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Review article

Effect of psychotherapy for depression on quality of life: meta-analysis

Spyros Kolovos, Annet Kleiboer and Pim Cuijpers

Background

Several meta-analyses have shown that psychotherapy is effective for reducing depressive symptom severity. However, the impact on quality of life (QoL) is as yet unknown.

Aims

To investigate the effectiveness of psychotherapy for depression on global QoL and on the mental health and physical health components of QoL.

Method

We conducted a meta-analysis of 44 randomised clinical trials comparing psychotherapy for adults experiencing clinical depression or elevated depressive symptoms with a control group. We used subgroup analyses to explore the influence of various study characteristics on the effectiveness of treatment.

Results

We detected a small to moderate effect size (Hedges' $g=0.33$, 95% CI 0.24–0.42) for global QoL, a moderate effect

size for the mental health component ($g=0.42$, 95% CI 0.33–0.51) and, after removing an outlier, a small but statistically significant effect size for the physical health component ($g=0.16$, 95% CI 0.05–0.27). Multivariate meta-regression analyses showed that the effect size of depressive symptoms was significantly related to the effect size of the mental health component of QoL. The effect size of depressive symptoms was not related to global QoL or the physical health component.

Conclusions

Psychotherapy for depression has a positive impact on the QoL of patients with depression. Improvements in QoL are not fully explained by improvements in depressive symptom severity.

Declaration of interest

None.

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Depression is one of the most common mental disorders among adults.^{1,2} Major depression in particular ranks currently fourth in disease burden worldwide, and is expected to rank first in high-income countries by 2030.³ In addition, depression is an enormous societal burden due to high healthcare use and reduced work performance.^{4–6} Furthermore, depression is associated with substantial impairments in quality of life.^{7,8} Quality of life (QoL) is a broad concept that comprises a range of life domains of the individual, such as social relationships, physical abilities, mental health functioning, role functioning and engagement in daily activities. Deficits in all these domains have been identified in people experiencing depressive symptoms.⁹ Several meta-analyses have shown the effectiveness of different psychotherapies in reducing depressive symptoms compared with control conditions.^{10,11} Even though it is often postulated that improvements in depressive symptoms during treatment coincide with improvements in QoL, evidence to support the effectiveness of depression treatment on QoL is limited.¹² Research indicates that QoL and depressive symptoms are moderately correlated at post-treatment assessment but suggest a weaker relationship in the long term.¹³ Additionally, research suggests that people in remission from depression experience persistent deficits in QoL.^{14,15} Therefore, clinical remission, as well as overall well-being, defined exclusively by the absence of depressive symptoms, may be insufficient. To date, no meta-analysis has quantified the effects of psychotherapy for depression on QoL. Therefore, we examined how the effects of psychotherapy for depression compared with control conditions on global QoL and on two specific domains of QoL, namely mental and physical health.

Method

Initially, we searched an existing database (www.evidencebasedpsychotherapies.org) that has previously been used in a series of

meta-analyses and contains 1476 randomised controlled trials (RCTs).¹⁶ This database was developed through a systematic literature search (from 1966 to 1 January 2013) and is periodically updated. Additionally, a systematic literature search was conducted in PubMed, EMBASE, PsycINFO and the Cochrane Central Register of Controlled Trials from 1 January 2013 to 1 January 2015.

Study selection

We included published RCTs in which psychotherapy for depression was compared with a control condition, for participants 18 years old or over, and which reported a measure of QoL at post-treatment assessment. Psychotherapy was defined as an intervention in which the core element was verbal communication between a participant and a therapist, or as a systematic psychological treatment in the form of a website or book which the participant worked through more or less independently but with personal support from a therapist.¹⁶ The control condition was defined as waiting list, care as usual, placebo or another minimal treatment. Studies were included if participants were diagnosed with a depressive disorder on the basis of a structured clinical interview or if they reported elevated depressive symptoms based on a standardised measurement of depressive symptom severity. A measure of QoL was defined as any patient-reported measure aiming to assess perceived health status, well-being or effective performance in daily life.¹⁷ These measures could provide a global (overall) score, or separate scores for different domains or components. We differentiated between two components, namely mental health and physical health. The mental health component of QoL was defined as personal satisfaction with the current psychological state, whereas the physical health component was defined as the perceived competence for performance and functioning in various everyday

activities.¹⁷ Our search was restricted to studies written in English and German. Studies regarding treatment maintenance were excluded. Comorbid psychiatric or medical disorders were not used as an exclusion criterion. Finally, we excluded studies for which we did not have sufficient statistics to perform the meta-analysis.

Quality assessment

The validity of the studies was assessed following the guidelines provided by the Cochrane Collaboration's tool for assessing risk of bias.¹⁸ Risk of bias was examined in four domains: random sequence generation, allocation concealment, blinding (masking) of outcome assessment and intention-to-treat analysis. Two authors (S.K. and A.K.) conducted the assessment. Disagreements between the two reviewers were resolved by discussion until consensus was reached.

Statistical analysis

For the univariate analyses we used the Comprehensive Meta-Analysis (CMA) software package.¹⁹ For the multivariate analyses we used the *metareg* module within Stata,²⁰ because these analyses cannot be performed with CMA. We calculated the effect size following the procedure described by Hedges & Olkin to correct for small sample size bias.²¹ We estimated the pooled effect sizes using the random effects model to account for heterogeneity among studies.²² Heterogeneity was examined with the I^2 statistic, where a value of 25% determines low heterogeneity, 50% moderate heterogeneity and 75% high heterogeneity.²³ We further calculated the 95% confidence intervals around I^2 statistic,²⁴ by using the non-central χ^2 -based approach within the *heterogi* module for Stata.²⁵ Finally, publication bias was examined by visual inspection of the funnel plot and by implementing Duval & Tweedie's trim and fill procedure, which is a test of symmetry of the funnel plot. In addition, the method developed by Duval & Tweedie yields an adjusted pooled effect size after accounting for missing studies due to publication bias.²⁶ We conducted a number of subgroup analyses to identify potential moderators of the outcome using the mixed effects model, in which studies within the subgroups are pooled with the random effects model and the tests for significant differences between the subgroups are carried out with the fixed effects model.²⁷ Subgroup analyses were performed when at least three studies were available for each group. Moreover, we conducted sensitivity analyses because a number of studies compared more than one experimental group with the same control condition, and therefore the assumption of independency was violated. In the sensitivity analyses we first included the largest effect size for each study and then the lowest effect size for each study.

