

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.

Copyright and usage

© The Royal College of Psychiatrists 2016.

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).

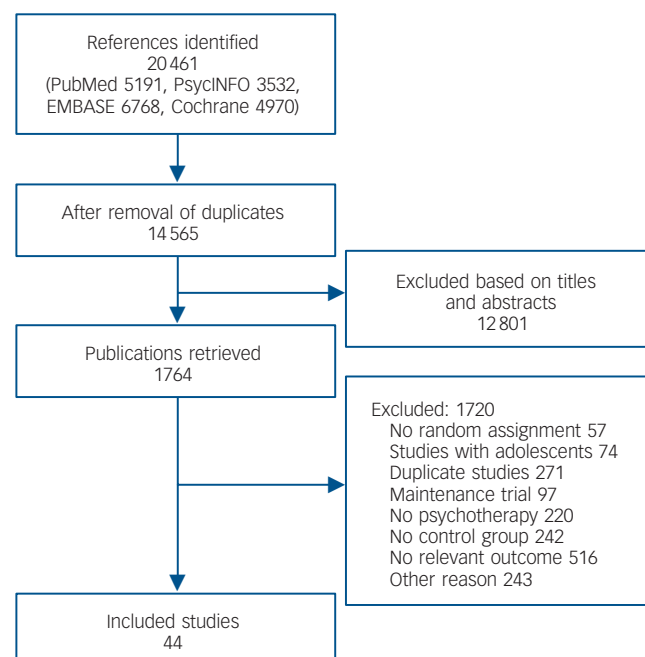


Fig. 1 Study selection.

