

# NKR 43 Angst PICO 3 Psykoterapi vs SSRI/SNRI

## Review information

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### What's new

Date / Event	Description
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## History

Date / Event	Description
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## Characteristics of studies

### Characteristics of included studies

#### *Beidel 2007*

<b>Methods</b>	<p><b>Study design:</b> Randomized controlled trial</p> <p><b>Study grouping:</b> Parallel group</p> <p><b>Open Label:</b></p> <p><b>Cluster RCT:</b></p>
<b>Participants</b>	<p><b>Baseline Characteristics</b></p> <p>Intervention</p> <ul style="list-style-type: none"> <li>● <i>Number with primary social phobia (n, %): 57, 100%</i></li> <li>● <i>Number with primary generalized anxiety disorder (n, %): 0,0</i></li> <li>● <i>Number with primary separation anxiety disorder (n, %): 0,0</i></li> <li>● <i>Number with other types of primary anxiety disorders (n, %): 0,0</i></li> <li>● <i>Age in years (mean, SD): Not reported by intervention. Total sample:11.61 (2.6)</i></li> <li>● <i>Age range and proportion of children and adolescents: Not reported by intervention. Total sample: 7-17</i></li> </ul> <p>Control</p> <ul style="list-style-type: none"> <li>● <i>Number with primary social phobia (n, %): 33, 100%</i></li> <li>● <i>Number with primary generalized anxiety disorder (n, %): 0,0</i></li> <li>● <i>Number with primary separation anxiety disorder (n, %): 0,0</i></li> <li>● <i>Number with other types of primary anxiety disorders (n, %): 0,0</i></li> <li>● <i>Age in years (mean, SD): Not reported by intervention. Total sample:11.61 (2.6)</i></li> <li>● <i>Age range and proportion of children and adolescents: Not reported by intervention. Total sample: 7-17</i></li> </ul> <p><b>Included criteria:</b> All of the youths were between the ages of 7 and 17 and had a primary diagnosis of social phobia. To be considered primary, social phobia symptoms were of at least moderate severity (4 or higher on an 8-point scale) and created functional impairment.</p>

	<p><b>Excluded criteria:</b> To ensure generalization of study findings, secondary comorbid diagnoses were allowed, with the exception of bipolar disorder, psychosis, conduct disorder, autism spectrum disorders, and mental retardation. Youths with moderate to severe depression who expressed active suicidal ideation or who had a previous unsuccessful trial of fluoxetine or behavior therapy were excluded.</p> <p><b>Pretreatment:</b></p>
<p><b>Interventions</b></p>	<p><b>Intervention Characteristics</b></p> <p>Intervention</p> <ul style="list-style-type: none"> <li>● <i>Description of type of intervention/control:</i> SET-C (Beidel et al., 2000) includes SST, peer generalization experiences, and in vivo exposure. (The SET-C treatment manual is available from Multi-Health Systems, Inc.) Social skills training/peer generalization is conducted in small groups (four to five youths), whereas in vivo exposure is conducted individually. Treatment consisted of one individual and one group session per week for 12 weeks. Group sessions are 150 minutes in length (60 minutes of SST and 90 minutes of peer generalization) and were constituted with no more than a 3-year age span (e.g., 8-11, 9-12) and individual sessions averaged 60 minutes. SST targeted seven major topic areas: greetings skills, initiating and maintaining conversations, listening skills, joining groups of peers, friendship establishment and maintenance, positive and negative assertion, and telephone skills. Content of the treatment sessions and role-play scenarios were always modified to be age appropriate. Non-verbal skills (eye contact, voice volume, vocal tone) were also addressed. Peer generalization experiences directly followed SST and allowed practice of social skills outside the clinic. Activities varied depending on group age but typically included roller skating, bowling, video arcades, pizza parlors, picnics, flying kites, and children's museums. Same-age non-anxious peers, recruited from the community, were trained as facilitators and participated in generalization sessions. In vivo exposure targeted unique fear patterns. Commonly used tasks included reading in front of others, writing on the blackboard, conversing with same-age peers, and asking questions of adults. Discontinued when self-reported anxiety returned to baseline (see Beidel et al., 2000), individual exposure sessions averaged 60 minutes but did not exceed 90 minutes</li> <li>● <i>Length of intervention/control (weeks and sessions):</i> 12 weeks, 2 sessions per week</li> <li>● <i>Length of follow-up (in months):</i> Only FU for treatment responders so no relevant data</li> </ul> <p>Control</p> <ul style="list-style-type: none"> <li>● <i>Description of type of intervention/control:</i> Fluoxetine was chosen based on its safety and efficacy profile. Identically appearing fluoxetine or placebo capsules were dispensed using the following titration schedule: weeks 1 and 2, one capsule (10 mg) per day; weeks 3 and 4, two capsules (20 mg) per day; and weeks 5 and 6, three capsules (30 mg) per day. At week 7, dose was increased to 40 mg (four capsules) and held constant throughout treatment (through week 12). Dose reduction/discontinuation was allowed in the case of moderate or severe side effects, but</li> </ul>

	<p>no adjustments were necessary. There were no reports of suicidal ideation or parasuicidal behaviors. Children continued to be seen weekly throughout the 12-week program. In addition to medication management, the psychiatrist offered general encouragement and support, but no specific exposure instructions during each 60-minute session</p> <ul style="list-style-type: none"> <li>● <i>Length of intervention/control (weeks and sessions):</i> 12 weeks, 12 sessions</li> <li>● <i>Length of follow-up (in months):</i></li> </ul>
<p><b>Outcomes</b></p>	<p><i>Remission of primary anxiety diagnosis (EoT)</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> Dichotomous Outcome</li> <li>● <b>Reporting:</b> Fully reported</li> <li>● <b>Direction:</b> Higher is better</li> <li>● <b>Data value:</b> Endpoint</li> </ul> <p><i>Youth reported anxiety symptoms (EoT)</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> Continuous Outcome</li> <li>● <b>Reporting:</b> Fully reported</li> <li>● <b>Scale:</b> SPAI-C</li> <li>● <b>Range:</b> 0-52</li> <li>● <b>Unit of measure:</b> Points</li> <li>● <b>Direction:</b> Lower is better</li> <li>● <b>Data value:</b> Endpoint</li> </ul> <p><i>Parent reported anxiety symptoms (EoT)</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> Continuous Outcome</li> <li>● <b>Reporting:</b> Fully reported</li> <li>● <b>Scale:</b> CBCL-internalizing</li> <li>● <b>Range:</b> 0 – 64</li> <li>● <b>Unit of measure:</b> Points</li> <li>● <b>Direction:</b> Lower is better</li> <li>● <b>Data value:</b> Endpoint</li> </ul> <p><i>Remission of primary anxiety diagnosis (longest FU, at least 3 months)</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> Dichotomous Outcome</li> <li>● <b>Reporting:</b> Fully reported</li> <li>● <b>Direction:</b> Higher is better</li> </ul>

- **Data value:** Endpoint
- **Notes:** 1-year follow-up. These 18 youths were matched by age (within 2 years) and sex to 18 youths treated with SET-C who also completed follow-up. Because most nonresponders self-selected out of followup, this selection bias negates the use of last observation carried forward as an appropriate analysis.

