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Multiple session early psychological interventions for the prevention of post-traumatic stress disorder (Review)
Roberts NP, Kitchiner NJ, Kenardy J, Robertson L, Lewis C, Bisson JI
Roberts NP, Kitchiner NJ, Kenardy J, Robertson L, Lewis C, Bisson JI. Multiple session early psychological interventions for the prevention of post-traumatic stress disorder.
Cochrane Database of Systematic Reviews 2019, Issue 8. Art. No.: CD006869. DOI: 10.1002/14651858.CD006869.pub3.

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[Intervention Review]

# Multiple session early psychological interventions for the prevention of post-traumatic stress disorder

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**Editorial group:** Cochrane Common Mental Disorders Group.

Publication status and date: New search for studies and content updated (conclusions changed), published in Issue 8, 2019.

**Citation:** Roberts NP, Kitchiner NJ, Kenardy J, Robertson L, Lewis C, Bisson JI. Multiple session early psychological interventions for the prevention of post-traumatic stress disorder. *Cochrane Database of Systematic Reviews* 2019, Issue 8. Art. No.: CD006869. DOI: 10.1002/14651858.CD006869.pub3.

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#### **ABSTRACT**

### **Background**

The prevention of long-term psychological distress following traumatic events is a major concern. Systematic reviews have suggested that individual psychological debriefing is not an effective intervention at preventing post-traumatic stress disorder (PTSD). Over the past 20 years, other forms of intervention have been developed with the aim of preventing PTSD.

### **Objectives**

To examine the efficacy of psychological interventions aimed at preventing PTSD in individuals exposed to a traumatic event but not identified as experiencing any specific psychological difficulties, in comparison with control conditions (e.g. usual care, waiting list and no treatment) and other psychological interventions.

### Search methods

We searched the Cochrane Central Register of Controlled Trials (CENTRAL), MEDLINE, Embase, PsycINFO and ProQuest's Published International Literature On Traumatic Stress (PILOTS) database to 3 March 2018. An earlier search of CENTRAL and the Ovid databases was conducted via the Cochrane Common Mental Disorders Controlled Trial Register (CCMD-CTR) (all years to May 2016). We handsearched reference lists of relevant guidelines, systematic reviews and included study reports. Identified studies were shared with key experts in the field.

We conducted an update search (15 March 2019) and placed any new trials in the 'awaiting classification' section. These will be incorporated into the next version of this review, as appropriate.

### **Selection criteria**

We searched for randomised controlled trials of any multiple session (two or more sessions) early psychological intervention or treatment designed to prevent symptoms of PTSD. We excluded single session individual/group psychological interventions. Comparator interventions included waiting list/usual care and active control condition. We included studies of adults who experienced a traumatic event which met the criterion A1 according to the Diagnostic and Statistical Manual (DSM-IV) for PTSD.



### **Data collection and analysis**

We entered data into Review Manager 5 software. We analysed categorical outcomes as risk ratios (RRs), and continuous outcomes as mean differences (MD) or standardised mean differences (SMDs), with 95% confidence intervals (CI). We pooled data with a fixed-effect meta-analysis, except where there was heterogeneity, in which case we used a random-effects model. Two review authors independently assessed the included studies for risk of bias and discussed any conflicts with a third review author.

#### **Main results**

This is an update of a previous review.

We included 27 studies with 3963 participants. The meta-analysis included 21 studies of 2721 participants. Seventeen studies compared multiple session early psychological intervention versus treatment as usual and four studies compared a multiple session early psychological intervention with active control condition.

Low-certainty evidence indicated that multiple session early psychological interventions may be more effective than usual care in reducing PTSD diagnosis at three to six months' follow-up (RR 0.62, 95% CI 0.41 to 0.93;  $I^2 = 34\%$ ; studies = 5; participants = 758). However, there was no statistically significant difference post-treatment (RR 1.06, 95% CI 0.85 to 1.32;  $I^2 = 0\%$ ; studies = 5; participants = 556; very low-certainty evidence) or at seven to 12 months (RR 0.94, 95% CI 0.20 to 4.49; studies = 1; participants = 132; very low-certainty evidence). Meta-analysis indicated that there was no statistical difference in dropouts compared with usual care (RR 1.34, 95% CI 0.91 to 1.95;  $I^2 = 34\%$ ; studies = 11; participants = 1154; low-certainty evidence). At the primary endpoint of three to six months, low-certainty evidence indicated no statistical difference between groups in reducing severity of PTSD (SMD -0.10, 95% CI -0.22 to 0.02;  $I^2 = 34\%$ ; studies = 15; participants = 1921), depression (SMD -0.04, 95% CI -0.19 to 0.10;  $I^2 = 6\%$ ; studies = 7; participants = 1009) or anxiety symptoms (SMD -0.05, 95% CI -0.19 to 0.10;  $I^2 = 2\%$ ; studies = 6; participants = 945).

No studies comparing an intervention and active control reported outcomes for PTSD diagnosis. Low-certainty evidence showed that interventions may be associated with a higher dropout rate than active control condition (RR 1.61, 95% CI 1.11 to 2.34; studies = 2; participants = 425). At three to six months, low-certainty evidence indicated no statistical difference between interventions in terms of severity of PTSD symptoms (SMD -0.02, 95% CI -0.31 to 0.26;  $I^2 = 43\%$ ; studies = 4; participants = 465), depression (SMD 0.04, 95% CI -0.16 to 0.23;  $I^2 = 0\%$ ; studies = 2; participants = 409), anxiety (SMD 0.00, 95% CI -0.19 to 0.19;  $I^2 = 0\%$ ; studies = 2; participants = 414) or quality of life (MD -0.03, 95% CI -0.06 to 0.00; studies = 1; participants = 239).

None of the included studies reported on adverse events or use of health-related resources.

### **Authors' conclusions**

While the review found some beneficial effects of multiple session early psychological interventions in the prevention of PTSD, the certainty of the evidence was low due to the high risk of bias in the included trials. The clear practice implication of this is that, at present, multiple session interventions aimed at everyone exposed to traumatic events cannot be recommended. There are a number of ongoing studies, demonstrating that this is a fast moving field of research. Future updates of this review will integrate the results of these new studies.

### PLAIN LANGUAGE SUMMARY

### Multiple session early psychological interventions for prevention of post-traumatic stress disorder

### Why was this review important?

Traumatic events can have a significant effect on the ability of individuals, families and communities to cope. In the past, single session interventions such as psychological debriefing were widely used with the aim of preventing continuing psychological difficulties. However, previous reviews have found that single session individual interventions have not been effective at preventing post-traumatic stress disorder (PTSD). A range of other forms of intervention have been developed to try to prevent people exposed to trauma from developing PTSD.

### Who will be interested in this review?

- People exposed to traumatic events and their loved ones.
- Professionals working in mental health services.
- General practitioners.
- · Commissioners.

### What questions did this review try to answer?



Are multiple session early psychological interventions (i.e. interventions over two or more sessions beginning within the first three months after the traumatic event) more effective than treatment as usual or another psychological intervention in:

- reducing the number of people diagnosed with PTSD;
- reducing the severity of PTSD symptoms;
- reducing the severity of depressive symptoms;
- reducing the severity of anxiety symptoms;
- improving the general functioning (e.g. social, psychological, occupational and functioning) of recipients of the intervention.

#### Which studies were included in the review?

We searched for randomised controlled trials (clinical studies where people are randomly put into one of two or more treatment groups) that examined multiple session early psychological interventions in the prevention of PTSD, published between 1970 and March 2018.

We included 27 studies with 3963 participants.

### What did the evidence from the review tell us?

- We found low-certainty evidence that multiple session early psychological interventions may be more effective than treatment as usual in preventing PTSD diagnosis three to six months after receiving the intervention.
- We found very low-certainty evidence that multiple session early psychological interventions may be neither more nor less effective than treatment as usual in preventing PTSD, either immediately after, or at seven to 12 months after, the intervention. We also found very low-certainty evidence that multiple session early psychological interventions may be neither more nor less effective than treatment as usual in reducing the severity of PTSD symptoms, either immediately or at subsequent points of follow-up.
- We found low-certainty evidence that multiple session early psychological interventions may be associated with a higher dropout rate than other psychological interventions.
- We found low-certainty evidence that multiple session early psychological interventions may be neither more nor less effective than other psychological interventions in diagnosing PTSD; reducing the severity of PTSD, depression and anxiety; or in maintaining the general functioning of participants receiving the intervention.
- We found no studies that measured adverse effects.
- We found no studies that measured use of health-related resources.

### What should happen next?

The current evidence base is small. However, new studies are being conducted and future updates of this review will incorporate the results of these.



Summary of findings for the main comparison. Any early psychological intervention compared to waiting list/usual care for the prevention of post-traumatic stress disorder

Any early psychological intervention compared to waiting list/usual care for the prevention of post-traumatic stress disorder

Patient or population: various

**Setting:** various

Intervention: any early psychological intervention

**Comparison:** waiting list/usual care

Outcomes  Nº of partici-  pants the eviden (studies)  (GRADE)	-	Certainty of	Relative ef- fect	Anticipated absolute effects* (95% CI)			
		(95% CI)	Risk with waiting list/usual care	Risk difference with any early psychological intervention			
PTSD diagnosis: post- treatment 556 ⊕⊙⊝⊝ Very lo		⊕⊝⊝⊝ Very low <sup>a,b</sup>	<b>RR 1.06</b> (0.85 to	Study population			
	very ton	1.32)	283 per 1000	17 more per 1000 (42 fewer to 91 more)			
PTSD diagnosis: 3-6	758 (5 RCTs)	⊕⊕⊝⊝ <b>Low</b> c,d	<b>RR 0.62</b> (0.41 to	Study population			
months (5 RC1s) Lowe, a	LOW	0.93)	215 per 1000	82 fewer per 1000 (127 fewer to 15 fewer)			
PTSD diagnosis: 7-12 months	_	⊕⊝⊝⊝ Very low <sup>e,f</sup>	<b>RR 0.94</b> (0.20 to	Study population			
months	(1 RCT)	very tow-	4.49)	47 per 1000	3 fewer per 1000 (38 fewer to 164 more)		
Dropouts from treatment	1154 (11 RCTs)	⊕⊕⊝⊝ <b>Low</b> d,k	RR 1.34	Study population			
	(11 KC15)	LOW <sup>u</sup> ,n	(0.91 to 1.95)	125 per 1000	43 more per 1000 (11 fewer to 119 more)		
Severity of PTSD symp- toms: 3-6 months	1921 (15 RCTs)	⊕⊕⊝⊝ <b>Low</b> g,h	-	The mean severity of PTSD symptoms: 3–6 months was 0	SMD 0.1 lower (0.22 lower to 0.02 higher)		

Severity of depressive symptoms: 3-6 months	1009 (7 RCTs)	⊕⊕⊝⊝ <b>Low</b> g,i	_	The mean severity of depressive symptoms at 3–6 months was 0	SMD 0.04 lower (0.19 lower to 0.1 higher)
Severity of anxiety symptoms: 3–6 months	945 (6 RCTs)	⊕⊕⊝⊝ <b>Low</b> g,j	_	The mean severity of anxiety symptoms at 3–6 months was 0	SMD 0.05 lower (0.19 lower to 0.10 higher)

<sup>\*</sup>The risk in the intervention group (and its 95% confidence interval) is based on the assumed risk in the comparison group and the relative effect of the intervention (and its 95% CI).

CI: confidence interval; PTSD: post-traumatic stress disorder; RCT: randomised controlled trial; RR: risk ratio; SMD: standardised mean difference.

### **GRADE Working Group grades of evidence**

**High certainty:** we are very confident that the true effect lies close to that of the estimate of the effect.

**Moderate certainty:** we are moderately confident in the effect estimate: the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different.

Low certainty: our confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the effect.

Very low certainty: we have very little confidence in the effect estimate: the true effect is likely to be substantially different from the estimate of effect.

<sup>a</sup>Three of the five studies were at high risk of bias (Brunet 2013; Mouthaan 2013; Rothbaum 2012).

bDowngraded two levels for imprecision as the total number of events was fewer than 300 and 95% CI included both little or no effect.

<sup>c</sup>Three of the five studies were at high risk of bias (Jones 2010; Mouthaan 2013; Rothbaum 2012).

<sup>d</sup>Downgraded one level for imprecision as the total number of events was fewer than 300.

eThe study was at high risk of bias (Mouthaan 2013).

Downgraded two levels as the total number of events was fewer than 300 and 95% CI included both little or no effect.

gDowngraded one level for imprecision as the 95% CI includes both little or no effect.

hTwelve of the 15 studies were at high risk of bias (Als 2015; Borghini 2014; Brom 1993; Brunet 2013; Curtis 2016; Holmes 2007; Jensen 2016; Jones 2010; Kazak 2005; Mouthaan 2013; Rothbaum 2012; Zatzick 2001).

Six of the seven studies were at high risk of bias (Als 2015; Curtis 2016; Holmes 2007; Jensen 2016; Mouthaan 2013; Zatzick 2001).

JFive of the six studies were at high risk of bias (Als 2015; Curtis 2016; Jensen 2016; Kazak 2005; Mouthaan 2013).

kEight of the 11 studies were at high risk of bias (Brom 1993; Brunet 2013; Holmes 2007; Kazak 2005; Rothbaum 2012; Ryding 1998; Ryding 2004; Zatzick 2001).

## Summary of findings 2. Any early psychological intervention compared to active control condition for the prevention of post-traumatic stress disorder

Any early psychological intervention compared to active control condition for the prevention of post-traumatic stress disorder

Patient or population: various

**Setting:** various

**Intervention:** any early psychological intervention

<b>Comparison:</b> active control of	condition
--------------------------------------	-----------

Outcomes	№ of partic-	№ of partic- Certainty ipants of the evi- (studies) dence (GRADE)	Relative effect (95% CI)	Anticipated absolute effects* (95% CI)				
	•			Risk with active control condition	Risk difference with any early psychological intervention			
Dropouts from treat-	Dropouts from treatment  425 (2 RCTs)  B⊕⊕⊙⊝ (1.11 to 2.34)		RR 1.61	Study population				
			•	168 per 1000	103 more per 1000 (19 more to 225 more)			
Severity of PTSD symp- toms: 3-6 months	465 (4 RCTs)	⊕⊕⊝⊝ <b>Low</b> <sup>c,d</sup>	_	The mean severity of PTSD symptoms at 3–6 months was 0	SMD 0.02 lower (0.31 lower to 0.26 higher)			
Severity of depressive symptoms: 3–6 months	409 (2 RCTs)	⊕⊕⊝⊝ <b>Low</b> d,e	_	The mean severity of depressive symptoms at 3–6 months was 0	SMD 0.04 higher (0.16 lower to 0.23 higher)			
Severity of anxiety symptoms: 3–6 months	414 (2 RCTs)	⊕⊕⊝⊝ <b>Low</b> d,e	_	The mean severity of anxiety symptoms at 3–6 months was 0	SMD 0 (0.19 lower to 0.19 higher)			
General functioning: 3-6 months	239 (1 RCT)	⊕⊕⊝⊝ <b>Low</b> g,h	_	The mean general functioning at 3–6 months was 0	MD 0.03 lower (0.06 lower to 0)			

<sup>\*</sup>The risk in the intervention group (and its 95% confidence interval) is based on the assumed risk in the comparison group and the relative effect of the intervention (and its 95% CI).

CI: confidence interval; MD: mean difference; PTSD: post-traumatic stress disorder; RCT: randomised controlled trial; RR: risk ratio; SMD: standardised mean difference.

#### **GRADE Working Group grades of evidence**

High certainty: we are very confident that the true effect lies close to that of the estimate of the effect.

**Moderate certainty:** we are moderately confident in the effect estimate: the true effect is likely to be close to the estimate of the effect, but there is a possibility that it is substantially different.

Low certainty: our confidence in the effect estimate is limited: the true effect may be substantially different from the estimate of the effect.

**Very low certainty:** we have very little confidence in the effect estimate: the true effect is likely to be substantially different from the estimate of effect.

<sup>&</sup>lt;sup>a</sup>Study was at high risk of bias (Gidron 2001).

bDowngraded two levels for imprecision as total number of events was fewer than 300 and 95% CI included both little or no effect.

cAll four studies were at high risk or bias (Cox 2018a; Gamble 2010; Gidron 2001; Gidron 2007).

<sup>&</sup>lt;sup>d</sup>Downgraded one level for imprecision as 95% CI included both little or no effect.

eBoth studies were at high risk of bias (Cox 2018a; Gamble 2010).

fDowngraded one level for imprecision as total number of events was fewer than 300.

gStudy was at high risk of bias (Gamble 2010).

<sup>&</sup>lt;sup>h</sup>Downgraded one level for imprecision as total number of participants was fewer than 400.





#### BACKGROUND

### **Description of the condition**

There is now a large body of literature to show that traumatic experience can cause significant psychological difficulties for large numbers of people, through events such as natural disasters (e.g. Berger 2012; Schulz 2013), human made disasters (e.g. Jenkins 2012), military combat (Brunet 2015; Richardson 2019; Stevelink 2018), rape (Dworkin 2017), violent crime (e.g. Lowe 2017; Wilson 2015), and road traffic accidents (Heron-Delaney 2013). Many individuals show great resilience in the face of such experiences and will manifest short-lived or subclinical stress reactions that diminish over time, although some will experience delayed onset of symptoms (Bryant 2013). Most people recover without medical or psychological assistance (McNally 2003). Nevertheless, a range of psychological difficulties may develop following trauma in some of those who have been exposed. These include depressive reactions; phobic reactions and other anxiety disorders; alcohol and other substance misuse and less frequently obsessive compulsive disorder, psychotic reactions and conversion symptoms. Some individuals display symptoms consistent with acute stress disorder (ASD) in the early phase after a traumatic event. Post-traumatic stress disorder (PTSD) is one of the most common enduring mental health problems to occur and has probably received most attention in the research literature.

PTSD is defined by the Diagnostic and Statistical Manual 5th edition (DSM5) as a syndrome which is comprised of four clusters of symptoms: repeated re-experiencing of the trauma; avoidance of internal and external reminders; negative alterations in cognition and mood; and alterations in arousal and reactivity (APA 2013). For a diagnosis of PTSD to be made, symptoms have to have been present for more than one month. The Diagnostic and Statistical Manual 4th edition (DSM-IV) used the term 'acute PTSD' to describe PTSD beginning before three months (APA 1994). While this term is no longer used in DSM5, the first three months continues to be seen as the priority period for early intervention. Reported rates of PTSD beginning within 12 months vary across different trauma exposed populations, with an estimated prevalence across studies at around 29% at one-month post-trauma and 17% at 12 months (Santiago 2013). Prevalence rates tend to be higher for people who have experienced intentional over non-intentional trauma (Santiago 2013). Epidemiological research suggests that around 40% of people who develop early onset PTSD go on to develop a chronic disorder (Santiago 2013). The impact on social, interpersonal and occupational functioning for people who develop chronic PTSD can be very significant across the life span (Litz 2004; Kearns 2012).