We used univariate and multivariate meta-regression analyses to examine the relationship between changes in QoL and depressive symptom severity. The meta-regressions were undertaken using the mixed effects model.²⁸ For the univariate meta-regressions the effect size of QoL was set as the dependent variable and the effect size of depressive symptom severity as the predictor. For the multivariate meta-regressions a number of potential confounder variables were added simultaneously as predictors alongside the effect size for depressive symptoms.

Power calculation

We presumed that a limited number of studies would have administered QoL measures. Thus, we carried out a power calculation to estimate whether the included studies would provide sufficient statistical power to detect small effect sizes,

according to the recommendations of Borenstein *et al.*²⁷ Although there is no consensus about a clear definition, we defined a small effect size as $g=0.20$.²⁹ We conservatively assumed a high level of between-study variance (τ^2), a statistical power of 0.80 and an alpha value of 0.05. The power calculation demonstrated that we would need 20 studies with a mean sample size of 40 participants or 15 studies with 54 participants.

Results

The databases search resulted in 20 461 titles. We retrieved the full text of 1764 studies, from which 44 studies were included in this meta-analysis (Fig. 1).

Study characteristics

The 44 studies included in total 5264 patients: 2907 in the intervention group and 2357 in the control group (see online Table DS1). More specifically, the meta-analysis of global QoL included 27 studies with 2448 patients, the meta-analysis of the mental health component included 18 studies with 2463 patients and that of the physical health component included 13 studies with 1561 patients. Psychotherapies that could be clustered in the cognitive-behavioural group (i.e. cognitive-behavioural therapy, mindfulness-based cognitive therapy and coping with depression) were provided in 25 studies (56%). Life review was offered in five studies (14%), problem-solving treatment in three studies (8%), acceptance and commitment therapy in three studies (8%) and interpersonal psychotherapy in two studies (5%). Care as usual was the most common control condition and was included in 20 studies (45%). It consisted mainly of psychotherapy, antidepressant medication or combination treatments, but was only superficially described in the published papers. Therefore, we did not have enough information to cluster usual treatments based on their modality. Waiting-list groups were included in 17 studies (39%). Other types of control conditions were included in 7 studies (16%), and consisted of discussion groups, psycho-education, a 20 min educational video or placebo pill. The mean number of treatment sessions was 10 (median 9, range 1–25).

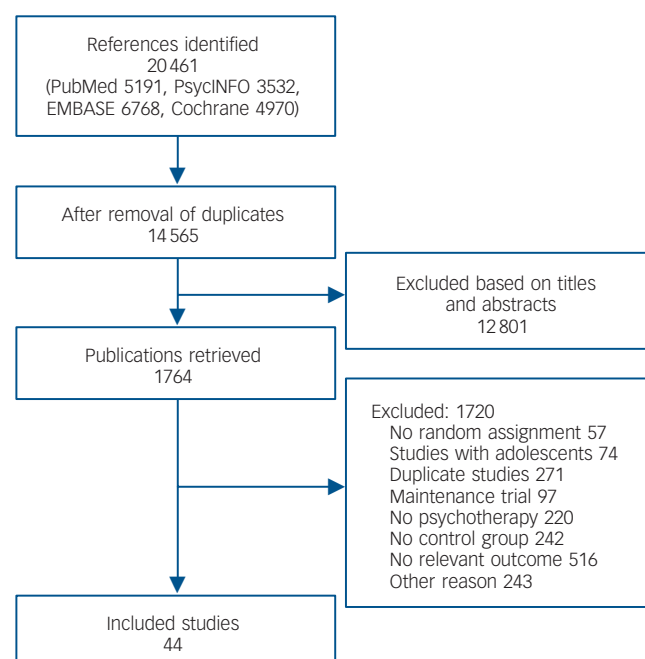


Fig. 1 Study selection.

All the quality criteria were met by 24 studies (55%) and at least three out of four criteria were met by 33 studies (75%). Finally, 36 studies (82%) reported a method of handling incomplete outcome data. The characteristics of the participants varied among the studies. Adult patients (both men and women) were included in 27 studies (61%), older adults in 11 studies (25%) and exclusively women in six studies (14%). Eighteen studies (41%) included participants diagnosed with major depressive disorder. Patients with comorbid physical or mental health symptoms were included in 15 studies (34%).

Global QoL

Thirty-one comparisons were included in the meta-analysis of global QoL (Fig. 2). The mean effect size (Hedges' *g*) was 0.33 (95% CI 0.24–0.42). We detected low between-study heterogeneity ($I^2 = 21$, 95% CI 0–49). After adjusting for publication bias using the trim and fill procedure the mean effect size was $g = 0.30$ (95% CI 0.21–0.40), with three imputed studies (Table 1). Moreover, meta-analysis including only the largest effect size of each study resulted in an overall effect size of $g = 0.35$ (95% CI 0.25–0.45). When only the smallest effect size was included the pooled effect size was $g = 0.34$ (95% CI 0.24–0.45). We also calculated the effect sizes separately for scores on the Quality of Life Inventory (QOLI; $g = 0.32$, 95% CI 0.15–0.48) and the EuroQol EQ-5D ($g = 0.19$, 95% CI 0.07–0.32). Studies including people with major depressive disorder resulted in larger effect sizes ($g = 0.49$, 95%

CI 0.36–0.61) than studies including people without such a diagnosis ($g = 0.23$, 95% CI 0.13–0.34, $P = 0.002$). Studies including adults reported larger effect sizes ($g = 0.39$, 95% CI 0.29–0.49) than studies including older adults ($g = 0.15$, 95% CI 0.00–0.30, $P = 0.009$). Other study characteristics were not significantly related to the effect size of global QoL.