*Youth reported anxiety symptoms (longest FU, at least 3 months)*

- **Outcome type:** ContinuousOutcome
- **Reporting:** Not reported
- **Unit of measure:** Points
- **Direction:** Lower is better
- **Data value:** Endpoint

*Parent reported anxiety symptoms (longest FU, at least 3 months)*

- **Outcome type:** ContinuousOutcome
- **Reporting:** Not reported
- **Unit of measure:** Points
- **Direction:** Lower is better
- **Data value:** Endpoint

*Youth reported functioning (EoT)*

- **Outcome type:** ContinuousOutcome
- **Reporting:** Not reported
- **Direction:** Lower is better
- **Data value:** Endpoint

*Observer reported functioning (EoT)*

- **Outcome type:** ContinuousOutcome
- **Reporting:** Fully reported
- **Scale:** Children's Global Assessment Scale (CGAS)
- **Range:** 1-100
- **Unit of measure:** Points
- **Direction:** Higher is better
- **Data value:** Endpoint
- **Notes:** Clinician rated

*Combined youth and observer reported functioning (EoT)*

- **Outcome type:** DichotomousOutcome
- **Reporting:** Fully reported
- **Scale:** "High-end state functioning"
- **Direction:** Higher is better
- **Data value:** Endpoint
- **Notes:** Children were designated to have high end-state functioning if they had both a score of less than 18 on the SPAI-C (a cutoff score previously documented as characteristic of youths without social phobia) and a rating of 8 or 9 on the CGAS, indicating no more than mild impairment on overall functioning.

*Number that discontinued treatment or control (EoT)*

- **Outcome type:** DichotomousOutcome
- **Reporting:** Fully reported
- **Direction:** Lower is better
- **Data value:** Endpoint
- **Notes:** Discontinued after randomization

*Suicidal thoughts (EoT)*

- **Outcome type:** AdverseEvent
- **Reporting:** Partially reported
- **Direction:** Lower is better
- **Data value:** Endpoint
- **Notes:** Not reported for intervention group

*Suicidal behavior (EoT)*

- **Outcome type:** AdverseEvent
- **Reporting:** Partially reported
- **Direction:** Lower is better
- **Data value:** Endpoint
- **Notes:** Not reported for intervention group

*Serious adverse events (EoT)*

- **Outcome type:** AdverseEvent
- **Reporting:** Partially reported
- **Direction:** Lower is better
- **Data value:** Endpoint

	<ul style="list-style-type: none"> <li>● <b>Notes:</b> Not reported for the Intervention group. For the control: In the fluoxetine group, no symptoms were rated as severe; eight symptoms (diarrhea, heartburn, facial pallor, fatigue, weakness, lethargy, anger outbursts, and difficulty concentrating) were rated as moderate.</li> </ul>
<b>Identification</b>	<p><b>Sponsorship source:</b> This research was supported in part by NIMH grant R01MH53703 to the first three authors. Lilly Corporation supplied the fluoxetine and matching placebo capsules. Disclosure: Drs. Beidel and Turner are the authors of the Social Phobia and Anxiety Inventory for Children and the Social Effectiveness Therapy for Children Treatment Manual and receive royalties from Multi-Health Systems, Inc. for the sale of these items. Dr. Sallee receives grant support from Shire and Bristol-Myers Squibb pharmaceutical companies; is a paid consultant to Shire and Abbott Laboratories and a speaker for Pfizer and Takeda Pharmaceuticals; and is on the board of directors of P2Dinc and Satiety Solutions LLC. Dr. Pathak is employed by Astra Zeneca and has received royalties from Forest Laboratories. The other authors have no financial relationships to disclose</p> <p><b>Country:</b> USA</p> <p><b>Setting:</b> Media announcements and referrals from mental health clinicians</p> <p><b>Comments:</b> Unique identifier: NCT00043537</p> <p><b>Authors name:</b> Beidel et al. 2007</p> <p><b>Institution:</b> Department of Psychology, University of Central Florida</p> <p><b>Email:</b> dbeidel@mail.ucf.edu</p> <p><b>Address:</b></p>
<b>Notes</b>	<p><i>Nkr 43 Angst on 07/04/2016 22:44</i></p> <p><b>Select</b></p> <p>Three conditions: Social skill training, SSRI, pill placebo. No combination</p>

Risk of bias table

Bias	Authors' judgement	Support for judgement
Sequence Generation	Low risk	Quote: "Using a computer-generated program, the remaining 139 subjects, ages 7 to 17, were randomized to one of three treatment groups: SET-C, fluoxetine, or pill placebo."
Allocation concealment	Unclear risk	Judgement Comment: Not described

Blinding of participants and personnel	High risk	Judgement Comment: Not blinded
Blinding of outcome assessors	Low risk	Quote: "Rating scales were completed by the diagnostic interviewer at pre- treatment and by an independent evaluator blinded to group status at all subsequent evaluations"
Incomplete outcome data	High risk	Judgement Comment: Between 21,2% to 30,2 % missing data
Selective outcome reporting	Low risk	Judgement Comment: Match to protocol
Other sources of bias	Low risk	Quote: "Finally, this study was conducted during the issuance of the black box warning by the U.S. Food and Drug Administration, which effectively stopped parental agreement to participate in the protocol. Therefore, the study was terminated prematurely, which would have resulted in more even cell sizes." Judgement Comment: Does likely not influence as it was stopped for general safety concerns regarding SSRI and not due to large treatment effect

### Walkup 2008

<b>Methods</b>	<p><b>Study design:</b> Randomized controlled trial</p> <p><b>Study grouping:</b> Parallel group</p> <p><b>Open Label:</b></p> <p><b>Cluster RCT:</b></p>
<b>Participants</b>	<p><b>Baseline Characteristics</b></p> <p>Intervention(psychotherapy)</p> <ul style="list-style-type: none"> <li>● <i>Number with primary social phobia (n, %):</i> Not reported specifically</li> <li>● <i>Number with primary generalized anxiety disorder (n, %):</i> Not reported specifically</li> <li>● <i>Number with primary separation anxiety disorder (n, %):</i> Not reported specifically</li> <li>● <i>Number with other types of primary anxiety disorders (n, %):</i> 0,0%</li> <li>● <i>Age in years (mean, SD):</i> 10.5 (2.9)</li> <li>● <i>Age range and proportion of children and adolescents:</i> 7-17 (77.7% children[7-12])</li> </ul> <p>Control</p> <ul style="list-style-type: none"> <li>● <i>Number with primary social phobia (n, %):</i> Not reported specifically</li> <li>● <i>Number with primary generalized anxiety disorder (n, %):</i> Not reported specifically</li> <li>● <i>Number with primary separation anxiety disorder (n, %):</i> Not reported specifically</li> <li>● <i>Number with other types of primary anxiety disorders (n, %):</i> 0,0%</li> </ul>