### **Description of the intervention**

To date, Cochrane Reviews have considered psychological intervention of PTSD (Bisson 2013), and pharmacological treatment of PTSD (Stein 2006). A large number of randomised controlled trials (RCTs) have demonstrated the effectiveness of some psychological interventions in treating chronic PTSD (Bisson 2013; NICE 2018). Trauma-focused cognitive behavioural therapy (CBT; see Bisson 2013; Bradley 2005), and eye movement desensitisation and reprocessing (EMDR) (NICE 2018), have the strongest evidence base. Evidence-based interventions are not effective for everyone and many people remain symptomatic, even after treatment is completed (Bradley 2005).

Since the 1990s, clinicians have been increasingly involved in attempts to develop interventions that might mitigate against the effects of trauma and prevent the onset of chronic PTSD. For several years, single session interventions such as psychological debriefing were a widely used and popular form of intervention. Debriefing came under increasing scrutiny in the 1990s and has been the subject of one Cochrane Review first published in 1998 and subsequently updated (Rose 2002). Other reviews reported similar findings (van Emmerik 2002; Bastos 2015). The lack of evidence for the efficacy of single session individual debriefing has therefore led many experts in the field to caution against its use (e.g. NICE 2018).

### How the intervention might work

Increasingly the field has turned its attention to other models of intervention (Kearns 2012; Qi 2016). These models have included multiple session interventions aimed at any individual exposed to a traumatic event with the aim of preventing the development of PTSD, interventions aimed at individuals with a known or suspected specific risk factor and interventions aimed at individuals who are clearly symptomatic. For example, psychological first aid has been increasingly prescribed as an initial form of intervention (NCTSN/NCP 2006). Psychological first aid refers to the provision of basic comfort, information, support and attendance to immediate practical and emotional needs. Brief forms of CBT offered from around two weeks' post incident have been proposed as interventions to prevent the onset of PTSD and to treat those who develop symptoms in the early stages after a trauma. Interventions aimed at enhancing social support have also been suggested (Litz 2002; Ormerod 2002). Several recent studies have been conducted to evaluate some of these forms of intervention.

### Why it is important to do this review

Some experts in the field (e.g. Bisson 2003; Brewin 2008; Qi 2016) advocate interventions that are targeted at those who are most at risk of continuing psychological difficulty. However, in the immediate aftermath of a traumatic event there is often a strong imperative from healthcare services, the public and politicians to provide psychological intervention to everyone who has been exposed regardless of symptomatology. The issues of who should be offered the intervention, timing of intervention and mode of intervention are at this time still contentious. We previously published a Cochrane Review of multiple session early psychological interventions for the prevention of PTSD (Roberts 2009) in which we found 11 studies of brief psychological interventions aimed at preventing PTSD in individuals exposed to a specific traumatic event. We found no evidence to support the use of these interventions. In a second review, we found evidence for trauma-focused CBT over a waiting list control and over supportive counselling for individuals who were displaying traumatic stress symptoms (Roberts 2010). Evidence was strongest when individuals met diagnosis for ASD or acute PTSD. This review aims to clarify the current evidence base by conducting an updated review of multiple session early interventions aimed at preventing PTSD in individuals who have been exposed to a traumatic event but have not been identified as experiencing any specific psychological difficulties.

### **OBJECTIVES**

To examine the efficacy of psychological interventions aimed at preventing PTSD in individuals exposed to a traumatic event



but not identified as experiencing any specific psychological difficulties, in comparison with control conditions (e.g. usual care, waiting list and no treatment) and other psychological interventions.

#### **METHODS**

### Criteria for considering studies for this review

#### Types of studies

RCTs. Sample size, language and publication status were not used to determine whether or not a study should be included.

### **Types of participants**

Any adult aged 18 years or older, exposed to a traumatic event. When a study included mixed adult and adolescent participants, we attempted to obtain separate data for adults when this was available. If this data was not available we required that at least 80% of the sample was aged 18 or over to include the study. For the purposes of the review, an event was considered traumatic if it was likely to meet criterion A1 of DSM-IV for PTSD (APA 1994). Therefore, the majority of participants in included studies were considered to have experienced, witnessed, or been confronted with an event or events that involved actual or threatened death or serious injury, or a threat to the physical integrity of self or others.

We excluded studies that enlisted participants who met a certain symptom profile (e.g. ASD, acute PTSD, depression) or recruited participants on the basis of responses to a screening measure.

### Types of interventions

This review considered any multiple session early psychological intervention designed to prevent symptoms of traumatic stress, and begun within three months of a traumatic incident. We excluded single session interventions because they are the subject of a separate Cochrane Review (Rose 2002). Early psychological interventions aimed at treating individuals who were identified as symptomatic (e.g. with ASD or acute PTSD) is subject to a separate review (Roberts 2010) conducted at the same time as this review.

For the purpose of the review, a psychological intervention included any specified non-pharmaceutical intervention aimed at preventing the onset of PTSD offered by one or more health professional or lay person, with contact between therapist and participant on at least two occasions. We decided a priori that eligible intervention categories would include forms of psychological therapy based on a specified theoretical model. Potential intervention categories were identified from previous PTSD-based reviews (Bisson 2013; NICE 2018). These were:

- trauma-focused cognitive behavioural therapy (TF-CBT) any psychological intervention that predominantly used trauma-focused cognitive, behavioural or cognitive-behavioural techniques. This category included exposure therapy;
- stress management/relaxation any psychological intervention that predominantly taught relaxation or anxiety/stress management techniques;
- TF-CBT group therapy any approach delivered in a group setting that predominantly used trauma-focused cognitive, behavioural or cognitive-behavioural techniques;

- CBT any psychological intervention that predominantly used non-trauma-focused cognitive, behavioural or cognitivebehavioural techniques. This category excluded the use of exposure therapy;
- 5. EMDR any psychological intervention that predominantly used FMDR.
- non-trauma-focused CBT group therapy any approach delivered in a group that predominantly used nontrauma-focused cognitive, behavioural or cognitive-behavioural techniques;
- 7. other psychological intervention any psychological intervention that predominantly used non-trauma-focused techniques that would not be considered cognitive, behavioural or cognitive-behavioural techniques. This category included non-directive counselling, psychodynamic therapy and hypnotherapy.

We also decided a priori that eligible interventions would include non-pharmaceutical interventions that were not based or only partially based on a specified theoretical model but that nevertheless aimed to reduce symptoms of traumatic stress, to include the following categories.

- Education or information giving intervention any intervention which predominantly provided only education or information about possible future difficulties or offered advice about constructive means of coping, or both.
- 2. Stepped care any a priori specified care plan which offered intervention in a stepped care manner based on the continuing needs of the included participants.
- 3. Interventions aimed at enhancing positive coping skills and improving overall well-being any non-pharmaceutical intervention which aimed to improve well-being such as an occupational therapy intervention, an exercise-based intervention or a guided self-help intervention.

We decided a priori that the trials considered would include:

- psychological intervention versus waiting list or usual care control;
- 2. psychological intervention versus other psychological intervention.

From prior knowledge of the literature, it was clear that a number of different forms of intervention had been evaluated on differing participant groups. Several studies were thought to have offered intervention to all individuals exposed. Others were known to have evaluated interventions for those who met inclusion based on predictors of future risk. We decided to undertake comparison of all interventions together initially and to undertake subanalysis on specific interventions and interventions targeted at individuals meeting specific risk factors as appropriate.

### Types of outcome measures

### **Primary outcomes**

- Rates of PTSD among those exposed to trauma as measured by a standard classificatory system, assessed using a standardised measure such as the Clinician-Administered PTSD Scale (CAPS; Blake 1995).
- 2. Dropout from treatment.



### Secondary outcomes

- Severity of traumatic stress symptoms using a standardised measure such as the CAPS (Blake 1995), Impact of Event Scale (Horowitz 1979), the Davidson Trauma Scale (Davidson 1997), or the Post-traumatic Diagnostic Scale (Foa 1995). In circumstances where an individual study utilised both a clinician-administered and a self-reported measure, primacy was given to outcomes using the clinician-administered measure, as such measures are considered to provide the 'gold standard' in the traumatic stress field (e.g. Foa 1997).
- Severity of self-reported depressive symptoms using a standardised measure such as the Beck Depression Inventory (Beck 1961).
- Severity of self-reported anxiety symptoms using a standardised measure such as the Beck Anxiety Inventory (Beck 1988), or the Spielberger State-Trait Anxiety Inventory (Spielberger 1970)
- 4. Adverse effects.
- 5. General functioning including quality of life measures such as the 36-item Short Form (SF-36; Ware 1993).
- 6. Use of health-related resources.

Comparisons involving follow-up data would only be made when outcome data were available for similar time points. These time points were decided a priori as post-treatment, three to six months post-trauma, seven to 12 months post-trauma, one to two years post-trauma, two years and beyond. Three to six months post-trauma was the primary outcome period.

### Search methods for identification of studies

### **Electronic searches**

## Cochrane Common Mental Disorders Controlled Trials Register (CCMD-CTR)

The Cochrane Common Mental Disorders (CCMD) Group maintains a specialised register of RCTs, the CCMD-CTR. This register contains over 40,000 reference records (reports of RCTs) for anxiety disorders, depression, bipolar disorder, eating disorders, self-harm and other mental disorders within the scope of this Group. The CCMD-CTR is a partially studies-based register with more than 50% of reference records tagged to approximately 12,500 individually PICO-coded study records. Reports of trials for inclusion in the register are collated from (weekly) generic searches of MEDLINE (from 1950), Embase (from 1974) and PsycINFO (from 1967); quarterly searches of the Cochrane Central Register of Controlled Trials (CENTRAL), and review specific searches of additional databases. Reports of trials are also sourced from international trial registries, drug companies, the handsearching of key journals, conference proceedings, and other (non-Cochrane) systematic reviews and meta-analyses. Details of CCMD's core search strategies (used to identify RCTs) can be found on the Group's website, with an example of the core MEDLINE search displayed in Appendix 1. The CCMD-CTR fell out of date in June 2016, with the move of the editorial base from the University of Bristol to York.

Searches for this review were conducted in August 2008, May 2016 and March 2018.

### Search one (1 August 2008)

CCMDCTR-Studies: Diagnosis = "stress disorder\*" or PTSD and Intervention = therapy or intervention or counsel\* or debriefing

and Age-group = adult or aged or "not stated" or unclear <u>and not</u> Duration of therapy = "1 session"

CCMDCTR-References: Keyword = "Stress Disorder\*" or "Stress-Disorder\*" or Free-text = PTSD and Free-text = debrief\* or \*therap\* or intervention\* or counsel\*

#### Search two (6 May 2016)

CCMDCTR-Studies Register: (PTSD or posttrauma\* or posttrauma\* or "post trauma\*" or "combat disorder\*" or "stress disorder\*"):sco,stc

CCDMDCTR-References Register: (PTSD or posttrauma\* or posttrauma\* or "post trauma\*" or "combat disorder\*" or "stress disorder\*"):ti,ab,kw,ky,emt,mh,mc

[Key to field tags. ti:title; ab:abstract; kw:keywords; ky:other keywords; mh:MeSH headings; mc:MeSH check words; emt:EMTREE headings; sco:healthcare condition; stc:target condition]

#### Search three (3 March 2018)

CCMD's information specialist conducted additional searches on the following bibliographic databases, using relevant subject headings (controlled vocabularies) and search syntax, appropriate to each resource.

The search was for a suite of PTSD reviews and the search terms were based on population <u>or</u> psychological debriefing (Appendix 2).

- The Cochrane Central Register of Controlled Trials (CENTRAL), Issue 2 of 12, February 2018;
- 2. Ovid MEDLINE (2014 to 3 March 2018);
- 3. Ovid Embase (2014 to 3 March 2018);
- 4. Ovid PsycINFO (2014 to 3 March 2018);
- 5. Ebsco PILOTS (2014 to 3 March 2018).

### Search four (15 March 2019)

In keeping with MECIR conduct standard c37 (searches to be within 12 months of publication), we ran an update search in March 2019. We screened the abstracts and placed any new trials matching our inclusion criteria as 'awaiting classification'. These will be incorporated into the next version of this review, as appropriate.

### **Searching other resources**

### Reference lists

We searched reference lists of the National Institute for Health and Care Excellence PTSD guidelines (NICE 2018), and studies identified in the search and of related review articles.

### **Personal communication**

We provided a list of included references on the website of the International Society for Traumatic Stress Studies and contacted the membership to ask them to identify any studies that they thought might be missing.



### Data collection and analysis

#### **Selection of studies**

Two review authors (NR and CL) independently read the abstracts of all potential trials. If an abstract appeared to represent an RCT, each review author independently read the full report to determine if the trial met the inclusion criteria. When agreement could not be reached about inclusion, we consulted a third review author. The studies excluded on further reading are listed in the Characteristics of excluded studies table, with reasons for their exclusion.

### **Data extraction and management**

We designed a data extraction sheet to capture data that was entered into Review Manager 5 software (Review Manager 2014). Information extracted included demographic details of participants, details of the traumatic event, the randomisation process, the interventions used, dropout rates and outcome data. Two review authors (of NR, NK and JK) independently extracted data. When agreement could not be reached, we discussed the issue a third review author.

#### Assessment of risk of bias in included studies

Two review authors (of NR, NK and JK) independently assessed the risk of bias for each study, using the criteria outlined in the *Cochrane Handbook for Systematic Reviews of Interventions* and listed below (Higgins 2011). We resolved conflicts through discussion with a third review author (JB).

- 1. Random sequence generation.
- 2. Allocation concealment.
- 3. Blinding of outcome assessment.
- 4. Incomplete outcome data.
- 5. Selective outcome reporting.
- 6. Other bias (including baseline imbalances, early termination of the trial, researcher allegiance).

We did not assess blinding of participants and personnel as a double-blind methodology for studies of psychological treatment is impossible as it is clear to participants what treatment they are receiving.

We judged each potential source of bias as high, low or unclear, and provided a supporting quotation from the study report, together with a justification for the judgement, in the 'Risk of bias' table. We summarised risk of bias judgements across different studies for each of the domains listed. We considered blinding separately for different key outcomes when necessary. When information on risk of bias related to unpublished data or correspondence with a trialist, we noted this in the 'Risk of bias' table.

When considering treatment effects, we took into account risk of bias for studies that contributed to that outcome.

### Measures of treatment effect

### Continuous data

We analysed continuous outcomes using mean difference (MD) when all trials had measured outcome on the same scale. When trials measured outcomes on different scales, we used the standardised mean difference (SMD).

#### Dichotomous data

We used the risk ratio (RR) as the main categorical outcome measure as this is more widely used than odds ratio (OR) in health-related practice. All outcomes were presented using 95% confidence intervals (CI).

### Unit of analysis issues

For trials which had a crossover design, we considered only results from the first randomisation period. If the trial was a three (or more) armed trial, consideration was given to undertaking pairwise meta-analysis with each arm, depending upon the nature of the intervention in each arm and the relevance to the review objectives. Management of cluster randomised trials followed guidance provided in the *Cochrane Handbook for Systematic Reviews of Interventions* (Higgins 2011).

### Dealing with missing data

When intention-to-treat (ITT) data were available, we reported these in the results. We attempted to access ITT data wherever possible. We used completer-only data when this was the only type of data available. In cases where there was inadequate information within a particular paper to undertake analysis, we attempted to compute missing data from other information available within the paper, using guidance provided in the *Cochrane Handbook for Systematic Reviews of Interventions* (Higgins 2011). For example, we calculated standard deviations (SDs) for continuous data when only the standard error (SE) or t statistics or P values were reported. When imputation was not possible or when further clarification was required, we attempted to contact the authors to request additional information. In cases where no further useable data were available, the study was not included in further analysis.

### Assessment of heterogeneity

We used a visual inspection of the forest plots initially to explore for possible statistical heterogeneity (variation in intervention effects or results). We measured heterogeneity between studies by observing the I² test and the Chi² test (P < 0.10). An I² of less than 30% was considered to indicate that statistical heterogeneity might not be important; an I² of 30% to 60% to indicate moderate heterogeneity; an I² of 50% to 90% to indicate substantial heterogeneity and an I² greater than 75% to indicate considerable heterogeneity. Clinical and methodological heterogeneity reflect issues such as differences in participant populations, intervention types, study design and methodological rigour. We anticipated that included studies would evaluate a range of different interventions in a wide variety of populations. Therefore, we used a random-effects model for all comparisons.

### **Assessment of reporting biases**

It was decided a priori that if a minimum of 10 studies were available in a meta-analysis, we would prepare funnel plots and examine them for signs of asymmetry. Where there was asymmetry, we planned to consider other possible reasons for this.

### **Data synthesis**

We pooled data from more than one study using a fixed-effect model, except where heterogeneity was considered to be present. In these cases, we used a random-effects model as described below.



### Subgroup analysis and investigation of heterogeneity

It was decided a priori that we would explore the following possible causes of clinical heterogeneity if sufficient data allowed.

- Number of treatment sessions taken (two to six versus seven or more).
- 2. Type of traumatic event (combat-related trauma versus rape and sexual assault versus other civilian trauma).
- 3. Participant characteristics (men versus women)

### Sensitivity analysis

It was decided a priori that sensitivity analysis would explore possible causes of methodological heterogeneity if sufficient data allowed. Analysis would be based on the following criteria.

- 1. Trials considered most susceptible to bias would be excluded based on the following quality assessment criteria:
  - a. those with unclear allocation concealment;
  - b. high levels of postrandomisation losses (more than 40%) or exclusions;
  - c. unblinded outcome assessment or blinding of outcome assessment uncertain.
- 2. Use of ITT analysis versus completer outcomes would be undertaken depending on available data.

### 'Summary of findings' tables

We evaluated the certainty of available evidence using the GRADE approach. We generated 'Summary of findings' tables using GRADEpro GDT software, which imports data from Review Manager 5 (GRADEpro GDT; Review Manager 2014). These tables provided outcome-specific information concerning the overall certainty of evidence from studies included in the comparison, the magnitude of effect of the interventions examined and the sum of available data on outcomes considered. We included information on the first seven outcomes of our review: PTSD diagnosis (on a clinician-administered scale), severity of traumatic stress symptoms, severity of self-reported depressive symptoms,

severity of self-reported anxiety symptoms, dropouts, adverse effects and general functioning. For the primary outcome of PTSD diagnosis, we reported all time points available. For the secondary outcomes, we prioritised the primary end point of three to six months' postintervention.

We assessed the certainty of evidence using five factors.

- 1. Limitations in study design and implementation of available studies.
- 2. Indirectness of evidence.
- 3. Unexplained heterogeneity or inconsistency of results.
- 4. Imprecision of effect estimates.
- 5. Potential publication bias.

For each outcome, we classified the certainty of evidence according to the following categories.

- 1. High certainty: further research is very unlikely to change our confidence in the estimate of effect.
- Moderate certainty: further research is likely to have an important impact on our confidence in the estimate of effect, and may change the estimate.
- 3. Low certainty: further research is very likely to have an important impact on our confidence in the estimate of effect, and is likely to change the estimate.
- 4. Very low certainty: we are very uncertain about the estimate.

We downgraded the evidence from high certainty by one level for serious (or by two for very serious) study limitations (risk of bias), indirectness of evidence, serious inconsistency, imprecision of effect estimates or potential publication bias.