The mean effect size for depressive symptoms was $g = 0.60$ (95% CI 0.50–0.70) and therefore considerably larger than that for global QoL. Heterogeneity was moderate ($I^2 = 32\%$, 95% CI 0–56). After adjusting for publication bias the mean effect size decreased to $g = 0.54$ (95% CI 0.44–0.64), with five imputed studies. The univariate meta-regression analysis indicated a significant relationship between the effect size of global QoL and the effect size of depressive symptoms (slope 0.52, 95% CI 0.21–0.84, $P = 0.002$), suggesting that with each increase in effect size of depressive symptom severity by 1 the effect size for global QoL increased by 0.52. The effect size of global QoL was not significantly associated with the number of treatment sessions (slope 0.00, 95% CI –0.03 to 0.03, $P = 0.803$). The multivariate meta-regression results showed that none of the predictors included in the model was significantly related to the effect of psychotherapy on global QoL at post-treatment assessment (see Table 4).

Mental health

Twenty-one comparisons were included in the meta-analysis of the effects of psychotherapy for depression on the mental health

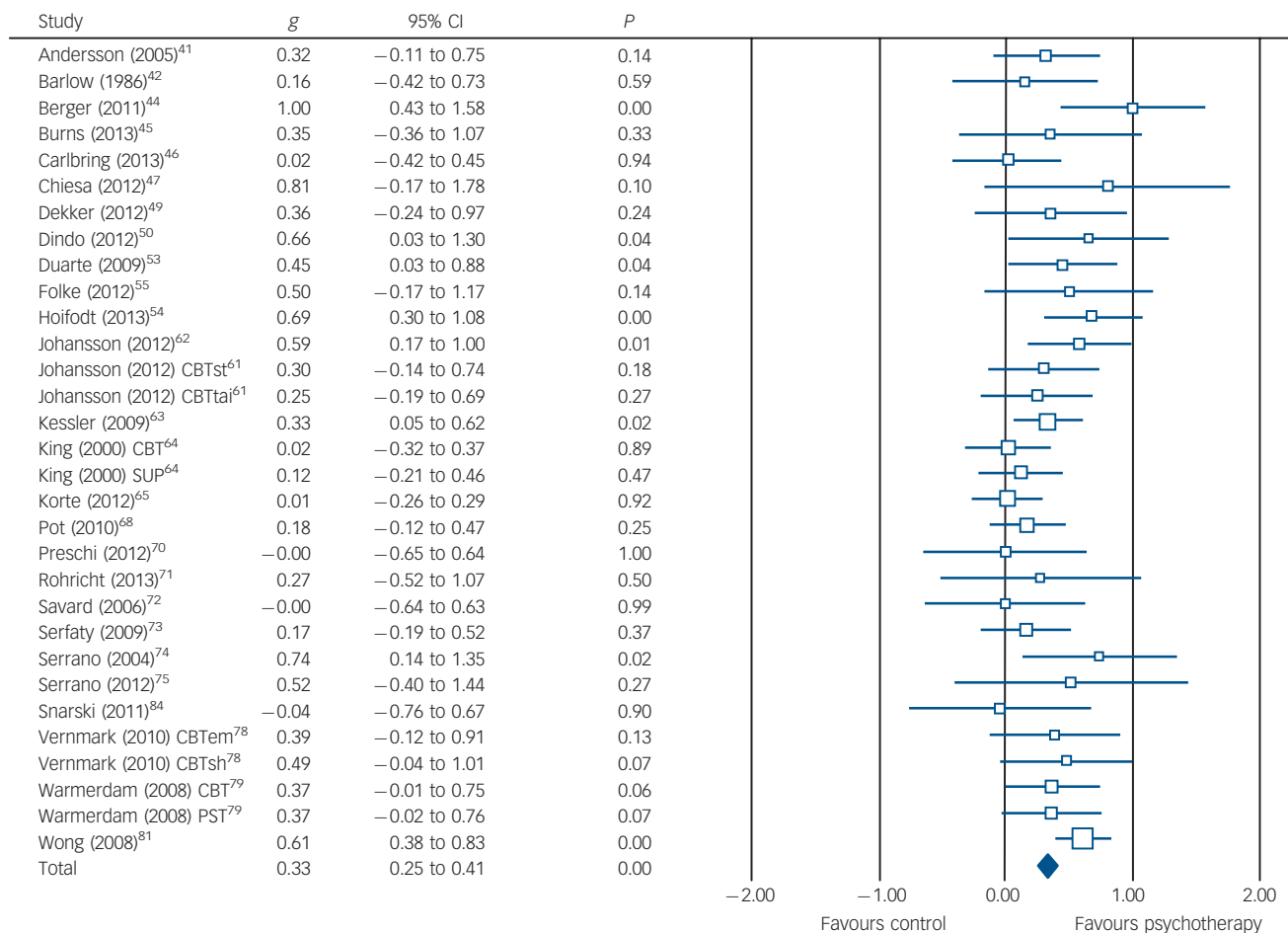


Fig. 2 Standardised effect sizes (Hedges' *g*) of psychotherapy for depression compared with control conditions on global quality of life. CBT, cognitive-behavioural therapy (ern, email therapy; sh, guided self-help; st, standard treatment; tai, tailored treatment); PST, problem-solving therapy; SUP, supportive therapy.