	<ul style="list-style-type: none"> <li>● <i>Age in years (mean, SD): 10.8 (2.8)</i></li> <li>● <i>Age range and proportion of children and adolescents: 7-17 (74.4% children[7-12])</i></li> </ul> <p>Intervention(SSRI + therapy)</p> <ul style="list-style-type: none"> <li>● <i>Number with primary social phobia (n, %): Not reported specifically</i></li> <li>● <i>Number with primary generalized anxiety disorder (n, %): Not reported specifically</i></li> <li>● <i>Number with primary separation anxiety disorder (n, %): Not reported specifically</i></li> <li>● <i>Number with other types of primary anxiety disorders (n, %): 0,0%</i></li> <li>● <i>Age in years (mean, SD): 10.7 (2.8)</i></li> <li>● <i>Age range and proportion of children and adolescents: 7-17 (72.1% children[7-12])</i></li> </ul> <p><b>Included criteria:</b> Children between the ages of 7 and 17 years with a primary diagnosis of separation or generalized anxiety disorder or social phobia (according to the criteria of the Diagnostic and Statistical Manual of Mental Disorders, fourth edition, text revision [DSM-IV-TR]16), substantial impairment, and an IQ of 80 or more were eligible to participate. Children with coexisting psychiatric diagnoses of lesser severity than the three target disorders were also allowed to participate; such diagnoses included attention deficit-hyperactivity disorder (ADHD) while receiving stable doses of stimulant and obsessive-compulsive, post-traumatic stress, oppositional-defiant, and conduct disorders</p> <p><b>Excluded criteria:</b> Children were excluded if they had an unstable medical condition, were refusing to attend school because of anxiety, or had tried but had not had a response to two adequate trials of SSRIs or an adequate trial of cognitive behavioral therapy. Girls who were pregnant or were sexually active and were not using an effective method of birth control were also excluded. Children who were receiving psychoactive medications other than stable doses of stimulants and who had psychiatric diagnoses that made participation in the study clinically inappropriate (i.e., current major depressive or substance-use disorder; unmedicated ADHD, combined type; or a lifetime history of bipolar, psychotic, or pervasive developmental disorders) or who presented an acute risk to themselves or others were also excluded.</p> <p><b>Pretreatment:</b> No group differences detected</p>
<p><b>Interventions</b></p>	<p><b>Intervention Characteristics</b></p> <p>Intervention(psychotherapy)</p> <ul style="list-style-type: none"> <li>● <i>Description of type of intervention/control:</i> Cognitive behavioral therapy involved fourteen 60-minute sessions, which included review and ratings of the severity of subjects' anxiety, response to treatment, and adverse events. Therapy was based on the Coping Cat program, which was adapted for the subjects' age and the duration of the study. Each subject who was assigned to receive cognitive behavioral therapy received training in anxiety-management skills, followed by behavioral exposure to anxiety-provoking situations. Parents attended weekly check-ins and two parent-only sessions. Experienced psychotherapists, certified in the Coping Cat protocol, received regular site-level and cross-site supervision</li> </ul>

	<ul style="list-style-type: none"> <li>● <i>Length of intervention/control (weeks and sessions):</i> 12 weeks, 14 sessions</li> <li>● <i>Length of follow-up (in months):</i> 6 month but only for responders. There is also a follow up study (CAMELS) that describes remission for a portion of the responders 6 years after randomization</li> </ul> <p>Control</p> <ul style="list-style-type: none"> <li>● <i>Description of type of intervention/control:</i> Pharmacotherapy involved eight sessions of 30 to 60 minutes each that included review and ratings of the severity of subjects' anxiety, their response to treatment, and adverse events. Ser-traline (Zoloft) and matching placebo were ad-ministered on a fixed-flexible schedule begin-ning with 25 mg per day and adjusted up to 200 mg per day by week 8. Through week 8, subjects who were considered to be mildly ill or worse and who had minimal side effects were eligible for dose increases. Psychiatrists and nurse clini-cians with experience in medicating children with anxiety disorders were certified in the study phar-macotherapy protocol and received regular site-level and cross-site supervision. Pill counts and medication diaries were used to facilitate and document adherence</li> <li>● <i>Length of intervention/control (weeks and sessions):</i> 12 weeks, 8 sessions of medication review and administration</li> <li>● <i>Length of follow-up (in months):</i></li> </ul> <p>Intervention(SSRI + therapy)</p> <ul style="list-style-type: none"> <li>● <i>Description of type of intervention/control:</i> Combination therapy consisted of the admin-istration of sertraline and cognitive behavioral therapy. Whenever possible, therapy and medica-tion sessions occurred on the same day for the convenience of subject</li> <li>● <i>Length of intervention/control (weeks and sessions):</i> 12 weeks, 14 sessions of Copiing Cat and 8 sessions of medication administration</li> <li>● <i>Length of follow-up (in months):</i></li> </ul>
<p><b>Outcomes</b></p>	<p><i>Remission of primary anxiety diagnosis (EoT)</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> DichotomousOutcome</li> <li>● <b>Reporting:</b> Not reported</li> <li>● <b>Scale:</b> ADIS-C/P</li> <li>● <b>Direction:</b> Higher is better</li> <li>● <b>Data value:</b> Endpoint</li> <li>● <b>Notes:</b> Reported in Piacentini et al., 2014</li> </ul> <p><i>Youth reported anxiety symptoms (EoT)</i></p> <ul style="list-style-type: none"> <li>● <b>Outcome type:</b> ContinuousOutcome</li> <li>● <b>Reporting:</b> Not reported</li> </ul>

- **Notes:** No self-report in study

*Parent reported anxiety symptoms (EoT)*

- **Outcome type:** ContinuousOutcome
- **Reporting:** Not reported
- **Notes:** No parent report in study

*Remission of primary anxiety diagnosis (longest FU, at least 3 months)*

- **Outcome type:** DichotomousOutcome
- **Scale:** ADIS
- **Direction:** Higher is better
- **Data value:** Endpoint
- **Notes:** 6 year follow-up (based on Ginsburg et al., 2014)

*Youth reported anxiety symptoms (longest FU, at least 3 months)*

- **Outcome type:** ContinuousOutcome
- **Reporting:** Not reported
- **Notes:** No self-report in study