### RESULTS

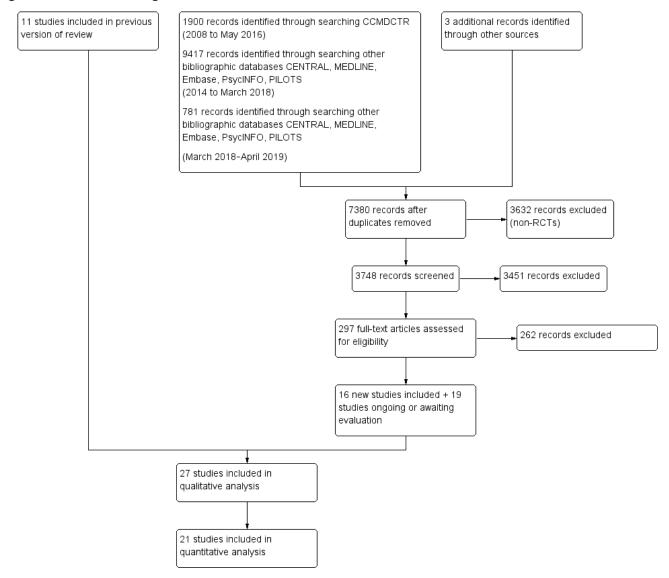
### **Description of studies**

### Results of the search

See Figure 1.



Figure 1. PRISMA flow diagram.



The search identified 7380 titles and abstracts and two review authors independently read 297 papers in detail to establish if they met the specified inclusion criteria.

An update search (15 March 2019) identified 781 RCT records. We screened the abstracts and have added 11 new studies to those awaiting classification.

### **Included studies**

Twenty-seven studies, including 11 identified in Roberts 2009, evaluated brief (two or more sessions) psychological interventions aimed at preventing PTSD in people exposed to a specific traumatic event. Twenty five studies were reported in English, one in French (Andre 1997), and one in Persian (Taghizadeh 2008). The studies are described in the Characteristics of included studies table.

### Study design

All included studies were two armed RCTs, apart from one study (Lindwall 2014), which included three arms, one of which evaluated intervention offered to children which was not eligible for inclusion

in this review. Participants would not have been blind to their allocation group. Sample size in the included studies varied from 17 (Gidron 2001) to 386 (Jensen 2016) participants. The total number of participants randomised in the 27 studies was 3963.

### **Participants**

Seven studies were conducted in the USA (Biggs 2016; Cox 2018a; Curtis 2016; Kazak 2005; Lindwall 2014; Rothbaum 2012; Zatzick 2001); three in Canada (Brunet 2013; Irvine 2011; Marchand 2006); three in Australia (Gamble 2005; Gamble 2010; Holmes 2007); two in the Netherlands (Brom 1993; Mouthaan 2013); two in Israel (Gidron 2001; Gidron 2007); two in Sweden (Ryding 1998; Ryding 2004); one in France (Andre 1997); one in Iran (Taghizadeh 2008); one in the UK (Als 2015); one in Switzerland (Borghini 2014); one in Denmark (Jensen 2016); one in China (Wang 2015); one in Sri Lanka (Wijesinghe 2015); and one study was conducted in various European countries including Denmark, Italy, Norway, Portugal, Sweden and the UK (Jones 2010).



Five studies evaluated interventions offered to mothers who had experienced traumatic births (Gamble 2005; Gamble 2010; Ryding 1998; Ryding 2004; Taghizadeh 2008), while another study was of mothers of babies born at less than 33 weeks' gestation (Borghini 2014). Five studies included individuals who had been involved in road traffic accidents (Brom 1993; Gidron 2001; Gidron 2007; Wang 2015; Zatzick 2001). In four studies, participants were family members (Als 2015; Curtis 2016; Kazak 2005; Lindwall 2014): one study was in relatives of patients admitted to an intensive care unit (Curtis 2016), one was in parents whose child was newly diagnosed with cancer (Kazak 2005), one was in parents whose child received a stem cell or bone marrow transplant (Lindwall 2014), and one was in parents of a child admitted to a paediatric intensive care unit (Als 2015). Three studies were in individuals who had experienced major physical trauma and had been admitted to a trauma centre (Holmes 2007; Mouthaan 2013; Rothbaum 2012), while another three were in individuals who had been admitted to an intensive care unit (Cox 2018a; Jensen 2016; Jones 2010). One study included individuals who had been exposed to armed robbery, involving acts of violence (Marchand 2006). Participants in this study had to have reported experiencing intense fear, helplessness or horror during or after the robbery for inclusion. One study was conducted in individuals who had been exposed to a life-threatening event (Brunet 2013), one included bus drivers who had been assaulted (Andre 1997), one was in people who had received an implantable cardioverter defibrillator (ICD) (Irvine 2011), one was on military mortuary workers (Biggs 2016), and one was in snakebite victims (Wijesinghe 2015).

### Interventions

Twenty-one studies compared multiple session early interventions versus treatment as usual. Five studies used an approach which we grouped as "brief individual trauma processing" (Brom 1993; Gamble 2005; Marchand 2006; Rothbaum 2012; Ryding 1998). This subgroup consisted of a number of brief therapies - lasting two or more sessions - that were theoretically diverse but shared similar core treatment components. These included: psychoeducation, therapist directed reliving of the index trauma to promote elaboration of the trauma memory and help to contextualise or reframe aspects of the experience. Two studies used CBT (Andre 1997; Irvine 2011), two used brief dyadic CBT (therapy involving a trauma exposed individual in conjunction with a significant other person, such as a partner, spouse or other family member) (Brunet 2013; Kazak 2005), one used a self-guided Internetbased intervention (Mouthaan 2013), one used brief interpersonal therapy (IPT) (Holmes 2007), one used a counselling intervention (Taghizadeh 2008), one used group counselling (Ryding 2004), one used psychological first aid group sessions (Biggs 2016), one used collaborative care (Zatzick 2001), one used intensive care diaries (Jones 2010), one used three-step early intervention (Borghini 2014), one used supported psychoeducation (Als 2015), one used communication facilitator in an intensive care setting (Curtis 2016), one used nurse-led intensive care recovery programme (Jensen 2016), and one used creative art (Wang 2015).

Six studies compared a psychological intervention with another intervention. Two studies compared memory structuring intervention with supportive listening (Gidron 2001; Gidron 2007), one study compared CBT with an education programme (Cox 2018a), one study compared a resilience programme with parenting support (Gamble 2010), one study compared a child-targeted intervention (massage and humour therapy) with child-targeted

plus a parent-targeted (massage, relaxation, imagery) intervention (Lindwall 2014), and one compared psychological first aid and psychoeducation and CBT versus psychological first aid and psychoeducation (Wijesinghe 2015).

Further information about specific interventions is provided in the Characteristics of included studies table.

#### **Outcomes**

Most of the included studies used well validated self-report measures of PTSD, depression or anxiety as key outcomes. Measures used are listed in the Characteristics of included studies table. The most commonly used tool for measuring PTSD was the Impact of Events Scale (IES) (Als 2015; Andre 1997; Brom 1993; Marchand 2006; Rothbaum 2012; Ryding 1998; Ryding 2004; Taghizadeh 2008), followed by the Impact of Event Scale - Revised (IES-R) (Brunet 2013; Cox 2018a; Irvine 2011; Kazak 2005; Lindwall 2014; Mouthaan 2013; Wang 2015), Post-Traumatic Stress Disorder Checklist (PCL) (Biggs 2016; Curtis 2016; Holmes 2007; Zatzick 2001), and CAPS (Brunet 2013; Mouthaan 2013; Wang 2015). Other PTSD scales used less commonly included the Mini-International Neuropsychiatric Interview for PTSD (MINI-PTSD) (Gamble 2005), Perinatal Posttraumatic Stress Disorder Questionnaire (PPQ) (Borghini 2014), Harvard Trauma Questionnaire (HTQ) (Jensen 2016), PTSD Symptom Scale -Interview Version (PSS-I) (Rothbaum 2012), and Post-traumatic Stress Symptom Scale – Self Report (PSS-SR) (Wijesinghe 2015).

Other outcomes such as anxiety, depression and quality of life were measured using Hospital Anxiety and Depression Scale (HADS) (Als 2015; Andre 1997; Cox 2018a; Holmes 2007; Irvine 2011; Jensen 2016; Mouthaan 2013; Wang 2015), Patient Health Questionnaire (PHQ-9) (Biggs 2016; Curtis 2016), SF-36 (Holmes 2007; Irvine 2011; Jensen 2016; Mouthaan 2013), Generalised Anxiety Disorder Assessment (GAD-7) (Curtis 2016), World Health Organization Quality of Life Assessment (Biggs 2016), Trauma Symptom Inventory (Brom 1993), Social Constraints Scale (SCS) (Brunet 2013), Social Adjustment Scale by Self-Report (SAS-SR) (Brunet 2013), EuroQol (EQ-5D) (Cox 2018a), Patient-Reported Outcomes Measurement Information System (PROMIS) (Cox 2018a), Brief Coping Orientation to Problems Experienced (COPE) (Cox 2018a), Edinburgh Postnatal Depression Scale (EPDS) (Gamble 2005; Gamble 2010; Ryding 2004), Depression Anxiety and Stress Scale-21 (DASS-21) (Gamble 2005; Gamble 2010), Maternity Social Support Scale (MSSS) (Gamble 2005), and health-related quality of life (HRQoL) (Gamble 2010).

We used data from a clinician-administered PTSD severity measure for six studies (Brunet 2013; Gamble 2005; Jones 2010; Marchand 2006; Mouthaan 2013; Rothbaum 2012). Data were only available from self-report measures from17 trials (Als 2015; Borghini 2014; Brom 1993; Cox 2018a; Curtis 2016; Gamble 2010; Gidron 2001; Gidron 2007; Holmes 2007; Irvine 2011; Jensen 2016; Kazak 2005; Lindwall 2014; Ryding 1998; Ryding 2004; Taghizadeh 2008; Zatzick 2001).

### **Excluded studies**

See Characteristics of excluded studies table.

In this update, we culled the list of excluded studies (previously reported in the first version of this review (Roberts 2009), to those



studies which narrowly missed the inclusion criteria, those readers may plausibly expect to see included.

We excluded 38 studies following access of full-text articles, as they did not satisfy the inclusion criteria. Of these, participants already had PTSD symptoms in 34 trials (Ben-Zion 2018; Bisson 2004; Bryant 1998; Bryant 1999; Bryant 2003; Bryant 2005; Bryant 2008; Bugg 2009; Cernvall 2015; Echeburua 1996; Ehlers 2003; Foa 2006; Freedman (in preparation); Freedman (submitted); Freyth 2010; Jarero 2011; Jarero 2015; Nixon 2012; Nixon 2016; O'Donnell (in preparation); O'Donnell 2012; Öst unpublished; Shalev 2012; Shapiro 2015; Shapiro 2018; Shaw 2013; Sijbrandij 2007; Skogstad 2015; van Emmerik 2008; Wagner 2007; Wu 2014; Zatzick 2004; Zatzick 2013; Zatzick 2015). Four studies evaluated single session interventions (Resnick 2005; Rose 1999; Rothbaum (submitted); Turpin 2005).

### Studies awaiting classification

One study could not be accessed and has been included in studies awaiting classification (Kilpatrick 1984). A further 11 newly completed studies were identified in the most recent search of March 2019 and are awaiting classification (see Characteristics of studies awaiting classification table).

#### **Ongoing studies**

For details of ongoing studies, see Characteristics of ongoing studies table.

#### Risk of bias in included studies

See Figure 2 and Figure 3.



Figure 2. Risk of bias summary: review authors' judgements about each risk of bias item for each included study.

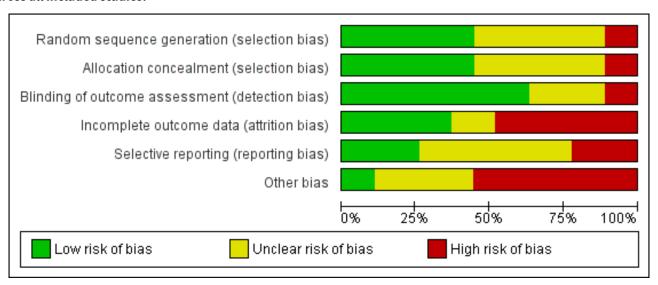
	Random sequence generation (selection bias)	Allocation concealment (selection bias)	Blinding of outcome assessment (detection bias)	Incomplete outcome data (attrition bias)	Selective reporting (reporting bias)	Other bias
Als 2015	•	•	•	•	•	•
Andre 1997	?	?	?			•
Biggs 2016	?	?	•	•	•	•
Borghini 2014	?	•	?	•	?	?
Brom 1993	?	?	•	•	?	•
Brunet 2013	•	?	?	•	?	•
Cox 2018a	•	•	•	•	•	•
Curtis 2016	•	•	•	•	•	?
Gamble 2005	•	•	?	•	?	?
Gamble 2010	•	?	?	•	•	•
Gidron 2001	?	?	•	•	?	•
Gidron 2007	?	?	•		•	•
Holmes 2007	•	•	•		?	?
Irvine 2011	•	?	•	•	•	?
Jensen 2016	•	•	?	•	•	•
Jones 2010	•	•	•	•	?	
Kazak 2005	?	•	•	•	?	•
Lindwall 2014	?	?	?	•	•	•
Marchand 2006	?	?	•	•	?	?
Mouthaan 2013	•	•	•	•	•	•
Rothbaum 2012	•	•	•	?	?	
Dudina 1000					2	<u></u>



Figure 2. (Continued)



Figure 3. Risk of bias graph: review authors' judgements about each risk of bias item presented as percentages across all included studies.



Methodological quality of many of the included studies was poor. The process of recruitment used by Brom 1993 was of particular concern as recruitment took place prior to invitation to potential participants to join the study. This may lead to significant difference in response rate, dropout rate, other dropout factors and baseline scores between the treatment and control group.

### **Random sequence generation**

Twelve studies provided an adequate description of the randomisation process and were at low risk of selection bias (Als 2015; Brunet 2013; Cox 2018a; Curtis 2016; Gamble 2005; Gamble 2010; Holmes 2007; Irvine 2011; Jensen 2016; Jones 2010; Mouthaan 2013; Rothbaum 2012). Twelve studies did not report the randomisation process and were therefore at unclear risk of selection bias (Andre 1997; Biggs 2016; Borghini 2014; Brom 1993; Gidron 2001; Gidron 2007; Kazak 2005; Lindwall 2014; Marchand 2006; Wang 2015; Wijesinghe 2015; Zatzick 2001). Three studies were at high risk of selection bias as the process was not truly random (Ryding 1998; Ryding 2004: Taghizadeh 2008). Ryding 2004 randomised women who gave birth on approximately 18 predetermined days of the month to the counselling group. Ryding 1998 selected every second emergency caesarean section

patient, according to the delivery ward register, for counselling, the remainder were in the comparison group. Taghizadeh 2008 randomised participants by day of the week.

### **Allocation concealment**

Twelve studies reported adequate concealment procedures and were at low risk of selection bias (Als 2015; Borghini 2014; Cox 2018a; Curtis 2016; Gamble 2005; Holmes 2007; Jensen 2016; Jones 2010; Kazak 2005; Mouthaan 2013; Rothbaum 2012; Zatzick 2001). In 12 studies, allocation concealment was unclear or inadequate (Andre 1997; Biggs 2016; Brom 1993; Brunet 2013; Gamble 2010; Gidron 2001; Gidron 2007; Irvine 2011; Lindwall 2014; Marchand 2006; Wang 2015; Wijesinghe 2015). Three studies made no attempt to hide allocation concealment and were at high risk of selection bias (Ryding 1998; Ryding 2004; Taghizadeh 2008).

### Blinding of participants and personnel

This was not assessed as a double-blind methodology for studies of psychological treatment is impossible as it is clear to participants what treatment they are receiving.



#### Blinding of outcome assessors

Seventeen studies blinded outcome assessors and were at low risk of detection bias (Als 2015; Cox 2018a; Curtis 2016; Gidron 2001; Gidron 2007; Holmes 2007; Irvine 2011; Kazak 2005; Marchand 2006; Mouthaan 2013; Rothbaum 2012; Ryding 1998; Ryding 2004; Taghizadeh 2008; Wang 2015; Wijesinghe 2015; Zatzick 2001). Seven studies did not report this (Andre 1997; Borghini 2014; Brunet 2013; Gamble 2005; Gamble 2010; Jensen 2016; Lindwall 2014), and three studies were at high risk of detection bias (Biggs 2016; Brom 1993; Jones 2010).

#### Incomplete outcome data

Ten studies fully reported loss to follow-up and adequately dealt with missing outcome data (Brunet 2013; Cox 2018a; Gamble 2005; Irvine 2011; Jones 2010; Marchand 2006; Mouthaan 2013; Ryding 1998; Wang 2015; Zatzick 2001). Gamble 2005 included followup data from all participants (one participant in the intervention group could not be contacted at initial follow-up). Marchand 2006 and Zatzick 2001 included withdrawals in analysis by estimation of outcome by the method of 'last observation carried forward'. Thirteen studies did not adequately report missing outcome data and loss to follow-up and were at high risk of bias (Als 2015; Andre 1997; Biggs 2016; Borghini 2014; Brom 1993; Curtis 2016; Gamble 2010; Gidron 2001; Gidron 2007; Holmes 2007; Jensen 2016; Kazak 2005; Lindwall 2014). Four studies provided data only for treatment completers and recorded withdrawals without reasons by group or the number of withdrawals was not specified and were at unclear risk of bias (Rothbaum 2012; Ryding 2004; Taghizadeh 2008; Wijesinghe 2015).

### **Selective reporting**

Seven studies published protocols and reported on prespecified outcomes and were at low risk of reporting bias (Als 2015; Cox 2018a; Curtis 2016; Gamble 2010; Irvine 2011; Jensen 2016; Mouthaan 2013). Fourteen studies did not publish protocols and therefore it was unclear if they were free from reporting bias (Borghini 2014; Brom 1993; Brunet 2013; Gamble 2005; Gidron 2001; Holmes 2007; Jones 2010; Kazak 2005; Marchand 2006; Rothbaum 2012; Ryding 1998; Ryding 2004; Taghizadeh 2008; Zatzick 2001). One study did not report PTSD diagnosis as a primary outcome and was at high risk of reporting bias (Gidron 2007). Five other studies with null findings were at high risk of selective reporting because they did not provide data that could be used in meta-analysis (Andre 1997; Biggs 2016; Lindwall 2014; Wang 2015; Wijesinghe 2015).

### Other bias

Three studies were at low risk of other bias (Andre 1997; Lindwall 2014; Taghizadeh 2008). Nine studies were at unclear risk of other bias (Borghini 2014; Curtis 2016; Gamble 2005; Holmes 2007; Irvine 2011; Marchand 2006; Ryding 1998; Ryding 2004; Wijesinghe 2015). Fifteen studies were at high risk of other bias (Als 2015; Biggs 2016; Brom 1993; Brunet 2013; Cox 2018a; Gamble 2010; Gidron 2001; Gidron 2007; Jensen 2016; Jones 2010; Kazak 2005; Mouthaan 2013; Rothbaum 2012; Wang 2015; Zatzick 2001). Reasons for studies being judged at high risk of other bias include small sample size (Als 2015; Gidron 2001; Gidron 2007; Kazak 2005; Wang 2015; Zatzick 2001), inadequate description of the intervention (Biggs 2016), evaluation of treatment adherence not reported (Biggs 2016; Gidron 2001; Gidron 2007; Jensen 2016), differences in treatment

groups at baseline (Brom 1993; Gamble 2010), study authors affiliated with intervention (Brunet 2013; Gidron 2001; Gidron 2007; Mouthaan 2013; Rothbaum 2012), higher-than expected attrition rate (Cox 2018a), PTSD diagnosis based on assessor administration of PDS (Jones 2010) and intervention not manualised (Wang 2015; Zatzick 2001).

