



The Effects of Patient Education on Psychological Status and Clinical Outcomes in Rheumatoid Arthritis: A Systematic Review and Meta-Analysis

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Wu Z, Zhu Y, Wang Y, Zhou R, Ye X, Chen Z, Li C, Li J, Ye Z, Wang Z, Liu W and Xu X (2022) The Effects of Patient Education on Psychological Status and Clinical Outcomes in Rheumatoid Arthritis: A Systematic Review and Meta-Analysis. Front. Psychiatry 13:848427. doi: 10.3389/fpsyt.2022.848427 **Background:** Rheumatoid arthritis (RA) is a common systemic inflammatory autoimmune disease. The disease has a serious impact on mental health and requires more effective non-pharmacological interventions.

Objective: This study aims to systematically evaluate the effectiveness of patient education on psychological status and clinical outcomes in rheumatoid arthritis.

Methods: This systematic review and meta-analysis was conducted based on the Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) guidelines. PubMed, Cochrane Library, EMBASE database, and Web of Science database were screened for articles published until November 2, 2021. Randomized controlled trials (RCTs) of patient education for RA were included. Outcomes measures included pain, physical function, disease activity, erythrocyte sedimentation rate (ESR), C-reactive protein (CRP), anxiety, depression, Arthritis Self-Efficacy (pain, other symptoms, total), and General health. For each outcome, standardized mean differences or mean differences and 95% confidence intervals (CIs) were calculated.

Results: A total of 24 RCTs (n = 2,276) were included according to the inclusion and exclusion criteria. Meta-analysis revealed a statistically significant overall effect in favor of patient education for physical function [SMD = -0.52, 95% CI (-0.96, -0.08), $l^2 = 93\%$, P = 0.02], disease activity [SMD = -1.97, 95% CI (-3.24, -0.71), $l^2 = 97\%$, P = 0.002], ASE (pain) [SMD = -1.24, 95% CI (-2.05, -0.43), $l^2 = 95\%$, P = 0.003], ASE (other symptoms) [SMD = -0.25, 95% CI (-0.41, -0.09), $l^2 = 25\%$, P = 0.002], ASE (total) [SMD = -0.67, 95% CI (-1.30, -0.05), $l^2 = 90\%$, P = 0.03], and general health [SMD = -1.11, 95% CI (-1.36, -0.86), $l^2 = 96\%$, P < 0.0001]. No effects were found for anxiety [SMD = 0.17, 95% CI (-0.64, 0.98), $l^2 = 82\%$, P = 0.68], depression [SMD = -0.18, 95% CI (-0.52, 0.15), $l^2 = 52\%$, P = 0.28], pain [SMD = -0.37, 95% CI (-0.80, 0.05), $l^2 = 89\%$, P = 0.08], and CRP [SMD = -0.27, 95% CI (-0.57, 0.02), $l^2 = 0\%$, P = 0.07].

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Conclusions: Patient education may be effective in improving clinical outcomes and psychological status in patients with rheumatoid arthritis. Considering the methodological limitations of the included RCTs, more high-quality and large-sample RCTs are needed to confirm this conclusion in the future.

Systematic Review Registration: http://www.crd.york.ac.uk/prospero, identifier: CRD42021250607.

Keywords: patient education, psychotherapy, rheumatoid arthritis, meta-analysis, systematic review

INTRODUCTION

Rheumatoid arthritis (RA) is an autoimmune disease characterized by chronic inflammation, which can cause joint destruction, deformity, pain, and dysfunction (1, 2). This chronic disease has led to a decline in patient's physical function, quality of life, and workability, as well as an increase in medical expenses (3, 4), which brings a heavy burden to individuals and society (5). At present, RA has become one of the main global public health problems, affecting nearly 1% of the world's population (6). RA is clinically incurable, but antirheumatic drugs and biological agents can control symptoms and improve inflammation (7). However, the efficacy of drugs was affected by patients' adherence with medications (8), and oral medications have brought many adverse reactions to patients (9). Studies have found that about 12%-17% of patients have adverse drug events after discharge (10). Therefore, doctors and patients are often looking for more beneficial non-pharmacological interventions. In addition, some studies have found that 30 to 80% of patients with rheumatic and chronic musculoskeletal diseases do not adhere to treatment plans, that poor patient adherence to treatment can affect treatment outcomes, and that poor adherence is associated with reduced functioning and health-related quality of life (11-13). Therefore, strategies to improve patient adherence are critical to improving the effectiveness of clinical interventions (14). Several studies have made recommendations to increase patient adherence, such as patient-centredness, the inclusion of patients in treatment decisions, and patient participation in shared decision-making may be critical factors in improving adherence (13). Some studies suggest that individualized patient education improves patient adherence (15, 16).

Patient education is a low-cost intervention with no side effects, and it has been accepted by patients, family members, and medical workers. At present, the educational intervention has become an effective supplement to traditional medical treatment, which aims to support and help patients with RA to strengthen their life and health management (17). Previous studies have found that educational interventions can increase awareness of patients with RA about the disease and treatment methods, thereby improving their medication adherence (17, 18). Other studies have reported that educational intervention may have a positive effect on the control of disease activity (19, 20). At the same time, educational intervention can improve the health, pain, swollen joint count, tender joint count, and physical function of patients with RA (21). However, other studies have shown that the effect of educational intervention on disease control was not yet clear, the short-term and long-term effects may be inconsistent (22, 23).

According to previously reported studies, the effectiveness of patient education interventions is still controversial. It is necessary to conduct a systematic review and meta-analysis to evaluate its efficacy. Although some previous studies have summarized the effect of patient education on rheumatoid arthritis using a systematic review approach, this study did not perform a meta-analysis (24, 25). Another meta-analysis summarized the impact of patient education on rheumatoid arthritis, reporting outcomes including disability, tender joint count, depression, general health, and psychological status. The study found that patient education had a short-term effect on rheumatoid arthritis and no long-term effects. The literature included in this study was mainly published before 2001, which was published a long time ago (26). In recent years, due to the growing interest of researchers in patient education, many randomized controlled trials of patient education in the treatment of rheumatoid arthritis have been published. It is necessary to recapitulate and update this evidence based on the latest published literature. This meta-analysis aims to review and analyze the effectiveness of patient education in the treatment of RA. Several variables were compared, including pain, physical function, disease activity, erythrocyte sedimentation rate (ESR), C-reactive protein (CRP), anxiety, depression, Arthritis Self-Efficacy (pain, other symptoms, total), and general health.

METHODS

This study was conducted according to the PRISMA guidelines and the recommendations of the Cochrane Collaboration (27). All analyses were based on previously published studies, and ethical approval was not required in this review. Systematic Review Registration: http://www.crd.york.ac.uk/prospero, identifier: CRD42021250607.

Search Strategy

We searched all clinical studies published in PubMed, Cochrane Library, Embase, and Web of Science database before November 2, 2021. Search terms such as the following were used: "Arthritis, Rheumatoid," "Rheumatoid Arthritis," "Education," "Educational Activities," "Training Programs," "Workshops," "Randomized Controlled Trial," "Clinical Trial," "Randomly," "Randomised," and "random allocation." We conducted a detailed search for each database according to the search methods of the different databases. The **Supplementary Appendix** described the search strategy of each database in detail. Four researchers screened the retrieved documents according to the inclusion criteria and exclusion criteria to read the title, abstract, and full text. The disagreements between the four researchers were discussed with the fifth researcher until a consensus was reached.

Selection Criteria

Patients

This study included patients with RA, and the diagnostic criteria included diagnosis based on doctors (Physician diagnosed), or the diagnostic criteria of the American College of Rheumatology (ACR). There were no restrictions on the patients' age, gender, course of the disease, and where RA occurs.

Interventions

Patient education is defined as a planned and systematic educational activity aimed at improving the health of patients, such as providing disease-related information, health consultation, behavior guidance, behavior modification and advice (24, 28). These methods are aimed at improving the patient's experience of their disease, raising awareness of the disease, promoting the patient's healthy behavior, and improving the patient's ability to deal with the disease. These educational activities can be carried out verbally, in writing, or remotely (such as by telephone). Patient education for patients with rheumatoid arthritis mainly includes providing patients with rheumatoid arthritis with disease-related information, treatment methods, coping strategies for disease symptoms, suggestions and guidance on daily activities, and other activities to improve patients' disease knowledge and health behavior. These educational activities are planned and systematic. We also excluded studies that systematically taught patients to exercise or exercise therapy with the primary goal of exercise or increasing their exercise adherence. However, some suggestions for exercise methods can be used as part of the intervention component of patient education. There were no restrictions on the duration, frequency, and specific methods of educational intervention.

Comparisons

In this study, the types of interventions in the intervention and control group included education vs. usual care, education + usual care vs. usual care, education + conventional treatment vs. conventional treatment, and education vs. waiting list (no intervention). Conventional treatment includes other treatments that patients used before entering the study, such as medications or other routine treatments. No intervention mainly refers to that the control group patients were in the waiting list group and did not receive any intervention before the end of the study, while the same interventions as the intervention group were used after the end of the study.

