

Best Evidence Summaries of Topics in Mental Healthcare

BEST in MH clinical question-answering service

Question

"In adults with a personality disorder how effective are brief dynamic psychotherapies, compared to any other intervention, for improving patient outcomes?"

Clarification of question using PICO structure

Patients: Adults with a personality disorder

Intervention: Brief dynamic psychotherapies / short-term psychodynamic psychotherapies (STPP)

Comparator: Any Outcome: Any



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Clinical and research implications

Two poor quality systematic reviews and one small open randomised controlled trial (RCT) with limitations in the reported analyses provided very limited information on the effectiveness of short-term psychodynamic psychotherapies for the treatment of people with personality disorders. The results reported in the systematic reviews were not sufficient to support any firm conclusions. The additional RCT provided some preliminary evidence that Psychic Representation-Focused Psychotherapy, used in addition to standard treatment, may improve a range of symptoms in people with borderline personality disorder. However, these results were for immediate, post-treatment effects only; longer-term follow-up from this study is ongoing. There is currently insufficient evidence to support the use of short-term psychodynamic psychotherapies in people with personality disorders and more research is needed.

What does the evidence say?

Number of included studies/reviews (number of participants)

We identified two systematic reviews, ^{1,2} and one additional RCT³ that reported data relevant to this evidence summary. The first systematic review assessed the effectiveness of intensive short-term dynamic psychotherapy (ISTDP), based on the Davanloo method, and included all patient groups and outcome measures. ¹ Three randomised controlled trials and one additional observational study included in this review were conducted in people with varying severities of personality disorder; the review results derived from these studies are included in this evidence summary. ¹ The second systematic review included RCTs of individual Short-Term Psychodynamic Psychotherapy (STPP) conducted in people with varying severities of personality disorder, but did not report any data for between group comparisons. ² The additional RCT was a report of preliminary, post-treatment results from a trial comparing Psychic Representation-Focused Psychotherapy (PRFP) plus conventional treatment to conventional treatment alone, for people with borderline personality disorder. ³

Main Findings

The first systematic review reported that studies in patients with personality disorders found that ISTDP was associated with pre-post-treatment improvements in symptoms and was significantly superior to waiting list control, but not significantly different from other active treatments. However, no numerical results were provided to support these statements. The second systematic review included data from seven RCTs of STPP, but did not report any between group comparisons, i.e. there were no reported estimates of the effectiveness of STPP compared to control or other active treatments. Reported meta-analyses indicated STPP was associated with pre- post-treatment improvements in measures of symptoms, Inventory of Interpersonal Problems and Global Assessment of Functioning. Finally, the additional RCT found that PRFP plus conventional treatment was associated with significant improvements in Severity Global Index (SCI-90), Zanarini score (total, relations and feeling of emptiness), Montgomery-Asberg Depression Rating Scale (MADRS) score (total and suicide), Rosemberg score, Barratt score and Social Adaptation Self-evaluation Scale (SASS) score, compared to conventional treatment alone. However, these results were for immediate post-treatment assessment only; 6-12 month follow-up is ongoing.

Authors Conclusions

One systematic review concluded that the limited evidence available supports the application of intensive short-term dynamic psychotherapy (ISTDP) across a broad range of populations. The second systematic review concluded that Short-Term Psychodynamic Psychotherapy (STPP) offers an efficacious treatment option for personality disorder, which is superior to waiting list controls, and comparable to psychodynamic and cognitive behavioural approaches. Both reviews concluded that further research is needed to support their findings. An additional RCT concluded that preliminary post-treatment results showed significantly better outcomes in people treated with Psychic Representation-Focused Psychotherapy (PRFP) plus conventional treatment than in those managed with conventional treatment alone, and that PRFP may represent an important contribution for the treatment of borderline personality disorder patients.

Reliability of conclusions/Strength of evidence

Two poor quality systematic reviews and one small open RCT with limitations in the reported analyses provided very limited information on the effectiveness of short-term psychodynamic psychotherapies for the treatment of people with personality disorders. There is currently insufficient evidence to support the use of these treatments in this population.

What do guidelines say?

Neither National Institute for Health and Clinical Excellence (NICE) and Scottish Intercollegiate Guidelines Network (SIGN) guidelines comment upon the use of STPP for treating personality disorders.