### **Effects of interventions**

See: Summary of findings for the main comparison Any early psychological intervention compared to waiting list/usual care for the prevention of post-traumatic stress disorder; Summary of findings 2 Any early psychological intervention compared to active control condition for the prevention of post-traumatic stress disorder

The meta-analyses included 21 studies with 2721 participants. Results were reported for all available outcome measures specified in the methodology. None of the studies identified reported data on adverse effects or use of health-related resources. Six studies could not be used in the meta-analysis as they reported no useable data (Andre 1997; Biggs 2016; Lindwall 2014; Taghizadeh 2008; Wang 2015; Wijesinghe 2015).

### Comparison 1: any intervention versus waiting list/usual care

Twenty-one studies compared a psychological intervention against a waiting list or treatment as usual condition (Als 2015; Andre 1997; Biggs 2016; Borghini 2014; Brom 1993; Brunet 2013; Curtis 2016; Gamble 2005; Holmes 2007; Irvine 2011; Jensen 2016; Jones 2010; Kazak 2005; Marchand 2006; Mouthaan 2013; Rothbaum 2012; Ryding 1998; Ryding 2004; Taghizadeh 2008; Wang 2015; Zatzick 2001).

### **Primary outcome**

### 1. PTSD diagnosis

Six studies used a clinician-administered scale to measure PTSD diagnosis and were entered into the meta-analyses (Brunet 2013; Gamble 2005; Jones 2010; Marchand 2006; Mouthaan 2013; Rothbaum 2012). The remaining 15 studies used a self-reported measure and were not included in the meta-analyses (Als 2015; Andre 1997; Biggs 2016; Borghini 2014; Brom 1993; Curtis 2016; Holmes 2007; Irvine 2011; Jensen 2016; Kazak 2005; Ryding 1998; Ryding 2004; Taghizadeh 2008; Wang 2015; Zatzick 2001).

### **Post-treatment**

Five studies provided data on clinician-administered diagnosis of PTSD (Brunet 2013; Gamble 2005; Marchand 2006; Mouthaan 2013; Rothbaum 2012). Post-treatment there was a lack of evidence for difference between intervention and control conditions (RR 1.06, 95% CI 0.85 to 1.32;  $I^2 = 0\%$ ; studies = 5; participants = 556; very low-certainty evidence; Analysis 1.1).

When analysed by type of psychological intervention, there was a lack of evidence for difference between treatment as usual and brief individual trauma processing therapy (RR 1.10, 95% CI 0.87 to 1.40;  $I^2 = 0\%$ ; studies = 3; participants = 262). There was uncertainty in estimating differences between treatment as usual and self-guided Internet-based interventions (RR 0.91, 95% CI 0.40 to 2.06; studies = 1; participants = 228), brief dyadic CBT (RR 0.59, 95% CI 0.21 to 1.61; studies = 1; participants = 66) reflected by the very wide confidence intervals.



#### Three to six months' follow-up

Five studies provided data on clinician-administered diagnosis of PTSD at three to six months' follow-up (Gamble 2005; Jones 2010; Marchand 2006; Mouthaan 2013; Rothbaum 2012). Multiple session early psychological interventions were associated with a reduction in PTSD symptoms compared to treatment as usual (RR 0.62, 95% CI 0.41 to 0.93; I<sup>2</sup> = 34%; studies = 5; participants = 758; low-certainty evidence; Analysis 1.2).

When analysed by type of intervention, meta-analysis of three studies identified uncertainty for the difference between treatment as usual and brief individual trauma processing therapy reflected in the wide confidence interval (RR 0.73, 95% CI 0.44 to 1.22; I² = 41%; studies = 3; participants = 251). There was considerable imprecision in estimating the difference between self-guided Internet-based interventions and treatment as usual (RR 0.50, 95% CI 0.18 to 1.45; studies = 1; participants = 185). Evidence from one study of 322 participants showed that intensive care diaries were more effective than delayed access to diaries (RR 0.38, 95% CI 0.17 to 0.82).

#### Seven to 12 months' follow-up

One study measured clinician-administered PTSD at seven to 12 months' follow-up (Mouthaan 2013). There was very high imprecision in estimating differences between self-guided Internet-based intervention and treatment as usual (RR 0.94, 95% CI 0.20 to 4.49; studies = 1; participants = 132; very low-certainty evidence; Analysis 1.3).

#### 2. Dropout from treatment

Eleven studies provided data on the number of participants who left the study early (Brom 1993; Brunet 2013; Gamble 2005; Holmes 2007; Irvine 2011; Kazak 2005; Marchand 2006; Rothbaum 2012; Ryding 1998; Ryding 2004; Zatzick 2001). Meta-analysis indicated that there was no significant difference in dropouts (RR 1.34, 95% CI 0.91 to 1.95; I<sup>2</sup> = 34%; studies = 11; participants = 1154; low-certainty evidence; Analysis 1.4).

When analysed by type of intervention, evidence from one study indicated brief IPT was associated with a higher dropout rate than TAU (RR 3.06, 95% CI 1.39 to 6.75; participants = 90). There was no significant difference, and wide confidence intervals, in the dropout rates between treatment as usual and brief individual trauma processing therapy (RR 1.06, 95% CI 0.67 to 1.68; I² = 26%; studies = 5; participants = 571), brief dyadic therapy (RR 2.09, 95% CI 0.76 to 5.75; I² = 0%; studies = 2; participants = 112), group therapy (RR 0.72, 95% CI 0.27 to 1.89; studies = 1; participants = 162), collaborative care (RR 0.75, 95% CI 0.14 to 3.94; studies = 1; participants = 34), or telephone-based CBT (RR 2.42, 95% CI 0.79 to 7.44; studies = 1; participants = 185).

### Secondary outcomes

### 1. Severity of PTSD symptoms

### **Post-treatment**

Nine studies provided data on the severity of PTSD symptoms post-treatment (Borghini 2014; Brom 1993; Gamble 2005; Jones 2010; Marchand 2006; Mouthaan 2013; Rothbaum 2012; Ryding 2004; Zatzick 2001). Meta-analysis showed no evidence of a difference between multiple session early psychological interventions and treatment as usual (SMD -0.09, 95% CI -0.29 to 0.12; studies =

9; participants = 1326; Analysis 1.5). There was a high degree of heterogeneity in this result ( $I^2 = 67\%$ ).

When analysed by type of intervention, evidence from one study of 300 participants indicated that self-guided Internet-based CBT may be more effective than treatment as usual (SMD -0.38,95% CI -0.61 to -0.15). Another study of 330 participants showed that intensive care diaries may be more effective than delayed access to diaries in reducing severity of PTSD symptoms postintervention (SMD -0.22,95% CI -0.44 to -0.01). There was very high imprecision (wide confidence intervals) in estimating differences between treatment as usual and brief individual trauma processing therapy (SMD 0.04,95% CI -0.34 to 0.425; I² = 76%; studies = 4; participants = 46), group counselling (SMD -0.09,95% CI -0.41 to 0.24; studies = 1; participants = 147), collaborative care (SMD -0.50,95% CI -1.24 to 0.25; studies = 1; participants = 29), three-step early intervention (SMD 0.33,95% CI -0.20 to 0.86; studies = 1; participants = 55).

It is evident that the heterogeneity in the group of participants who received brief individual trauma processing therapy contributed to the overall heterogeneity observed. It was not possible to investigate heterogeneity by number of sessions of the intervention, time between trauma exposure and intervention or type of traumatic event. However, it was possible to undertake a sensitivity analysis to look into gender of participants. Of the four studies on brief individual trauma processing therapy, one was done in women only (Gamble 2005). However, removing this study from the meta-analysis had little effect on the heterogeneity within this group ( $I^2 = 82\%$ ). In the subgroup analysis, there was no evidence (X2=0.89, P=0.35) of difference between studies of brief individual trauma processing therapy targeting women only (SMD -0.19, 95% CI -0.58 to 0.20; studies = 1; participants = 102) and studies where gender was mixed (SMD 0.12, 95% CI -0.39 to 0.62; studies = 3; participants = 363). But the small number of studies limits the ability to identify differences between subgroups.

We planned to perform a sensitivity analysis excluding studies with unclear allocation concealment, high levels of post-randomisation losses (greater than 40%) and unblinded outcome assessment or blinding of outcome assessment uncertain. However, this was not possible as three studies were at unclear risk of selection bias (Brom 1993; Gamble 2005; Marchand 2006), two studies were at high risk of attrition bias (Brom 1993; Rothbaum 2012), and two studies failed to blind or failed to report the blinding of outcome assessors (Brom 1993; Gamble 2005).

One study measured severity of PTSD symptoms using the PCL-17 (Biggs 2016). Data were not in a usable format for meta-analysis. Authors reported that the severity of PTSD symptoms did not differ significantly between the intervention and control group at one or two months' post-treatment.

### Three to six months' follow-up

Fifteen studies provided data on the severity of PTSD symptoms at three to six months' follow-up (Als 2015; Borghini 2014; Brom 1993; Brunet 2013; Curtis 2016; Gamble 2005; Holmes 2007; Irvine 2011; Jensen 2016; Jones 2010; Kazak 2005; Marchand 2006; Mouthaan 2013; Rothbaum 2012; Zatzick 2001). There was no significant difference in severity of PTSD symptoms between multiple session early psychological interventions and treatment as usual (SMD – 0.10,95% CI –0.22 to 0.02;  $I^2=34\%$ ; studies = 15; participants = 1921; low-certainty evidence; Analysis 1.6).



When analysed by type of intervention, evidence from one study (300 participants) indicated that self-guided Internet-based CBT may be more effective than treatment as usual (SMD –0.27, 95% CI –0.50 to –0.04). Two studies on a total of 103 participants showed that brief dyadic therapy may be more effective in reducing severity of PTSD symptoms three to six months' post-treatment (SMD –0.41, 95% CI –0.81 to –0.02). None of the other interventions (brief individual trauma processing therapy, collaborative care, brief IPT, intensive care diaries, three-step early intervention, telephone-based CBT, supported psychoeducation, communication facilitation in an ICU setting and nurse-led ICU recovery programme), showed any evidence of effectiveness over treatment as usual in the severity of PTSD symptoms three to six months' postintervention.

One study measured severity of PTSD at six months but could not be entered in the meta-analysis as authors did not report SDs (Andre 1997). The mean IES in the group receiving CBT decreased from 21.7 to 17.2, while the mean IES in the control group decreased from 15.6 to 12.0. However, data were only available for 39/65 participants in the intervention group and 45/67 in the control group.

One study measured severity of PTSD symptoms using the PCL-17 (Biggs 2016). Data were not in a usable format for meta-analysis. Authors reported that the severity of PTSD symptoms did not differ significantly between the intervention and control group at three or four months' post-treatment.

### Seven to 12 months' follow-up

Four studies provided data on the severity of PTSD symptoms at seven to 12 months' follow-up (Borghini 2014; Irvine 2011; Jensen 2016; Mouthaan 2013). There was no evidence of difference in severity of PTSD symptoms between multiple session early psychological interventions and treatment as usual (SMD -0.09, 95% CI -0.32 to 0.14; I<sup>2</sup> = 57%; studies = 4; participants = 765; Analysis 1.7).

It was not possible to investigate heterogeneity by number of sessions of the intervention, time between trauma exposure and intervention, or type of traumatic event. One study was conducted in women only, but excluding this from the meta-analysis had little effect on the heterogeneity (I<sup>2</sup> = 70%) (Borghini 2014). We planned to perform a sensitivity analysis excluding studies at high risk of bias. However, this was not possible as three of the four studies which measured this outcome at seven to 12 months were at unclear risk of selection bias (Irvine 2011), unclear risk of detection bias (Borghini 2014; Jensen 2016), and high risk of attrition bias (Borghini 2014; Jensen 2016).

When analysed by type of intervention, evidence from one study (185 participants) indicated that telephone-based CBT may be more effective than treatment as usual (SMD -0.38, 95% CI -0.67 to -0.09). For all other comparisons, confidence intervals were wide reflecting uncertainty in differences between treatment as usual and self-guided Internet-based CBT (SMD 0.00, 95% CI -0.23 to 0.23; studies = 1; participants = 300), three-step early intervention (SMD -0.18, 95% CI -0.71 to 0.35; studies = 1; participants = 55), or nurseled intensive care recovery programme (SMD 0.12, 95% CI -0.14 to 0.38; studies = 1; participants = 225).

The study by Wang 2015 measured severity of PTSD symptoms at 12 months' follow-up. Authors did not report SDs and, therefore,

it could not be used in a meta-analysis. There were no significant differences between group in terms of PTSD severity as measured by the CAPS score (P = 0.74) or the IES-R score (P = 0.68).

One study measured severity of PTSD symptoms using the PCL-17 (Biggs 2016). Data were not in a usable format for meta-analysis. Authors reported that the severity of PTSD symptoms did not differ significantly between the intervention and control group at seven or 10 months' post-treatment.

#### 2. Severity of depressive symptoms

#### Post-treatment

Five studies reported data on severity of depressive symptoms post-treatment (Holmes 2007; Mouthaan 2013; Rothbaum 2012; Ryding 2004; Zatzick 2001). Meta-analysis indicated a lack of evidence of difference between multiple session early psychological interventions and treatment as usual at postintervention (SMD –0.19, 95% CI –0.40 to 0.01; I<sup>2</sup> = 35%; studies = 5; participants = 671; Analysis 1.8).

When analysed by type of intervention, evidence from one study (137 participants) indicated that brief individual trauma processing therapy may be more effective than treatment as usual (SMD – 0.46, 95% CI –0.80 to –0.12). There was a lack of evidence of a difference between treatment as usual and self-guided Internet-based CBT (SMD –0.17, 95% CI –0.39 to 0.06; studies = 1; participants = 300), group therapy (SMD –0.22, 95% CI –0.55 to 0.10; studies = 1; participants = 147). There were very wide confidence intervals when comparing treatment as usual with collaborative care (SMD –0.26, 95% CI –0.99 to 0.48; studies = 1; participants = 29), or brief IPT (SMD 0.32, 95% CI –0.20 to 0.84; studies = 1; participants = 58).

One study measured severity of depression using the PHQ-9 (Biggs 2016). Data were not in a usable format for meta-analysis. Authors reported that the severity of PTSD symptoms did not differ significantly between the intervention and control group at one or two months' post-treatment.

### Three to six months' follow-up

Seven studies reported data on severity of depressive symptoms at three to six months' follow-up (Als 2015; Curtis 2016; Holmes 2007; Irvine 2011; Jensen 2016; Mouthaan 2013; Zatzick 2001). Meta-analysis indicated no difference between multiple session early psychological interventions and treatment as usual (SMD -0.04, 95% CI -0.19 to 0.10; I $^2$  = 6%; studies = 7; participants = 1009; low-certainty evidence; Analysis 1.9).

When analysed by type of intervention, there was no statistically significant difference between any specific intervention and treatment as usual.

One study measured depression at six months (Andre 1997). However, SDs were not reported and the study could not be entered in the meta-analysis. Authors reported that the mean change in HAD score was 3.2 in the intervention group and 3.3 in the control group. There was a large loss to follow-up in this study; data were only available for 39/65 participants in the intervention group and 45/67 in the control group.

One study measured severity of depression using the PHQ-9 (Biggs 2016). Data were not in a usable format for meta-analysis. Authors reported that the severity of PTSD symptoms did not differ



significantly between the intervention and control groups at three or four months' post-treatment.

#### Seven to 12 months' follow-up

Three studies reported data on severity of depressive symptoms at seven to 12 months' follow-up (Irvine 2011; Jensen 2016; Mouthaan 2013). Meta-analysis indicated no evidence of difference between multiple session early psychological interventions and treatment as usual (SMD 0.01, 95% CI -0.14 to 0.15;  $I^2 = 0\%$ ; studies = 3; participants = 745; Analysis 1.10).

When analysed by type of intervention, there was no difference between any specific intervention and treatment as usual.

One study measured severity of depression at 12 months' follow-up (Wang 2015). Authors did not report SDs and, therefore, it could not be used in a meta-analysis. The mean change in depression according to HADS was 3.29 in the treatment and 3.15 in the control group (P = 0.64).

One study measured severity of depression using the PHQ-9 (Biggs 2016). Data were not in a usable format for meta-analysis. Authors reported that the severity of PTSD symptoms did not differ significantly between the intervention and control group at seven or 10 months' post-treatment.

#### 3. Severity of anxiety symptoms

#### Post-treatment

Three studies provided data on the severity of anxiety symptoms post-treatment (Holmes 2007; Kazak 2005; Mouthaan 2013). Meta-analysis indicated no statistically significant difference between groups in the severity of anxiety symptoms postintervention (SMD -0.41, 95% CI -0.98 to 0.16; I<sup>2</sup> = 65%; studies = 3; participants = 358; Analysis 1.11).

It was not possible to investigate heterogeneity by number of sessions of the intervention, time between trauma exposure and intervention, type of traumatic event or gender of participants. We planned to perform a sensitivity analysis excluding studies at high risk of bias but this was not possible as two studies were at high risk of attrition bias (Holmes 2007; Kazak 2005), and two were at high risk of other bias (Kazak 2005; Mouthaan 2013).

When analysed by type of intervention, there was no statistically significant difference between treatment as usual and self-guided Internet-based therapy (SMD –0.03, 95% CI –0.25 to 0.20;  $I^2$  = 0%; studies = 1; participants = 300). Although compared with treatment as usual there was evidence of potential benefit for brief dyadic therapy (SMD –0.76, 95% CI –1.54 to 0.02; studies = 1; participants = 29) or brief IPT (SMD –0.76, 95% CI –1.54 to 0.02; studies = 1; participants = 29) confidence intervals were too wide to rule out no difference.

### Three to six months' follow-up

Six studies provided data on the severity of anxiety symptoms at three to six months' follow-up (Als 2015; Curtis 2016; Irvine 2011; Jensen 2016; Kazak 2005; Mouthaan 2013). Meta-analysis indicated no difference between multiple session early psychological interventions and treatment as usual (SMD -0.05, 95% CI -0.19 to 0.10; I $^2$  = 2%; studies = 6; participants = 945; low-certainty evidence; Analysis 1.12).

When analysed by type of intervention, there was no significant difference between any multiple session early psychological treatment and treatment as usual.

One study measured anxiety at six months (Andre 1997). However, SDs were not reported and the study could not be entered in the meta-analysis. Authors reported that the mean change in HAD score was 6.2 in the intervention group and 6.8 in the control group. There was a large loss to follow-up in this study; data were only available for 39/65 participants in the intervention group and 45/67 in the control group.

#### Seven to 12 months' follow-up

Three studies provided data on the severity of anxiety symptoms at seven to 12 months' follow-up (Irvine 2011; Jensen 2016; Mouthaan 2013). Meta-analysis indicated no statistically significant difference between multiple session early psychological interventions and treatment as usual (SMD -0.04, 95% CI -0.27 to 0.18; I<sup>2</sup> = 58%; studies = 3; participants = 746; Analysis 1.13).