Outcomes

For inclusion in this review, RCTs had to assess at least one outcome, and the outcome parameters in the respective studies had to be the primary outcomes:

1. Pain. Pain was measured using the visual analog scale (VAS), or Arthritis Impact Measurement Scales 2 (AIMS 2).

- 2. Physical function. Physical function was measured using the Health Assessment Questionnaire (HAQ), Arthritis Impact Measurement Scales 2 (AIMS2), or the Short-Form Health Survey (SF-36).
- 3. Disease activity. Disease activity was measured using the disease activity score 28 (DAS-28) or Ritchie Articular Index (RAI).
- 4. Erythrocyte sedimentation rate (ESR).
- 5. C-reactive protein (CRP).
- 6. Anxiety. Anxiety was measured using the Hospital Anxiety and Depression Scale (HADS) or State-Trait Anxiety Inventory (STAI).
- 7. Depression. Depression was measured using the Hospital Anxiety and Depression Scale (HADS), Beck Depression Inventory (BDI), or Center for Epidemiologic Studies Depression Scale (CES-D).
- 8. Arthritis Self-Efficacy (pain, other symptoms, total). ASE was measured using the Arthritis Self-Efficacy scale (ASE).
- 9. General health. General health was measured using the Short-Form Health Survey (SF-36), EuroQol Five Dimensions Questionnaire (EQ5D), or Arthritis Impact Measurement Scales 2 (AIMS2).

Studies Types

This study included only randomized controlled trials (RCTs). Observational studies, non-randomized controlled trials, and reviews were not included. The language of the included studies was limited to English.

Data Extraction and Quality Assessment

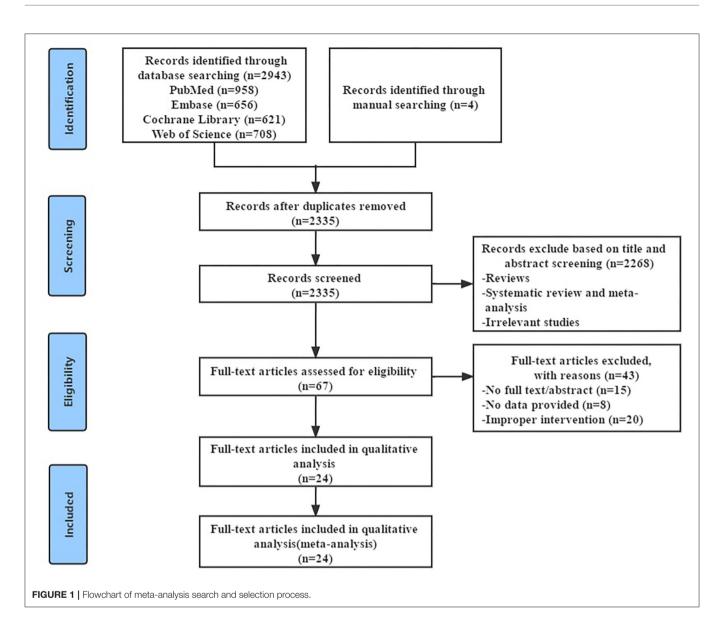
Four reviewers independently extracted study data from eligible studies according to a prespecified study protocol, including the characteristics of the researchers (e.g., name and country), patient characteristics (e.g., age, gender, and duration of disease), research characteristics (e.g., study design, publication years, sample size, the frequency and duration of intervention), and study outcomes. When the follow-up time was inconsistent between studies, we chose to include the final follow-up time. The disagreements between the four researchers were discussed with the fifth researcher until a consensus was reached.

Assessment of Risk of Bias in Included Studies

Two reviewers independently used the Cochrane risk-of-bias tool to assess the quality and risk of bias of the included studies, which included the following domains: selection bias (random sequence generation and allocation concealment), performance bias (blinding of participants and personnel), detection bias (blinding of outcome assessment), attrition bias (incomplete outcome data), reporting bias (selective reporting), and other bias (29). The evaluation results were examined by a third reviewer, and the disagreed evaluations were further discussed until a consensus was reached.

Rating Quality of Evidence

The Grading of Recommendations, Assessment, Development and Evaluation (GRADE) system was used to evaluate the quality of evidence for each outcome. The strength of the evidence was



categorized as high, moderate, low, or very low. Two reviewers independently used the GRADE system to assess the quality of evidence. The disagreements between the two researchers were discussed with the third researcher until a consensus was reached.

Statistical Analysis

We conducted this meta-analysis of the included literature by Review Manager 5.3 software (Cochrane Collaboration, Oxford, UK) and illustrated the results of data merging intuitively with a forest map. The mean differences (MDs), standard mean differences (SMDs), and 95% confidence intervals (CIs) were calculated by random-effects models or fixedeffects models. The heterogeneity between various studies was statistically analyzed by I^2 and chi-square tests. Significant heterogeneity was indicated when $I^2 \ge 50\%$ or P < 0.1, and the random-effects models were used. When $I^2 < 50\%$ or P > 0.1 showed no significant heterogeneity, the fixedeffects models were used. When there was heterogeneity among various studies, a subgroup analysis was conducted according to the type of interventions. We used metaregression and sensitivity analysis to explore the sources of heterogeneity. In addition, publication bias was assessed using Egger's and Begg's tests (30). *P*-values <0.05 were considered statistically significant.

RESULTS

Study Selection

The literature search strategy was detailed in the **Supplementary Appendix**, and the screening process was summarized in **Figure 1**. We searched the four English databases (PubMed, Cochrane Library, Embase, and Web of Science database). Initially, we retrieved a total of 2,947 potentially

TABLE 1 | Study characteristics.

References	Population	Country	Years	Study design	Mean age	(SD), years	Samp	le size	Male/fe	emale	Symptom dura	ition in years (SI
					EG	CG	EG	CG	EG	CG	EG	CG
Helewa et al. (31)	RAc	Canada	1991	RCT	52.7 (12.6)	55.3 (11.8)	53	52	Not rep	orted	Not	reported
Barlow et al. (32)	RA ^c	The U.K.	1997	RCT	58.62 (11.25)	60.04 (10.82)	53	55	9/44	11/44	14.62 (11.49)	17.04 (12.29)
Scholten et al. (33)	RA ^b	Austria	1999	RCT	48.3	(5.6)	38	30	14/5	54	8.9	9 (1.2)
Hill et al. (34)	RA ^b	The U.K.	2001	RCT	63 (Not reported)	62 (Not reported)	51	49	17/34	10/39	13 (Not reported)	12 (Not reported)
Riemsma et al. (35)	RA ^b	Netherlands	2003	RCT	55.1 (10.3)	57.0 (8.3)	71	76	24/47	29/47	11.7 (11.1)	11.4 (8.9)
Hammond et al. (36)	RA ^c	The U.K.	2004	RCT	53.9 (13.9)	57.1 (13.5)	162	164	41/121	49/115	9.0 (7.7)	9.9 (8.8)
Kirwan et al. (37)	RA ^b	The U.K.	2005	RCT	56.4 (10.17)	57.1 (10.71)	30	28	11/19	7/21	13.2 (12.72)	16.7 (12.18)
Montserrat et al. (38)	RA ^b	Spain	2006	RCT	55.40 (16.32)	51.09 (16.62)	22	21	14/8	15/6	21.50 (15.30)	19.47 (16.09)
Masiero et al. (39)	RA ^b	Italy	2007	RCT	54.2 (9.8)	52.5 (11.9)	36	34	7/29	6/28	14.8 (8.8)	16.1 (8.3)
Giraudet-Le Quintrec et al. (40)	RA ^b	France	2007	RCT	55.32 (11.08)	54.31 (14.37)	104	104	15/89	16/88	11.85 (9.44)	14.25 (10.27)
Lovisi et al. (41)	RA ^b	Brazil	2009	RCT	45.71 (10.50)	46.20 (9.52)	28	30	2/26	5/25	9.43 (9.10)	9.41 (7.95)
Macedo et al. (42)	RA ^b	Australia	2009	RCT	48.63 (11.56)	52.56 (7.65)	16	16	1/15	1/15	11.63 (9.95)	8.38
Mathieux et al. (43)	RA ^b	France	2009	RCT	48.3 (13.0)	47.0 (13.2)	30	30	8/22	9/21	Noti	reported
Conn et al. (44)	RA ^b	The U.S.	2013	RCT	54.2 (8.2)	52.9 (10.2)	52	52	11/41	11/41	9.1 (Not reported)	6.4 (Not reported)
Shigaki et al. (45)	RAc	The U.S.	2013	RCT	50.3 (11.6)	49.3 (12.3)	54	54	4/50	6/48	7.4 (8.6)	8.5 (10.3)
Yousefi et al. (46)	RA ^b	Iran	2015	RCT	42.9 (13.24)	46.6 (10.97)	100	106	14/86	11/95	7.6 (5.18)	6.51 (5.28)
Pot-Vaucel et al. (47)	RA ^d	France	2016	RCT	58.2 (10.7)	62.4 (9.8)	28	26	Not rep	orted	11.6 (9.4)	14.5 (3.0)
Anvar et al. (48)	RA ^c	Iran	2018	RCT	69.03 (No	t reported)	37	39	0/37	0/39	Not	reported
Hosseini Moghadam et al. (49)	RA ^a	Iran	2018	RCT	48.06 (10.51)	48.87 (9.24)	32	32	Not reported	Not	reported	

(Continued)

References	Population	Country	Years	Study design	Mean age (SD), years	SD), years	Samp	Sample size	Male/female	male	Symptom dura	Symptom duration in years (SD)
					Ð	g	ß	g	Ð	90	G	GO
Saeedifar et al. (50)	RA°	Iran	2018	RCT	44.27 (11.35)	11.35)	30	30	Not reported	orted	9.1 (Not	9.1 (Not reported)
Zhao et al. (51)	RA ^a	China	2019	RCT	56.93 (11.14)	54.15 (10.06)	46	46	12/34	14/32	Not r	Not reported
Shao et al. (52)	RA°	China	2020	RCT	60.40 (12.04)	56.59 (7.12)	15	17	5/10	4/13	9.57 (9.74)	10.88 (7.32)
Song et al. (53)	RA ^a	China	2020	RCT	57.05 (11.31)	53.22 (10.04)	41	36	11/30	11/25	Not re	Not reported
Shao et al. (54)	RA°	China	2021	RCT	58.2 (11.3)	59.5 (11.9)	112	112	18/94	14/98	10.2 (8.1)	11.1 (8.9)

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relevant records and excluded 612 duplicate records. After reading the title and abstract, we excluded 2,268 obviously irrelevant records, leaving 67 research need to read the full text for further confirmation. After reading the full text, 43 studies that did not meet the inclusion criteria were excluded, 24 RCTs were retained. Finally, 24 RCTs (31–54) were included, with a total of 2,276 patients with RA.