*Parent reported anxiety symptoms (longest FU, at least 3 months)*

- **Outcome type:** ContinuousOutcome
- **Reporting:** Not reported
- **Notes:** No parent report in study

*Youth reported functioning (EoT)*

- **Outcome type:** ContinuousOutcome
- **Reporting:** Not reported
- **Notes:** No self-report in study

*Observer reported functioning (EoT)*

- **Outcome type:** ContinuousOutcome
- **Reporting:** Fully reported
- **Scale:** Children's Global Assessment Scale (CGAS)
- **Range:** 1-100
- **Unit of measure:** Points
- **Direction:** Higher is better

- **Data value:** Endpoint

*Combined youth and observer reported functioning (EoT)*

- **Outcome type:** DichotomousOutcome
- **Reporting:** Not reported

*Number that discontinued treatment or control (EoT)*

- **Outcome type:** DichotomousOutcome
- **Reporting:** Fully reported
- **Direction:** Lower is better
- **Data value:** Endpoint

*Suicidal thoughts (EoT)*

- **Outcome type:** AdverseEvent
- **Reporting:** Fully reported
- **Direction:** Lower is better
- **Data value:** Endpoint

*Suicidal behavior (EoT)*

- **Outcome type:** AdverseEvent
- **Reporting:** Fully reported
- **Direction:** Lower is better
- **Data value:** Endpoint

*Serious adverse events (EoT)*

- **Outcome type:** AdverseEvent
- **Reporting:** Fully reported
- **Direction:** Lower is better
- **Data value:** Endpoint
- **Notes:** Moderate to severe adverse events: SSRI: Physical = 50.4%, Psychiatric = 17.3%, Harm-related = 2..3%, Medical or surgical = 0.8%. Sum = 70.8% CBT: Physical = 36.7%, Psychiatric = 9.4%, Harm-related = 5.8%, Medical or surgical = 0.7%. Sum = 52.6%

**Identification**

**Sponsorship source:** Supported by grants (U01 MH064089, to Dr. Walkup; U01 MH64092, to Dr. Albano; U01 MH64003, to Dr. Birmaher; U01 MH63747, to Dr. Kendall; U01 MH64107, to Dr. March; U01 MH64088, to Dr. Piacentini; and U01 MH064003, to Dr. Compton) from the National Institute of Mental Health (NIMH). Sertraline and matching placebo were supplied free of charge by Pfizer. Dr. Walkup reports receiving consulting fees from Eli Lilly and Jazz Pharmaceuticals and fees for legal consultation to de-fense counsel and submission of written reports in litigation involving GlaxoSmithKline, receiving lecture fees from CMP Media, Medical Education Reviews, McMahon Group, and Di-Medix, and receiving support in the form of free medication and matching placebo from Eli Lilly and free medication from Ab-bott for clinical trials funded by the NIMH; Dr. Albano, receiv-ing royalties from Oxford University Press for the Anxiety Disor-ders Interview Schedule for DSM-IV, Child and Parent Versions, but not for interviews used in this study, and royalties from the Guilford Press; Dr. Piacentini, receiving royalties from Oxford University Press for treatment manuals on childhood obsessive-compulsive disorder and tic disorders and from the Guilford Press and APA Books for other books on child mental health and receiving lecture fees from Janssen-Cilag; Dr. Birmaher, receiv-ing consulting fees from Jazz Pharmaceuticals, Solvay Pharma-ceuticals, and Abcomm, lecture fees from Solvay, and royalties from Random House for a book on children with bipolar disor-der; Dr. Rynn, receiving grant support from Neuropharm, Boeh-ringer Ingelheim Pharmaceuticals, and Wyeth Pharmaceuticals, consulting fees from Wyeth, and royalties from APPI for a book chapter on pediatric anxiety disorders; Dr. McCracken, receiving consulting fees from Sanofi-Aventis and Wyeth, lecture fees from Shire and UCB, and grant support from Aspect, Johnson & Johnson, Bristol-Myers Squibb, and Eli Lilly; Dr. Waslick, receiv-ing grant support from Baystate Health, Somerset Pharmaceuti-cals, and GlaxoSmithKline; Dr. Iyengar, receiving consulting fees from Westinghouse for statistical consultation; Dr. March, receiving study medications from Eli Lilly for an NIMH-funded clinical trial and receiving royalties from Pearson for being the author of the Multidimensional Anxiety Scale for Children, re-ceiving consulting fees from Eli Lilly, Pfizer, Wyeth, and Glaxo-SmithKline, having an equity interest in MedAvante, and serving on an advisory board for AstraZeneca and Johnson & Johnson; and Dr. Kendall, receiving royalties from Workbook Publishing for anxiety-treatment materials. No other potential conflict of interest relevant to this article was reported.

**Country:** USA

**Setting:** Recruited from Duke University Medical Cen-ter, New York State Psychiatric Institute-Colum-bia University Medical Center-New York Univer-sity, Johns Hopkins Medical Institutions, Temple University, University of California, Los Angeles, and Western Psychiatric Institute and Clinic-University of Pittsburgh Medical Center.

**Comments:** ClinicalTrials.gov number, NCT00052078

**Authors name:** Walkup 2008

**Institution:** Johns Hopkins Medical Institutions, Baltimore, New York

**Email:** not stated

**Address:**

<b>Notes</b>	Nkr 43 Angst on 07/04/2016 22:35 <b>Select</b> The CAMS study end of treatment
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### Risk of bias table

Bias	Authors' judgement	Support for judgement
Sequence Generation	Low risk	Quote: "The randomization sequence in a 2:2:2:1 ratio was determined by a computer-generated algorithm"
Allocation concealment	Low risk	Quote: "and maintained by the central pharmacy, with stratification according to age, sex, and study center."
Blinding of participants and personnel	High risk	Judgement Comment: Not blinded for CBT
Blinding of outcome assessors	Low risk	Quote: "The study protocol called for independent evaluators who completed assessments to be unaware of all treatment assignments."
Incomplete outcome data	Low risk	Judgement Comment: Attrition between 4.32 % to 17.29 %
Selective outcome reporting	Low risk	Judgement Comment: Match to protocol
Other sources of bias	Low risk	Judgement Comment: No other sources detected

*Footnotes*

### Characteristics of excluded studies

*Footnotes*

## Characteristics of studies awaiting classification

*Footnotes*

## Characteristics of ongoing studies

*Footnotes*

## Summary of findings tables

## Additional tables

## References to studies

### Included studies

#### ***Beidel 2007***

Beidel,D. C.; Turner,S. M.; Sallee,F. R.; Ammerman,R. T.; Crosby,L. A.; Pathak,S.. SET-C versus fluoxetine in the treatment of childhood social phobia. Journal of the American Academy of Child and Adolescent Psychiatry 2007;46(12):1622-1632. [DOI: 10.1097/chi.0b013e318154bb57 [doi]]