It was not possible to investigate heterogeneity by number of sessions of the intervention, time between trauma exposure and intervention, type of traumatic event or gender of participants. We planned to perform a sensitivity analysis excluding studies at high risk of bias. However, this was not possible as two of the three studies which measured this outcome at seven to 12 months, were at unclear risk of selection bias (Irvine 2011), unclear risk of detection bias (Jensen 2016), and high risk of attrition bias (Jensen 2016).

When analysed by type of intervention, evidence from one study of 185 participants indicated that telephone-based CBT may be more effective than treatment as usual (SMD -0.30, 95% CI -0.59 to -0.01). It was unclear if self-guided Internet-based CBT (SMD 0.09, 95% CI -0.13 to 0.32; studies = 1; participants = 300), or nurse-led intensive care diaries (SMD 0.04, 95% CI -0.21 to 0.28; studies = 1; participants = 261) were more or less effective than treatment as usual.

One study measured severity of anxiety at 12 months' follow-up (Wang 2015). Authors did not report SDs and, therefore, it could not be used in a meta-analysis. The mean change in anxiety according to HADS was 4.10 in the treatment group and 3.85 in the control group (P = 0.36).

### 4. Adverse effects

No studies provided data on adverse effects.

### 5. General functioning

One study measured quality of life using the WHOQOL-BREF (Biggs 2016). Data were not in a usable format for meta-analysis. Authors reported that the severity of PTSD symptoms did not differ significantly between the intervention and control groups at one, two, four, seven or 10 months' post-treatment.

### 6. Use of health-related resources

No studies provided data on use of health-related resources.

## Comparison 2: any intervention versus other psychological intervention

Six studies compared a psychological intervention against another active intervention (Cox 2018a; Gamble 2010; Gidron 2001; Gidron



2007; Lindwall 2014; Wijesinghe 2015). However, two of the studies provided no useable data and, therefore, could not be included in the meta-analyses (Lindwall 2014; Wijesinghe 2015).

### **Primary outcome**

#### 1. PTSD diagnosis

One study measured PTSD diagnosis using the Post-traumatic Stress Symptom Scale. As this is a self-reported rather than clinician-administered tool, this study was not included in the meta-analysis (Wijesinghe 2015).

#### **Post-treatment**

No study measured PTSD diagnosis post-treatment.

#### Three to six months' follow-up

No study measured PTSD diagnosis post-treatment using a clinician-administered tool.

#### Seven to 12 months' follow-up

No study measured PTSD diagnosis at seven to 12 months' followup.

### 2. Dropouts from treatment

Two studies provided data on number of dropouts for any reason, showing participants in early intervention groups were more likely to drop out than active controls (RR 1.61, 95% CI 1.11 to 2.34; I $^2$  = 0%; studies = 2; participants = 425; low-certainty evidence; Analysis 2.1) (Cox 2018a; Gamble 2010). When analysed by type of intervention, there may be higher drop out for brief individual trauma processing therapy compared with parenting support (RR 1.78, 95% CI 1.12 to 2.84; studies = 1; participants = 262). It was uncertain if there was a difference in drop out between guided self-help and physical education (RR 1.33, 95% CI 0.71 to 2.50; studies = 1; participants = 163).

### Secondary outcomes

### 1. Severity of PTSD symptoms

Five studies provided data on severity of PTSD symptoms (Cox 2018a; Gamble 2010; Gidron 2001; Gidron 2007; Lindwall 2014), but only four could be entered in a meta-analysis (Cox 2018a; Gamble 2010; Gidron 2001; Gidron 2007).

### Post-treatment

Two studies provided data on severity of PTSD symptoms post-treatment (Cox 2018a; Gamble 2010). Meta-analysis showed a lack of evidence for difference between treatments (SMD 0.13, 95% CI -0.06 to 0.33; I<sup>2</sup> = 0%; studies = 2; participants = 392; Analysis 2.2). When analysed by type of intervention, there was imprecision (wide confidence intervals) in estimating differences between brief individual trauma processing therapy and parenting support (SMD 0.14, 95% CI -0.13 to 0.41; studies = 1; participants = 217), or guided self-help and physical education (SMD 0.13, 95% CI -0.17 to 0.42; studies = 1; participants = 175).

### Three to six months' follow-up

Four studies provided data on severity of PTSD symptoms at three to six months' follow-up (Cox 2018a; Gamble 2010; Gidron 2001; Gidron 2007). There was a wide confidence interval indicating uncertainty on differences between treatments (SMD –0.02, 95% CI

-0.31 to 0.26;  $I^2 = 43\%$ ; studies = 4; participants = 465; low-certainty evidence; Analysis 2.3).

It was not possible to investigate heterogeneity by number of sessions of the intervention, time between trauma exposure and intervention, type of traumatic event or gender of participants. We planned to perform a sensitivity analysis excluding studies at high risk of bias. However, this was not possible as three of the four studies which measured this outcome at three to six months were at unclear risk of selection bias (Gamble 2010; Gidron 2001; Gidron 2007), unclear risk of detection bias (Gamble 2010), and high risk of attrition bias (Gamble 2010; Gidron 2007).

When analysed by type of intervention, confidence intervals were wide reflecting a lack of certainty on differences between brief individual trauma processing therapy and supportive listening (SMD -0.54, 95% CI -1.42 to 0.34; I<sup>2</sup> = 52%; studies = 2; participants = 51), brief individual trauma processing therapy and parenting support (SMD 0.06, 95% CI -0.19 to 0.31; studies = 1; participants = 239), or guided self-help and physical education (SMD 0.13, 95% CI -0.16 to 0.43; studies = 1; participants = 175).

One study measured severity of PTSD symptoms in parents of children undergoing stem cell transplantation (Lindwall 2014). A child-targeted intervention (massage and humour therapy) was compared with a child-targeted plus a parent-targeted intervention (massage, relaxation, imagery). Data could not be included in the meta-analysis. Authors reported no significant differences between interventions (P = 0.45), but that PTSD did decrease significantly from baseline to six months for the two intervention groups as a whole (P < 0.001).

### Seven to 12 months' follow-up

One study provided data on severity of PTSD symptoms at seven to 12 months' follow-up (Gamble 2010). There was no evidence of a significant difference between brief individual trauma processing therapy and parenting support (MD 1.27, 95% CI –0.60 to 3.14; studies = 1; participants = 200; Analysis 2.4),

### 2. Severity of depressive symptoms

Three studies measured severity of depressive symptoms (Cox 2018a; Gamble 2010; Lindwall 2014), but only two provided data that could be used in a meta-analysis (Cox 2018a; Gamble 2010).

### **Post-treatment**

Two studies provided data on severity of depressive symptoms post-treatment (Cox 2018a; Gamble 2010). There was no evidence of a significant difference between treatments (SMD 0.12, 95% CI -0.08 to 0.32; I<sup>2</sup> = 0%; studies = 2; participants = 392; Analysis 2.5). When analysed by type of intervention, there was a wide confidence interval when comparing brief individual trauma processing therapy and parenting support (SMD 0.06, 95% CI -0.21 to 0.33; studies = 1; participants = 217) suggesting a lack of certainty on group differences. Although the direction of effect favoured physical education over guided self help the difference wasn't statistically significant (SMD 0.19, 95% CI -0.11 to 0.49; studies = 1; participants = 175).

### Three to six months' follow-up

Two studies provided data on severity of depressive symptoms at three to six months' follow-up (Cox 2018a; Gamble 2010). There was



no evidence of a difference between treatments (SMD 0.04, 95% CI -0.16 to 0.23; I<sup>2</sup> = 0%; studies = 2; participants = 409; low-certainty evidence; Analysis 2.6). When analysed by type of intervention, there was no evidence of a difference between brief individual trauma processing therapy and parenting support (SMD 0.09, 95% CI -0.17 to 0.34; studies = 1; participants = 234) or guided self-help and physical education (SMD -0.04, 95% CI -0.33 to 0.26; studies = 1; participants = 175).

One study measured depression in parents of children undergoing stem cell transplantation using the CES-D scale (Lindwall 2014). Data could not be entered into a meta-analysis but authors reported no significant differences between groups a child-targeted intervention (massage and humour therapy) with child-targeted plus a parent-targeted (massage, relaxation, imagery) intervention (P = 0.63).

#### Seven to 12 months' follow-up

One study provided data on severity of depressive symptoms at seven to 12 months' follow-up (Gamble 2010). There was no evidence of a difference between brief individual trauma processing therapy and parenting support (MD 0.79, 95% CI –0.66 to 2.24; studies = 1; participants = 198; Analysis 2.7).

#### 3. Severity of anxiety symptoms

#### Post-treatment

Two studies provided data on severity of anxiety symptoms post-treatment (Cox 2018a; Gamble 2010). There was no evidence of a significant difference between treatments (SMD 0.08, 95% CI – 0.12 to 0.28;  $I^2$  = 0%; studies = 2; participants = 392; Analysis 2.8). When analysed by type of intervention, there was no evidence of a significant difference between brief individual trauma processing therapy and parenting support (SMD 0.10, 95% CI –0.17 to 0.37; studies = 1; participants = 217) or guided self-help and physical education (SMD 0.05, 95% CI –0.24 to 0.35; studies = 1; participants = 175).

### Three to six months' follow-up

Two studies provided data on severity of anxiety symptoms at three to six months' follow-up (Cox 2018a; Gamble 2010). There was no evidence of a difference between treatments (SMD 0.00, 95% CI – 0.19 to 0.19;  $I^2$  = 0%; studies = 2; participants = 414; low-certainty evidence; Analysis 2.9). When analysed by type of intervention, there was no evidence of a difference between brief individual trauma processing therapy and parenting support (SMD 0.03, 95% CI –0.22 to 0.28; studies = 1; participants = 239) or guided self-help

and physical education (SMD -0.04, 95% CI -0.33 to 0.26; studies = 1; participants = 175).

#### Seven to 12 months' follow-up

One study provided data on severity of anxiety symptoms at seven to 12 months' follow-up (Gamble 2010). There was a very wide confidence interval when comparing brief individual trauma processing therapy with parenting support (MD -0.07, 95% CI -1.58 to 1.44; studies = 1; participants = 199; Analysis 2.10).

#### 4. Adverse effects

No studies provided data on adverse effects. One study reported interactions between initial traumatic stress symptoms, psychiatric history, past psychiatric diagnosis and IPT intervention in predicting worse PTSD symptoms at six months (Holmes 2007).

#### 5. General functioning

#### Post-treatment

One study measured general functioning using the HRQoL (Gamble 2010). There was no difference between counselling aimed at promoting resilience and parenting support (MD -0.02, 95% CI -0.05 to 0.01; studies = 1; participants = 218; Analysis 2.11).

#### Three to six months' follow-up

One study measured general functioning at three to six months' post-treatment (Gamble 2010). There was no significant difference between counselling aimed at promoting resilience and parenting support (MD -0.03, 95% CI -0.06 to 0.00; studies = 1; participants = 239; low-certainty evidence; Analysis 2.12).

### Seven to 12 months' follow-up

One study found no difference in general functioning at seven to 12 months' post-treatment (MD 0.01, 95% CI -0.03 to 0.05; studies = 1; participants = 199; Analysis 2.13) (Gamble 2010).

### 6. Use of health-related resources

No studies provided data on use of health-related resources.

### **Publication bias**

All of the studies identified for this review were published, apart from one (Gamble 2010). Only two comparisons were sufficiently large for us to be able to conduct funnel plots (Analysis 1.6; Analysis 1.4). Visual inspection of these plots suggested that there was no indication of asymmetry (Figure 4; Figure 5). We were unable to investigate the plots by intervention.



Figure 4. Funnel plot of comparison: 1 Any early psychological intervention versus waiting list/usual care, outcome: 1.5 Severity of PTSD symptoms: 3-6 months. TAU: treatment as usual.

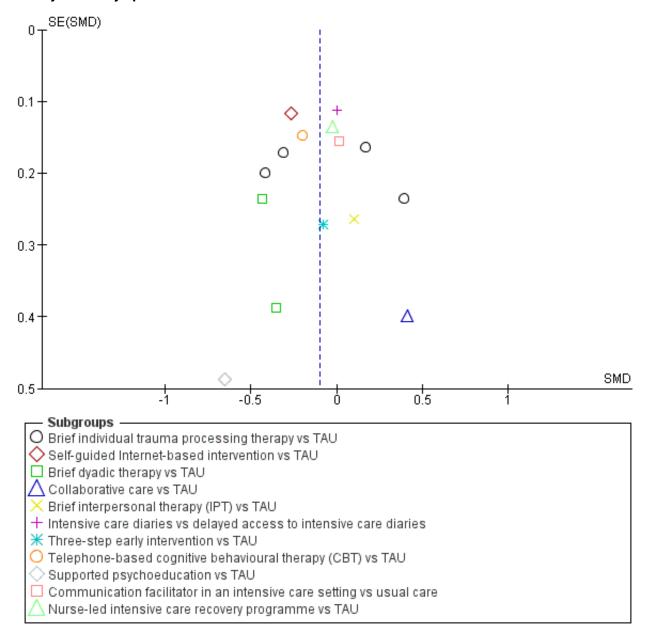
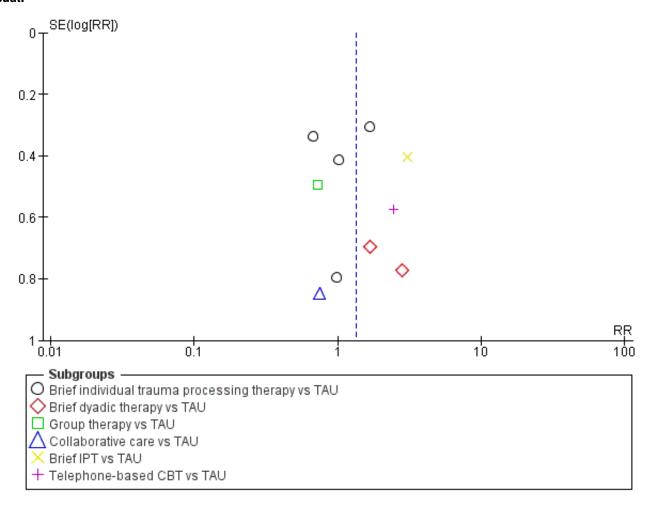




Figure 5. Funnel plot of comparison: 1 Any early psychological intervention versus waiting list/usual care, outcome: 1.13 Dropouts for any reason. CBT: cognitive behavioural therapy; IPT: interpersonal therapy; TAU: treatment as usual.



### DISCUSSION

### **Summary of main results**

We identified 27 RCTs (3963 participants) of early multiple session psychological interventions starting within three months of a traumatic event that were designed to prevent traumatic stress symptoms. Twenty-one studies of 2721 participants provided usable data and were included in a meta-analysis.

Multiple session early psychological interventions may be more effective than treatment as usual in reducing the number of people diagnosed with PTSD three to six months after receiving the intervention. However, there was no significant difference between the two groups immediately post-treatment or at seven to 12 months' follow-up. Evidence was not strong enough to suggest that early psychological interventions may be more effective than treatment as usual in reducing severity of PTSD symptoms, depression and anxiety at any time point. Differences in dropouts were not significant. In terms of specific intervention types, intensive care diaries may be more effective than treatment as usual in reducing the number of people diagnosed with PTSD based on data from one study (Jones 2010), but we found no difference

in terms of PTSD severity; self-guided Internet-based intervention may be more effective than treatment as usual at reducing PTSD severity based on data from one study (Mouthaan 2013); brief dyadic therapy may be more effective than treatment as usual at reducing PTSD severity based on data from two studies (Brunet 2013; Mouthaan 2013). We found no evidence of effect for any other intervention.

There were no significant differences between multiple session early psychological interventions and active control interventions in terms of PTSD diagnosis, reducing the severity of depression and anxiety, or improving general functioning of the recipient either postintervention or at three to 12 months' follow-up. However, participants who received multiple session interventions were more likely to stop the intervention early.

### Overall completeness and applicability of evidence

The studies included in this review directly addressed the primary review question. It was possible to perform meta-analyses of RCTs of multiple session psychological interventions aimed at preventing PTSD in individuals who had been exposed to a traumatic event but had not been identified as experiencing any



specific psychological difficulties. However, meta-analyses could not be performed for all outcomes and the limited number of studies, their small sample sizes and heterogeneity (see below) complicated interpretation. The question of how interventions fare against another psychological interventions could only be answered for memory structuring and not for other interventions. The number of new studies identified in our most recent search (see Characteristics of studies awaiting classification table) suggests that this is a quickly moving field, particularly in terms of early psychological intervention in medical settings and that a further updated review will be required in the near future.

Given the prominence of cognitive behavioural interventions in the evidence base for psychological therapies it is surprising that only three of the studies identified in this review evaluated a CBT intervention (Andre 1997; Irvine 2011; Rothbaum 2012), although another seven studies included CBT components in their interventions (Brunet 2013; Cox 2018a; Kazak 2005; Marchand 2006; Mouthaan 2013; Wijesinghe 2015; Zatzick 2001). Andre 1997 reported a significant decrease in intrusive symptoms and anxiety in their treatment group. Unfortunately, data provided in their paper did not permit inclusion in a meta-analysis. Brief individual trauma processing therapies which were based on several different theoretical models but shared active treatment components were the most frequently evaluated interventions. The evidence reviewed did not provide any support for this approach as a preventive intervention.

Participants in the studies included in the review were exposed to single individual traumatic events and there were no studies of larger-scale traumatic events such as disasters or wars, which limits the external validity of the results across the full range of traumatic events.

Unfortunately, other than the studies where participants were critically ill and at high risk of mortality (Irvine 2011; Jensen 2016; Jones 2010), only one study considered adverse effects (Holmes 2007), and it was unclear whether or not any adverse effects occurred in the other studies. The absence of tolerability assessment (an evaluation of the extent to which patients can endure the unwanted adverse effects of an intervention) is a key shortcoming in the RCTs identified and one that has previously been noted in psychological treatment studies of chronic PTSD (Bisson 2007).

### Quality of the evidence

### Heterogeneity

There was evidence of both clinical and statistical heterogeneity in the included studies. Although all the trials attempted to prevent PTSD symptoms, the nature of the interventions included in the meta-analyses was quite diverse. The interventions included in the primary meta-analysis were two sessions of brief dyadic CBT (Brunet 2013), two sessions of counselling (one face-to-face and one by telephone) (Gamble 2005), two sessions of adapted critical incident stress debriefing (Marchand 2006), sharing of an intensive care diary (Jones 2010), brief self-guided Internet-based intervention (Mouthaan 2013), and three sessions of modified prolonged exposure (Rothbaum 2012). It was very difficult to compare such trials and there did appear to be some differences in outcomes, for example Jones 2010 appeared to be more effective than Marchand 2006.