Study Characteristics

Overview of Included Studies

The study characteristics of the 24 RCTs can be observed in Table 1. These studies were published from 1991 to 2021. The 24 RCTs included were conducted in different countries, with two in the United States (44, 45), three in France (40, 43, 47), four in China (51-54), four in United Kingdom (32, 34, 36, 37), four in Iran (46, 48-50), one each in the Canada (31), Austria (33), Netherlands (35) Spain (38), Italy (39), Brazil (41), and Australia (42). Of the 24 studies included, 2,276 patients with RA were analyzed. The number of patients with RA in each study ranged from 32 to 258, with a total of 1,155 patients in the education group and 1,121 in the control group. Twentyfour studies reported the age of patients with RA. Their mean age ranged from 44.27 to 69.03 years. Eighteen studies reported the duration of symptoms, with the mean duration ranged from 7.05 to 20.48 years. Sixteen studies were based on the diagnosis of KOA based on clinical and radiographic features by a physician (33-35, 37-44, 46, 47, 49, 51, 53), while the remaining eight studies were based on the diagnosis criteria of the American College of Rheumatology (31, 32, 36, 45, 48, 50, 52, 54).

Intervention Characteristics and Outcome Measures

Table 2 shows the characteristics of interventions in these 24 RCTs, including the specific methods of education, duration of intervention, and outcomes. To compare interventions between the education group and the control group, eight studies used education vs. usual care (32, 35, 37, 44, 48, 51, 52, 54), and five studies used education + usual care vs. usual care (36, 40, 42, 46, 53), five studies used education + conventional treatment vs. conventional treatment (34, 38, 39, 49, 50), and six studies used education vs. waiting list (no intervention) (31, 33, 41, 43, 45, 47). The outcomes of these 24 RCTs include pain, physical function, disease activity, erythrocyte sedimentation rate (ESR), C-reactive protein (CRP), anxiety, depression, ASE (pain), ASE (other symptoms), ASE(total), and general health. Twelve studies that assessed pain used VAS (10 cm) scores (32, 34, 37, 38, 41, 50), VAS (100 mm) scores (36, 39, 42, 46), and AIMS2 scores (35, 45), respectively. Fifteen studies assessed physical function using HAQ (31, 33, 36-44, 51, 54), SF-36 (46) and AIMS2 (35), respectively. Eight studies assessed disease activity using DAS-28 (35, 40, 42, 51-54) and RAI (39), respectively. Five studies assessed the anxiety using HADS (32, 37, 40) and STAI (41, 47), respectively. Five studies assessed the depression using HADS (32, 37, 40), BDI (31, 33, 41, 47), and CES-D (45), respectively. Ten studies assessed ASE using Arthritis Self-Efficacy Questionnaire (32, 35-37, 45, 48, 49, 51, 52, 54). Four studies assessed the general health using SF-36 (41, 46), AIMS2 (45) and EQ5D (42), respectively.

TABLE 2 | Intervention, main measures, and results.

References	Specific methods of education	Intervention length, fr	equency, and duration	Main outcomes and results
		Education group	Control group	
Helewa et al. (31)	Patients were provided with joint protection education, daily living skills and coping strategies	Education (6 weeks)	Waiting list	1. Physical function (HAQ); 2. Depression (Beck)
Barlow et al. (32)	Patients were educated through leaflets that include information about rheumatoid arthritis disease, disease management and medication	Education (3 weeks)	Usual care	1. Pain (VAS-10 cm); 2. Anxiety (HADS); 3. Depression (HADS); 4. ASE (pain); 5. ASE (other symptoms)
Scholten et al. (33)	Multidisciplinary education of patients through lectures provides information on disease and medication, pain management and joint protection, and coping skills for disease symptoms and sequelae	Education (6 weeks)	Waiting list	1. Physical function (HAQ); 2. Depression (BDI)
Hill et al. (34)	Educational curricula have been developed for patients, including information on rheumatoid arthritis, treatment medications, pain management strategies, joint protection skills, exercise advice and other daily coping strategies	Education (30 min each; 7 times in total; 24 weeks) + Conventional treatment	Conventional treatment	1. Pain (VAS-10 cm); 2. CRP
Riemsma et al. (35)	Group education was used to provide educational courses for patients, and education-related pamphlets and tapes were distributed to provide patients with information about diseases and treatments, as well as various coping strategies related to diseases, so as to promote patients to strengthen the management of diseases	Education (2.5-h each; Five times a week; 6 weeks)	Usual care	1. Pain (AIMS); 2. Physical function (AIMS); 3. ASE (pain); 4. ASE (other symptoms); 5. Disease activity (DAS-28)
Hammond et al. (36)	Basic information on rheumatoid arthritis and its treatment and management, coping strategies for daily life, exercise and advice on joint protection were provided to patients through education	Education (1 times per week; 5 weeks) + Usual care	Usual care	1. Pain (VAS-100 mm); 2. Physical function (HAQ); 3. ASE (total)
Kirwan et al. (37)	Education was provided to patients through courses, including knowledge of the disease, advice and guidance on coping strategies in daily life, pain management, emotional management, joint protection, drug use, and so on	Education (2.5-h each; The first, second, third, fourth and eighth weeks; Five times in total)	Usual care	1. Pain (VAS-10 cm); 2. Physical function (HAQ); 3. Anxiety (HADS); 4. Depression (HADS); 5. ASE (pain); 6. ASE (other symptoms)
Montserrat et al. (38)	Through a combination of individual and group education, rheumatoid arthritis knowledge, joint protection methods, exercise advice and pain management strategies were taught to patients	Education (30 min each; Once every 3 months; 12 weeks) + Conventional treatment	Conventional treatment	1. Pain (VAS-10 cm); 2. Physical function (HAQ); 3. ESR; 4. CRP
Masiero et al. (39)	Through lectures, conferences and pamphlets, patients were taught about the disease, the mechanism of pain and other symptoms, as well as information about exercise and pain management	Education (6-h each; Once every 3 weeks; 12 weeks) + Conventional treatment	Conventional treatment	 Pain (VAS-100 mm); 2. Physical function (HAQ); Disease activity (RAI)
Giraudet-Le Quintrec et al. (40)	Through multidisciplinary education, patients were taught information about the disease, diet, exercise and treatment methods, as well as coping strategies for certain diseases	Education (6-h each; Eight times a week; 72weeks) + Usual care	Usual care	1. Physical function (HAQ); 2. Anxiety (HADS); 3. Depression (HADS); 4. Disease activity (DAS-28)

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TABLE 2 | Continued

References	Specific methods of education	Intervention length, fr	equency, and duration	Main outcomes and results
		Education group	Control group	
Lovisi et al. (41)	Comprehensive information on rheumatoid arthritis, including etiology, pathogenesis, disease management, drug therapy and rehabilitation, was taught to patients through meetings	Education (1-h each; Once a week; 6 weeks)	Waiting list	1. Pain (VAS-10 cm); 2. Physical function (HAQ); 3 Anxiety (STAI); 4. Depression (BDI); 5. General health (SF36)
Macedo et al. (42)	Provide patients with information about rheumatoid arthritis, medications, and coping strategies for their daily lives	Education (1 times per week; 5 weeks) + Usual care	Usual care	 Pain (VAS-100 mm); 2. Physical function (HAQ); Disease activity (DAS-28); 4. General health (EQ5D); 5. ESR
Mathieux et al. (43)	Provide multidisciplinary education to patients, including information on disease and treatment, advice and guidance on joint protection	Education (24 weeks)	Waiting list	1. Physical function (HAQ)
Conn et al. (44)	Provide educational manuals for patients to increase their understanding of the disease and treatment, provide coping strategies for the disease, and increase patients' management of the disease	Education (2-h each; 1 times per week; 6 weeks)	Usual care	1. Physical function (HAQ)
Shigaki et al. (45)	Provide educational courses for patients, including the causes of illness, treatment of illness, and pain management	Education (6 weeks)	Waiting list	1. Pain (AIMS2); 2. Depression (CES-D); 3. ASE (total); 4. General health (AIMS2)
Yousefi et al. (46)	Provide multidisciplinary education and educational brochures to patients to increase their awareness of rheumatoid arthritis, provide pain management methods and joint protection methods, and enhance patients' disease management skills	Education (5-h each; 1 times per week; 6 weeks) + Usual care	Usual care	1. Pain (VAS-100 mm); 2. Physical function (SF36) 3. General health (SF36)
Pot-Vaucel et al. (47)	To provide patients with information about the disease and treatment, and to increase their understanding of the disease and treatment, and to provide disease management skills and medication information	Education (24 weeks)	Waiting list	1. Anxiety (STAI); 2. Depression (Beck)
Anvar et al. (48)	A multidisciplinary group education approach was used to improve patient awareness of all aspects of the disease, coping strategies for pain management and daily activities, and information on diet and exercise	Education (1–1.5 h each; 6 times per week; 6 weeks)	Usual care	1. ASE (pain);
Hosseini Moghadam et al. (49)	Group education for patients with rheumatoid arthritis includes providing coping strategies for the disease, guidance on daily activities and precautions to improve their knowledge of RA	Education (30 min each; Twice a week; 8 weeks) + Conventional treatment	Conventional treatment	1. ASE (pain); 2. ASE (other symptoms); 3. ASE (total)
Saeedifar et al. (50)	Patients were provided with information about rheumatoid arthritis, joint protection methods, pain management strategies, methods to prevent a recurrence, appropriate exercise patterns, and enhanced management of the disease	Education (1–2 h each; 1 times per week; 24 weeks) + Conventional treatment	Conventional treatment	1. Pain (VAS-10 cm)