Date question received: 26/02/2008

Date searches conducted: 07/11/2014, updated from 26/02/2008

Date answer completed: 08/12/2014

References

Systematic reviews

- 1. Abbass, A., Town, J., & Driessen, E. (2012). Intensive short-term dynamic psychotherapy: A systematic review and meta-analysis of outcome research. *Harvard Review of Psychiatry*, 20(2), 97-108.
- 2. Town, J. M., Abbass, A., & Hardy, G. (2011). Short-term psychodynamic psychotherapy for personality disorders: A critical review of randomized controlled trials. *Journal of Personality Disorders*, *25*(6), 723-740.

Randomised controlled trials

Renesis, B., Galian, M., Serrano, R., Figuera, D., del Moral, A. F., López-Ibor, J. J., ... & Trujillo, M. (2013). A new Time Limited Psychotherapy for BPD: Preliminary Results of a Randomized and Controlled Trial. *Actas Esp Psiquiatr*, 41(3), 139-48.

Results

Systematic Reviews

Author	Search	Inclusion criteria	Number of	Summary of results	Risk of bias
(year)	Date		included studies		
Abbass et	Not	Participants: Any patient group.	n = 21	This review aimed to assess the	The article reports a
al. (2012)	reported	Intervention: Intensive short-term	publications, of	effectiveness of intensive short-term	broad survey of the
		dynamic psychotherapy, based upon	which 6 were	dynamic psychotherapy (ISTDP).	literature with no
		Davanloo's books or articles, either	considered		clear research
		individual or group-based.	relevant to this	The article listed four randomised controlled	objective and no
		Comparator: Any comparator.	evidence	trials (RCTs) conducted in people with	specific inclusion
		Outcome: Any outcome.	summary	personality disorder, however, results were	criteria, beyond the
		Study design : Any study design.	(conducted in	only described for three of these (all three	intervention of
			people with	were also included in Town et al. 2011, see	interest.
			personality	below); study details indicate that the fourth	
			disorders)	publication may have been an earlier	Studies were
				publication of one of the three trials	retrieved from four
				described, but this is not clear. Two further	previous meta-
				publications reported the results of an	analyses conducted
				observational study with long-term follow-	by the authors,
				up.	supplemented by up-
					date searches in
				One RCT (n=81) compared ISTDP, comprising	three bibliographic
				a mean of 40.3 one hour sessions, (n=15) to	databases (limited
				brief adaptational psychotherapy, not	search terms
				described, (n not reported) and a waiting list	reported), screening
				control (n not reported). This study was	the references of
				described as having found that ISTDP	identified articles

significantly outperformed waiting list and contact with control on patients' self-rated target experts. Only complaints (Target Complaints published studies Questionnaire), symptoms (Symptom were included. Checklist [SCL]-90) and the Social Adjustment Scale, however, no numerical Two reviewers results were reported. No differences in assessed articles for performance between ISTDP and the brief eligibility, but no adaptational psychotherapy were reported. details of the data A second RCT (n=49) compared ISTDP, extraction process of comprising a mean of 28.5 one hour any assessment of sessions, (n=25) to brief supportive the methodological psychotherapy, not described, (n=24). This quality of included study was described as having reported a studies were significant decrease in symptoms following reported. treatment (no details reported), but no significant reduction on the Inventory of The meta-analyses Interpersonal Problems (IIP) and no (not used in this differences in performance between ISTDP evidence summary) and the brief supportive psychotherapy. pooled outcomes Both studies were conducted in people with data across studies a DSM-III diagnosis of personality disorder, that included people described as "largely cluster C personality with a wide range of disorders." Follow-up was described as 6 to physical and psychiatric 18 months for these two studies combined. conditions. No The third RCT (n=27) compared ISTDP, mean numerical results of 27.7 sessions, to a minimal contact control. individual studies The study included participants with more were reported and