There were also differences in the clinical populations which included motor vehicle accident victims, victims of armed robbery or aggression, military mortuary attendants, people in ICUs, women who had emergency caesarean sections or had premature babies, parents of children who were in ICU or had recently been diagnosed with cancer. Unfortunately, the limited number of trials meant that sensitivity analyses could not be performed in a meaningful way to explore these issues further. Statistical heterogeneity was apparent in several analyses, the 12 value demonstrating inconsistencies in the outcomes of some trials that were grouped. When there was statistical heterogeneity, we used a random-effects model as opposed to a fixed-effect model to calculate more conservative CIs. We concluded that all trials were essentially trying to measure the same outcome and that it was worthwhile summarising their combined results, but the variation meant that caution should be applied when interpreting the results (Fletcher 2007).

### **Methodological certainty**

We used the five GRADE considerations (study limitations, consistency of effect, imprecision, indirectness and publication bias) to assess the certainty of the body of evidence for each outcome, and to draw conclusions about the certainty of evidence within the text of the review. One of the main considerations for downgrading GRADE judgements was risk of bias. Concerns over the certainty of the evidence also limit the extent to which conclusions can be generalised. For details of the risk of bias judgements for each study, see the Characteristics of included studies table and the graphical representation of risk of bias presented in Figure 2 and Figure 3.

There were several issues that were problematic in several studies including the randomisation process, incomplete reporting of dropouts and absence of a manualised, replicable specific treatment. As with all psychological treatment trials there are issues with the control groups. This is particularly important in early intervention research where a reduction in symptoms over the duration of the trial would be expected, given the natural course of traumatic stress reactions. The development of a psychological treatment placebo is very difficult, if not impossible, as is blinding of participants and therapists. Some of the wait list/usual care groups may have received some form of intervention by virtue of contact through symptom monitoring, but this was not properly evaluated and it is not possible to determine what, if any, impact on outcomes this would have had. The four studies that did have an active control group showed no difference between that and multiple session early psychological interventions (Cox 2018a; Gamble 2010; Gidron 2001; Gidron 2007).

The sample sizes of most of the studies were also an important limitation. However, the intervention and control groups in most studies appeared well matched at baseline, reducing the risk of the reported unadjusted mean outcomes being influenced by baseline differences. We had intended to undertake subgroup analyses investigating heterogeneity in terms of the type of trauma and the number of sessions of intervention, but there were not enough studies for us to do this.

### Potential biases in the review process

This review adhered strictly to the Cochrane guidelines (Higgins 2011). Two review authors independently screened the abstracts



identified by the literature search; read all potentially relevant studies; assessed each study against the inclusion criteria; extracted data from the written reports and rated each study for risk of bias. We discussed any disagreements with a third review author, and reached unanimous decisions for inclusion and classification. We carefully followed guidelines set out by Cochrane on statistical methods and used GRADE to assess the certainty of evidence (Higgins 2011). Following these procedures minimised the potential for bias, but some unavoidable issues remained.

All studies included in the review apart from Gamble 2010 were published, which led to the possibility of publication bias. We were unable to undertake investigate for possible publication bias for our primary outcome, but we were able to generate funnel plots for two comparisons and found no evidence to suggest that publication bias was indicated.

We systematically searched numerous online databases for potentially relevant studies, and scrutinised reference lists of included studies. We also contacted experts in the field requesting help to identify missed studies or ongoing work. Nonetheless, we could not fully eliminate the possibility of overlooked RCTs.

There was considerable statistical heterogeneity in four of the pooled comparisons. In circumstances where heterogeneity was thought to be potentially problematic, we used a random-effects model. Heterogeneity was also a factor that was taken into consideration in downgrading the certainty of the evidence with GRADE.

## Agreements and disagreements with other studies or reviews

To our knowledge, this is the only specific systematic review of multiple session interventions to prevent PTSD. The results are consistent with the findings of the previous version of this review (Roberts 2009). Our findings are also consistent with other systematic reviews that have included studies in this area (NICE 2018), and also with prevailing guidance regarding how best to respond following a traumatic event (ACPMH 2007; NICE 2018).

### **AUTHORS' CONCLUSIONS**

### Implications for practice

These results suggest that at this time there is very limited evidence to support the use of psychological intervention for routine use following traumatic events. We found some contradictory evidence for the use of intensive care diaries in intensive care units and some evidence to suggest the benefit of brief dyadic cognitive behavioural therapy (CBT)-based approaches and self-guided Internet-based intervention, but we did not feel that the available evidence was strong enough to recommend regular use of these interventions without further evaluation.

### Implications for research

The number of ongoing studies reflects that this a fast moving field of research. Future updates of this review will incorporate the

results of these studies currently underway. Further well-designed randomised controlled trials of interventions that appeared to show promise could be subjected to further evaluation (e.g. Brunet 2013; Jones 2010; Kazak 2005; Mouthaan 2013). However, it may be that these interventions would demonstrate greater effect when targeted at individuals who are symptomatic (Roberts 2010), given that many individuals will gradually recover without the need of any intervention (Bryant 2013). We note that preventive cognitive behavioural interventions have not been adequately investigated. However, the absence of effect for brief individual trauma processing therapies, which share some features of traumafocused cognitive behavioural therapies, suggests that models that are not targeted on those with traumatic stress symptoms may not be the most fruitful means of intervention delivery. Current evidence favours approaches that focus intervention on people who are symptomatic or have been diagnosed with acute stress disorder or acute post-traumatic stress disorder (PTSD) (Roberts 2010). Given the positive findings for brief dyadic therapy, it would be of interest to evaluate forms of familial and community intervention and interventions aimed at improving coping skills and enhancing positive and helpful behaviours (Ruzek 2007). Internet- and app-based interventions potentially offer relatively cheap and accessible ways of offering interventions to large numbers of people and should be investigated further. The findings of Mouthaan 2013 demonstrate some support for this approach and there are now a number of studies that have demonstrated the effectiveness of Internet-based CBT for PTSD (Lewis 2018). Future research should also explore the optimal time to intervene, consider adverse events and tolerability of treatment, and carefully control for additional intervention.

### ACKNOWLEDGEMENTS

We wish to express our thanks to Melissa Alderfer, Randall Curtis, Jessica Ford, Jenny Gamble, Elena Garralda, Janet Jensen, Christina Jones, Joanne Mouthaan, Elsa Lena Ryding and Doug Zatzick for providing additional unpublished data. We also thank Sarah Dawson for her help with searches. Finally, thanks to those experts who responded with additional suggestions and contacts during the search process.

The authors and the Cochrane Common Mental Disorders Editorial Team, are grateful to the following peer reviewers and editors for their time and comments: Bradley Belsher, Nick Meader, Karen Morley, and Barbara Rothbaum. They would also like to thank copy editor, Anne Lawson.

CRG Funding Acknowledgement: The National Institute for Health Research (NIHR) is the largest single funder of the Cochrane Common Mental Disorders Group.

Disclaimer: views and opinions expressed herein are those of the review authors and do not necessarily reflect those of the NIHR, National Health Service (NHS) or the Department of Health and Social Care.



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\* Indicates the major publication for the study



## CHARACTERISTICS OF STUDIES

## **Characteristics of included studies** [ordered by study ID]

## Als 2015

Methods	Design: RCT			
Participants	Setting: PICU			
	Type of trauma exposure: parents of children aged 4–16 years admitted to PICU.			
	Inclusion criteria: parents with a child aged 4–16 years admitted to the PICU for ≥ 12 hours.			
	<b>Exclusion criteria:</b> child death prior to discharge; discharge to palliative care; planned admissions; history of PICU admission; overseas address or insufficient English to complete study questionnaires.			
	Sample size: 442 individuals assessed for eligibility; 31 randomised.			
	Mean age: 41.1 (SD not reported) years where reported.			
	<b>Gender:</b> 5 (16.1%) men; 26 (83.9%) women			
	<b>Ethnicity:</b> 12 participants (38.7%) were identified as "white UK". The ethnicity of other participants was not reported.			
	Country: UK			
Interventions	Group 1: supported psychoeducational intervention: n = 22			
	2 phases: the first phase (receipt of the psychoeducational tool), was planned to occur within 7 days of discharge from hospital and the second phase (receipt of the telephone call), within 14 days of receiving the tool. The psychoeducational tool consisted of a handbook developed by mental health and paediatric experts and parents with experience of having a child in PICU. The handbook covered 3 main areas: emotional recovery, behavioural recovery and getting back to normal learning. The first section included a description of common emotional reactions in children, their siblings and parents following discharge from PICU, with advice regarding their management. It also included an outline of when recovery became stalled by the development of PTSD, its manifestations, what treatments are available and their rationale. The second section gave more detailed advice to parents about managing behavioural problems in children following hospital discharge. The third section addressed possible learning difficulties (e.g. slowed information processing, memory and attention problems) in the aftermath of the child's admission and provided guidance on how to support affected children. There was an additional section containing a list of contacts of possible sources of further support and advice. The telephone call, conducted by the researcher, was used to discuss each family's post-PICU experience, reinforce the material in the handbook (thus ensuring all families were exposed to the information), and support families in putting into practice the advice given, if appropriate. <b>Group 2: TAU:</b> n = 9			
Outcomes	PTSD: IES			
	Other: HADS; PSS-PICU			
	Follow-up: 3–6 months' postdischarge from PICU			
Notes	Primary aim was to evaluate the feasibility of the intervention and procedures to evaluate it through a larger trial.			
Risk of bias				



Als 2015 (Continued)		
Random sequence generation (selection bias)	Low risk	Participants were randomised to the intervention or TAU using a computer-generated list of random numbers prepared by an independent statistician.
Allocation concealment (selection bias)	Low risk	Allocation sequence was concealed from the researcher enrolling and assessing participants and was stored with an administrator who had no other involvement in the trial.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Baseline and 3- to 6-month follow-up questionnaires were posted to families and returned using stamped addressed envelopes.
Incomplete outcome data (attrition bias) All outcomes	High risk	Outcome data were assessed on an ITT basis and involved all parent–child pairs randomly assigned and providing follow-up data. However, outcome data were only available for 17 (77.8%) of the intervention group and 6 (66.6%) of the TAU group.
Selective reporting (reporting bias)	Low risk	A study protocol was published at the beginning of the trial. The reported plan was to assess changes in mental health outcomes from baseline to follow-up across both groups. However, the research group was unable to collect baseline data within the specified time frame, and they, therefore, focused solely on the 3- to 6-month outcome data. We did not judge this to constitute a significant reporting bias.
Other bias	High risk	Small sample size

## **Andre 1997**

Methods	Design: RCT		
Participants	Setting: community		
	Type of trauma exposure: victims of aggression		
	Inclusion criteria: bus drivers who were victims of aggression		
	Exclusion criteria: not reported		
	Sample size: 132 randomised: 65 to intervention, 67 to control		
	Mean age: 35.2 (SD not reported) years		
	Gender: not reported		
	Ethnicity: not reported		
	Country: France		
Interventions	Group 1:		
	CBT intervention with 1–6 sessions, 45–60 min each. Mean 2.3 sessions per participant		
	Group 2:		
	Standard care (medical/social care provided by company)		
Outcomes	PTSD: IES		
	Other: HAD anxiety; number of days off work		



#### Andre 1997 (Continued)

Follow-up: 6 months' postintervention

Notes

#### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Not reported
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Not reported
Incomplete outcome data	High risk	High attrition: intervention (40%), control (33%)
(attrition bias) All outcomes		Adherence to intervention not reported
Selective reporting (reporting bias)	High risk	No protocol identified. The study authors reported no statistically significant differences between intervention and control groups, which meant that it was not possible to include this study in meta-analyses. This has the potential to lead to an overestimate of effectiveness for those outcomes.
Other bias	Low risk	Appeared free from other sources of bias.

## **Biggs 2016**

Methods	Design: longitudinal RCT
Participants	Setting: participants were recruited from military MAs returning from deployment to the Middle East.
	Type of trauma exposure: military mortuary work
	Inclusion criteria: all returning MAs were offered the opportunity to take part.
	Exclusion criteria: none reported
	Sample size: 362 soldiers assessed for eligibility, 126 randomised
	Mean age: 28.1 (SD not reported) years
	<b>Gender:</b> 85 (67.5%) men; 41 (32.5%) women
	<b>Ethnicity:</b> 57.6% white, the ethnicity of other participants not reported.
	Country: USA
Interventions	Group 1: Troop Education for Army Morale (TEAM): n = 68
	Intervention based on the principles of PEA and included 4 x 2-hour interactive group sessions held a

Intervention based on the principles of PFA and included 4 × 2-hour interactive group sessions held approximately 2, 3, 4 and 7 months' postdeployment, informational handouts, access to a dedicated website, e-mail and telephone line service. 2 trained psychologists or psychiatrists facilitated intervention sessions.



<b>Bigg</b>	s 2016	(Continued)
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**Group 2: assessment only:** n = 58

Outcomes

PTSD: PCL

Other: PHQ-9 depression; WHOQOLBREF

Follow-up: 2, 3, 4, 7 and 10 months' postdeployment

Notes

#### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Not reported
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	High risk	Intervention group completed surveys before the start of each intervention session. We judged that this may have influenced responses.
Incomplete outcome data (attrition bias) All outcomes	High risk	The longitudinal effect of treatment on outcome measures was derived from linear mixed models using total score as the dependent variable and time (baseline and 5 follow-ups), treatment (intervention vs comparison), and the interaction between time and treatment as independent variables. The way in which missing data were handled was not reported and dropout was high.
Selective reporting (reporting bias)	High risk	No study protocol identified. The study authors reported no statistically significant differences between intervention and control groups, which meant that it was not possible to include this study in meta-analyses. This has the potential to lead to an overestimate of effectiveness for those outcomes.
Other bias	High risk	Intervention only described very briefly. No description of an intervention manual and evaluation of treatment adherence was not reported.

#### Borghini 2014

Methods	Design: RCT

Participants

**Setting:** participants recruited from a NICU.

Type of trauma exposure: mothers of preterm infants born at < 33 weeks' gestation.

**Inclusion criteria:** mothers of infants born at < 33 weeks' gestation.

**Exclusion criteria:** death of the neonate, parents who were not fluent in French, infant malformation, periventricular leukomalacia grade ≥ 2 or intraventricular haemorrhage, parental mental illness and neonatal abstinence syndrome. Participants were also excluded at 6 months' corrected age (6 months after 40 weeks' gestation), if an infant had severe mental or psychomotor delays during the paediatric appointment.

**Sample size:** 242 mothers eligible for inclusion; 62 met initial inclusion criteria, agreed to take part and were randomised; 2 infants developed severe mental or psychomotor delays during the paediatric appointment at 6 months and the mother was removed from the study, leaving a final sample of 60.



#### Borghini 2014 (Continued)

Mean age: 33 (SD 4.1) years Gender: 60 (100%) women

Ethnicity: not reported. 45/55 completing participants were reported to be of Swiss nationality.

Country: Switzerland

#### Interventions

#### **Group 1: 3-step early intervention:** n = 30

Aimed at 3 main dimensions: parental support, parent–infant relationship support and infant development support. Intervention had 3 phases occurring at 33 weeks after conception, 42 weeks after conception and 4 months after the theoretical term of 40 weeks after conception. At 33 weeks after conception, the intervention consisted of a joint observation, including the mother, a nurse and a therapist, of the infant's reactions and adjustments to various stimuli and his/her interaction abilities during a standard care procedure in the NICU. The observation lasted 30–60 min and was videotaped. Parents were encouraged to comment on their infants' behaviours and on their own emotions. At 42 weeks after conception, the intervention consisted of an assessment using the NBAS, which identified infants' stress reactions and self-regulation capacities. Each NBAS was videotaped. At the infant's 4-month corrected age, the intervention consisted of 3 sessions, 1 week apart, of a 10-min mother–infant free play. The interactions were videotaped and used later by the examiner in the interaction guidance phase with the mothers. Each session could last 40–60 min and aimed at promoting the mother's caregiving qualities and sensitivity to the infant through the careful observation of the infant's reactions, needs, competences and vulnerabilities.

## **Group 2: TAU:** n = 30

At 33 weeks' conception, families in the preterm group without intervention met in the NICU to obtain information about their infant and their relationship with him or her. There was no joint observation of the infant. At 42 weeks after conception the preterm group without intervention also had the NBAS assessment, but the NBAS was followed by a semi-structured interview with the parents based on the Clinical Interview for Parents of High-Risk Infants which allowed mothers to express much of their emotional experiences with their infants at birth and during their subsequent hospitalisation. At the infant's 4-month corrected age, the preterm group without intervention met for a single appointment. Mothers were asked to play freely with their infant for 10 min and the interaction was videotaped, but there was no interaction guidance afterwards.

#### Outcomes

PTSD: PPQ

**Other:** CARE-Index (3rd revision) was used to code the quality of video-recorded mother–infant interactions

Follow-up: 42 weeks' postconception, and 4 and 12 months' corrected infant birth.

#### Notes

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Not specified
Allocation concealment (selection bias)	Low risk	Randomised using a sealed envelope procedure.
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	PTSD outcomes assessed via a self-report measure which appeared to have been administered by the treating clinician.



Borghini 2014 (Continued)			
Incomplete outcome data (attrition bias) All outcomes	High risk	Analysis was based on completers. 4 (13.3%) participants dropped out of the intervention group; 1 (3.3%) dropped out of the TAU group. There was no explanation of the causes of dropout or a consort diagram. It was unclear how many participants contributed to each follow-up point.	
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.	
Other bias	Unclear risk	No indication that treatment adherence was assessed.	

## **Brom 1993**

Methods	Design: RCT		
Participants	Setting: Participants were recruited from police records of MVA and seen on an outpatient basis.		
	Type of trauma exposure: MVA		
	Inclusion criteria: exposure to an MVAs of moderately serious-to-serious severity		
	Exclusion criteria: not specified		
	<b>Sample size:</b> 738 people randomised and written to by the police and invited to take part in study; 151 agreed to take part in the study; 83 (36%) agreed to take part in the monitoring group and 68 (13%) took up the offer of intervention.		
	<b>Mean age:</b> 37.6 (SD 16.6) years		
	<b>Gender:</b> 89 (58.9%) men; 62 (41.1%) women		
	Ethnicity: not reported		
	Country: Netherlands		
Interventions	Group 1: counselling intervention: n = 68		
	Counselling intervention lasted 3–6 sessions depending on need. Components of the intervention included: practical help and information on procedures following an MVA and general information about psychological reactions; support to recognise that the event was over, to explore the experience and related emotions and to mobilise support networks; reality testing and confronting emotional reactions; confronting the memory of the experience; ongoing contact at least 2–3 months after the accident; referral for psychotherapeutic treatment where indicated.		
	<b>Group 2: monitoring group:</b> n = 83		
	Monitoring group were unaware that they were part of a comparison study and received assessments only.		
Outcomes	PTSD: IES		
	Other: Trauma Symptom Inventory		
	Follow-up: 1 and 6 months		
Notes			
Risk of bias			
Bias	Authors' judgement Support for judgement		



Brom 1993 (Continued)		
Random sequence generation (selection bias)	Unclear risk	Not reported
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	High risk	Not reported but outcomes were likely to be measured by the therapists.
Incomplete outcome data (attrition bias) All outcomes	High risk	The plan of analysis and management of dropout is not reported.
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.
Other bias	High risk	Randomisation occurred prior to potential participants agreeing to take part in the study and participants were only aware that they were either invited to participate in a secondary prevention programme or that they were invited to participate in a research project. The response rate for the monitoring group was 36%, for the intervention group 13%. Symptom scores on the IES were not matched at baseline. Groups were not matched for marital status and income. Individuals in the monitoring group were more likely to be married and had a statistically significant higher level of income.