(Continued)

References	Specific methods of education	Intervention length, fre	Intervention length, frequency, and duration	Main outcomes and results
		Education group	Control group	
Zhao et al. (51)	The clinical guidelines on rheumatoid arthritis were taught by telephone, including basic information on rheumatoid arthritis, as well as medication, diet and exercise	Education (20–40 min each; Once every 3 weeks; 12 weeks)	Usual care	 Physical function (HAQ); 2. ASE (total); 3. Disease activity (DAS-28)
Shao et al. (52)	Through the course, patients will be provided with disease-related knowledge and information, coping skills and joint protection strategies to enhance patients' management of disease	Education (6 weeks)	Usual care	1. Disease activity (DAS-28); 2. ASE (pain); 3. ASE (other symptoms)
Song et al. (53)	Educational courses were provided to patients over the phone, which included knowledge and skills in rheumatoid arthritis, treatment methods, information on medication, exercise and diet	Education (20-40 min each; Once every 3 weeks; 12 weeks) + Usual care	Usual care	1. Disease activity (DAS-28); 2. ESR
Shao et al. (54)	Through the course, patients are provided with disease-related knowledge and information, disease-related coping skills and joint protection strategies to enhance their disease management	Education (25-40 min a day, 8 weeks)	Usual care	 Physical function (HAQ); 2.ASE (pain); 3. ASE (other symptoms); 4. Disease activity (DAS-28)

Risk of Bias

The details of the risk of bias for each study can be observed in Figure 2. Sixteen studies reported specific randomization methods (31, 34, 36-38, 40-42, 44, 46, 50-54), eight RCTs did not specify the specific randomization method (32, 33, 35, 39, 43, 45, 47, 48). The allocation concealment of nine RCTs remained unclear (32, 33, 35, 38, 43, 45, 47, 48, 52). The performance bias of five RCTs was judged to be high (41, 42, 50, 51, 53), and nine RCTs was judged to be unclear (32, 33, 35, 37, 43-45, 47, 48). The detection bias of one RCT was judged to be high (53), and fourteen RCTs remained unclear (32, 33, 35, 37-40, 43-46, 49, 50, 52). Two RCT was judged to be high in incomplete outcome data (attrition bias) (36, 46), and two RCTs remained unclear (32, 48). The other bias of the thirteen studies was unclear (32, 33, 35, 39, 42, 43, 45-48, 50, 52, 53). Among the twenty-four RCTs included, only three RCTs had a low risk of publication bias (31, 34, 54).

Quality of Evidence

The results of the quality of evidence assessed by the GRADE system can be found in **Table 3**. The quality of evidence for pain, physical function, disease activity, and depression was classified as moderate. The quality of evidence for anxiety, ASE (pain), ASE (other symptoms), and general health was classified as low. The quality of evidence for ESR, CRP, and ASE (total) was classified as very low.

Assessment of Overall Effect Size Pain

Twelve studies evaluated pain and included 1,160 participants. Twelve studies that assessed pain used VAS (10 cm) scores (32, 34, 37, 38, 41, 50), VAS (100 mm) scores (36, 39, 42, 46), and AIMS2 scores (35, 45), respectively. The lower the VAS score, and AIMS2 score, the less painful. Of the 12 RCTs, five studies reported that patient education improved pain in patients with RA (P <0.05) (38, 42, 46, 50), while the other seven studies reported that patient education did not improve pain in patients with RA (P > 0.05) (32, 34-37, 39, 41, 45). Four RCTs was not included in the meta-analysis (35, 38, 41, 42). Eight of the twelve RCTs were included in the meta-analysis (32, 34, 36, 37, 39, 45, 46, 50). The pooled results showed no significant improvement in pain in the education group compared to the control group [SMD = -0.37, 95% CI $(-0.80, 0.05), I^2 = 89\%, P = 0.08]$. When education was compared with usual care, the subgroup analysis showed no significant difference in pain improvement in the education group [SMD = 0.07, 95% CI (-0.29, 0.42), $I^2 = 13\%$, P = 0.71]. Similarly, when education + usual care was compared with usual care alone, subgroup analysis showed no difference in pain improvement in the education group [SMD = -0.78, 95% CI (-2.27, 0.71), $I^2 = 98\%$, P = 0.30]. In addition, when education was compared with waiting list (no intervention), subgroup analysis showed no difference in pain improvement in the education group [SMD = -0.08, 95% CI (-0.50, 0.34), P = 0.72]. However, when education + conventional treatment was compared with conventional treatment alone, subgroup analysis showed a significant difference in pain improvement in

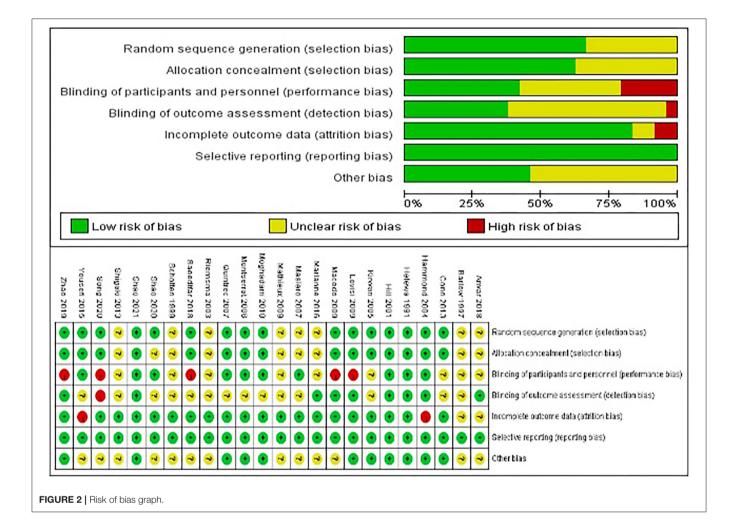
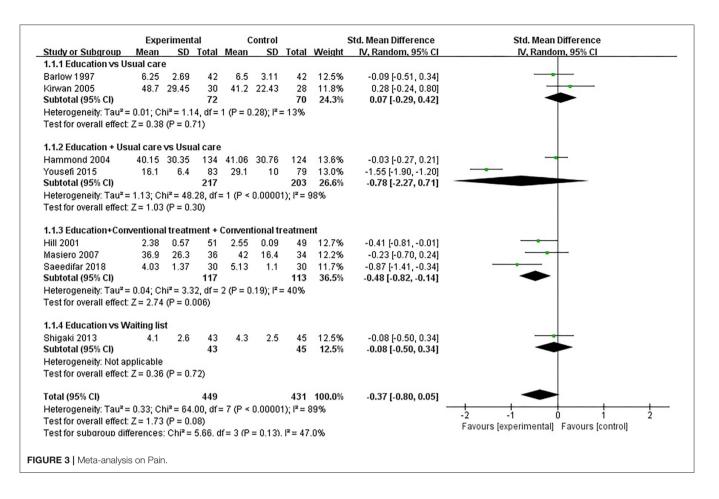


TABLE 3 | Evidence quality rated using the GRADE approach.

Outcomes	No. of studies	Sample size	Risk of bias	Inconsistency	Indirectness	Imprecision	Publication bias	Evider	nce quality
Pain	12	1,160	Not serious	Serious	Not serious	Not serious	Not serious	$\oplus \oplus \oplus \ominus$	Moderate
Physical function	15	1,641	Not serious	Serious	Not serious	Not serious	Not serious	$\oplus \oplus \oplus \ominus \ominus$	Moderate
Disease activity	8	838	Not serious	Serious	Not serious	Not serious	Not serious	$\oplus \oplus \oplus \ominus \ominus$	Moderate
ESR	3	152	Serious	Not serious	Not serious	Very serious	Not serious	$\oplus \ominus \ominus \ominus$	Very low
CRP	3	220	Serious	Not serious	Not serious	Very serious	Not serious	$\oplus \ominus \ominus \ominus$	Very low
Anxiety	5	443	Not serious	Serious	Not serious	Serious	Not serious	$\oplus \oplus \ominus \ominus$	Low
Depression	8	700	Not serious	Serious	Not serious	Not serious	Not serious	$\oplus \oplus \oplus \ominus$	Moderate
ASE (pain)	7	675	Serious	Serious	Not serious	Not serious	Not serious	$\oplus \oplus \ominus \ominus$	Low
ASE (other symptoms)	6	599	Serious	Serious	Not serious	Not serious	Not serious	$\oplus\oplus\ominus\ominus\ominus$	Low
ASE (total)	4	487	Serious	Serious	Not serious	Serious	Not serious	$\oplus \ominus \ominus \ominus$	Very low
General health	4	340	Serious	Not serious	Not serious	Serious	Not serious	$\oplus\oplus\ominus\ominus$	Low

ESR, erythrocyte sedimentation rate; CRP, C-reactive protein; ASE, Arthritis Self-Efficacy.



the education group [SMD = -0.48, 95% CI (-0.82, -0.14), $I^2 = 40\%$, P = 0.006] (Figure 3).