Table 1 Global quality of life: effect sizes in meta-analysis of studies comparing psychotherapy with a control group

Comparison ^a	Number of comparisons	Effect size		Heterogeneity ^b		
		<i>g</i>	95% CI	<i>I</i> ²	95% CI	<i>P</i>
All studies	31	0.33***	0.24–0.42	21	0–49	
Adjusted values	34	0.30	0.21–0.40			
Effect size for depression	31	0.60***	0.50–0.70	32	0–56	
Adjusted values	36	0.54	0.44–0.64			
One effect size per study (highest)	27	0.35***	0.25–0.45	25	0–53	
One effect size per study (lowest)	27	0.34***	0.24–0.45	28	0–55	
QOLI	8	0.32***	0.15–0.48	0	0–56	
EQ-5D	8	0.19**	0.07–0.32	0	0–56	
Subgroup analyses						
MDD						
Yes	15	0.49***	0.36–0.61	0	0–46	0.002
No	16	0.23***	0.13–0.34	6	0–48	
Comorbidity						
Yes	10	0.23**	0.08–0.38	0	0–53	0.148
No	21	0.37***	0.26–0.49	31	0–59	
Control group						
Care as usual	10	0.18**	0.05–0.32	1	0–53	0.053
Waiting list	14	0.42***	0.28–0.56	23	0–59	
Other	7	0.33***	0.16–0.50	0	0–58	
Intent-to-treat analysis						
Yes	27	0.34***	0.24–0.43	25	0–53	0.895
No	4	0.32*	0.03–0.60	5	0–69	
Treatment type						
Individual	10	0.17*	0.01–0.33	0	0–53	0.258
Group	8	0.35**	0.13–0.57	52	0–77	
Internet-based treatment						
Yes	12	0.41***	0.28–0.53	3	0–51	0.331
No	19	0.27***	0.15–0.40	26	0–57	
Target group						
Adults in general	23	0.39***	0.29–0.49	11	0–47	0.009
Older adults	8	0.15	–0.00–0.30	0	0–56	
CBT v. other						
CBT	20	0.36***	0.25–0.47	21	0–54	0.429
Other	11	0.28***	0.13–0.44	18	0–60	
Life review v. other						
Life review	5	0.19	–0.05–0.42	27	0–73	0.171
Other	26	0.37***	0.26–0.43	11	0–45	

CBT, cognitive-behavioural therapy; MDD, major depressive disorder; QoL, quality of life; QOLI, Quality of Life Inventory.
a. The data presented here are from analysis using the random effects model.
b. Variance between studies as a proportion of the total variance; heterogeneity tested using the *I*² statistic.
P*<0.05, *P*<0.01, ****P*<0.001.

component of QoL compared with a control condition. The mean effect size was $g=0.42$ (95% CI 0.33–0.51). We detected low between-study heterogeneity ($I^2=3$, 95% CI 0–54). After adjustment for publication bias the mean effect size was $g=0.37$ (95% CI 0.28–0.47), with five imputed studies (Table 2). Additionally, we performed a meta-analysis including the largest effect size of each study, which resulted in an overall effect size of $g=0.43$ (95% CI 0.32–0.53). When the smallest effect size was included the pooled effect size was $g=0.40$ (95% CI 0.31–0.50). Finally we conducted a series of subgroup analyses, but none of the included moderators was significantly related to the effect size of the mental health component (Table 2). For this group of studies the mean effect size for depressive symptoms was $g=0.48$ (95% CI 0.36–0.60). Between-study heterogeneity was moderate to high ($I^2=55$, 95% CI 17–72). After adjusting for publication bias the effect size decreased ($g=0.41$, 95% CI 0.28–0.54), with four missing studies.

Meta-regression analysis indicated a significant association between the effect size of the mental health component of QoL and the effect size of depressive symptoms at post-treatment measurement (slope 0.49, 95% CI 0.19–0.80, $P=0.003$). The results suggested that with each increase in effect size of depressive symptom severity by 1 the effect size for the mental health

component of QoL increased by 0.49 (Fig. 3). The effect size of the mental health component was not significantly related to the number of treatment sessions (slope 0.01, 95% CI –0.01 to 0.03, $P=0.544$). The multivariate meta-regression analysis demonstrated that only the effect size of depression severity was a significant predictor of the effect of psychotherapy on the mental health component QoL ($b=0.50$, 95% CI 0.16–0.83, $P=0.007$; see Table 4).

Physical health

Fourteen comparisons were included in the meta-analysis of the physical health component of QoL (Table 3). The mean effect size was $g=0.27$ (95% CI 0.07–0.46). We detected high between-study heterogeneity ($I^2=70$, 95% CI 41–81). We repeated the analysis after removing an outlier with an effect size of $g=2.16$.³⁰ The pooled effect size decreased ($g=0.16$, 95% CI 0.05–0.27); in addition heterogeneity was low ($I^2=4$, 95% CI 0–49). We adjusted for publication bias and the mean effect size decreased to $g=0.13$ (95% CI 0.01–0.25), with two studies missing. The meta-analysis including the largest effect sizes of each study resulted in an overall effect size of $g=0.16$ (95% CI 0.04–0.28). Next we included the smallest effect sizes, resulting in an overall

Table 2 Mental health component of quality of life: effect sizes in meta-analysis of studies comparing psychotherapy with a control group