#### ***Walkup 2008***

Ginsburg G.S.; Becker E.M.; Keeton C.P.; Sakolsky D.; Piacentini J.; Albano A.M.; Compton S.N.; Iyengar S.; Sullivan K.; Caporino N.; Peris T.; Birmaher B.; Rynn M.; March J.; Kendall,P. C.. Naturalistic follow-up of youths treated for pediatric anxiety disorders.. JAMA Psychiatry 2014;71(3):310-318. [DOI: ]

Piacentini,John; Bennett,Shannon; Compton,Scott N.; Kendall,Phillip C.; Birmaher,Boris; Albano,Anne Marie; March,John; Sherrill,Joel; Sakolsky,Dara; Ginsburg,Golda; Rynn,Moira; Bergman,R. Lindsey; Gosch,Elizabeth; Waslick,Bruce; Iyengar,Satish; McCracken,James; Walkup,John. 24- and 36-week outcomes for the Child/Adolescent Anxiety Multimodal Study (CAMS).. Journal of the American Academy of Child & Adolescent Psychiatry 2014;53(3):297-310. [DOI: ]

Rynn,Moira A.; Walkup,John T.; Compton,Scott N.; Sakolsky,Dara J.; Sherrill,Joel T.; Shen,Sa; Kendall,Philip C.; McCracken,James; Albano,Anne Marie; Piacentini,John; Riddle,Mark A.; Keeton,Courtney; Waslick,Bruce; Chrisman,Allan; Iyengar,Satish; March,John S.; Birmaher,Boris. Child/adolescent anxiety multimodal study: Evaluating safety.. Journal of the American Academy of Child & Adolescent Psychiatry 2015;54(3):180-190. [DOI: ]

Walkup, J. T.; Albano, A. M.; Piacentini, J.; Birmaher, B.; Compton, S. N.; Sherrill, J. T.; Ginsburg, G. S.; Rynn, M. A.; McCracken, J.; Waslick, B.; Iyengar, S.; March, J. S.; Kendall, P. C.. Cognitive behavioral therapy, sertraline, or a combination in childhood anxiety. *The New England journal of medicine* 2008;359(26):2753-2766. [DOI: 10.1056/NEJMoa0804633 [doi]]

## Excluded studies

## Studies awaiting classification

## Ongoing studies

## Other references

## Additional references

## Other published versions of this review

## Data and analyses

### 1 Psychotherapy vs SSRI

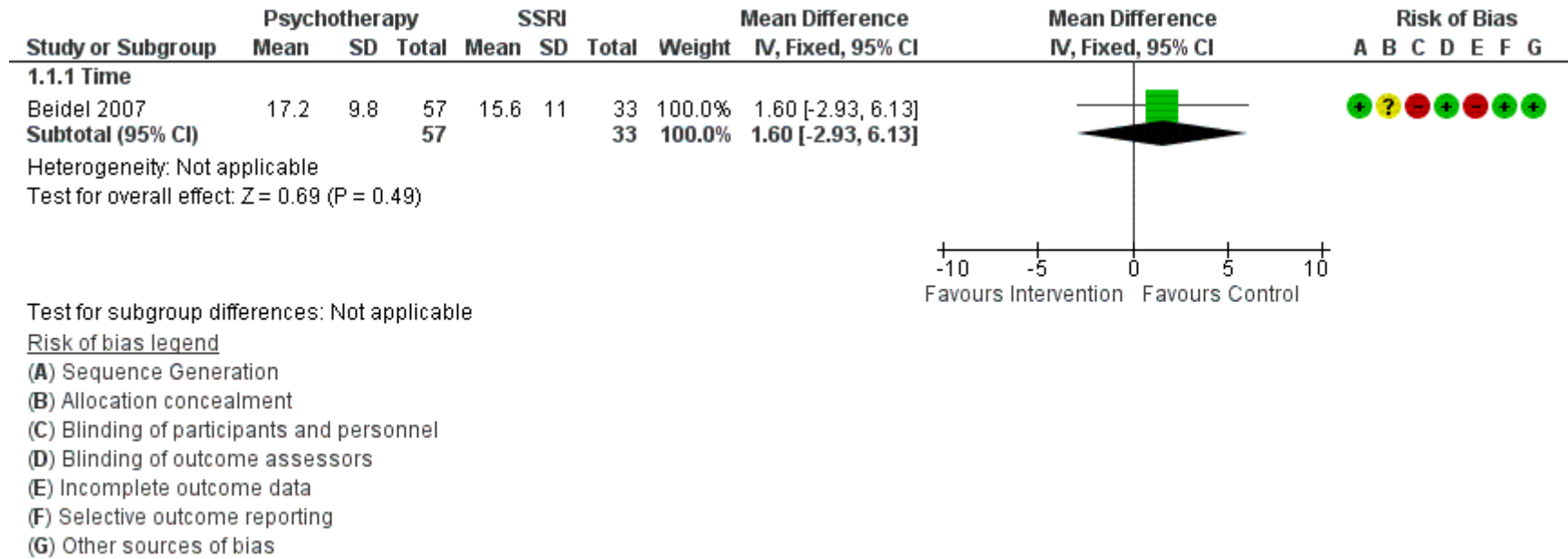
Outcome or Subgroup	Studies	Participants	Statistical Method	Effect Estimate
1.1 Youth reported anxiety symptoms (EoT)	1		Mean Difference (IV, Fixed, 95% CI)	Subtotals only
1.1.1 Time	1	90	Mean Difference (IV, Fixed, 95% CI)	1.60 [-2.93, 6.13]
1.2 Parent reported anxiety symptoms (EoT)	1		Mean Difference (IV, Fixed, 95% CI)	Subtotals only
1.2.1 Time	1	90	Mean Difference (IV, Fixed, 95% CI)	1.20 [-2.52, 4.92]
1.3 Youth reported anxiety symptoms (longest FU, at least 3 months)	0	0	Mean Difference (IV, Fixed, 95% CI)	Not estimable
1.4 Parent reported anxiety symptoms (longest FU, at least 3 months)	0	0	Mean Difference (IV, Fixed, 95% CI)	Not estimable
1.5 Youth reported functioning (EoT)	0	0	Mean Difference (IV, Fixed, 95% CI)	Not estimable