Physical Function

Physical function was assessed in fifteen studies involving 1,641 participants. Fifteen studies assessed physical function using HAQ (31, 33, 36-44, 51, 54), SF-36 (46) and AIMS2 (35), respectively. The lower the HAQ score and AIMS2 score, the better the physical function, while the higher the SF-36 score, the better the physical function. Of the fifteen RCTs, six studies reported that patient education improved physical function in patients with RA (P < 0.05) (33, 38, 39, 43, 46, 54), while the other nine studies reported that patient education did not improve physical function in patients with RA (P > 0.05) (31, 35–37, 40– 42, 44, 51). Five RCTs was not included in the meta-analysis (31, 35, 37, 41, 42). Ten of the 15 RCTs were included in the meta-analysis (33, 36, 38-40, 43, 44, 46, 51, 54). Pooled results showed a significant improvement in physical function in the education group compared to the control group [SMD = -0.52, 95% CI (-0.96, -0.08), $I^2 = 93\%$, P = 0.02]. When education was compared with usual care, the subgroup analysis showed no significant difference in physical function improvement in the education group [SMD = -0.21, 95% CI (-0.61, 0.19), I^2 = 72%, P = 0.30]. Similarly, when education + usual care was compared with usual care alone, subgroup analysis showed no difference in physical function improvement in the education group [SMD = -0.66, 95% CI (-1.95, 0.63), $I^2 = 98\%$, P = 0.32]. However, when education was compared with waiting list (no intervention), subgroup analysis showed a significant difference in physical function improvement in the education group [SMD = -0.69, 95% CI (-1.05, -0.33), $I^2 = 0\%$, P = 0.0002]. In addition, when education + conventional treatment was compared with conventional treatment alone, subgroup analysis showed a significant difference in physical function improvement in the education group [SMD = -0.66, 95% CI (-1.04, -0.28), $I^2 = 0\%$, P = 0.0007] (Figure 4).

Disease Activity

Eight studies assessed disease activity, involving a total of 838 participants. Eight studies assessed disease activity using DAS-28 (35, 40, 42, 51–54) and RAI (39), respectively. The lower the DAS-28 score and RAI score, the better the disease activity. Of the eight RCTs, four studies reported that patient education improved disease activity in patients with RA (P < 0.05) (35, 38, 51, 53), while the other four studies reported that patient education did not improve disease activity in patients with RA (P > 0.05) (40, 42, 52, 54). Three RCTs was not included in the meta-analysis (35, 40, 42). Five of the eight RCTs were included in the meta-analysis (39, 51–54). Pooled results showed a significant improvement in disease activity in the education group compared to the

	Expe	erimenta			ontrol			Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean		Total	Mean	SD	Total	Weight	IV, Random, 95% Cl	IV, Random, 95% Cl
1.1 Education vs Us	sual care								
Conn 2013	1.57		52	1.5	0.8		10.2%	0.09 [-0.29, 0.48]	
Shao 2021	21.44	2.99	108	24.75		106	10.6%	-0.53 [-0.80, -0.26]	
Zhao 2019	0.13	0.75	41	0.25	1	36	9.9%	-0.14 [-0.58, 0.31]	
Subtotal (95% CI)	Section .	ini. miadr	201	in a state		194	30.6%	-0.21 [-0.61, 0.19]	
Heterogeneity: Tau ² =				? (P = 0.0)3); l² =	= 72%			
Test for overall effect:	Z=1.04	(P = 0.3	0)						
2.1.2 Education + Us	ual care	vs Usua	l care						
Hammond 2004	0.94	0.85	134	0.97	0.62	124	10.7%	-0.04 [-0.28, 0.20]	
Quintrec 2007	1.18	0.46	96	1.06	0.47	93	10.5%	0.26 [-0.03, 0.54]	++
Yousefi 2015	-75.05	10.64	83	-53.58	8.45	79	10.1%	-2.22 [-2.61, -1.82]	
Subtotal (95% CI)			313			296	31.3%	-0.66 [-1.95, 0.63]	
Heterogeneity: Tau ² =	1.27; Ch	ni² = 110	.07, df	= 2 (P <	0.000	01); I ² =	98%		
Test for overall effect:	Z = 1.00	(P = 0.3	2)						
2.1.3 Education + Co	vention	al treatn	nent vs	Conver	ntional	treatn	nent		
Masiero 2007	0.93	0.44	36	1.24	0.54	34	9.7%	-0.62 [-1.10, -0.14]	
Montserrat 2006	0.5	0.63	22	1.1	0.98	21	9.1%	-0.72 [-1.34, -0.10]	
Subtotal (95% CI)			58			55	18.8%	-0.66 [-1.04, -0.28]	◆
Heterogeneity: Tau ² =	0.00; Ch	ni² = 0.08	6, df = 1	(P = 0.8	31); l² =	= 0%			
Test for overall effect:	Z = 3.41	(P = 0.0	007)						
2.1.4 Education vs W	aiting lis	t							
Mathieux 2009	0.19	0.19	30	0.35	0.32	30	9.6%	-0.60 [-1.12, -0.08]	
Scholten 1999	2.2	0.32	38	2.6	0.69	30	9.7%	-0.77 [-1.26, -0.27]	
			68			60	19.2%	-0.69 [-1.05, -0.33]	◆
Subtotal (95% CI)	0.00.04	ni² = 0.20), df = 1	(P = 0.8	65); l² =	= 0%			
	0.00, Cr		002)						
Subtotal (95% CI)	•	(P = 0.0	002)						
Subtotal (95% Cl) Heterogeneity: Tau² =	•	(P = 0.0	640			605	100.0%	-0.52 [-0.96, -0.08]	•
Subtotal (95% Cl) Heterogeneity: Tau² = Test for overall effect:	Z= 3.75		640	= 9 (P <	0.000			-0.52 [-0.96, -0.08]	
Subtotal (95% Cl) Heterogeneity: Tau² = Test for overall effect: Total (95% Cl)	Z = 3.75	ni² = 123.	640 .60, df	= 9 (P <	0.000			-0.52 [-0.96, -0.08]	-2 -1 0 1 2 Favours [control]

FIGURE 4 | Meta-analysis on Physical function.

Study or Subarcum	Mean	riment			ontrol	Total	Weight	Std. Mean Difference IV, Random, 95% Cl	Std. Mean Difference IV, Random, 95% Cl
Study or Subgroup 3.1.1 Education vs Us			TUTAL	mean	50	TUI	weight	IV, Rahuom, 95% CI	IV, Kalluolli, 95% Cl
		-							
Shao 2020		0.95	15	3.37		17		-0.18 [-0.88, 0.51]	Ţ
Shao 2021		1.15	108		1.26	106	22.2%	-0.20 [-0.47, 0.07]	22
Zhao 2019	3.82	1.49	41	4.58	1.79	36	21.8%	-0.46 [-0.91, -0.01]	
Subtotal (95% CI)			164			159	64.9%	-0.26 [-0.48, -0.04]	•
Heterogeneity: Tau ² =	0.00; CI	hi² = 0.	99, df =	2 (P =	0.61);	l² = 0%			
Test for overall effect:	Z = 2.30	(P = 0	.02)						
3.1.2 Education + Us	ual care	vs Usi	ual care	е					
Song 2020	3.23	1.38	41	4.1	1.92	36	21.7%	-0.52 [-0.98, -0.07]	
Subtotal (95% CI)			41			36	21.7%	-0.52 [-0.98, -0.07]	◆
Heterogeneity: Not ap	plicable								
Test for overall effect:	Z = 2.24	(P = 0	.03)						
3.1.3 Education + Co	nvention	al trea	tment	+ Conve	ention	al treat	ment		
Masiero 2007	17.9	0.29	36	22.1	0.37	34	13.4%	-12.54 [-14.73, -10.35]	
Subtotal (95% CI)			36			34	13.4%	-12.54 [-14.73, -10.35]	◆
Heterogeneity: Not ap	oplicable								
Test for overall effect:			0.0000	1)					
Total (95% Cl)			241			229	100.0%	-1.97 [-3.24, -0.71]	•
Heterogeneity: Tau ² =	1 06. 01	hiz - 11		f - 1 /D	~ 0.00			-1.57 [-5.24, -0.71]	
Test for overall effect:				n – 4 (r	~ 0.00	,001),1	- 57 %		-10 -5 0 5 10
Test for subgroup diff				0 46- 0	0 /0 - /		12 - 00	201	Favours [experimental] Favours [control]
Testior subdroub diff	erences	: Uni*=	= 120.0	3. $uf = 2$	((P < (1.00001	D. F= 98.	570	

control group [SMD = -1.97, 95% CI (-3.24, -0.71), $I^2 = 97\%$, P = 0.002]. When education was compared with usual care, subgroup analysis showed that the education group had significantly improved disease activity [SMD = -0.26, 95% CI (-0.48, -0.04), $I^2 = 0\%$, P = 0.02]. Similarly, when education + usual care was compared with usual care alone, subgroup analysis showed that the education group had significantly improved disease activity [SMD = -0.52, 95% CI (-0.98, -0.07), P = 0.03]. In addition, when education + conventional treatment was compared with conventional treatment alone, subgroup analysis showed a significant difference in disease activity improvement in the education group [SMD = -12.54, 95% CI (-14.73, -10.35), P < 0.00001] (**Figure 5**).

