore which aspects of smartphone interventions are relevant to the greater or lesser efficacy of treating symptoms.

Research results

We identified ten articles, including seven randomized clinical trials (985 participants) and three single-group trials (169 participants). Analysis of the between-group study showed that smartphone-based interventions had positive effects in reducing manic ($g = -0.19$, 95%CI: -0.33 to -0.04, $P = 0.01$) and depressive ($g = -0.28$, 95%CI: -0.55 to -0.01, $P < 0.05$) symptoms. In within-group analysis, smartphone-based interventions significantly reduced manic ($g = 0.17$, 95%CI: 0.04 to 0.30, $P < 0.01$) and depressive ($g = 0.48$, 95%CI: 0.18 to 0.78) symptoms compared to the baseline. Nevertheless, smartphone-based monitoring systems significantly reduced manic ($g = 0.27$, 95%CI: 0.02 to 0.51, $P < 0.05$) but not depressive symptoms. Subgroup analysis indicated that the interventions with psychoeducation were effective for depressive ($g = -0.62$, 95%CI: -0.81 to -0.43, $P < 0.01$) and manic ($g = -0.24$, 95%CI: -0.43 to -0.06, $P = 0.01$) symptoms compared to the controlled conditions, while the interventions without psychoeducation did not ($P > 0.05$). The contacts between therapists and patients that contributed to the implementation of psychological therapy reduced depression symptoms ($g = -0.47$, 95%CI: -0.75 to -0.18, $P = 0.01$).

Research conclusions

Smartphone-based interventions and monitoring have a significant positive impact on depressive and manic symptoms of BD patients in between-group and within-group analysis.

Research perspectives

The current meta-analysis suggests that smartphone-based interventions provide evidence of any reduction in manic and depressive symptoms. Nevertheless, smartphone-based monitoring systems are only effective for participants with manic but not depressive symptoms. The findings have implied that these digital tools can be used as the clinically future treatments for symptoms of BD. However, future trials need to keep pace with the development of these apps and a better understanding of the numerous factors that influence outcomes of smartphone interventions for BD are also required.

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