1.6 Observer reported functioning C-GAS(EoT)	2		Mean Difference (IV, Random, 95% CI)	Subtotals only
1.6.1 Time	2	362	Mean Difference (IV, Random, 95% CI)	1.73 [-4.34, 7.80]
1.7 Remission of primary anxiety diagnosis (EoT)	2		Risk Ratio (IV, Random, 95% CI)	Subtotals only
1.7.1 Time	2	362	Risk Ratio (IV, Random, 95% CI)	1.61 [0.58, 4.44]
1.8 Remission of primary anxiety diagnosis (longest FU, 6 years)	1		Risk Ratio (IV, Fixed, 95% CI)	Subtotals only
1.8.1 Time	1	162	Risk Ratio (IV, Fixed, 95% CI)	0.95 [0.70, 1.29]
1.9 Remission of primary anxiety diagnosis (longest FU, 6 months)	1		Risk Ratio (IV, Fixed, 95% CI)	Subtotals only
1.9.1 Time	1	272	Risk Ratio (IV, Fixed, 95% CI)	1.03 [0.81, 1.30]
1.10 Number that discontinued treatment or control (EoT)	2		Risk Ratio (IV, Random, 95% CI)	Subtotals only
1.10.1 Time	2	374	Risk Ratio (IV, Random, 95% CI)	0.45 [0.17, 1.23]
1.11 Combined youth and observer reported functioning (EoT)	1		Risk Ratio (IV, Fixed, 95% CI)	Subtotals only
1.11.1 Time	1	90	Risk Ratio (IV, Fixed, 95% CI)	2.51 [1.15, 5.46]
1.12 Suicidal ideation (EoT)	1	272	Risk Ratio (M-H, Fixed, 95% CI)	10.53 [0.59, 188.57]
1.13 Suicide attempt (EoT)	1	272	Risk Ratio (M-H, Fixed, 95% CI)	Not estimable
1.14 Serious adverse events (EoT)	1	272	Risk Ratio (M-H, Fixed, 95% CI)	0.19 [0.01, 3.95]

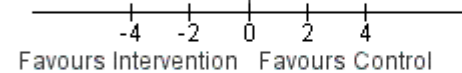
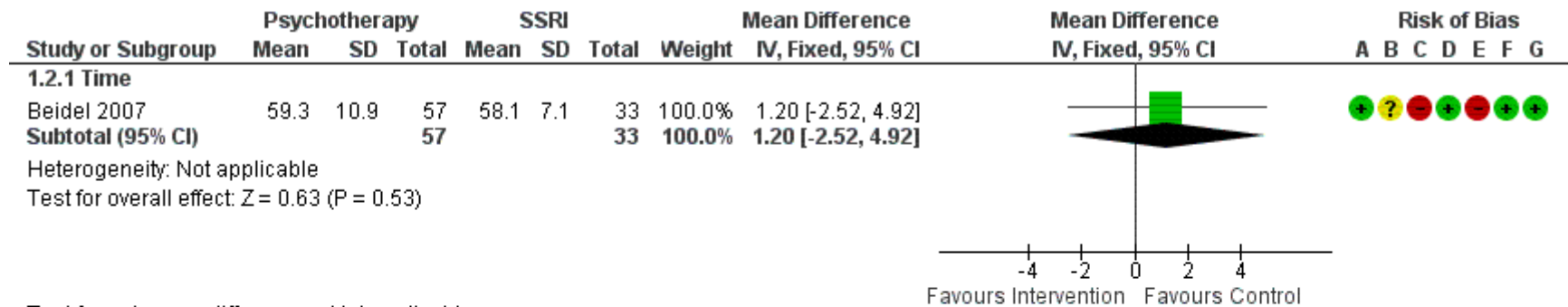
# Figures

Figure 1 (Analysis 1.1)



Forest plot of comparison: 1 Psychotherapy vs SSRI, outcome: 1.1 Youth reported anxiety symptoms (EoT).

Figure 2 (Analysis 1.2)

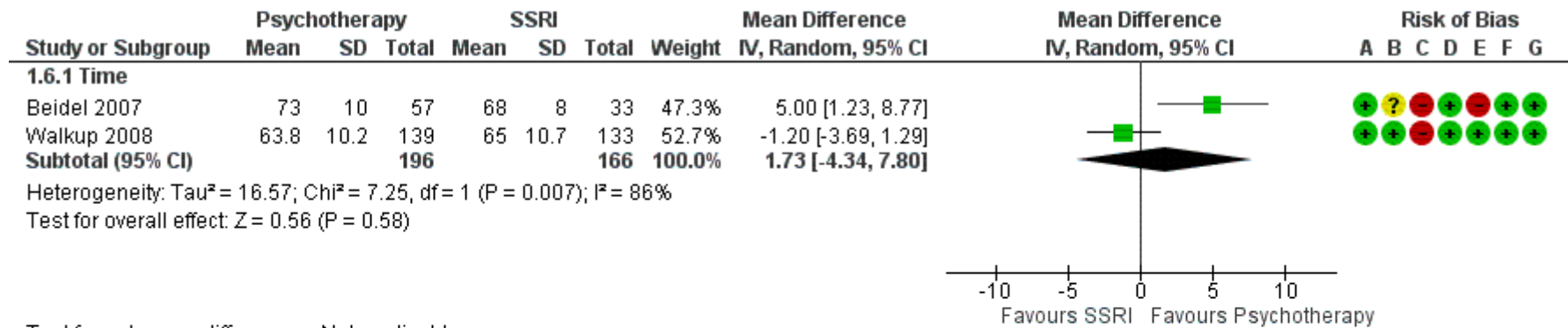


Risk of bias legend

- (A) Sequence Generation
- (B) Allocation concealment
- (C) Blinding of participants and personnel
- (D) Blinding of outcome assessors
- (E) Incomplete outcome data
- (F) Selective outcome reporting
- (G) Other sources of bias

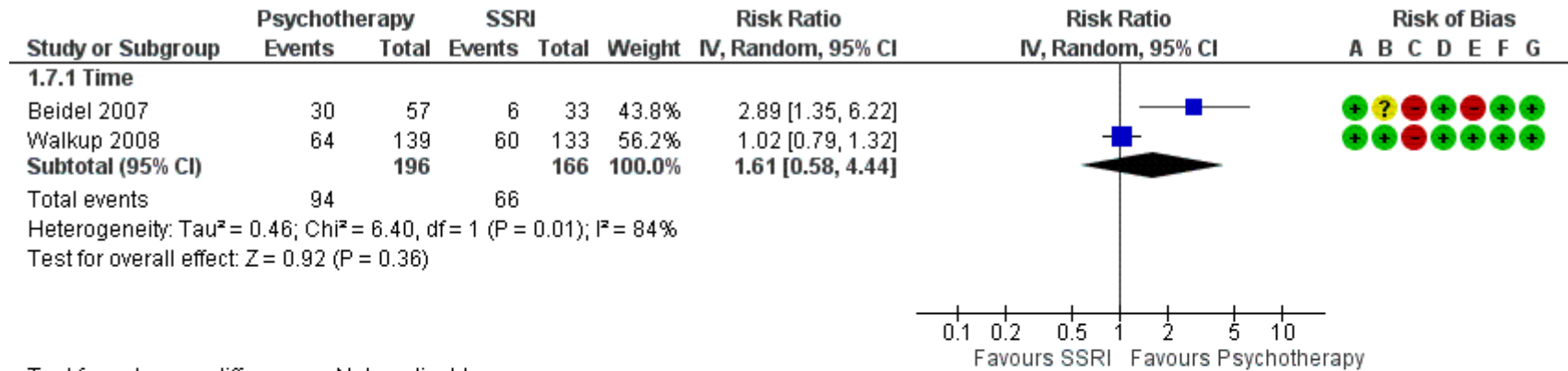
Forest plot of comparison: 1 Psychotherapy vs SSRI, outcome: 1.2 Parent reported anxiety symptoms (EoT).