				severe personality disorders. It was stated that participants in the ISTDP group had significantly improved IIP, and functional measures compared to controls, and that participants in the ISTDP group had significant reductions in medication usage and an increase in employment rate and work hours, whereas controls did not. No numerical results were reported. The observational study reported results from a residential treatment program for personality disorders, in which all patients received individual ISTDP sessions in combination with group psychotherapy and different forms of non-verbal therapy. Patients' self-reported quality of interpersonal relationships improved at discharge and increased further both at 1-year and 3-year follow-up. No numerical results were reported.	some studies were omitted from the description of results altogether.
Town et al. (2011)	Not reported	Participants: Those who met specific criteria for a psychiatric diagnosis of personality disorder (not described). Intervention: Individual STPP, based upon the principles of Mann, Sifneos, Malan, Davanloo, Luborsky, with sessions lasting between 45-60 minutes, with	n = 8 RCTs (total 211 participants from 7 reported studies)	This article aimed to provide a "critical review of the literature" on Short-Term Psychodynamic Psychotherapy (STPP) for the treatment of people with personality disorders. The authors reported that eight studies were	The article reports a "critical review" of the literature. No clear research question was stated, but inclusion criteria were defined for

overall treatment length less than 40 weeks.

Comparator: Any comparator.

Outcome: Not defined. Study design: RCTs

included in the review, but details were only presented for seven studies. The types of personality disorder in the included studies varied: four studies included people with cluster types A, B and C (one of these also included people with not otherwise specified personality disorder), two studies included only people with cluster type C, and one study included only people with borderline personality disorder. The mean number of intervention sessions in the included studies ranged from 12 to 40, and the mean follow-up duration was 19 ± 7.3 months.

The mean pre- post treatment effect of STTP on symptomatic measures (Symptom Checklist (SCL-90), Brief Symptom Inventory (BSI), or Global Symptom Index (GSI)) was Cohen's d 0.92 ± 0.49 , seven studies. The mean pre- post treatment effect of STTP on IIP was Cohen's d 0.86 ± 0.47 , five studies. The mean pre- post treatment effect of STTP on Global Assessment of Functioning (GAF) was Cohen's d 1.47 ± 1.12 , two studies. Reductions in personality disorder diagnoses at follow-up ranged from 38% to 83.3%, four studies.

participants, intervention and study design.

Studies were retrieved from a previous Cochrane review, supplemented by update searches of four bibliographic databases and screening of the reference lists of retrieved articles.

Two reviewers independently assessed studies for inclusion, but no details of the data extraction or quality assessment processes were reported.

The Cochrane Collaboration Depression Anxiety

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		and Neurosis quality
		rating scale was used
		to rate the quality of
		included studies, but
		only overall quality
		scores were
		reported.
		Although the review
		included RCTs, only
		pre- to post-
		treatment effect
		sizes for the
		intervention were
		reported (no
		comparative
		effectiveness data).
		Overall Cohen's d
		pre- to post-
		treatment effect
		sizes appear to have
		been derived by
		calculating a simple,
		un-weighted mean
		across studies which
		included participants
		with varying
		severities of
		personality disorder.

			Further, Cohen's d
			effect sizes were
			used when this was
			not necessary (i.e.
			when results of
			studies with the
			same outcome
			measure were being
			combined.
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Randomised controlled trials

Author	Inclusion criteria	Number of	Summary of results	Risk of bias
(year)		participants		
Reneses	Participants: DSM-IV diagnosis of	n = 53	This study aimed to assess the effectiveness of Psychic	Randomisation
et al.	Borderline Personality Disorder, attending	(intervention	Representation-Focused Psychotherapy (PRFP) plus	used a
(2013)	outpatient services, aged 18-50. Exclusion	group = 25;	conventional treatment, compared to conventional	computer
	criteria: having active suicide risk	comparator	treatment alone, for people with borderline personality	generated,
	symptoms, showing violent behaviours,	group = 28)	disorder.	random
	comorbid diagnosis of eating behaviour			number
	disorder or toxic dependence disorder, or		There were no significant differences between the	sequence.
	current severe physical disease.		intervention and control group at baseline in demographic or	
	Intervention: Psychic Representation-		socioeconomic characteristics, pharmacological treatments	No details of
	focused psychotherapy (PRFP) - based on		or disease severity. Approximately half of study participants	allocation
	classic psychoanalytic principles and		had a concomitant psychiatric condition on Axis I of the DSM-	concealment
	focuses on distorted psychic		IV classification. All participants were receiving	were
	representations and their link with affect		pharmacological treatment (90% anti-depressants, >40%	reported.

and emotions. 20 face-to-face, 45 minute consecutive weekly sessions, plus conventional treatment.

Comparator: Conventional treatment - psychopharmacological treatment (antidepressants, mood stabilizers, antipsychotics, depending on patients' symptoms), for 6 months.