ESR

ESR were reported in three RCTs (38, 42, 53) with a total of 152 patients with RA. Three RCTs was not included in the metaanalysis (38, 42, 53). In three RCTs, the comparisons involved (1) education + conventional treatment was compared with conventional treatment alone, and (2) education + usual care was compared with usual care alone. When education + conventional treatment was compared with conventional treatment alone, the results showed that there was difference in improved ESR (P < 0.05). However, when education + usual care was compared with usual care alone, the results showed that there was no difference in improved ESR (P > 0.05).

CRP

Three studies assessed CRP and included a total of 220 participants. Of the three RCTs, three studies reported that patient education did not improve CRP in patients with RA (P >0.05) (34, 38, 53). One RCT was not included in the meta-analysis (38). Two of the three RCTs were included in the meta-analysis (34, 53). The pooled results showed no significant improvement in CRP in the education group compared to the control group [SMD = -0.27, 95% CI (-0.57, 0.02), $I^2 = 0\%$, P = 0.07]. When education + conventional treatment was compared with conventional treatment alone, the subgroup analysis revealed that the education group showed no significant difference in improved CRP [SMD = -0.34, 95% CI (-0.73, 0.06), P = 0.09]. Similarly, when education was compared with usual care, subgroup analysis also showed no statistical difference in CRP improvement between the education group and usual care [SMD = -0.19,95% CI (-0.63, 0.26), P = 0.42] (Figure 6).

Anxiety

Anxiety was evaluated in five studies, and 443 participants were included. Five studies assessed anxiety using HADS (32, 37, 40) and STAI (41, 47), respectively. The lower the HADS score and STAI score, the better the anxiety. Of the five RCTs, five studies reported that patient education did not improve anxiety in patients with RA (P > 0.05) (32, 37, 40, 41, 47). Three RCTs was not included in the meta-analysis (37, 40, 41). Two of the five RCTs were included in the meta-analysis (32, 47). The pooled results showed no significant improvement in anxiety in the education group compared to the control group [SMD = 0.17, 95% CI (-0.64, 0.98), $I^2 = 82\%$, P = 0.68]. When education

was compared with usual care, the subgroup analysis showed no significant improvement in the anxiety of the education group [SMD = -0.23, 95% CI (-0.66, 0.20), P = 0.30]. However, when education was compared with waiting list (no intervention), the subgroup analysis revealed that the control group showed a significant difference in improved anxiety [SMD = 0.60, 95% CI (0.06, 1.15), P = 0.03] (Figure 7).

Depression

Depression was evaluated in eight studies, and 700 participants were included. Eight studies assessed depression using HADS (32, 37, 40), BDI (31, 33, 41, 47), or CES-D (45), respectively. The lower the HADS score, BDI score and CES-D score, the better the depression. Of the eight RCTs, eight studies reported that patient education did not improve depression in patients with RA (P > 0.05) (31–33, 37, 40, 41, 45, 47). Four RCTs was not included in the meta-analysis (31, 37, 40, 41). Four of the eight RCTs were included in the meta-analysis (32, 33, 45, 47). The pooled results showed no significant improvement in depression in the education group compared to the control group [SMD = -0.18, 95% CI (-0.52, 0.15), $I^2 = 52\%, P = 0.28$]. When education was compared with usual care, the subgroup analysis showed no significant improvement in the depression of the education group [SMD = -0.37,95% CI (-0.80,0.07), P = 0.10]. In addition, When education was compared with waiting list (no intervention), the subgroup analysis showed that there was no significant difference in improved depression [SMD = -0.11, 95% CI (-0.56, 0.35), $I^2 = 63\%, P = 0.64$] (Figure 8).

ASE (Pain)

ASE (pain) was evaluated in seven studies involving a total of 675 participants. Seven studies assessed ASE (pain) using the ASE scale (32, 35, 37, 48, 49, 52, 54). Of the seven RCTs, three studies reported that patient education improved ASE (pain) in patients with RA (P < 0.05) (48, 49, 54), while the other four studies reported that patient education did not improve ASE (pain) in patients with RA (*P* > 0.05) (32, 35, 37, 52). Seven RCTs were included in the meta-analysis (32, 35, 37, 48, 49, 52, 54). Pooled results showed a significant improvement in ASE(pain) in the education group compared to the control group [SMD = -1.24, 95% CI (-2.05, -0.43), $I^2 = 95\%$, P = 0.003]. When education was compared with usual care, the subgroup analysis showed a significant improvement in ASE (pain) in the education group [SMD = -1.27, 95% CI (-2.20, -0.34), $I^2 = 96\%$, P =0.007]. Similarly, when education + conventional treatment was compared with conventional treatment alone, subgroup analysis showed a significant improvement in ASE (pain) [SMD = -1.16, 95% CI (−1.69, −0.63), *P* < 0.0001] (**Figure 9**).

ASE (Other Symptoms)

ASE (other symptoms) was evaluated in six studies involving a total of 599 participants. Six studies assessed ASE (other symptoms) using the ASE scale (32, 35, 37, 49, 52, 54). Of the six RCTs, one studies reported that patient education improved ASE (other symptoms) in patients with RA (P < 0.05) (49), while the other five studies reported that patient education did not improve ASE (other symptoms) in patients with RA (P

	Expe	eriment	al	C	Control		1	Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl
5.1.1 Education + Cor	vention	al treat	ment v	s Conv	entional	treatn	nent		
Hill 2001	24.8	28.75	51	39	51.5	49	56.3%	-0.34 [-0.73, 0.06]	
Subtotal (95% CI)			51			49	56.3%	-0.34 [-0.73, 0.06]	
Heterogeneity: Not ap	plicable								
Test for overall effect:	Z=1.69	(P = 0.0	09)						
5.1.2 Education + Usu	ial care	vs Usu	al care						
Song 2020	6.09	15.05	41	12.7	48.94	36	43.7%	-0.19 [-0.63, 0.26]	
Subtotal (95% CI)			41			36	43.7%	-0.19 [-0.63, 0.26]	
Heterogeneity: Not ap	plicable								
Test for overall effect:	Z = 0.81	(P = 0.4	42)						
Total (95% CI)			92			85	100.0%	-0.27 [-0.57, 0.02]	
Heterogeneity: Tau ² =	0.00: CI	hi ² = 0.2	5. df =	1 (P = 0)	.61): I ² =	= 0%			
Test for overall effect:									-0.5 -0.25 0 0.25 0.5
Test for subaroup diff		•	'	f = 1 (P	= 0.61)	$l^{2} = 0.9$	6		Favours [experimental] Favours [control]
							-		
URE 6 Meta-analysis		۰ ۲							

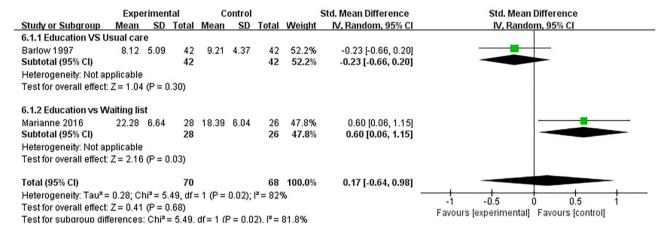


FIGURE 7 | Meta-analysis on Anxiety.

	Expe	rimen	tal	Co	ontrol		:	Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% CI	IV, Random, 95% Cl
7.1.1 Education vs Us	sual care	е							
Barlow 1997	6.24	3.94	42	7.6	3.4	42	26.9%	-0.37 [-0.80, 0.07]	
Subtotal (95% CI)			42			42	26.9%	-0.37 [-0.80, 0.07]	
Heterogeneity: Not ap	plicable								
Test for overall effect:	Z=1.66	(P = 0	1.10)						
7.1.2 Education vs W	aiting lis	st							
Marianne 2016	13.68	9.77	28	10.44	5.5	26	21.5%	0.40 [-0.14, 0.94]	
Scholten 1999	10.8	2	38	12.7	6.2	30	24.1%	-0.43 [-0.91, 0.06]	
Shigaki 2013	10.8	8.2	43	13.2	11.2	45	27.6%	-0.24 [-0.66, 0.18]	
Subtotal (95% CI)			109			101	73.1%	-0.11 [-0.56, 0.35]	
Heterogeneity: Tau ² =	0.10; CI	hi² = 5.	43, df=	= 2 (P = 1	0.07);	² = 639	Хо		
Test for overall effect:	Z = 0.47	(P = 0	1.64)						
Total (95% CI)			151			143	100.0%	-0.18 [-0.52, 0.15]	
Heterogeneity: Tau ² =	0.06; CI	hi² = 6.	20, df =	= 3 (P = 1	0.10);	² = 529	λ.		-1 -0.5 0 0.5 1
Test for overall effect:	Z = 1.07	(P = 0	.28)						Favours [experimental] Favours [control]
Test for subaroup diff	ferences	: Chi ² :	= 0.64.	df = 1 (F	= 0.4	2), ² =	0%		r avours lexperimentalj Favours (controlj
URE 8 Meta-analysis	s on Dep	ressior	۱.						