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

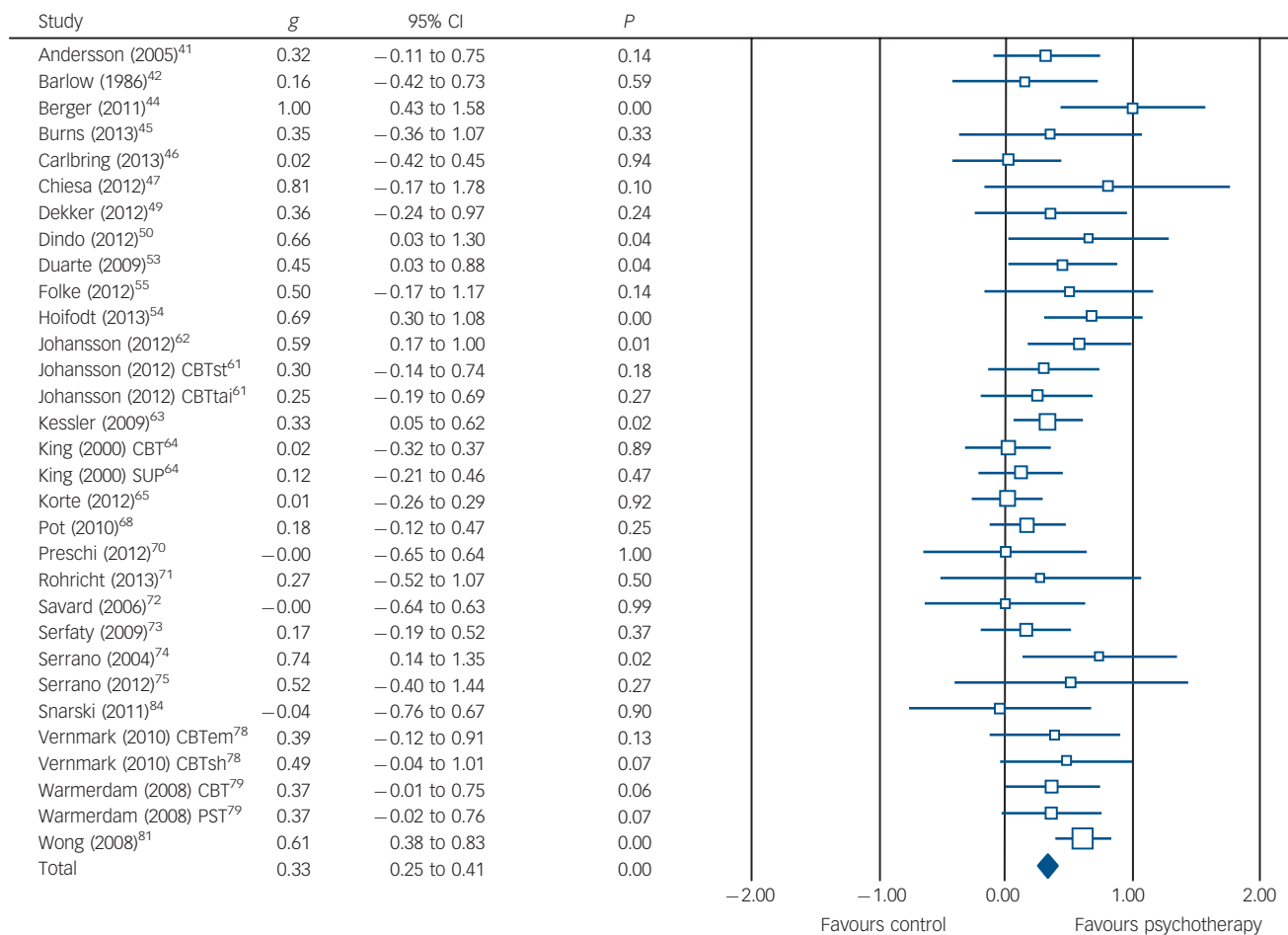


Fig. 2 Standardised effect sizes (Hedges' *g*) of psychotherapy for depression compared with control conditions on global quality of life. CBT, cognitive-behavioural therapy (ern, email therapy; sh, guided self-help; st, standard treatment; tai, tailored treatment); PST, problem-solving therapy; SUP,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

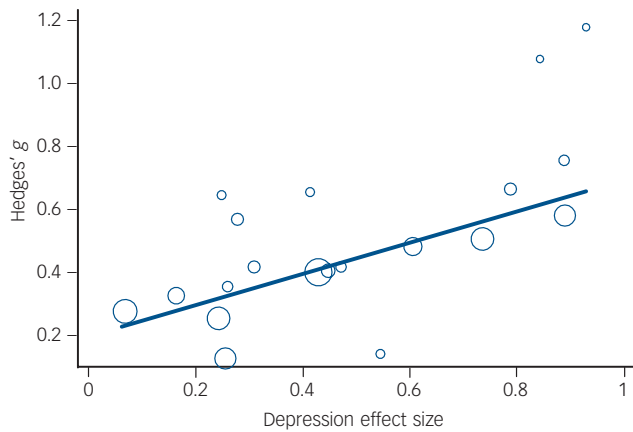


Fig. 3 Relationship between effect sizes for depressive symptom severity and the mental health component of quality of life.

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.

Spyros Kolovos, MSc, Department of Health Sciences, VU University Amsterdam, EMGO Institute for Health and Care Research, Amsterdam, The Netherlands; **Annet Kleiboer**, PhD, Department of Clinical Psychology, VU University Amsterdam, EMGO Institute for Health and Care Research, VU University, Department of Clinical and Health Psychology, Utrecht University, The Netherlands; **Pim Cuijpers**, PhD, Department of Clinical Psychology, VU University Amsterdam, EMGO Institute for Health and Care Research, VU University and VU University Medical Centre Amsterdam, The Netherlands, and Leuphana University, Lüneburg, Germany

Correspondence: Spyros Kolovos, Department of Health Sciences, EMGO Institute for Health and Care Research, De Boelelaan 1085, 1081 HV Amsterdam, The Netherlands. Email: s.kolovos@vu.nl

First received 2 Sep 2015, final revision 14 Jan 2016, accepted 27 Apr 2016

Acknowledgements

We thank Eirini Karyotaki, David Turner and Erica Weitz for their contribution in various parts of this meta-analysis.