Brunet 2013	
Methods	Design: RCT
Participants	Setting: emergency departments of 2 public hospitals
	Type of trauma exposure: exposure to a life-threatening event
	<b>Inclusion criteria:</b> experienced a life-threatening event in the last 10 days that elicited a peritraumatic reaction of fear, helplessness or horror.
	<b>Exclusion criteria:</b> did not speak either French or English; had or were suspected of having a traumatic brain injury; had a lifetime diagnosis of psychosis, substance or alcohol dependence, bipolar disorder or mental retardation; had been clinically depressed in the last 2 years; were taking psychotropic medication at the onset of the study; were injured to the extent that they could not participate in the study; lived outside the Montreal metropolitan area; did not have a significant other (a friend, spouse or another family member) to bring to the therapy session; or did not succeed in making an appointment with the therapist within 30 days after trauma exposure.
	<b>Sample size:</b> 90 people eligible to take part; 83 randomised; 74 completed baseline measures and were identified as the ITT sample; 5 individuals were subsequently withdrawn for protocol violation.
	<b>Mean age:</b> 36.29 (SD 11.05) years
	<b>Gender:</b> 40 (54.1%) men; 34 (45.9%) women
	Ethnicity: 85.1% white; 14.9% other
	Country: Canada
Interventions	Group 1: brief dyadic CBT intervention: n = 37



#### **Brunet 2013** (Continued)

2-session dyadic intervention included elements of psychoeducation and motivational interviewing, and targeted communication between the patient and significant other, aiming to facilitate support, promote bidirectional disclosure, reduce disclosure constraining behaviours and improve coping. It promoted the disclosure of thoughts and emotions about the trauma in the natural environment of the dyad while attempting to reduce social constraints on disclosure and negative social support interactions.

**Group 2: waiting list:** n = 37

Waiting list group filled out and returned the questionnaires.

Outcomes PTSD: IES-R; CAPS

Other: SCS; SAS-SR

Follow-up: postintervention

#### Notes

#### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Allocation undertaken through randomly permuted blocks.
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	IES-R (self-report measure) was the primary outcome. We judged the risk of detection bias to be low using this measure. PTSD diagnosis was made using the CAPS. It was unclear whether assessor were blind to allocation.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Analysis undertaken using ITT principles. 4 (5.4%) individuals were removed from the study for protocol violation and not included in follow-up analyses. A further 3 (4.1%) participants did not complete outcomes.
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.
Other bias	High risk	Authors affiliated with experimental intervention.

#### Cox 2018a

Methods	Design: multisite RCT
Participants	Setting: hospital ICUs
	<b>Type of trauma exposure:</b> patients admitted to an ICU who had been in receipt of mechanical ventilation for > 48 consecutive hours.
	<b>Inclusion criteria:</b> aged $\geq$ 18 years of age and in receipt of mechanical ventilation for > 48 consecutive hours, and successful extubation before discharge.
	<b>Exclusion criteria:</b> pre-existing or current cognitive impairment; treatment for severe mental illness during the 6 months preceding admission (e.g. active psychosis, suicidality); residence at a location other than home immediately before admission,; poor English fluency; ICU attending physician's expectation of patient survival < 3 months; inability to complete study procedures as determined by



#### Cox 2018a (Continued)

study staff; and failure to return home from either a hospital or postacute care facility within 3 months after discharge. Exclusion criteria for family members included history of cognitive impairment and poor English fluency.

Sample size: 2002 patients assessed for eligibility; 175 randomised

Mean age: 51.73 (SD 13.76) years

Gender: 100 (57.1%) men; 75 (42.9%) women

**Ethnicity:** 21.7% African-American; 70.1% white; 1.1% Asian; 1.1% American Indian/Alaska Native; 0.6% Native Hawaiian or other Pacific Islander; 1.1% Hispanic; 0.6% did not know or did not wish to an-

swer.

Country: USA

#### Interventions

## Group 1: telephone- and web-based CBT-based coping skills training: n = 86

CBT intervention delivered in 6 weekly telephone sessions, each lasting approximately 30 min, that addressed the following: introduction and relaxation exercise; progressive muscle relaxation; pleasant activities and activity-rest cycle; communication; cognitive restructuring and pleasant imagery; review and planning for sustainability. Psychologists taught each skill by providing a description and rationale for its use, leading participants through practice with feedback in the context of any self-reported ongoing stressors, helping participants plan how to apply the skill in real life, and highlighting relevant web-based content. In addition to learning skills themselves, family members coached patients in applying skills and using the web content on a day-to-day basis.

#### **Group 2: education programme:** n = 89

An education programme was designed with the help of a stakeholder group to address poor comprehension of critical illness, omitting any mention of post-ICU psychological distress. Programme consisted of 6 informational videos with accompanying web-based content. Study staff with content expertise in critical illness conducted 2 × 30-min telephone calls with participants during the 6-week intervention period to review materials and answer related questions.

## Outcomes

PTSD: IES-R

**Other:** HADS; EQ-5D; global mental and physical health status assessed with the PROMIS; adaptive coping behaviours measured by the Brief COPE.

**Follow-up:** 3 months' postbaseline (immediately post-treatment), and 6 months' postbaseline (3 months' post-treatment).

#### Notes

Family members of participants were also included and randomised to both interventions. The data reported here are for patient participants only. Eight participants died during the study. Patients were, on average, middle-aged and severely ill, with an expected hospital mortality of approximately 50%. Approximately one-third of patients had been treated for depression, anxiety or PTSD in the 3 months preceding admission. Readmission occurred among 43 (25%) patients during follow-up.

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation was undertaken via a password protected computerised system allocated patient–family member dyads at a 1:1 treatment group ratio with blocks of 4, stratifying by 3 factors to ensure balance.
Allocation concealment (selection bias)	Low risk	Not reported
Blinding of outcome assessment (detection bias)	Low risk	Trained research coordinators blinded to treatment group collected clinical data. Participants completed study surveys by telephone with treatment



Cox 2018a (Continued) All outcomes		group-blinded University coordinators or by a password-protected electronic patient-reported outcomes system at baseline and follow-up.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Analyses were conducted according to the ITT principle. Primary and secondary survey outcome analyses were conducted with full likelihood methods.
Selective reporting (reporting bias)	Low risk	Results reported as described in preregistered protocol.
Other bias	High risk	Study conducted with a severely ill population with high levels of premorbid mental health problems. The authors observed higher-than-expected attrition, attributable primarily to patients' serious illnesses and occurring after consent but before randomisation. Adherence to the interventions was fairly low and partly attributed to illness.

#### Curtis 2016

Methods	Design: RCT

#### **Participants**

Setting: 5 ICUs in 2 hospitals

Type of trauma exposure: family members of patients in ICU with high risk of mortality.

**Inclusion criteria:** predicted mortality  $\geq$  30% and a surrogate decision maker. Eligibility criteria for patients included: in ICU for > 24 hours; aged > 18 years; mechanically ventilated at enrolment; SOFA score  $\geq$  6 or diagnostic criteria predicting a  $\geq$  30% risk of hospital mortality; legal surrogate decision maker to consent for patient participation; and a family member able to come to the hospital. Eligibility criteria for family members included aged > 18 years and able to complete consent process and questionnaires in English.

Exclusion criteria: none.

**Sample size:** 2209 ICU patients screened for eligibility. Family members for 488 patients approached for participation and 170 enrolled; 2 family members withdrew before randomisation. The 168 randomised patients had 268 family members who participated.

Mean age of family member participants where available: 50.85 (SD 13.12) years

Gender (family members): 79 (29.5%) men; 189 (70.5%) women.

**Ethnicity where available:** 4.8% African-American; 84.8% white; 2.2% Asian; 3.9% Native American; 0.4% Native Hawaiian; 3.9% other or mixed

Country: USA

## Interventions

## **Group 1: communication facilitator:** n = 137

Communication facilitators assisted families of patients by providing communication support during the ICU stay with the aim to increase families' and clinicians' self-efficacy expectations about communication in the ICU. Intervention included: interviews by facilitators with family to understand the family's concerns, needs and communication characteristics; meetings by facilitators with physicians, nurses or other clinicians offering brief summary of family concerns, needs and communication characteristics; provision of communication and emotional support adapted to the family member's attachment style; facilitator participation in family conferences and 24-hour follow-up with the family after discharge to acute care.

**Group 2: usual care:** n = 131

Outcomes

PTSD: PCL



#### Curtis 2016 (Continued)

Other: PHQ-9; GAD-7; length of stay in ICU; cost of care

**Follow-up:** symptoms were assessed 3 and 6 months after the patient died in, or was discharged from, the ICU.

#### Notes

#### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Quote: "Randomization was stratified by hospital in block sizes of six, with results provided to study staff in sealed, opaque, consecutively numbered envelopes".
		Comment: information in the study paper implied computer-based randomisation.
Allocation concealment (selection bias)	Low risk	As above
Blinding of outcome as- sessment (detection bias) All outcomes	Low risk	Follow-up surveys assessing depression, anxiety and PTSD were mailed to family members' homes at follow-up points.
Incomplete outcome data (attrition bias) All outcomes	High risk	Loss to follow-up was high. Analysis was based on 133 (49.6%) of those randomised at 3 months and 122 (45.5%) at 6 months.
Selective reporting (re- porting bias)	Low risk	Results reported as described in preregistered protocol.
Other bias	Unclear risk	Eligibility criteria were changed during the trial to improve recruitment: the required SOFA score was lowered from ≥ 10 to ≥ 6.

## Gamble 2005

Methods	Design: RCT

Participants Setting: antenatal clinics of 3 maternity teaching hospitals.

Type of trauma exposure: postpartum women at risk of developing psychological trauma symptoms.

**Inclusion criteria:** aged > 18 years, in the last trimester of pregnancy, expected to give birth to a live infant, and able to complete questionnaires and interviews in English and met criterion A of DSM-IV for PTSD.

**Exclusion criteria:** women experiencing stillbirth or neonatal death.

**Sample size:** 348 women screened for birth-related trauma exposure, 103 met inclusion criteria and

were randomised.

Mean age: 28 (SD 6.04) years Gender: 103 (100%) women

Ethnicity: 93.2% white/European; 0.97% Asian; 2.9% other



#### Gamble 2005 (Continued)

Country: Australia

#### Interventions

#### **Group 1: midwife-led brief counselling intervention:** n = 50

Counselling began within 72 hours of birth on the postnatal ward and again by telephone at 4–6 weeks' postpartum. Counselling duration 40–60 min. The counselling process incorporated elements of critical stress debriefing and issues pertinent to the childbearing context. Intervention included building a therapeutic relationship, accepting and working with the mother's perception of their experience, supporting expression of feelings, filling in missing pieces of the account, connecting the event with the mother's emotions and behaviour, reviewing labour management, enhancing social support, reviewing positive approaches to coping, exploring further solutions.

**Group 2: TAU:** n = 53

Standard postnatal care

Outcomes

PTSD: MINI-PTSD

Other: EPDS; DASS-21; MSSS

Follow-up: 4-6 weeks' postpartum and 3 months' postpartum

#### Notes

#### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Participants randomised using sealed, opaque envelopes containing computer-generated, random allocations.
Allocation concealment (selection bias)	Low risk	Participants randomised using sealed, opaque envelopes containing computer-generated, random allocations.
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Appeared that assessor who completed outcome assessments at 4–6 weeks was also providing intervention and was not blind to allocation. A second assessor who completed assessments at 3 months was reported to be blind to allocation. Outcomes at 4–6 weeks appeared at high risk of detection bias and outcomes at 3 months appeared at low risk of bias.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Study retention rate was 100% at 3 months. 1 individual could not be contacted at 4–6 weeks.
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.
Other bias	Unclear risk	Treatment adherence not reported.

## Gamble 2010

Methods	Design: RCT
Participants	Setting: 4 hospital-based antenatal clinics
	Type of trauma exposure: women with traumatic birth/EmCS.



#### Gamble 2010 (Continued)

**Inclusion criteria:** women in third trimester of pregnancy expecting a full term, normal birth, able to understand sufficient English to discuss their experiences meeting DSM-IV criterion A for PTSD.

**Exclusion criteria:** pre-existing mental illness, or expecting an adverse birth outcome.

Sample size: 890 women screened for inclusion and 262 randomised

Mean age: 30 (SD not specified) years

**Gender:** 262 (100%) women **Ethnicity:** not specified

Country: Australia

#### Interventions

## **Group 1: PRIME:** n = 137

PRIME refers to a treatment condition for distressed postpartum women. Counselling was delivered at 72 hours after birth in person and at 6 weeks' postpartum by telephone. PRIME aims to support the expression of feelings and provide a framework for women to identify and work through distressing elements of childbirth (Gamble 2009). Women were provided with an opportunity to review the birth and gain a realistic perception of events. Focussed on developing individual situational supports for the present and near future, affirming negative things can be managed and developing a simple plan for achieving this.

#### **Group 2: parenting support:** n = 125)

Support focused on parenting concerns at 72 hours after birth and 6 weeks' postpartum by telephone. Midwives provided information on managing crying, feeding, sleeping and other parenting issues.

Both intervention were delivered in 2 sessions at 72 hours' and 6 weeks' postpartum by telephone.

**Group 3: control:** n = 138

Matched control group who were not traumatised were matched for age, parity and education. This group received standard maternity care as provided by their nominated facility. We have not included data from this group.

#### Outcomes

#### PTSD: PDS

**Other:** EPDS; DASS; HRQoL; mother-child relationship; parenting satisfaction; sense of competence; parenting stress

Follow-up: 6 weeks', 6 months', 12 months' postpartum

#### Notes

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Mothers screening positive were randomly allocated using computer-generated numbers.
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Not reported
Incomplete outcome data (attrition bias)	High risk	Analysis based on responses available at each follow-up point. Proportion of experimental group available at 6 and 12 months lower than control group.



Gamble 2010 (Continued)	
All outcomes	

Selective reporting (reporting bias)	Low risk	Outcomes reported as specified in protocol.
Other bias	High risk	Numerically, more participants indicating they received previous mental health help were in PRIME (40%) compared to the parenting group (29%; P = 0.054). No indication that this was controlled for in analyses.

#### Gidron 2001

Methods	Design: pilot RCT	
Participants	Setting: hospital emergency department	
	Type of trauma exposure: MVA	
	<b>Inclusion criteria:</b> survived an MVA within the last 24 hours; discharged from hospital within 24 hour after admission, indicating minor injury only and had heart rate > 95 beats/min upon admission into the emergency department	
	Exclusion criteria: brain damage	
	<b>Sample size:</b> number of patients approached to take part not reported; outcomes for 17 participants who completed the study were reported.	
	Mean age: 38.0 (SD not specified) years	
	<b>Gender:</b> 9 (52.9%) men; 8 (47.1%) women	
	Ethnicity: not specified	
	Country: Israel	
Interventions	<b>Group 1: MSI:</b> n = 8	

2-session intervention delivered over the telephone based on research about the nature of memory processing following trauma, specifically research suggesting that trauma memories are recalled as sensory, affective and fragmented information. Features included focus on time sections of the traumatic event; listening and clarifying details; memory structuring by having the therapist repeat the trauma narrative in an organised, labelled and logical manner, adding initial implications for the patient's life; patient described the traumatic event in the same structured, labelled and logical manner as the therapist did; between session practice of structured description; repeated practice of the new structure in the second session. Finally, the patient was taught about the importance of, and asked about, his/her social support.

## **Group 2: supportive listening:** n = 9

Patients telephoned twice, and invited to describe the event to the counsellor. The counsellor provided supportive listening and informed controls about the availability of treatment from the PTSD unit in the hospital. This condition attempted to control for therapist contact, attention to the problem, and simple disclosure and ventilation without guidance.

## Outcomes

PTSD: PDS
Other: none

Follow-up: 3-4 months

## Notes



#### Gidron 2001 (Continued)

#### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Not reported
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Low risk	At outcome assessment, participants were contacted by telephone by a researcher blind to their group status.
Incomplete outcome data (attrition bias) All outcomes	High risk	Study did not include a participant flowchart and the recruitment procedure and process of analysis was only briefly described.
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.
Other bias	High risk	Authors affiliated with experimental intervention. Small sample size. Study report not detailed. Therapist training and treatment adherence not reported.

#### Gidron 2007

Methods	Design: RCT		
Participants	Setting: university medical centre		
	Type of trauma exposure: traffic accident		
	<b>Inclusion criteria:</b> traffic accident 24–48 hours before entry to study; mild injuries; spent up to 24 hours in the hospital; aged 18–60 years and pulse rate of at least 95 beats/min.		
	Exclusion criteria: brain injury or loss of consciousness		
	<b>Sample size:</b> number screened and randomised not reported. Outcomes for 34 participants who completed the study reported.		
	<b>Mean age:</b> 28.3 (SD 10.0) years		
	<b>Gender:</b> 16 (%) men; 18 (%) women		
	Ethnicity: not specified		
	Country: Israel		
Interventions	<b>Group 1: MSI:</b> n = 19		
	2-session intervention delivered by telephone which is based on research about the nature of memo-		

2-session intervention delivered by telephone which is based on research about the nature of memory processing following trauma, specifically research suggesting that trauma memories are recalled as sensory, affective and fragmented information. Features of intervention included focus on time sections of the traumatic event; listening and clarifying details; memory structuring by having the therapist repeats the trauma narrative in an organised, labelled and logical manner, adding initial implications for the patient's life; patient described the traumatic event in the same structured, labelled and logical manner as the therapist; between session practice of structured description; repeated practice



#### Gidron 2007 (Continued)

of the new structure in the second session. Finally, the patient was taught about the importance of, and asked about, his/her social support.

## **Group 2: supportive listening:** n = 15

2 telephone calls that provided supportive listening. Included asking patients how they felt, asking whether they wanted to share the details of the event, listening to their story and telling them how to obtain further treatment should they feel such a need. The therapist provided only empathy without structuring patients' reports and without formal debriefing.

Outcomes PTSD: PDS

Follow-up: 3 months

Other: none

Notes Study reported that it was hypothesised that the MSI would result in fewer symptoms of PTSD than a supportive-listening control condition, and that these effects may depend on participants' gender. The

authors reported positive outcomes for PTSD for the experimental intervention for women only.

#### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Not reported
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Low risk	At follow-up, participants evaluated by telephone for their PTSD symptoms by a researcher blind to their group status.
Incomplete outcome data (attrition bias) All outcomes	High risk	Study did not include a participant flowchart and the recruitment procedure and process of analysis was only briefly described. The findings reported only fully participating patients and dropouts were not reported.
Selective reporting (reporting bias)	High risk	No study protocol identified. Study aimed to replicate findings from Gidron 2001. Gidron 2001 reported PTSD diagnosis as an outcome and it would have been reasonable to expect this study to do the same.
Other bias	High risk	Authors affiliated with experimental intervention. Small sample size. Study report not detailed. Therapist training and treatment adherence not reported.

## Holmes 2007

Methods	Design: RCT	
Participants	Setting: 2 level 1 trauma centres	
	<b>Type of trauma exposure:</b> major physical trauma. Approximately two-thirds of patients (43.9%) had received their injuries in road traffic accidents, falls or collisions (12.3%) or through non-accidental injury (9.6%). The cause of trauma for the remaining participants was not reported.	
	Inclusion criteria: aged > 18 years, had experienced major physical trauma.	
	Exclusion criteria: major head injury or injury was due to self-harm; current psychotic illness.	