Outcome: General symptoms (Severity Global Index of SCL-90-R; SGI), impulsivity (Barrat Impulsivity Scale; BIS), and social adaptation (Social Adaptation Scale; SAS). Secondary outcomes: BPD symptoms (Zanarini Rating Scale for Borderline Personality Disorder; ZRS), general clinical symptoms (Clinical Global Impression Scale; CGI), depressive symptoms (Montgomery-Asberg Depression Rating Scale; MADRS), anxiety symptoms (State-Trait Anxiety Inventory; STAI), and self-esteem (Rosemberg Self-Esteem Scale).

mood stabilizers, 30% antipsychotics).

There were seven drop-outs in the intervention period: 4 in the control group and 3 in the intervention group.

All outcome measures, with the exception of Zanarini rage score, showed statistically significant pre- to post-treatment improvements in the intervention group. CGI, Zanarini total score and the relations, identity, suicidality, effective instability and feeling of emptiness subgroups of the Zanarini scale also showed statistically significant pre- to post-treatment improvements in the control group.

There was a statistically significant treatment effect, favouring PRFP, for SCL-90, Zanarini score (total, relations and feeling of emptiness), MADRS score (total and suicide), Rosemberg score, Barratt score and SASS score. An effect size was reported, but it was not clear how this was estimated and between group mean differences in change from baseline were not reported.

The study was described as "open".

The methods stated that analyses were conducted on an intention-to-treat basis, but data were only reported for the 44 completers.

Results were reported for all specified outcomes.

Risk of Bias

Systematic reviews

Author (year)	Risk of Bias							
	Inclusion criteria	Searches	Quality	Synthesis				
				assessment				
Abbass et al.	(3)	(()	2	2	(i)			
(2012)	O	V		:	0			
Town et al.	8	⊗	2	8	(2)			
(2011)				O				

Randomised controlled trials

Study	RISK OF BIAS					
	Random allocation	Allocation concealment	Blinding of participants and personnel	Blinding of outcome assessment	Incomplete outcome data	Selective Reporting
Reneses et al. (2013)	©	?	\odot	(S)	\odot	©





? Unclear Risk

Search Details

Source	Search Strategy	Number of	Relevant
		hits	evidence
60 10			identified
SRs and G	· · · · · · · · · · · · · · · · · · ·		
NICE	psychodynamic personality disorder	17	0
DARE	(brief ADJ3 (psychosocial OR therap* OR psychotherap* OR psychoanaly* OR psychodynamic* OR analytic* OR dynamic*)) IN DARE WHERE LPD FROM 10/09/2006 TO 15/09/2014 34 Delete	86	2
	2 MeSH DESCRIPTOR Psychoanalysis EXPLODE ALL TREES 3 Delete		
	3 MeSH DESCRIPTOR Psychotherapy, Brief EXPLODE ALL TREES 59 Delete		
	4 MeSH DESCRIPTOR Psychotherapy, Psychodynamic EXPLODE ALL TREES 0 Delete		
	5 #1 OR #2 OR #3 OR #4		
Primary s	tudies		
CENTRAL	#1 MeSH descriptor: [Personality Disorders] explode all trees 836 #2 MeSH descriptor: [Psychotherapy, Brief] this term only 704 #3 "brief dynamic psychotherap*" 23 #4 "short term psychotherap*" 59 #5 "dynamic psychotherap*" 96 #6 #2 or #3 or #4 or #5 807 #7 #1 or #6 1598 #8 2008 or 2009 or 2010 or 2011 or 2012 or 2013 or 2014 or 2015 331260 #9 #7 and #8 714 Central only	548	
PsycINFO	 PsycINFO; exp PERSONALITY DISORDERS/; 21748 results. PsycINFO; BRIEF PSYCHOTHERAPY/; 4892 results. PsycINFO; "brief dynamic psychotherap*".ti,ab; 155 results. PsycINFO; "dynamic psychotherap*".ti,ab; 1210 results. PsycINFO; 2 OR 3 OR 4; 5591 results. PsycINFO; 1 AND 5; 198 results. 	20	