References

- 1 Hasin DS, Goodwin RD, Stinson FS, Grant BF. Epidemiology of major depressive disorder: results from the National Epidemiologic Survey on Alcoholism and Related Conditions. *Arch Gen Psychiatry* 2005; **62**: 1097–106.
- 2 Waraich P, Goldner EM, Somers JM, Hsu L. Prevalence and incidence studies of mood disorders: a systematic review of the literature. *Can J Psychiatry* 2004; **49**: 124–38.

- 3 Mathers CD, Loncar D. Projections of global mortality and burden of disease from 2002 to 2030. *PLoS Med* 2006; **3**: e442.
- 4 Olesen J, Gustavsson A, Svensson M, Wittchen HU, Jonsson B. The economic cost of brain disorders in Europe. *Eur J Neurol* 2012; **19**: 155–62.
- 5 Sobocki P, Jonsson B, Angst J, Rehnberg C. Cost of depression in Europe. *J Ment Health Policy Econ* 2006; **9**: 87–98.
- 6 Greenberg PE, Kessler RC, Birnbaum HG, Leong SA, Lowe SW, Berglund PA, et al. The economic burden of depression in the United States: how did it change between 1990 and 2000? *J Clin Psychiatry* 2003; **64**: 1465–75.
- 7 Saarni SI, Suvisaari J, Sintonen H, Pirkola S, Koskinen S, Aromaa A, et al. Impact of psychiatric disorders on health-related quality of life: general population survey. *Br J Psychiatry* 2007; **190**: 326–32.
- 8 Ustun TB, Ayuso-Mateos JL, Chatterji S, Mathers C, Murray CJL. Global burden of depressive disorders in the year 2000. *Br J Psychiatry* 2004; **184**: 386–92.
- 9 Papakostas GI, Petersen T, Mahal Y, Mischoulon D, Nierenberg AA, Fava M. Quality of life assessments in major depressive disorder: a review of the literature. *Gen Hosp Psychiatry* 2004; **26**: 13–7.
- 10 Cuijpers P, Geraedts AS, van Oppen P, Andersson G, Markowitz JC, van Straten A. Interpersonal psychotherapy for depression: a meta-analysis. *Am J Psychiatry* 2011; **168**: 581–92.
- 11 Cuijpers P, Smit F, Bohlmeijer E, Hollon SD, Andersson G. Efficacy of cognitive-behavioural therapy and other psychological treatments for adult depression: meta-analytic study of publication bias. *Br J Psychiatry* 2010; **196**: 173–8.
- 12 Ezquiaga E, Garcia-Lopez A, de Dios C, Leiva A, Bravo M, Montejó J. Clinical and psychosocial factors associated with the outcome of unipolar major depression: a one year prospective study. *J Affect Disord* 2004; **79**: 63–70.
- 13 McKnight PE, Kashdan TB. The importance of functional impairment to mental health outcomes: a case for reassessing our goals in depression treatment research. *Clin Psychol Rev* 2009; **29**: 243–59.
- 14 Ishak WW, Greenberg JM, Balayan K, Kapitanski N, Jeffrey J, Fathy H, et al. Quality of life: the ultimate outcome measure of interventions in major depressive disorder. *Harv Rev Psychiatry* 2011; **19**: 229–39.
- 15 Zimmerman M, Chelminski I, McGlinchey JB, Young D. Diagnosing major depressive disorder VI: performance of an objective test as a diagnostic criterion. *J Nerv Ment Dis* 2006; **194**: 565–9.
- 16 Cuijpers P, van Straten A, Warmerdam L, Andersson G. Psychological treatment of depression: a meta-analytic database of randomized studies. *BMC Psychiatry* 2008; **8**: 36.
- 17 Higgins J, Green S. *Cochrane Handbook for Systematic Reviews of Interventions Version 5.1.0*. Cochrane Collaboration, 2011.