	Expe	riment	al	С	ontrol			Std. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Random, 95% Cl	IV, Random, 95% Cl
8.1.1 Education vs Us	sual care								
Anvar 2018	-66.31	2.98	37	-45.56	2.66	39	11.3%	-7.28 [-8.55, -6.01]	
Barlow 1997	-23.1	11	42	-19.6	11	42	14.9%	-0.32 [-0.75, 0.12]	
Kirwan 2005	-62.4	17.4	30	-57.8	15.98	28	14.7%	-0.27 [-0.79, 0.25]	
Riemsma 2003	-3.5	0.7	71	-3.4	0.7	76	15.2%	-0.14 [-0.47, 0.18]	+
Shao 2020	-37.47	10.74	15	-37.47	12.58	17	14.0%	0.00 [-0.69, 0.69]	
Shao 2021	-41.76	6.23	108	-35.95	6.51	106	15.3%	-0.91 [-1.19, -0.63]	*
Subtotal (95% CI)			303			308	85.4%	-1.27 [-2.20, -0.34]	\bullet
Heterogeneity: Tau ² =	1.23; Ch	i ² = 124	.54, df	= 5 (P < 1	0.0000	1); $l^2 = 9$	96%		
Test for overall effect:	Z = 2.69	(P = 0.0)	07)						
8.1.2 Education + Cor	rventiona	al treatr	nent vs	Conver	tional t	reatme	ent		
Moghadam 2018	-8.9	2.4	32	-5.87	2.74	32	14.6%	-1.16 [-1.69, -0.63]	-
Subtotal (95% CI)			32			32	14.6%	-1.16 [-1.69, -0.63]	◆
Heterogeneity: Not ap	plicable								
Test for overall effect:	Z = 4.28	(P < 0.0	001)						
Total (95% CI)			335			340	100.0%	-1.24 [-2.05, -0.43]	\bullet
Heterogeneity: Tau ² =	1.09; Ch	i ² = 128	.56, df	= 6 (P <	0.0000	1); $ ^2 = 9$	95%		
Test for overall effect:									-4 -2 U 2 4
Test for subaroup diff				= 1 (P =	0.84).1	² = 0%			Favours [experimental] Favours [control]
FIGURE 9 Meta-analysi	s on ASE	(pain).							

> 0.05) (32, 35, 37, 52, 54). Six RCTs were included in the meta-analysis (32, 35, 37, 49, 52, 54). Pooled results showed a significant improvement in ASE (other symptoms) in the education group compared to the control group [SMD = -0.25, 95% CI (-0.41, -0.09), $I^2 = 25\%$, P = 0.002]. When education was compared with usual care, the subgroup analysis showed a significant improvement in ASE (other symptoms) in the education group [SMD = -0.19, 95% CI (-0.36, -0.02), $I^2 = 0\%$, P = 0.03]. Similarly, when education + conventional treatment was compared with conventional treatment alone, subgroup analysis showed a significant improvement in ASE (other symptoms) [SMD = -0.83, 95% CI (-1.34, -0.32), P = 0.001] (Figure 10).

ASE (Total)

ASE (total) was evaluated in four studies involving a total of 487 participants. Four studies assessed ASE (total) using the ASE scale (36, 45, 49, 51). Of the four RCTs, three studies reported that patient education improved ASE (total) in patients with RA (P < 0.05) (45, 49, 51), while the other study reported that patient education did not improve ASE (total) in patients with RA (P > 0.05) (36). Four RCTs were included in the meta-analysis (36, 45, 49, 51). Pooled results showed a significant improvement in ASE (total) in the education group compared to the control group [SMD = -0.67, 95% CI (-1.30, -0.05), $I^2 = 90\%$, P = 0.03]. When education was compared with usual care, the subgroup analysis showed a significant improvement in ASE (total) in the education group [SMD = -0.87, 95% CI (-1.33, -0.40), P = 0.0003]. When education + conventional treatment was compared with conventional treatment alone, the subgroup analysis showed a significant improvement in ASE (total) in the education group [SMD = -1.23, 95% CI (-1.76, -0.69), P < 0.00001]. Similarly, When education was compared waiting list (no intervention), the subgroup analysis showed a significant improvement in ASE (total) in the education group [SMD =

-0.76, 95% CI (-1.19, -0.33), P = 0.0006]. However, when education + usual care was compared with usual care alone, the subgroup analysis revealed that the education group showed no significant difference in improved ASE (total) [SMD = -0.05, 95% CI (-0.30, 0.19), P = 0.68] (**Figure 11**).

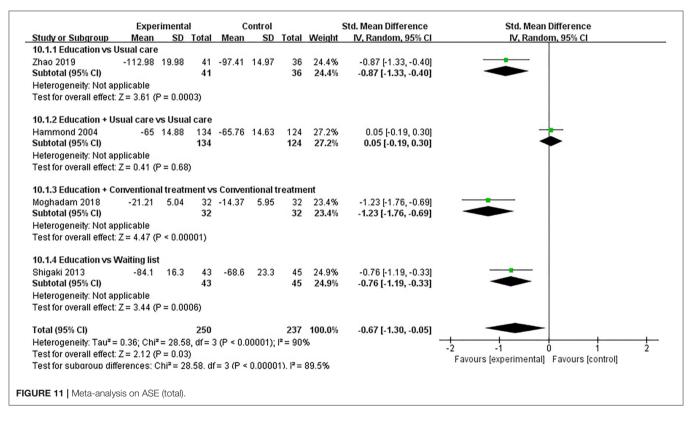
General Health

General health was evaluated in four studies involving a total of 340 participants. Four studies assessed general health using SF-36 (41, 46), EQ5D (42), or AIMS2 (45), respectively. The lower the AIMS2 score, the better the general health, and the higher the SF-36 and EQ5D, the better the general health. Of the four RCTs, two studies reported that patient education improved general health in patients with RA (P < 0.05) (42, 46), while the other two studies reported that patient education did not improve general health in patients with RA (P > 0.05) (41, 45). One RCT was not included in the meta-analysis (42). Three of the four RCTs were included in the meta-analysis (41, 45, 46). Pooled results showed a significant improvement in general health in the education group compared to the control group $[SMD = -1.11, 95\% CI (-1.36, -0.86), I^2 = 96\%, P < 0.00001].$ When education was compared waiting list (no intervention), the subgroup analysis showed a significant improvement in general health in the education group [SMD = -0.37, 95% CI (-0.69, -0.04), $I^2 = 0\%$, P = 0.03]. Similarly, when education + usual care was compared with usual care alone, subgroup analysis showed a significant improvement in general health [SMD = -2.17,95% CI (-2.55, -1.78), P < 0.00001] (Figure 12).

Publication Bias

When the meta-analysis includes more than ten studies, the possibility of publication bias should be reported (55). We used the Egger's and Begg's test to assess the publication bias of each outcome (30). The evaluation results showed that there was no publication bias in the outcomes (**Table 4**).

	Expe	erimenta	al	C	ontrol		S	td. Mean Difference	Std. Mean Difference
Study or Subgroup	Mean	SD	Total	Mean	SD	Total	Weight	IV, Fixed, 95% Cl	IV, Fixed, 95% CI
9.1.1 Education vs Us	sual care								
Barlow 1997	-31.8	13.9	42	-29	11.5	42	14.1%	-0.22 [-0.65, 0.21]	
Kirwan 2005	-61.1	15.13	30	-61.7	21.92	28	9.8%	0.03 [-0.48, 0.55]	
Riemsma 2003	-3.9	0.6	71	-3.8	0.6	76	24.8%	-0.17 [-0.49, 0.16]	
Shao 2020	-55.33	7.75	15	-51.12	11.53	17	5.3%	-0.41 [-1.12, 0.29]	
Shao 2021	-55.59	6.89	108	-53.86	8.56	106	36.0%	-0.22 [-0.49, 0.05]	
Subtotal (95% CI)			266			269	90.1%	-0.19 [-0.36, -0.02]	•
Heterogeneity: Chi ² =	1.19, df=	= 4 (P =	0.88); I	² = 0%					
Test for overall effect:	Z=2.18	(P = 0.0)	3)						
9.1.2 Education + Co									
Moghadam 2018	-11.31	3.08	32	-8.5	3.57	32	9.9%	-0.83 [-1.34, -0.32]	
Subtotal (95% CI)			32			32	9.9%	-0.83 [-1.34, -0.32]	
Heterogeneity: Not ap	plicable								
Test for overall effect:	Z = 3.19	(P = 0.0	01)						
Total (95% CI)			298			301	100.0%	-0.25 [-0.41, -0.09]	
Total (95% CI) Heterogeneity: Chi² =	6.65, df=	= 5 (P =		²= 25%		301	100.0%	-0.25 [-0.41, -0.09]	
		•	0.25); 1	²= 25%		301	100.0%	-0.25 [-0.41, -0.09] -	-1 -0.5 0 0.5 1 Favours [experimental] Favours [control]

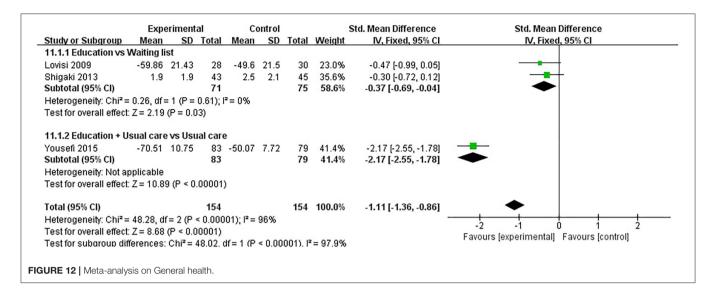


Meta-Regression Analysis and Sensitivity Analysis

In this study, we evaluated a total of 10 outcomes. The outcomes with high heterogeneity were pain, physical function, disease activity, anxiety, depression, ASE (pain), ASE (total), and general health. We used metaregression analysis and sensitivity analysis to explore the sources of heterogeneity. Results of meta-regression analysis and sensitivity analysis were presented in **Supplementary Appendix.**

DISCUSSION

We conducted this review to evaluate evidence for the benefits of patient education interventions in people with RA. The outcomes



include pain, physical function, disease activity, erythrocyte sedimentation rate (ESR), C-reactive protein (CRP), anxiety, depression, ASE (pain, other symptoms, total), and general health. The results of this meta-analysis suggest that patient education may be beneficial for improving physical function, disease activity, increasing ASE (pain), ASE (other symptoms), and ASE (total), and general health in patients with RA. However, there was no significant effect on anxiety, depression, pain, and CRP in patients with RA.