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8. PsycINFO; random*.ti,ab; 135095 results.		
9. PsycINFO; groups.ti,ab; 377354 results.		
10. PsycINFO; (double adj3 blind).ti,ab; 18245 results.		
11. PsycINFO; (single adj3 blind).ti,ab; 1458 results.		
12. PsycINFO; EXPERIMENTAL DESIGN/; 9340 results.		
13. PsycINFO; controlled.ti,ab; 83711 results.		
14. PsycINFO; (clinical adj3 study).ti,ab; 8183 results.		
15. PsycINFO; trial.ti,ab; 71017 results.		
16. PsycINFO; "treatment outcome clinical trial".md; 28168 results.		
17. PsycINFO; 7 OR 8 OR 9 OR 10 OR 11 OR 12 OR 13 OR 14 OR 15 OR 16; 585266 results.		
18. PsycINFO; 6 AND 17; 59 results.		
19. PsycINFO; 18 [Limit to: Publication Year 2008-2014]; 20 results.		
7. EMBASE; exp PERSONALITY DISORDERS/; 46359 results.	201	
8. EMBASE; BRIEF PSYCHOTHERAPY/; 2 results.		
9. EMBASE; "brief dynamic psychotherap*".ti,ab; 93 results.		
10. EMBASE; "dynamic psychotherap*".ti,ab; 531 results.		
11. EMBASE; 8 OR 9 OR 10; 533 results.		
12. EMBASE; 7 AND 11; 104 results.		
13. EMBASE; PSYCHOTHERAPY, BRIEF/; 75430 results.		
14. EMBASE; 9 OR 10 OR 13; 75486 results.		
15. EMBASE; 7 AND 14; 4680 results.		
16. EMBASE; random*.ti,ab; 907780 results.		
17. EMBASE; factorial*.ti,ab; 23517 results.		
18. EMBASE; (crossover* OR cross-over*).ti,ab; 70178 results.		
19. EMBASE; placebo*.ti,ab; 203202 results.		
20. EMBASE; (doubl* ADJ blind*).ti,ab; 144123 results.		
21. EMBASE; (singl* ADJ blind*).ti,ab; 14753 results.		
22. EMBASE; assign*.ti,ab; 243780 results.		
	9. PsycINFO; groups.ti,ab; 377354 results. 10. PsycINFO; (double adj3 blind).ti,ab; 18245 results. 11. PsycINFO; (single adj3 blind).ti,ab; 1458 results. 12. PsycINFO; EXPERIMENTAL DESIGN/; 9340 results. 13. PsycINFO; controlled.ti,ab; 83711 results. 14. PsycINFO; (clinical adj3 study).ti,ab; 8183 results. 15. PsycINFO; trial.ti,ab; 71017 results. 16. PsycINFO; "treatment outcome clinical trial".md; 28168 results. 17. PsycINFO; 7 OR 8 OR 9 OR 10 OR 11 OR 12 OR 13 OR 14 OR 15 OR 16; 585266 results. 18. PsycINFO; 6 AND 17; 59 results. 19. PsycINFO; 18 [Limit to: Publication Year 2008-2014]; 20 results. 19. PsycINFO; 18 [Limit to: Publication Year 2008-2014]; 20 results. 10. EMBASE; exp PERSONALITY DISORDERS/; 46359 results. 10. EMBASE; "brief dynamic psychotherap*".ti,ab; 93 results. 11. EMBASE; "dynamic psychotherap*".ti,ab; 531 results. 12. EMBASE; "dynamic psychotherap*".ti,ab; 531 results. 13. EMBASE; 7 AND 11; 104 results. 14. EMBASE; 9 OR 10 OR 13; 75486 results. 15. EMBASE; 9 OR 10 OR 13; 75486 results. 16. EMBASE; 7 and 14; 4680 results. 17. EMBASE; random*.ti,ab; 907780 results. 18. EMBASE; random*.ti,ab; 23517 results. 19. EMBASE; factorial*.ti,ab; 23517 results. 19. EMBASE; placebo*.ti,ab; 203202 results. 20. EMBASE; (doubl* ADJ blind*).ti,ab; 144123 results. 21. EMBASE; (singl* ADJ blind*).ti,ab; 14753 results.	8. PsycINFO; random*.ti,ab; 135095 results. 9. PsycINFO; groups.ti,ab; 377354 results. 10. PsycINFO; (double adj3 blind).ti,ab; 18245 results. 11. PsycINFO; (single adj3 blind).ti,ab; 1458 results. 12. PsycINFO; EXPERIMENTAL DESIGN/; 9340 results. 13. PsycINFO; controlled.ti,ab; 83711 results. 14. PsycINFO; controlled.ti,ab; 83711 results. 15. PsycINFO; clinical adj3 study).ti,ab; 8183 results. 16. PsycINFO; trial.ti,ab; 71017 results. 16. PsycINFO; "treatment outcome clinical trial".md; 28168 results. 17. PsycINFO; 7 OR 8 OR 9 OR 10 OR 11 OR 12 OR 13 OR 14 OR 15 OR 16; 585266 results. 18. PsycINFO; 6 AND 17; 59 results. 19. PsycINFO; 18 [Limit to: Publication Year 2008-2014]; 20 results. 201 8. EMBASE; exp PERSONALITY DISORDERS/; 46359 results. 201 8. EMBASE; "brief dynamic psychotherap*".ti,ab; 93 results. 10. EMBASE; "dynamic psychotherap*".ti,ab; 93 results. 11. EMBASE; 9 OR 9 OR 10; 533 results. 12. EMBASE; 7 AND 11; 104 results. 13. EMBASE; 9 SYCHOTHERAPY, BRIEF/; 75430 results. 14. EMBASE; 9 OR 10 OR 13; 75486 results. 15. EMBASE; 7 AND 14; 4680 results. 16. EMBASE; random*.ti,ab; 907780 results. 17. EMBASE; factorial*.ti,ab; 23517 results. 18. EMBASE; factorial*.ti,ab; 23517 results. 29. EMBASE; factorial*.ti,ab; 23517 results. 20. EMBASE; placebo*.ti,ab; 203202 results. 20. EMBASE; (crossover* OR cross-over*).ti,ab; 70178 results. 21. EMBASE; (singl* ADJ blind*).ti,ab; 144123 results. 22. EMBASE; (singl* ADJ blind*).ti,ab; 144737 results.