- 18 Higgins JP, Altman DG, Gotzsche PC, Juni P, Moher D, Oxman AD, et al. The Cochrane Collaboration's tool for assessing risk of bias in randomised trials. *BMJ* 2011; **343**: d5928.
- 19 Borenstein M, Hedges L, Higgins J, Rothstein H. *Comprehensive Meta-analysis Version 2*: 104. Biostat, 2005.
- 20 Harbord R, Higgins J. *Metareg: Stata Module to Perform Meta-analysis Regression*. Statistical Software Components, 2009.
- 21 Hedges LV, Olkin I. *Statistical Method for Meta-analysis*. Academic Press, 2014.
- 22 Hedges LV, Vevea JL. Fixed-and random-effects models in meta-analysis. *Psychol Methods* 1998; **3**: 486.
- 23 Higgins, Thompson SG, Deeks JJ, Altman DG. Measuring inconsistency in meta-analyses. *BMJ* 2003; **327**: 557–60.
- 24 Ioannidis JPA, Patsopoulos NA, Evangelou E. Uncertainty in heterogeneity estimates in meta-analyses. *BMJ* 2007; **335**: 914–6.
- 25 Orsini N, Bottai M, Higgins J, Buchan I. *Heterogi: Stata Module to Quantify Heterogeneity in a Meta-analysis*. Statistical Software Components, 2006.
- 26 Duval S, Tweedie R. Trim and fill: a simple funnel-plot-based method of testing and adjusting for publication bias in meta-analysis. *Biometrics* 2000; **56**: 455–63.
- 27 Borenstein M, Hedges LV, Higgins JP, Rothstein HR. *Introduction to Meta-analysis*. Wiley, 2011.
- 28 Viechtbauer W. Bias and efficiency of meta-analytic variance estimators in the random-effects model. *J Educ Behav Stat* 2005; **30**: 261–93.
- 29 Cohen J. Statistical power analysis. *Curr Dir Psychol Sci* 1992; **1**: 98–101.
- 30 Scheidt CE, Waller E, Endorf K, Schmidt S, König R, Zeeck A, et al. Is brief psychodynamic psychotherapy in primary fibromyalgia syndrome with concurrent depression an effective treatment? A randomized controlled trial. *Gen Hosp Psychiatry* 2013; **35**: 160–7.
- 31 Renner F, Cuijpers P, Huibers MJ. The effect of psychotherapy for depression on improvements in social functioning: a meta-analysis. *Psychol Med* 2014; **44**: 2913–26.
- 32 Eaton WW, Martins SS, Nestadt G, Bienvenu OJ, Clarke D, Alexandre P. The burden of mental disorders. *Epidemiol Rev* 2008; **30**: 1–14.
- 33 Kazdin AE. Mediators and mechanisms of change in psychotherapy research. *Annu Rev Clin Psychol* 2007; **3**: 1–27.
- 34 Demyttenaere K, De Fruyt J, Huygens R. Measuring quality of life in depression. *Curr Opin Psychiatry* 2002; **15**: 89–92.
- 35 Berlim M, Fleck MA. Quality of life and major depression. In *Quality of Life Impairment in Schizophrenia, Mood and Anxiety Disorders* (eds M Ritsner, AG Awad): 241–52. Springer Netherlands, 2007.
- 36 Fiske A, Wetherell JL, Gatz M. Depression in older adults. *Annu Rev Clin Psychol* 2009; **5**: 363–89.
- 37 Rejeski WJ, Mihalko SL. Physical activity and quality of life in older adults. *J Gerontol A Biol Sci Med Sci* 2001; **56**: 23–35.
- 38 Schulz KF, Altman DG, Moher D. CONSORT 2010 statement: updated guidelines for reporting parallel group randomized trials. *Ann Intern Med* 2010; **152**: 726–32.
- 39 Sterne JA, Gavaghan D, Egger M. Publication and related bias in meta-analysis: power of statistical tests and prevalence in the literature. *J Clin Epidemiol* 2000; **53**: 1119–29.
- 40 Cuijpers P, Hollon SD, van Straten A, Bockting C, Berking M, Andersson G. Does cognitive behaviour therapy have an enduring effect that is superior to keeping patients on continuation pharmacotherapy? A meta-analysis. *BMJ Open* 2013; **3**: e002542.