**Figure 3 (Analysis 1.6)**



Forest plot of comparison: 1 Psychotherapy vs SSRI, outcome: 1.6 Observer reported functioning C-GAS(EoT).

**Figure 4 (Analysis 1.7)**



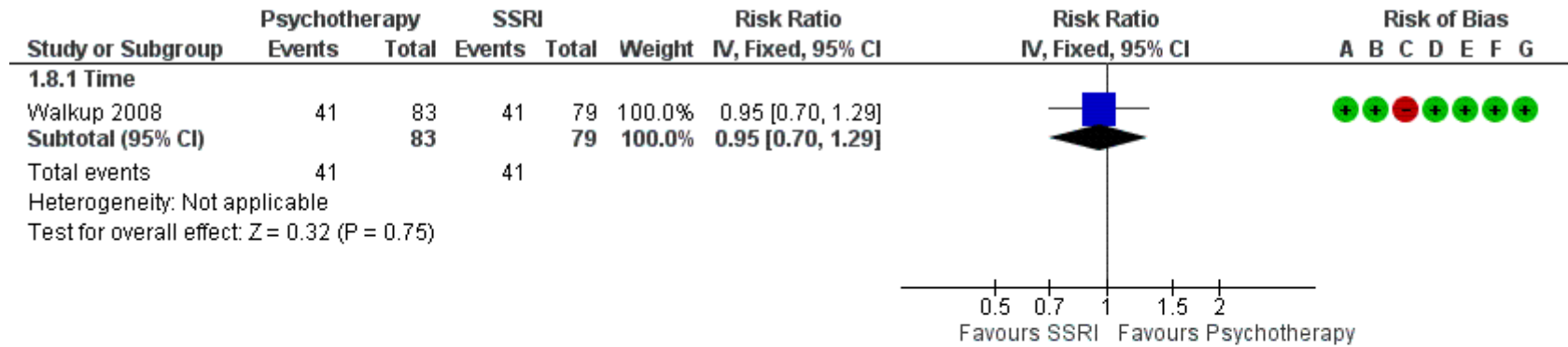
Test for subgroup differences: Not applicable

Risk of bias legend

- (A) Sequence Generation
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Forest plot of comparison: 1 Psychotherapy vs SSRI, outcome: 1.7 Remission of primary anxiety diagnosis (EoT).

**Figure 5 (Analysis 1.8)**

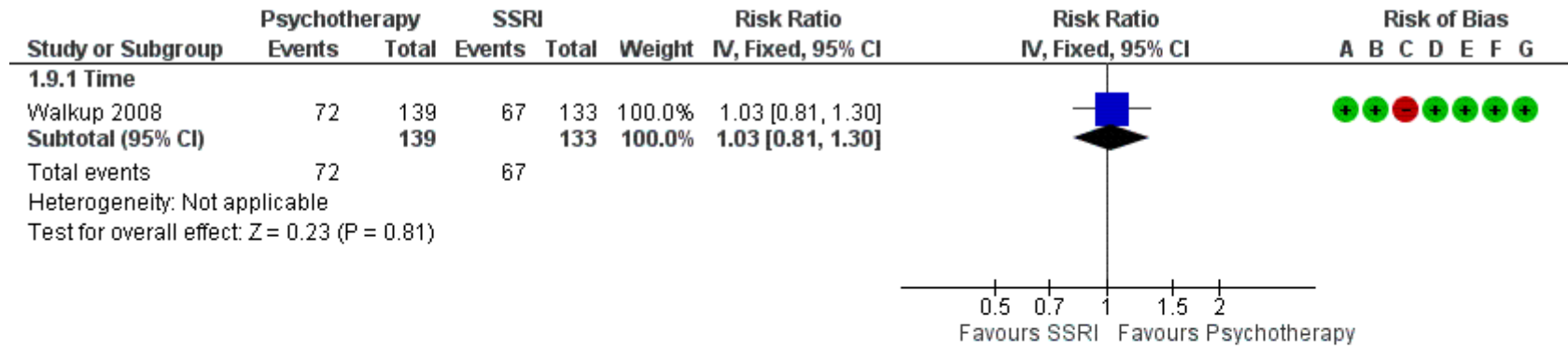


Risk of bias legend

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- (G) Other sources of bias

Forest plot of comparison: 1 Psychotherapy vs SSRI, outcome: 1.8 Remission of primary anxiety diagnosis (longest FU, 6 years).

**Figure 6 (Analysis 1.9)**

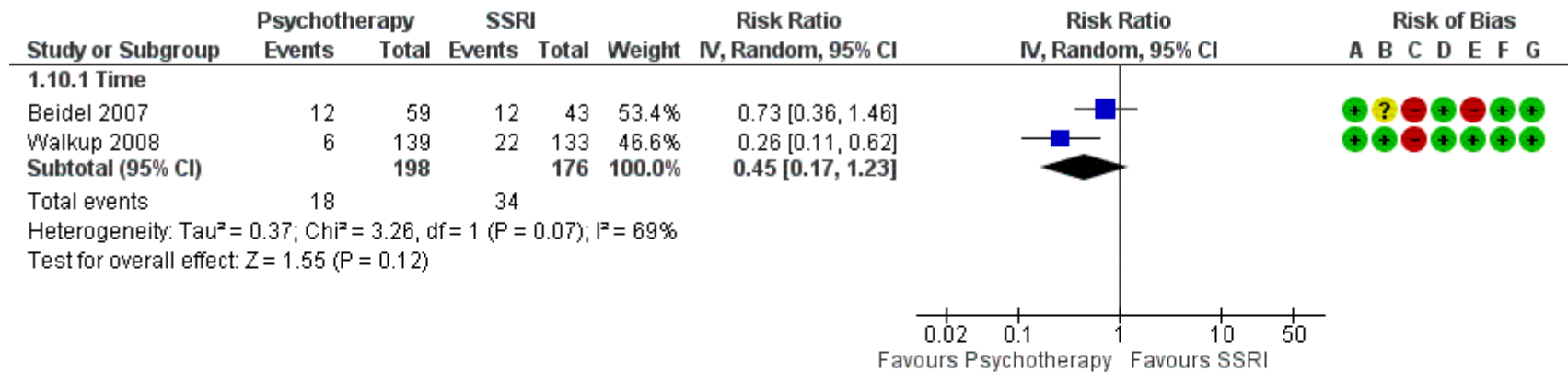


Risk of bias legend

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- (B) Allocation concealment
- (C) Blinding of participants and personnel
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Forest plot of comparison: 1 Psychotherapy vs SSRI, outcome: 1.9 Remission of primary anxiety diagnosis (longest FU, 6 months).