Comparison ^a	Number of comparisons	Effect size		Heterogeneity ^b		P
		g	95% CI	I ²	95% CI	
All studies	21	0.42***	0.33–0.51	23	0–54	
Adjusted values	26	0.37	0.28–0.47			
Effect size for depression	20	0.48***	0.36–0.60	55	17–72	
Adjusted values	24	0.41	0.28–0.54			
One effect size per study (highest)	18	0.43***	0.32–0.53	29	0–59	
One effect size per study (lowest)	18	0.40***	0.31–0.50	17	0–53	
Subgroup analyses						
MDD						
Yes	6	0.49***	0.34–0.64	0	0–61	0.277
No	15	0.39***	0.28–0.50	34	0–63	
Comorbidity						
Yes	7	0.34**	0.12–0.56	46	0–76	0.428
No	14	0.43***	0.35–0.52	1	0–48	
Control group						
Care as usual	13	0.38***	0.25–0.51	37	0–66	0.242
Waiting list	7	0.48***	0.36–0.61	0	0–58	
Target group						
Adults in general	13	0.43***	0.33–0.53	24	0–60	0.350
Women	4	0.54**	0.18–0.90	37	0–78	
Older adults	4	0.29**	0.09–0.48	4	0–69	
CBT v. other						
CBT	13	0.40***	0.28–0.53	35	0–65	0.585
Other	8	0.45***	0.33–0.57	0	0–56	

CBT, cognitive-behavioural therapy; MDD, major depressive disorder.

a. The data presented here are from analysis using the random effects model.

b. Variance between studies as a proportion of the total variance; heterogeneity tested using the I² statistic.

*P<0.05, **P<0.01, ***P<0.001.

Table 3 Physical health component of quality of life: effect sizes in meta-analysis of studies comparing psychotherapy with a control group

Comparison ^a	Number of comparisons	Effect size		Heterogeneity ^b		P
		g	95% CI	I ²	95% CI	
All studies	14	0.27**	0.07–0.46	70	41–81	
Outlier removed ^c	13	0.16**	0.05–0.27	4	0–49	
Adjusted values	15	0.13	0.01–0.25			
Effect size for depression	14	0.52***	0.38–0.66	38	0–66	
Adjusted values	17	0.44	0.28–0.59			
One effect size per study						
Highest	12	0.16	0.04–0.28	18	0–58	
Lowest	12	0.16	0.04–0.27	16	0–57	
Subgroup analyses						
MDD						
Yes	4	0.11	–0.11–0.32	13	0–72	0.528
No	9	0.19**	0.05–0.32	18	0–62	
Comorbidity						
Yes	6	0.18*	0.02–0.33	13	0–66	0.843
No	7	0.15	–0.02–0.33	22	0–67	
Treatment type						
Individual	10	0.17*	0.03–0.31	32	0–66	0.934
Group	3	0.18	–0.07–0.43	0	0–90	
Target group						
Adults in general	7	0.16*	0.04–0.28	0	0–58	0.952
Women	3	0.07	–0.49–0.62	65	0–88	
Older adults	3	0.16	–0.05–0.36	0	0–72	
CBT v. other						
CBT	10	0.18**	0.07–0.30	1	0–53	0.700
Other	3	0.12	–0.18–0.41	46	0–84	

CBT, cognitive-behavioural therapy; MDD, major depressive disorder.

a. The data presented here are from analysis using the random effects model.

b. Variance between studies as a proportion of the total variance; heterogeneity tested using the I² statistic.c. Outlier's effect size g=2.16 (Scheidt *et al*).³⁰

*P<0.05, **P<0.01, ***P<0.001.

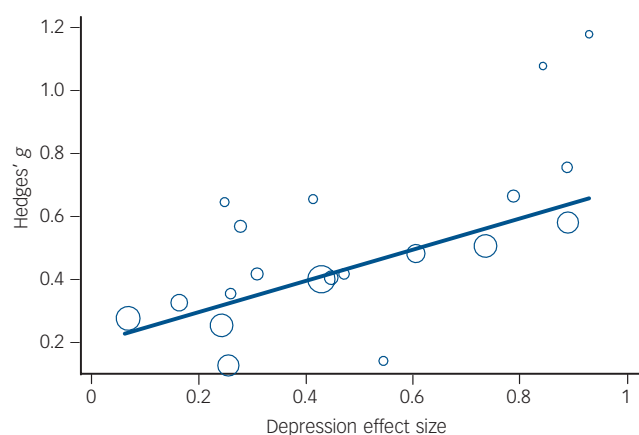


Fig. 3 Relationship between effect sizes for depressive symptom severity and the mental health component of quality of life.

effect size of $g=0.16$ (95% CI 0.04–0.27). Finally we conducted a number of subgroup analyses, but none of the included moderators was significantly associated with the effect size of the physical health component of QoL. The mean effect size of depression severity was $g=0.52$ (95% CI 0.38–0.66). Heterogeneity was moderate ($I^2=38$, 95% CI 0–66). After adjustment for publication bias the mean effect size decreased to $g=0.44$ (95% CI 0.28–0.59), with three missing studies.

Univariate meta-regressions showed no significant association between the effect size of physical health component and the effect

size of depressive symptoms (slope 0.35, 95% CI -0.12 to 0.82, $P=0.129$) or the number of treatment sessions (slope 0.00, 95% CI -0.03 to 0.03, $P=0.968$). Similarly, the multivariate meta-regression demonstrated that none of the predictors was significantly related to the effect of psychotherapy for depression on the physical health component of QoL at post-treatment assessment (Table 4).