- 41 Andersson G, Bergstrom J, Hollandare F, Carlbring P, Kalso V, Ekselius L. Internet-based self-help for depression: randomised controlled trial. *Br J Psychiatry* 2005; **187**: 456–61.
- 42 Barlow JP. *A Group Treatment for Depression in the Elderly*. University of Houston, 1986.
- 43 Beeber LS, Holditch-Davis D, Perreira K, Schwartz TA, Lewis V, Blanchard H, et al. Short-term in-home intervention reduces depressive symptoms in early Head Start Latina mothers of infants and toddlers. *Res Nurs Health* 2010; **33**: 60–76.
- 44 Berger T, Hammerli K, Gubser N, Andersson G, Caspar F. Internet-based treatment of depression: a randomized controlled trial comparing guided with unguided self-help. *Cogn Behav Ther* 2011; **40**: 251–66.
- 45 Burns A, O'Mahen H, Baxter H, Bennert K, Wiles N, Ramchandani P, et al. A pilot randomised controlled trial of cognitive behavioural therapy for antenatal depression. *BMC Psychiatry* 2013; **13**: 33.
- 46 Carlbring P, Hagglund M, Luthstrom A, Dahlin M, Kadowaki A, Vernmark K, et al. Internet-based behavioral activation and acceptance-based treatment for depression: a randomized controlled trial. *J Affect Disord* 2013; **148**: 331–7.
- 47 Chiesa A, Mandelli L, Serretti A. Mindfulness-based cognitive therapy versus psycho-education for patients with major depression who did not achieve remission following antidepressant treatment: a preliminary analysis. *J Altern Complement Med* 2012; **18**: 756–60.
- 48 Cramer H, Salisbury C, Conrad J, Eldred J, Araya R. Group cognitive behavioural therapy for women with depression: pilot and feasibility study for a randomised controlled trial using mixed methods. *BMC Psychiatry* 2011; **11**: 82.
- 49 Dekker RL, Moser DK, Peden AR, Lennie TA. Cognitive therapy improves three-month outcomes in hospitalized patients with heart failure. *J Card Fail* 2012; **18**: 10–20.
- 50 Dindo L, Recober A, Marchman JN, Turvey C, O'Hara MW. One-day behavioral treatment for patients with comorbid depression and migraine: a pilot study. *Behav Res Ther* 2012; **50**: 537–43.
- 51 Dobkin RD, Menza M, Allen LA, Gara MA, Mark MH, Tiu J, et al. Cognitive-behavioral therapy for depression in Parkinson's disease: a randomized, controlled trial. *Am J Psychiatry* 2011; **168**: 1066–74.
- 52 Dowrick C, Dunn G, Ayuso-Mateos JL, Dalgard OS, Page H, Lehtinen V, et al. Problem solving treatment and group psychoeducation for depression: multicentre randomised controlled trial. Outcomes of Depression International Network (ODIN) Group. *BMJ* 2000; **321**: 1450–4.
- 53 Duarte PS, Miyazaki MC, Blay SL, Sesso R. Cognitive-behavioral group therapy is an effective treatment for major depression in hemodialysis patients. *Kidney Int* 2009; **76**: 414–21.
- 54 Fledderus M, Bohlmeijer ET, Pieterse ME, Schreurs KM. Acceptance and commitment therapy as guided self-help for psychological distress and positive mental health: a randomized controlled trial. *Psychol Med* 2012; **42**: 485–95.
- 55 Folke F, Parling T, Melin L. Acceptance and commitment therapy for depression: a preliminary randomized clinical trial for unemployed on long-term sick leave. *Cogn Behav Pract* 2012; **19**: 583–94.