**Figure 7 (Analysis 1.10)**



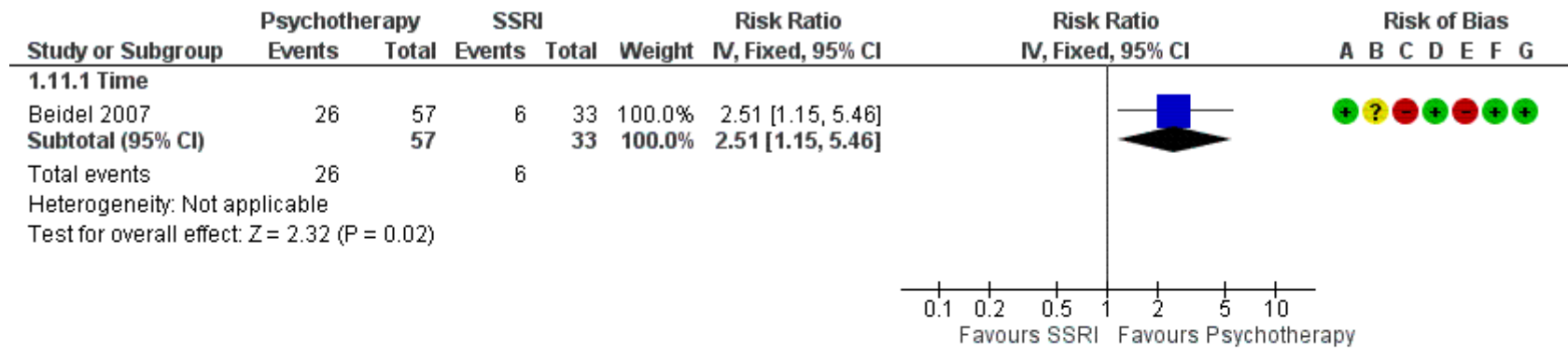
Risk of bias legend

- (A) Sequence Generation
- (B) Allocation concealment
- (C) Blinding of participants and personnel
- (D) Blinding of outcome assessors
- (E) Incomplete outcome data
- (F) Selective outcome reporting
- (G) Other sources of bias

Forest plot of comparison: 1 Psychotherapy vs SSRI, outcome: 1.10 Number that discontinued treatment or control (EoT).

**Figure 8 (Analysis 1.11)**



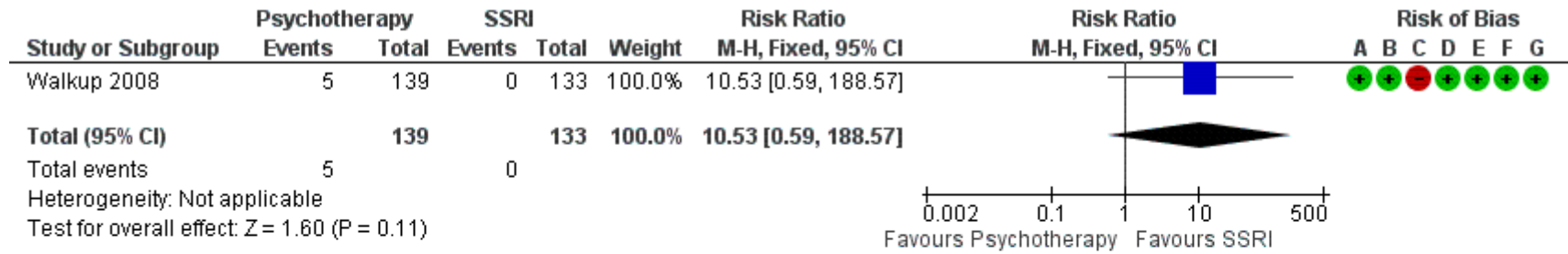


Risk of bias legend

- (A) Sequence Generation
- (B) Allocation concealment
- (C) Blinding of participants and personnel
- (D) Blinding of outcome assessors
- (E) Incomplete outcome data
- (F) Selective outcome reporting
- (G) Other sources of bias

Forest plot of comparison: 1 Psychotherapy vs SSRI, outcome: 1.11 Combined youth and observer reported functioning (EoT).

**Figure 9 (Analysis 1.12)**

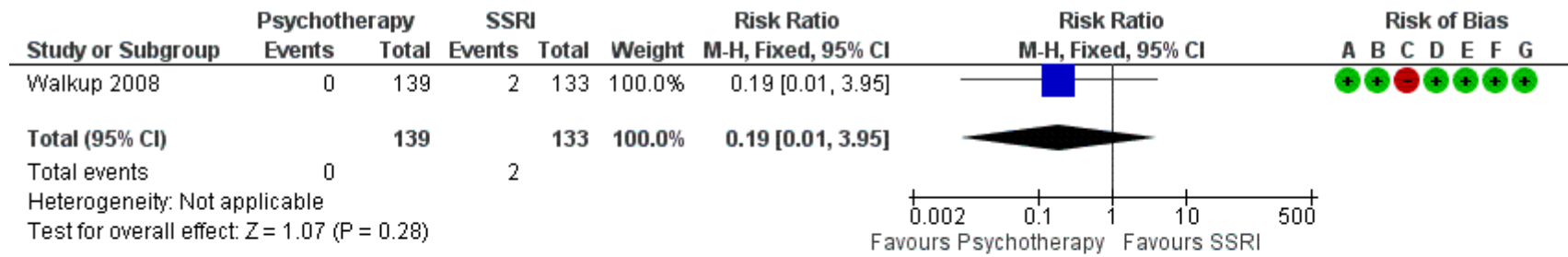


Risk of bias legend

- (A) Sequence Generation
- (B) Allocation concealment
- (C) Blinding of participants and personnel
- (D) Blinding of outcome assessors
- (E) Incomplete outcome data
- (F) Selective outcome reporting
- (G) Other sources of bias

Forest plot of comparison: 1 Psychotherapy vs SSRI, outcome: 1.12 Suicidal ideation (EoT).

**Figure 10 (Analysis 1.14)**



Risk of bias legend

- (A) Sequence Generation
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- (E) Incomplete outcome data
- (F) Selective outcome reporting
- (G) Other sources of bias

Forest plot of comparison: 1 Psychotherapy vs SSRI, outcome: 1.14 Serious adverse events (EoT).