	22 FAADACE, allowed to also 0FOAC months		
	23. EMBASE; allocat*.ti,ab; 85946 results.		
	24. EMBASE; volunteer*.ti,ab; 178672 results.		
	25. EMBASE; CROSSOVER PROCEDURE/; 40446 results.		
	26. EMBASE; DOUBLE BLIND PROCEDURE/; 115879 results.		
	27. EMBASE; RANDOMIZED CONTROLLED TRIAL/; 352001 results.		
	28. EMBASE; SINGLE BLIND PROCEDURE/; 18953 results.		
	29. EMBASE; 16 OR 17 OR 18 OR 19 OR 20 OR 21 OR 22 OR 23 OR 24 OR 25 OR 26 OR 27 OR 28;		
	1442862 results.		
	30. EMBASE; 15 AND 29; 390 results.		
	31. EMBASE; 30 [Limit to: Publication Year 2008-2014]; 218 results.		
	32. EMBASE; (juvenile OR adolescent*).ti,ab [Limit to: Publication Year 2008-2014]; 116905 results.		
	33. EMBASE; 31 not 32 [Limit to: Publication Year 2008-2014]; 201 results.		
Medline	7. MEDLINE; exp PERSONALITY DISORDERS/; 32896 results.	35	
	8. MEDLINE; BRIEF PSYCHOTHERAPY/; 2883 results.		
	9. MEDLINE; "brief dynamic psychotherap*".ti,ab; 70 results.		
	10. MEDLINE; "dynamic psychotherap*".ti,ab; 405 results.		
	11. MEDLINE; 8 OR 9 OR 10; 3143 results.		
	12. MEDLINE; 7 AND 11; 197 results.		
	13. MEDLINE; "randomized controlled trial".pt; 399438 results.		
	14. MEDLINE; "controlled clinical trial".pt; 90638 results.		
	15. MEDLINE; randomized.ab; 318530 results.		
	16. MEDLINE; placebo.ab; 163546 results.		
	17. MEDLINE; "drug therapy".fs; 1782347 results.		
	18. MEDLINE; randomly.ab; 228322 results.		
	19. MEDLINE; trial.ab; 332229 results.		
	20. MEDLINE; groups.ab; 1435618 results.		
	21. MEDLINE; 13 OR 14 OR 15 OR 16 OR 17 OR 18 OR 19 OR 20; 3523823 results.		
	22. MEDLINE; 12 AND 21; 66 results.		
	23. MEDLINE; 22 [Limit to: Publication Year 2008-2014]; 35 results.		
Summary	NA	NA	

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