- 56 Freedland KE, Skala JA, Carney RM, Rubin EH, Lustman PJ, Davila-Roman VG, et al. Treatment of depression after coronary artery bypass surgery: a randomized controlled trial. *Arch Gen Psychiatry* 2009; **66**: 387–96.
- 57 Harley R, Sprich S, Safren S, Jacobo M, Fava M. Adaptation of dialectical behavior therapy skills training group for treatment-resistant depression. *J Nerv Ment Dis* 2008; **196**: 136–43.
- 58 Haringsma R, Engels GI, Cuijpers P, Spinhoven P. Effectiveness of the Coping With Depression (CWD) course for older adults provided by the community-based mental health care system in the Netherlands: a randomized controlled field trial. *Int Psychogeriatr* 2006; **18**: 307–25.
- 59 Hoifodt RS, Lillevoll KR, Griffiths KM, Wilsgaard T, Eisemann M, Waterloo K, et al. The clinical effectiveness of web-based cognitive behavioral therapy with face-to-face therapist support for depressed primary care patients: randomized controlled trial. *J Med Internet Res* 2013; **15**: e153.
- 60 Hunter SB, Watkins KE, Hepner KA, Paddock SM, Ewing BA, Osilla KC, et al. Treating depression and substance use: a randomized controlled trial. *J Subst Abuse Treat* 2012; **43**: 137–51.
- 61 Johansson R, Sjöberg E, Sjögren M, Johnsson E, Carlbring P, Andersson T, et al. Tailored vs. standardized internet-based cognitive behavior therapy for depression and comorbid symptoms: a randomized controlled trial. *PLoS One* 2012; **7**: e36905.
- 62 Johansson R, Ekbladh S, Hebert A, Lindstrom M, Moller S, Pettitt E, et al. Psychodynamic guided self-help for adult depression through the internet: a randomised controlled trial. *PLoS One* 2012; **7**: e38021.
- 63 Kessler D, Lewis G, Kaur S, Wiles N, King M, Weich S, et al. Therapist-delivered Internet psychotherapy for depression in primary care: a randomised controlled trial. *Lancet* 2009; **374**: 628–34.
- 64 King M, Sibbald B, Ward E, Bower P, Lloyd M, Gabbay M, et al. Randomised controlled trial of non-directive counselling, cognitive-behaviour therapy and usual general practitioner care in the management of depression as well as mixed anxiety and depression in primary care. *Health Technol Assess* 2000; **4**: 1–83.
- 65 Korte J, Bohlmeijer ET, Cappeliez P, Smit F, Westerhof GJ. Life review therapy for older adults with moderate depressive symptomatology: a pragmatic randomized controlled trial. *Psychol Med* 2012; **42**: 1163–73.
- 66 Laidlaw K, Davidson K, Toner H, Jackson G, Clark S, Law J, et al. A randomised controlled trial of cognitive behaviour therapy vs treatment as usual in the treatment of mild to moderate late life depression. *Int J Geriatr Psychiatry* 2008; **23**: 843–50.
- 67 Lamers F, Jonkers CC, Bosma H, Kempen GI, Meijer JA, Penninx BW, et al. A minimal psychological intervention in chronically ill elderly patients with depression: a randomized trial. *Psychother Psychosom* 2010; **79**: 217–26.
- 68 Pot AM, Bohlmeijer ET, Onrust S, Melenhorst AS, Veerbeek M, De Vries W. The impact of life review on depression in older adults: a randomized controlled trial. *Int Psychogeriatr* 2010; **22**: 572–81.
- 69 Pots WT, Meulenbeek PA, Veehof MM, Klungers J, Bohlmeijer ET. The efficacy of mindfulness-based cognitive therapy as a public mental health intervention for adults with mild to moderate depressive symptomatology: a randomized controlled trial. *PLoS One* 2014; **9**: e109789.