Discussion

We examined the effects of psychotherapy on QoL of people with depression, separately for global QoL and for its mental and physical health components. The results were in line with previous findings, suggesting that psychotherapy for depression is beneficial not only for depressive symptoms but also for quality of life.^{9,13} Psychotherapy resulted in larger improvements in QoL than control conditions. The largest effect size was identified for the mental health component, whereas the effect size for global QoL was moderate. The smallest effect size was detected for the physical health component, which, however, included a limited number of comparisons. Nevertheless, even after excluding an outlier and adjusting for publication bias, the effect size for the physical health component remained statistically significant. Overall, it can be concluded that psychotherapy has a positive impact on various domains of a patient's life, such as mental functioning, social and work-related relationships, level of discomfort and engagement in everyday activities. Our findings, in conjunction with previous work, demonstrate the efficacy of psychotherapy for outcomes associated with depression.³¹ Particularly, the magnitude of the improvement in QoL is comparable – even though somewhat smaller – to that in social

Table 4 Study characteristics predicting the effect size of quality of life: multivariate meta-regression

	B	95% CI	s.e.	P
Global component				
Depression effect size	0.37	−0.11 to 0.85	0.23	0.122
Number of sessions	−0.01	−0.05 to 0.02	0.02	0.380
Target group: adults v. elderly	0.00	−0.48 to 0.48	0.23	0.999
Control group: care as usual v. waiting list	−0.05	−0.34 to 0.24	0.14	0.732
Control group: other v. waiting list	0.03	−0.26 to 0.28	0.13	0.824
CBT v. other	0.00	−0.24 to 0.22	0.11	0.994
Life review v. other	−0.23	−0.71 to 0.25	0.23	0.326
MDD	0.12	−0.08 to 0.32	0.11	0.272
Comorbidity	−0.22	−0.53 to 0.08	0.15	0.143
Internet-based treatment	−0.11	−0.38 to 0.15	0.13	0.370
Constant	0.34	−0.17 to 0.85	0.24	0.177
Mental health component				
Depression effect size	0.50	0.16 to 0.83	0.15	0.007
Number of sessions	−0.01	−0.04 to 0.02	0.01	0.466
Target group: adults v. elderly	0.07	−0.24 to 0.38	0.14	0.621
Target group: women v. elderly	0.22	−0.27 to 0.70	0.22	0.345
CBT v. other	0.03	−0.23 to 0.20	0.09	0.783
MDD	0.18	−0.17 to 0.22	0.18	0.319
Comorbidity	−0.06	−0.30 to 0.17	0.11	0.580
Constant	0.21	−0.15 to 0.57	0.17	0.223
Physical health component				
Depression effect size	0.34	−1.05 to 1.73	0.50	0.535
Number of sessions	0.01	−0.30 to 0.25	0.09	0.902
Target group: adults v. elderly	−0.18	−2.12 to 1.17	0.70	0.815
Target group: women v. elderly	−0.29	−1.66 to 1.07	0.49	0.587
CBT v. other	−0.06	−1.25 to 1.14	0.43	0.903
MDD	−0.05	−1.08 to 0.99	0.37	0.902
Comorbidity	0.05	−1.61 to 1.71	0.60	0.935
Individual treatment	−0.04	−0.91 to 0.84	0.32	0.912
Constant	0.07	−3.54 to 3.68	1.30	0.980

CBT, cognitive-behavioural therapy; MDD, major depressive disorder.

functioning of patients with depression.³¹ The influence of psychotherapy on domains of patients' lives other than the depressive symptoms is important because it can reduce the overall burden caused by the disease and decrease the risk of future depressive episodes.^{12,32}

We examined the association between QoL and depressive symptom severity by conducting various meta-regression analyses. The results were different for the global QoL and each of the two specific components of QoL. We found a significant positive relationship between the effect sizes for the mental health component and the effect sizes for depressive symptoms in the multivariate model. Nevertheless, the effect sizes for the physical health component were not related to the effect sizes for depressive symptoms. Finally, the relationship between the effect sizes for global QoL and depressive symptom severity were not statistically significant in the multivariate model. Overall, changes in QoL were not fully explained by changes in depressive symptoms. We can thus infer that decreased depressive symptom severity at the end of the treatment is not necessarily a manifestation of improvement in QoL of the patient or *vice versa*. This is an indication that QoL and depressive symptoms are two different constructs and that it is informative to use both as treatment outcomes. It should be highlighted that the applied research design did not allow us to determine causal or temporal relationships between QoL and depressive symptoms. A longitudinal design with repeated measurements of QoL and depressive symptoms is needed to examine whether changes in depressive symptoms lead to changes in QoL or *vice versa*.³³

The effect sizes for depressive symptom severity were larger than the effect sizes for QoL. This finding is in line with previous studies demonstrating that impairments in QoL persist even after patients reach remission.^{14,15} As a result, distortion in daily life may endure even when deficits related to depressive symptoms have ceased.⁹ A possible explanation is that improvements in QoL follow a slower pace than those in depressive symptoms.¹³ In addition, participants were eligible for the clinical trials because they experienced depressive symptoms and not because they reported low QoL. Therefore, they may not all have had low QoL to start with and thus had less to gain in terms of improvement in QoL. Finally, the detected difference in the effect sizes may also reflect the target of psychotherapy on reducing depressive symptom severity.

We found that the effects of psychotherapy on global QoL were larger for studies including exclusively participants with a diagnosis of major depressive disorder at baseline. Previous research has shown that deterioration of QoL is proportional to the severity of depression.^{34,35} Psychotherapy is therefore an effective treatment for people who experience both severe depressive symptoms and serious distress in their QoL. Furthermore, studies including adult patients yielded larger effect sizes than those including older adults. Older patients with depression demonstrate extensive age-related needs that may obstruct the efficacy of psychotherapy.³⁶ In addition, debilitated QoL in older adults may be related to risk factors other than depression, which are not the explicit target of psychotherapy.³⁷

Limitations of the study

The concept of QoL is inherently subjective and consequently is hard to measure with precision. It is thus possible that the different measures of QoL assessed slightly different constructs or parts of life. This limitation, however, applies mainly to the meta-analysis of global QoL where various instruments were included. The meta-analyses of the mental and physical health components were measured predominantly with the Medical