This study shows that patient education is generally beneficial for rheumatoid arthritis. Based on our previous work, we expected that patient education would contribute to clinical outcomes and psychological status in rheumatoid arthritis. Several previous studies have explored the impact of patient education on diseases such as low back pain and rheumatoid arthritis (26, 56). These studies found that patient education improved pain and general health in patients with acute low back pain, and improved general health, mental health, and depression in rheumatoid arthritis but did not significantly improve anxiety or disease activity. Our study found that patient education can help improve clinical outcomes in rheumatoid arthritis, such as physical function, disease activity, increasing ASE (pain), ASE (other symptoms), and ASE (total), and general health. The positive effect of patient education may be closely related to the active participation of patients, and it is very important for patients to adhere to rheumatoid arthritis-related interventions. However, our study found that patient education did not significantly improve anxiety, depression, pain, and CRP in patients with rheumatoid arthritis, and further research in this part is needed in the future. Overall, our results are in line with our prior expectations.

RA is a chronic inflammatory joint disease that can occur at any age, causing disability, loss of workability, and even death in patients with RA (57). Many patients with RA continue to experience joint pain, deformities, disability, and poor quality of life despite professional treatment (24). There are some interventions that can enhance the effectiveness of RA, and

patient education is one of them, this may be related to patient participation and shared decision making. It can help patients understand RA and strengthen the management of the disease (58, 59). Patient education has been defined as "any set of planned educational activities designed to improve patients' health behaviors and health status" (24). EULAR recommends that patient education should be part of the treatment of RA (17). Studies have shown that patients may play an important role in the disease management of RA (39), but this important role may require active participation and shared decision making by patients, and patients must contribute actively to adhere to interventions for their RA. Health education for patients with RA can have a positive impact on the perception of pain and the management of the disease, and they transform the knowledge of the disease and the methods of preventing pain into changes in health behavior, which not only reduces the symptoms of pain and disability but also it will also improve body function (39, 60, 61). Antirheumatoid medications and biological agents can effectively control the symptoms and inflammatory response of RA, but the effectiveness of the medications is severely limited by patient adherence (8), and patient education can enhance the adherence of patients with RA and improve the effectiveness of medications (53), this may have a positive effect on improving their pain, physical function and reducing inflammatory cytokines (such as ESR, CRP, and RF). For patients with RA, self-efficacy is an important influencing factor to comply with health advice and health outcomes (62). The study found that the self-efficacy of patients with RA is significantly related to their health status (63, 64) and that patient education has a positive impact on clinical efficacy through self-efficacy (65). In addition, the effect of patient education on patients with RA was also affected by the education content. Some education content includes guiding patients to take family exercise, and family exercise has been proved to promote the maintenance of muscle strength, increase handgrip strength and flexibility (66-68), which can further improve the physical function of patients with RA. Clinical studies have shown that patient

TABLE 4	Assessment	of	publication	bias.
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Outcomes	No. of studies	Egger's test (P-values)	Begg's test (P-values)	
Pain	12	0.595#	0.451#	
Physical function	15	0.278#	0.075#	

 $^{\#}P > 0.05.$

education was closely related to patients' mental health, especially in anxiety and depression, and patient education can improve patients' anxiety and depression (32, 69). The symptoms of RA are the result of the interaction between the body and the mind, and the symptoms of RA and psychological factors are closely related (69). Health education for patients with RA can increase the understanding of the disease and its treatment methods, which is beneficial to improve the anxiety and depression of patients with RA, and may also have a significant impact on the control of the disease. The study found that long-term behavior change of patients with RA through education can have a positive impact on disease control and also significantly reduce the disease activity of RA (19). Other studies have found that patient education may not have a direct impact on disease activity but maybe in an indirect way (17).

Patient education is increasing of interest to healthcare workers as a complementary intervention. Patient education has been recognized by many clinicians and is widely used in clinical practice. Some medical, orthopedic, and other diseases use patient education as an intervention, such as knee osteoarthritis, cancer, stroke, etc. (70-72). However, the mechanisms by which patient education works are complex and integrative. Some studies have found that patient education can reduce patient pain and function, and the mechanism by which patient education affects health outcomes may be related to self-efficacy (73). Patient education can improve patient self-efficacy, and some information, including exercise and health management, can promote healthy behavior changes (70, 73). Increased patient awareness of the disease can promote patient self-motivation, change their behavior, enhance disease management further, as well as increase patient adherence (74). In addition, patient education can improve psychosocial support, which contributes to disease management to a certain extent and promotes changes in patient behavior (75). Several studies suggest that patient education interventions should not be limited to treatment adherence alone but should also focus on the patient's mental health, which is valuable by combining psychosocial factors, clinical outcomes, and self-reported adherence (75, 76). Patient education is a long-term ongoing process and should be implemented with periodic assessment of its benefits and updating its interventions as necessary to accommodate changes in the patient's disease (76). The effectiveness of patient education may be affected by multiple factors, such as the patient's learning ability, literacy level, cultural environment, etc. Therefore, careful and comprehensive consideration should be given to developing interventions for patient education (77). In addition, some studies suggest that effective patient education should encourage patients to enhance self-efficacy and self-management, improve

Limitations

This systematic review and meta-analysis had several limitations. Firstly, although all the included RCTs were randomized, it was not clear whether some included RCTs had a bias in allocation concealment and bias in performance bias (blinding of participants and personnel) and detection bias (blinding of outcome assessment). Therefore, the quality of evidence for outcomes had been reduced to low levels. Secondly, the sample size included in the study was small. Although a small sample size can be used for meta-analysis, the results might be biased, and the conclusions drawn should be considered preliminary (81). Thirdly, the RCTs varied in terms of their patient populations (such as disease duration), comparative treatments (specific methods, the length of intervention time, and frequency of interventions), and outcome measures. Therefore, the results of the meta-analysis showed that there was a high degree of heterogeneity. However, due to the small number of included RCTs and the fact that some RCTs did not specifically report these differences, we could not conduct a subgroup analysis to check whether these factors had an impact on the outcome of the study.

Implications for Further Research and Practice

Considering that patient education is an intervention without side effects and helps to improve the curative effect of rheumatoid arthritis and control the development of rheumatoid arthritis, it is recommended that patient education be used as a treatment strategy for rheumatoid arthritis. The use of patient education as an intervention for RA is of great significance for reducing the medical cost of RA, especially in developing countries. In addition, this study also provides data for future clinical research on RA. However, due to the low level of evidence in this study and the high heterogeneity of the study, more highquality RCTs should be conducted in the future to verify these conclusions. We acknowledge the difficulty of conducting RCTs in this study, but some methods can make future study designs more comprehensive and rigorous. Firstly, the implementation of RCTs on patient education in the future should strictly follow the Consolidated Standards of Reporting Trials (CONSORT) statement to improve the quality of research (82). Secondly, researchers should complete registration at a standard clinical study center before starting a clinical study, and complete study protocols should be published to reduce publication bias (83). Finally, clinical studies should extend the duration of follow-up and increase the frequency of follow-up to assess the long-term efficacy of interventions.

Patient education is a broad-based intervention that may serve as a foundational intervention for many diseases. However, patient education is effective on clinical outcomes remains inconclusive. Studies have found that it is difficult to compare the effects of patient education on different chronic diseases and also among different types of arthritis (84). Therefore, we are only studying rheumatoid arthritis and not simultaneously studying and comparing larger diseases, such as rheumatic diseases, osteoarthritis, or other chronic musculoskeletal diseases. In the future, our research should be extended to other arthritis or chronic musculoskeletal diseases to explore further the impact of patient education on other arthritic or chronic musculoskeletal diseases. At the same time, patient education can be considered as a disease-based intervention, combining patient education with other interventions.

CONCLUSION

Patient education may be effective in improving clinical outcomes and psychological status in patients with rheumatoid arthritis. Considering the methodological limitations of the included RCTs, more high-quality and large-sample RCTs are needed to confirm this conclusion in the future.

DATA AVAILABILITY STATEMENT

The original contributions presented in the study are included in the article/**Supplementary Material**, further inquiries can be directed to the corresponding author/s.

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AUTHOR CONTRIBUTIONS

ZGW, WGL, and XMX designed the study. ZGW, YZ, YW, and RZ conducted a literature search and screening and extracted data from the literature. XLY, ZHC, CCL, and JYL checked the extracted data. ZGW, ZBW, and ZXY conducted the statistical analysis. ZGW, YZ, YW, and RZ wrote the first draft. WGL and XMX corrected the manuscript. ZXY did the language editing. WGL and XMX supervised the conduct of the study. All authors have read and approved the final submitted version.

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SUPPLEMENTARY MATERIAL

The Supplementary Material for this article can be found online at: https://www.frontiersin.org/articles/10.3389/fpsyt. 2022.848427/full#supplementary-material

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