- 70 Preschl B, Maercker A, Wagner B, Forstmeier S, Banos RM, Alcaniz M, et al. Life-review therapy with computer supplements for depression in the elderly: a randomized controlled trial. *Aging Ment Health* 2012; **16**: 964–74.
- 71 Rohricht F, Papadopoulos N, Priebe S. An exploratory randomized controlled trial of body psychotherapy for patients with chronic depression. *J Affect Disord* 2013; **151**: 85–91.
- 72 Savard J, Simard S, Giguere I, Ivers H, Morin CM, Maunsell E, et al. Randomized clinical trial on cognitive therapy for depression in women with metastatic breast cancer: psychological and immunological effects. *Palliat Support Care* 2006; **4**: 219–37.
- 73 Serfaty MA, Haworth D, Blanchard M, Buszewicz M, Murad S, King M. Clinical effectiveness of individual cognitive behavioral therapy for depressed older people in primary care: a randomized controlled trial. *Arch Gen Psychiatry* 2009; **66**: 1332–40.
- 74 Serrano JP, Latorre JM, Gatz M, Montanes J. Life review therapy using autobiographical retrieval practice for older adults with depressive symptomatology. *Psychol Aging* 2004; **19**: 270–7.
- 75 Serrano Selva JP, Latorre Postigo JM, Ros Segura L, Navarro Bravo B, Aguilar Corcoles MJ, Nieto Lopez M, et al. Life review therapy using autobiographical retrieval practice for older adults with clinical depression. *Psicothema* 2012; **24**: 224–9.
- 76 Strong V, Waters R, Hibberd C, Murray G, Wall L, Walker J, et al. Management of depression for people with cancer (SMArT oncology 1): a randomised trial. *Lancet* 2008; **372**: 40–8.
- 77 Talbot NL, Chaudron LH, Ward EA, Duberstein PR, Conwell Y, O'Hara MW, et al. A randomized effectiveness trial of interpersonal psychotherapy for depressed women with sexual abuse histories. *Psychiatr Serv* 2011; **62**: 374–80.
- 78 Vernmark K, Lenndin J, Bjarehed J, Carlsson M, Karlsson J, Oberg J, et al. Internet administered guided self-help versus individualized e-mail therapy: a randomized trial of two versions of CBT for major depression. *Behav Res Ther* 2010; **48**: 368–76.
- 79 Warmerdam L, van Straten A, Twisk J, Riper H, Cuijpers P. Internet-based treatment for adults with depressive symptoms: randomized controlled trial. *J Med Internet Res* 2008; **10**: e44.
- 80 Wiles N, Thomas L, Abel A, Ridgway N, Turner N, Campbell J, et al. Cognitive behavioural therapy as an adjunct to pharmacotherapy for primary care based patients with treatment resistant depression: results of the CoBaIT randomised controlled trial. *Lancet* 2013; **381**: 375–84.
- 81 Wong DF. Cognitive behavioral treatment groups for people with chronic depression in Hong Kong: a randomized wait-list control design. *Depress Anxiety* 2008; **25**: 142–8.
- 82 Wuthrich VM, Rapee RM. Randomised controlled trial of group cognitive behavioural therapy for comorbid anxiety and depression in older adults. *Behav Res Ther* 2013; **51**: 779–86.
- 83 Zilcha-Mano S, Dinger U, McCarthy KS, Barrett MS, Barber JP. Changes in well-being and quality of life in a randomized trial comparing dynamic psychotherapy and pharmacotherapy for major depressive disorder. *J Affect Disord* 2014; **152–154**: 538–42.
- 84 Snarski M, Scogin F, DiNapoli E, Presnell A, McAlpine J, Marcinkak J. The effects of behavioral activation therapy with inpatient geriatric psychiatry patients. *Behav Ther* 2011; **42**: 100–8.