Outcome Study Short Form. In addition, the number of studies in the meta-analysis of the physical health component was limited. Thus, we may have lacked adequate power to detect small effect sizes. A larger number of studies would allow us to interpret the results with more confidence, and therefore we strongly recommend the administration of measures of QoL in future clinical trials.¹⁴ Another limitation relates to the quality of the included studies, which was not ideal. Researchers are encouraged to follow precisely the recommended guidelines for conducting and reporting randomised trials.³⁸ Moreover, a concern for every meta-analysis is the prevalence of publication bias. The test of publication bias that we performed examines only whether the funnel plot is symmetrical or whether studies with small sizes are missing. This procedure may have limited power to detect moderate publication bias and accordingly our results may overestimate the true effect size of psychotherapy on QoL.³⁹ Furthermore, we examined only the short-term effects of psychotherapy; there is evidence that psychotherapy has long-term effects on depressive symptoms,⁴⁰ and it is therefore important to examine the possible long-term effects on QoL as well. Finally, we examined the effects of psychotherapy on QoL in comparison with control conditions. A meta-analysis focusing on the respective comparison between psychotherapy and pharmacotherapy for depression would be of importance.

Implications

Overall, this meta-analysis demonstrates that psychotherapy for depression is related to improvements in QoL. This evidence amplifies the notion that psychotherapy is beneficial not only for reducing depressive symptoms but also for improving additional outcomes related to depression. These effects are expected to reduce the enormous burden caused by depression and improve the lives of people with the disorder. Finally, the results indicate that QoL and depressive symptoms are two different constructs, and thus QoL could be assessed as an additional treatment outcome. Since the effects of psychotherapy are different for each component of QoL, it is informative to use specific scores for each domain, and not only an overall score for global QoL.

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Search (("Affective Symptoms"[MESH] OR "depression"[MESH] OR "mood disorders"[MESH] AND "randomized controlled trial") AND Clinical Trial[ptyp] AND ("2013/01/01"[PDat] : "2014/12/31"[PDat]) AND Humans[Mesh] AND adult[MeSH])

Filters: Publication date from 2013/01/01 to 2014/12/31; Humans; Adult: 19+ years

First author, year	Format ^a	Target group	Sessions	Intervention; control ^b (n)	HR-QoL measures ^c	Depression measures ^d	Trial quality ^e	Country ^f
Andersson, 2005 ⁴¹	GSH	Adults	5	CBT+DG (36); DG (49)	QOLI	MADRS; BDI	+++ -	SW
Barlow, 1986 ⁴²	GRP	Older adults	6	REM (22); WL (23)	LSI	CES-D	- - + -	USA
Beeber, 2010 ⁴³	GSH	Women	16	IPT (34); CAU (37)	SF-12	CES-D	- + + +	USA
Berger, 2011 ⁴⁴	GSH	Adults with MDD	10	CBT (25); WL(26)	WHOQOL	BDI-II	+ + + +	SL+GE
Burns, 2013 ⁴⁵	IND	Women	12	CBT (16); CAU (13)	EQ-5D	EPDS; PHQ-9	+ + - +	UK
Carlbring, 2013 ⁴⁶	GSH	Adults with MDD	7	BAT (40); WL (40)	QOLI	BDI-II; MADRS	+ + + +	SW
Chiesa, 2012 ⁴⁷	GRP	Adults with MDD	8	MBCT (9); PE (7)	PGWBI	HRSD	+ + + -	IT
Cramer, 2011 ⁴⁸	GRP	Women	12	CBT (48); CAU (19)	SF-12	PHQ-9	+ + + +	UK
Dekker, 2012 ⁴⁹	GSH	Adults with MDD and heart failure	2	CBT (20); CAU (21)	MLHFQ	BDI-II	+ + + +	US
Dindo, 2012 ⁵⁰	GRP	Adults with MDD and migraine	1	ACT (31); WL (14)	SF-36	HRSD; IDAS-D	+ - - -	US
Dobkin, 2011 ⁵¹	IND	Adults with Parkinson's disease	10	CBT (41); CAU (39)	SF-36	HRSD; BDI	+ + + +	US
Dowrick, 2000 ⁵²	GRP; GSH	Adults	12; 6	CWD (80); PST (128); CAU (139)	SF-36	BDI	+ + + +	UK

Duarte, 2009 ⁵³	GRP	Adults with MDD and haemodialysis	12	CT (41); CAU (40)	KDQOL-SF	BDI	++++	BR
Fledderus, 2012 ⁵⁴	GSH	Adults	9	ACT (125); WL (126)	SF-36	CES-D	++++	NL
Folke, 2012 ⁵⁵	GSH/GRP	Adults with MDD and unemployed	6	ACT (18); CAU (16)	WHOQOL	BDI	+ - + -	
Freedland, 2009 ⁵⁶	IND	Adults with coronary artery bypass surgery	12	CBT (41); SUP (42); CAU (40)	SF-36	HRSD; BDI	++++	
Harley, 2008 ⁵⁷	GRP	Adults with MDD	16	DBT (10); WL (9)	SOS	HRSD; BDI	- + + -	
Haringsma, 2006 ⁵⁸	GRP	Older adults	10	CWD (52); WL (58)	SF-20	CES-D; HRSD	- - + +	
Høifødt, 2013 ⁵⁹	IND/GSH	Adults	6	CBT (52); WL (54)	EQ-5D	BDI-II; HRSD	+ + - -	
Hunter, 2012 ⁶⁰	GRP	Adults with substance use disorders	18	CBT (47); CAU (26)	SF-12	BDI-II	- - + +	
Johansson, 2012 ⁶¹	GSH	Adults with MDD and comorbid symptoms	8	CBTst (37); CBTtai (36); DG (42)	QOLI	BDI-II; MADRS-S	++++	
Johansson, 2012 ⁶²	GSH	Adults with MDD	9	DYN (46); DG (46)	QOLI	BDI-II; PHQ-9; MADRS	++++	
Kessler, 2009 ⁶³	GSH	Adults	10	CBT (103); WL (91)	EQ-5D; SF-12	BDI	++++	
King, 2000 ⁶⁴	IND	Adults with comorbid anxiety	6	CBT (63); SUP (67); CAU (67)	EQ-5D	BDI	++++	
Korte, 2012 ⁶⁵	GRP	Older adults	8	LR (100); CAU (102)	EQ-5D	CES-D	++++	
Laidlaw, 2008 ⁶⁶	IND	Older adults with MDD	8	CBT (20); CAU (20)	WHOQOL	BDI; GDS; HRSD	+ + + -	
Lamers, 2010 ⁶⁷	IND	Chronically ill older adults	6	MPI (183); CAU (178)	SF-36	BDI	++++	
Pot, 2010 ⁶⁸	GRP	Older adults	12	LR (83); EV (88)	EQ-5D; MANSA	CES-D	+ + + -	NL
Pots, 2014 ⁶⁹	GRP	Mild and	11	MBCT (76);	SF-36	CES-D	+ + - +	NL

		moderate depression		WL (75)				
Preschl, 2012 ⁷⁰	IND	Older adults	8	LR (20); WL (16)	LSI; WHO-5	BDI-II	++++	SL
Röhricht, 2013 ⁷¹	GRP	Recurrent MDD	20	BPT (11); WL (12)	MANSA	HRSD	++++	UK
Savard, 2006 ⁷²	IND	Women with breast cancer	8	CT (21); WL (16)	QLQ	HRSD; BDI	+ + - +	CA
Scheidt, 2013 ³⁰	IND	Women with fibromyalgia syndrome	25	DYN (23); CAU (23)	SF-36	HRSD	++++	GE
Serfaty, 2009 ⁷³	IND	Older adults	12	CBT (64); CAU (55)	EQ-5D	BDI-II	++++	UK
Serrano, 2004 ⁷⁴	IND	Older adults	4	LR (20); CAU (9)	LSI	CES-D	- - + -	SP
Serrano, 2012 ⁷⁵	IND	Older adults with MDD	6	LR (23); PL (8)	LSI; QLSD	GDS	+ - + -	SP
Strong, 2008 ⁷⁶	IND	Adults with MDD and cancer	10	PST (91); CAU (92)	EQ-5D	SCL-20	++++	UK
Talbot, 2011 ⁷⁷	IND	Women with MDD and history of sexual abuse	16	IPT (37); CAU (33)	SF-36	BDI-II; HRSD	+ - - -	US
Vernmark, 2010 ⁷⁸	GSH	Adults with MDD	7	CBTem (27); CBTsh (29); WL (29)	QOLI	MADRS; BDI	++++	NL
Warmerdam, 2008 ⁷⁹	GSH	Adults	8; 5	CBT (46); PST (42); WL (63)	EQ-5D	CES-D	++++	NL
Wiles, 2013 ⁸⁰	IND	Treatment resistant depression	18	CBT+CAU (201); CAU (209)	SF-12	BDI; PHQ-9	++++	UK
Wong, 2008 ⁸¹	GRP	Adults with MDD	10	CBT (163); WL (159)	Q-LES	C-BDI	- + + +	HK
Wuthrich, 2013 ⁸²	GRP	Older adults with anxiety	12	CBT (27); WL (35)	SF-12	GDS	++++	AU
Zilcha-Mano, 2014 ⁸³	IND	Adults with MDD	20	SET (51); PL (50)	SF-36	BDI	++++	US

a. GSH, Guided self-help via internet (except Fledderus *et al*);⁵⁴ GRP, group treatment; IND, individual treatment.

b. ACT, acceptance and commitment therapy; BAT, behavioural activation treatment; BPT, body psychotherapy; CAU, care as usual; CBT, cognitive-behavioural therapy; CT, cognitive therapy; CWD, Coping With Depression; DBT, dialectical behaviour therapy; DG, discussion groups; DYN, psychodynamic psychotherapy; EV, educational video; IPT, interpersonal psychotherapy; LR, life review; MBCT, mindfulness-based cognitive therapy; MPI, minimal psychological intervention; PE, psychoeducation; PL, placebo; PST, problem solving treatment; REM, reminiscence therapy; SET, supportive expressive therapy; SUP, supportive therapy; WL, waiting list.

c. EQ-5D, EuroQol; KDQOL-SF, Kidney Disease and Quality of Life-Short Form; LSI, Life Satisfaction Index; MANSA, Manchester Short Assessment of Quality of Life; MLHFQ, Minnesota Living with Heart Failure Questionnaire; PGWB, Psychological General Well-being Index; QLQ, Quality of Life Questionnaire; Q-LES, Quality of Life Enjoyment and Satisfaction; QOLI, Quality of Life Inventory; SF, Short-Form Health Survey; WHO-5, Well-Being Index; WHOQOL, World Health Organization Quality of Life.

d. BDI, Beck Depression Inventory; BDI-II, Beck Depression Inventory-II; C-BDI, Beck Depression Inventory Chinese edition; CES-D, Center for Epidemiologic Studies Depression Scale; EPDS, Edinburgh Postnatal Depression Scale; GDS, Geriatric Depression Scale; HRSD, Hamilton Rating Scale for Depression; IDAS-D, Inventory of Anxiety and Depression Symptoms- General Depression Scale; MADRS, Montgomery-Åsberg Depression Rating Scale; PHQ-9, Patient Health Questionnaire; SCL-20, Symptom Checklist – Depression Scale.

e. Trial quality: intention to treat analysis, allocation concealment, blinding of outcome assessment, random sequence generation; + low risk of bias, – high risk of bias.

f. AU, Australia; BR, Brazil; CA, Canada; GE, Germany; IT, Italy; HK, Hong Kong; NL, Netherlands; SL, Switzerland; SP, Spain; SW, Sweden; UK, United Kingdom; US, United States.

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**Supplementary
Material**

Supplementary material can be found at:
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References

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