#### Holmes 2007 (Continued)

**Sample size:** 964 cases of major trauma screened; 146 patients eligible to take part in study; 114 recruited and 90 completed initial assessments and were randomised.

Mean age: 37.0 (SD 14.7) years of those recruited

Gender: 63 (70%) men; 27 (30%) women

Ethnicity: not reported

Country: Australia

#### Interventions

#### Group 1: IPC: n = 51

IPC is a brief universal psychological intervention aimed at addressing the physical and functional consequences of serious injury. The aim of the intervention was to promote adaptation to injury, reduce depressive and PTSD and prevent the emergence of new episodes of psychiatric disorder. As part of the therapy, the impact of the injury on interpersonal issues predating the injury were identified and explored. Interpersonal issues arising after the trauma that were addressed included the degree to which needs were communicated with and responded to by family, health professionals and, in some cases, insurers. The domain of role transition was expanded to include the transition from health to injury. Issues of grief and loss arising from the injury were explored and related to preinjury issues where appropriate. Strategies to enhance adaptation to these issues were outlined in therapy and practised by the patient between sessions. Number of sessions available not reported. Mean number of sessions for those completing therapy 5.9 (SD 1.1).

**Group 2: TAU:** n = 39

Participants were informed following randomisation that they would not be receiving the IPC and would be reassessed at 3 and 6 months. In the case of psychological distress, they were recommended to seek assessment through their primary practitioner, but were also able to contact the study co-ordinator.

## Outcomes

PTSD: PCL

Other: SCID; BDI; HADS; AUDIT; SF-36

Follow-up: 3 and 6 months

#### Notes

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation involved the initial assessor contacting a research officer who made a blinded selection from a box of mixed envelopes.
Allocation concealment (selection bias)	Low risk	Randomisation involved the initial assessor contacting a research officer who made a blinded selection from a box of mixed envelopes.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Measures completed at 3 months were self-reported. At 6 months, a repeat SCID was conducted by an assessor blinded to the treatment condition and the previous assessment results.
Incomplete outcome data (attrition bias) All outcomes	High risk	Study only presented data for those who completed the IPC intervention (52.9% of those allocated to this arm), although the participant flowchart suggests that 92.2% of those randomised to IPC completed the 3-month assessment and 90.2% completed the 6-month assessment.
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.



Holmes 2007 (Continued)

Other bias Unclear risk Treatment adherence was only evaluated for cases of completed therapy.

## Irvine 2011

Risk of bias

Methods Design: multisite RCT **Participants** Setting: 2 hospitals following ICD transplant Type of trauma exposure: surgery for ICD transplant Inclusion criteria: patients who received their first ICD implant for secondary prevention of sudden cardiac death or primary prevention of sudden cardiac death and if their underlying heart disease was hypertrophic cardiomyopathy. Exclusion criteria: insufficient English fluency; medical chart documented evidence of dementia, cognitive impairment or psychosis; ICD implant for primary prevention of sudden cardiac death due to coronary heart disease, Ischaemic cardiomyopathy and congestive heart failure. The latter were excluded because of concerns that their poor and declining heart function might confound evaluation of the CBT. Sample size: 292 patients were eligible to take part in this study, 193 consented and were randomised. Mean age: 64.4 (SD 14.3) years Gender: 159 (82.4%) men; 34 (17.6%) women Ethnicity: not reported Country: Canada Interventions Group 1: telephone-based CBT: n = 96 Based on the cognitive theory of anxiety (Salkovskis 1996). Content tailored to specific misconceptions that have been known to cause distress in cardiac and ICD patients and it was informed by our clinical work. It included a therapist manual, 8 telephone counselling sessions, a psychoeducational booklet for the participants and a CD with mindfulness-based exercises and a progressive muscle relaxation exercise. Ad hoc counselling sessions were offered to participants who experienced an ICD shock. **Group 2: TAU:** n = 97 Whatever the respective ICD treatment sites routinely offered their patients. All patients received standard educational materials explaining their heart disease and the ICD device. Follow-up appointments included device interrogation (i.e. to extract arrhythmia events and ICD therapies) and troubleshooting at 6-month intervals, cardiac care as necessary and non-systematic supportive reassurance delivered informally in the clinic. Each centre also had access to a cardiac rehabilitation programme and psychiatric consultation as needed. PTSD: IES-R Outcomes Other: HADS; phobic anxiety sub scale of the Crown-Crisp Experiential Index; Physical Component Summary and the Mental Component Summary of the SF-36 Follow-up: 6 and 12 months' postbaseline Notes 8 participants died during the study. Missing values were not imputed by the study authors for participants who died over follow-up, as the cause of death was assumed to be non-random and potentially related to the phenomenon under study.



#### Irvine 2011 (Continued)

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Computer-generated random numbers program used to randomise participants to the CBT or TAU condition within 4 blocking variables.
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Follow-up questionnaires were mailed to participants 6 and 12 months after baseline and included a prestamped return envelope.
Incomplete outcome data (attrition bias) All outcomes	Low risk	ITT approach employed for the treatment outcome analyses.
Selective reporting (reporting bias)	Low risk	Results reported as described in preregistered protocol.
Other bias	Unclear risk	Evaluation of treatment adherence not reported.

#### Jensen 2016

Methods	Design: RCT		
Participants	Setting: 10 ICUs		
	Type of trauma exposure: patients admitted to ICU and required mechanical ventilation		
	<b>Inclusion criteria:</b> Danish-speaking adults aged ≥ 18 years who had been mechanically ventilated ≥ 48 hours.		
	<b>Exclusion criteria:</b> meeting baseline criteria for dementia; patients, who were not oriented in personal data according to the verbal response in Glasgow Coma Score, with detected delirium using the Confusion Assessment Methods for the ICU at randomisation, or enrolled in other follow-up studies.		
	Sample size: 2105 patients assessed for eligibility; 386 randomised		
	Mean age: 66.8 (SD not reported) years		
	<b>Gender:</b> 229 (59.3%) men; 157 (40.7%) women		
	Ethnicity: not reported		
	Country: Denmark		
Interventions	Group 1: nurse-led intensive care recovery programme: n = 235		
	Individualised ICU recovery programme based on several theoretical approaches towards psycho-		

Individualised ICU recovery programme based on several theoretical approaches towards psychological recovery including Antonovsky's salutogenic model (Antonovsky 1987), illness narratives, person-centred communication and elements from guided self-determination and trauma-focused CBT. Programme consisted of 3 consultations conducted by trained study nurses. Nurse training included 10 workshop days of theory and practice with experts in their field. Included intervention patients received an information pamphlet *Life after ICU* at randomisation. First consultation conducted at the clinic with the patient and close relative at 1–3 months' post-ICU. Dialogue focused on past and present as the patient was supported in constructing an illness narrative. A prerequisite for dialogue was the provision of patient photographs taken by ICU nurses during ICU recovery. Second and third consultations at 5 and 10 months' post-ICU conducted by telephone. Patients prepared by completing "Reflection sheets" indicating issues of importance to the individual.



#### Jensen 2016 (Continued)

#### **Group 2: standard care:** n = 235

Standard care included light sedation, early mobilisation, daily Confusion Assessment Methods for the ICU delirium assessment, written information for visitors, and ICU discharge without follow-up. ICU diaries were not used, but unplanned ICU visits and access to the medical record after discharge were permitted. Physical training was initiated in the ICU and physical rehabilitation was offered to all patients.

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Outcomes PTSD: HTQ-IV

Other: HADS; SF-36; sense of coherence measured by the Orientation to Life Questionnaire

Follow-up: 3 and 12 months' postdischarge

Notes Among randomised patients, 36 (19 %) in intervention group vs 43 (22 %) in standard care group died

within the first year post-ICU.

#### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Information in the study paper implied computer-based randomisation as treatment allocation was concealed by random selection of opaque sealed envelopes in permuted blocks of 6.
Allocation concealment (selection bias)	Low risk	Information in the study paper implied computer-based randomisation as treatment allocation was concealed by random selection of opaque sealed envelopes in permuted blocks of 6.
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Self-reported questionnaire packages were sent by post at 3 and 12 months' post-ICU. To increase the response rate, a few patients were assisted by telephone in completing the questionnaires if unable on their own.
Incomplete outcome data (attrition bias) All outcomes	High risk	Primary analysis was reported to be based on ITT. However, data were only analysed on an ITT basis for 235 participants (67.5 % of participants still alive at 3 months and 76.5% of the participants still alive at 12 months).
Selective reporting (reporting bias)	Low risk	Results reported as described in preregistered protocol.
Other bias	High risk	Description of an intervention manual and evaluation of treatment adherence not reported.

## **Jones 2010**

Methods	Design: RCT	
Participants	Setting: 6 general district hospitals and 6 university hospitals in 6 European countries.	
	Type of trauma exposure: ICU patients with an ICU stay > 72 hours	
	Inclusion criteria: patients who had been admitted to the ICU and ventilated.	
	<b>Exclusion criteria:</b> patients stayed in the ICU for < 72 hours; ventilated for < 24 hours; were too confused to give informed consent (including severe traumatic brain injury) and had pre-existing psychotic illness such as schizophrenia and manic depression (a confounding factor for psychological recovery) or diagnosed PTSD.	



Jones 2010 (Continued)

Sample size: 1164 people screened for inclusion and 352 were randomised.

Mean age: intensive care diaries: 60 (SD 15.6) years; delayed intensive care diaries: 59 (SD 16) years.

**Gender:** 227 (64.5%) men; 125 (35.5%) women

Ethnicity: not reported

Country: Denmark, Italy, Norway, Portugal, Sweden, UK

#### Interventions

## **Group 1: intensive care diaries:** n = 177

All patients had an ICU diary written for them while they were in critical care, which the healthcare staff wrote and the family contributed to if they considered they could. Diary was a daily record of the patients' ICU stay, written in everyday language and accompanied by photographs. Memories of the ICU were assessed at 1-week post-ICU. Patients received their diary as soon as they wanted following randomisation at 1-month postdischarge. The diary was introduced to the patient by a research nurse or doctor who ensured that they understood its contents but did not give any advice on what to do with it. Most of the discussions took place in an outpatient setting in a hospital but a small number were in the patients' own homes. In units where the travelling distance was too great for the patient to return to the hospital the discussion of the diary took place over the telephone.

## Group 2: delayed access to ICU diary: n = 175

Control patients received their diaries after they completed the final follow-up questionnaires at 3 months

#### Outcomes

PTSD: PDS administered as a diagnostic interview

Other: PTSS-14, a 14-item questionnaire that has been validated with ICU patients

Follow-up: 3 months following discharge from ICU

#### Notes

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Patients were assigned to treatment or control group at 1 month using a closed, non-transparent envelope technique, randomised in blocks of 6 through computerised random number generation.
Allocation concealment (selection bias)	Low risk	Patients were assigned to treatment or control group at 1 month using a closed, non-transparent envelope technique, randomised in blocks of 6 through computerised random number generation.
Blinding of outcome assessment (detection bias) All outcomes	High risk	It was considered impractical to guarantee blinding of the allocation of the diary as patients would volunteer their use. In order to reduce bias and ensure blinding of the diagnosis of PTSD at the 3-month follow-up, the researchers were only trained to interview and administer the PDS but were not made aware of the scoring calculation or in what way each question contributed to the score and final diagnosis.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Outcomes based on available data at 3-month follow-up. Dropout from both groups was balanced and low.
Selective reporting (reporting bias)	Unclear risk	Study protocol registered after completion of study. It would make sense to report PTSD severity outcomes (e.g. on the PDS) but these were not reported.



Jones 2010 (Continued)

Other bias

High risk

The PTSS-14 has only been subject to preliminary validation in a small UK-only sample. PTSD diagnosis was based on assessor administration of the PDS. Although the PDS can be used to generate a provisional PTSD diagnosis it is not intended to replace a structured diagnostic interview. A number of participants were excluded retrospectively based on identification of PTSD with onset prior to admission at the 3-month follow-up point.

#### Kazak 2005

Methods **Design:** pilot RCT

**Participants** 

**Setting:** oncology service within a children's hospital.

Type of trauma exposure: parents of children newly diagnosed with cancer.

Inclusion criteria: primary caregivers and partner of children aged 0–17 years newly diagnosed with

cancer.

Exclusion criteria: single caregivers

Sample size: 88 families assessed for eligibility; 19 families (38 individuals were randomised)

Median age: 37 year for primary caregivers; 42 years for partners

Gender: 18 (47.4%) men; 20 (52.6%) women

Ethnicity: not reported. Ethnicity for participants' children: 36.8% African-American; 52.6% white; 5.3%

Asian; 5.3% mixed race

Country: USA

Interventions

**Group 1: SCCIP-ND:** n = 9 families, n = 18 individuals

SCCIP-ND is an adaptation of an integrated cognitive behavioural and family therapy intervention developed and tested with adolescent survivors of childhood cancer and their families. SCCIP-ND is intended for 2 caregivers of a child newly diagnosed with cancer. Caregivers work conjointly to identify beliefs about their experiences during the initial month of treatment, a time in which potentially traumatic events may occur. The focus is on understanding how beliefs about cancer and its treatment influence caregivers and to help family members anticipate the impact of cancer on the family over time. Intervention consisted of 3 × 45-min sessions of a manualised family intervention, SCCIP-ND. The goal was to deliver the intervention within the first month after the child's cancer diagnosis. Intervention was facilitated by the use of other families who had experienced cancer. The first session focused on identifying beliefs about cancer, its treatment and the impact on the family using the A-B-C Model to examine the relationships between perceptions of cancer-related events and their feelings, actions and relationships. The second session focused on changing beliefs to enhance family functioning. Caregivers learn how to use reframing to modify their beliefs and subsequently, their emotional, behavioural and interpersonal consequences. Participants are then coached to identify new beliefs that accept the uncontrollable; focus on the controllable; acknowledge their own strengths and use the positive session. The third session addressed family growth and the future through the use of 2 metaphors, "The Family Survival Roadmap" and "Putting Cancer in its Place," to help caregivers recognise their beliefs about the family's future, and share these beliefs with each other.

**Group2: TAU:** n = 10 families, n = 20 individuals

Usual psychosocial care. Each family was assigned a social worker who attended the initial family meeting, provided resources and supplemental information about the diagnosis and treatment, and offered support.

Outcomes

PTSD: IES-R



Kazak 2005 (Continued)

Other: STAI

Follow-up: 2 months' post-treatment

Notes

Participants were randomised by family dyad, rather than individually. All 19 primary caregivers were mothers. Of the partner caregivers, 18 were fathers and 1 was a grandmother.

### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Randomisation used a predetermined concealed random assignment list maintained by a staff member unaware of patient identity.
Allocation concealment (selection bias)	Low risk	Randomisation used a predetermined concealed random assignment list maintained by a staff member unaware of patient identity.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Outcome measures were self-report questionnaires administered by research assistants.
Incomplete outcome data (attrition bias) All outcomes	High risk	The authors did not undertake detailed statistical analysis due to the relatively small sample size. Outcome data were reported only for those available to follow-up, with a larger number of dropouts (including 4 participants withdrawals) from the intervention group.
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.
Other bias	High risk	Small sample size

#### Lindwall 2014

Methods	Design: RCT
Participants	Setting: 4 major paediatric stem cell transplantation centres
	<b>Type of trauma exposure:</b> participants were parents of children undergoing stem cell or bone marrow transplantation.
	<b>Inclusion criteria:</b> parent of children aged 6–18 years undergoing stem cell or bone marrow transplantation with an expected stay of 3 weeks were eligible if they were primarily responsible for caring for the child during his/her hospital stay; available to participate throughout the duration of the child's hospitalisation for transplantation; and able to speak and read English fluently.
	Exclusion criteria: none
	Sample size: 242 child-parent dyads eligible to take part; 171 were randomised.
	Mean age: not reported
	Gender: 59.1% of child-parent dyad men; (40.9%) women
	Ethnicity: 4.3% Asian; 14.6% black; 5.3% Hispanic; 70.7% white; 5.3% other/unknown
	Country: USA
Interventions	Group 1: child-targeted intervention involving massage and humour therapy: n = 58



#### Lindwall 2014 (Continued)

Patients were provided with psychoeducation about the benefits of both massage and humour therapy. Patients were scheduled for 3 massage sessions per week over the course of 4 weeks (i.e. admission through week +3). Humour therapy consisted of providing a "humor cart" that contained enjoyable items such as videos, books and games. Scheduled humour sessions involving an interventionist and the patient occurred once per week. In addition, the humour cart was made available to families ≥ 3 times per week over the course of 4 weeks (i.e. admission through week +3).

# Group 2: child-targeted intervention plus a parent-targeted intervention involving massage and relaxation or imagery (or both): n = 57

Child-targeted intervention described above plus an additional parent intervention. Parents were provided with psychoeducation about how promoting their own well-being (i.e. engaging in massage and relaxation training) may also positively benefit their children. Parents were scheduled for a massage session 3 times per week for 4 weeks (i.e. admission through week +3). In addition, parents participated in weekly relaxation training sessions (i.e. admission through week +3) with a member of the research team, which promoted strategies such as muscle relaxation, breathing exercises and guided imagery. Parents were provided with a relaxation tape and player, and they were encouraged to engage in relaxation exercises 15–20 min daily.

**Group 3: usual care:** n = 56

Patient-parent dyads in the standard care arm of the study did not receive any additional intervention beyond the routine, comprehensive services that are provided for families during the SCT process at these major paediatric SCT centres.

Outcomes

PTSD: IES-R

Other: CES-D for depression

Follow-up: 24 weeks' postadmission

Notes

Sample characteristics were reported for parents and children in combination and it was not possible to separate them. Data from group 1 was not used in meta-analyses.

Bias	Authors' judgement	Support for judgement
	- Judgement	
Random sequence generation (selection bias)	Unclear risk	Not reported
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Unclear risk	Not reported
Incomplete outcome data (attrition bias) All outcomes	High risk	Outcomes were completed for 97/167 (58.1%) participants for whom there was available data at baseline. 25 patients died, 11 withdrew (3 withdrew immediately after being randomised to the standard care arm; the remaining 8 withdrew after a period of noncompliance indicating they were no longer interested or felt study procedures were too burdensome), 8 were taken off study for medical reasons (relapse, second transplantation) and 22 failed the week +24 assessment.
Selective reporting (reporting bias)	High risk	No study protocol identified. The study authors reported no statistically significant differences between intervention and controls, which meant that it was not possible to include this study in meta-analyses. This has the potential to lead to an overestimate of effectiveness for those outcomes.



## Lindwall 2014 (Continued)

Other bias Low risk No other biases indicated.

## Marchand 2006

Methods	Design: RCT		
Participants	<b>Setting:</b> with the collaboration of a major convenience store chain in the metropolitan area of Montreal.		
	Type of trauma exposure: victims of armed robbery.		
	<b>Inclusion criteria:</b> victim of an armed robbery that included acts of violence ranging from threat of death or injury to physical assault and threat with a weapon. They also had to have reported to the screening interviewer that they experienced intense fear, helplessness or horror during or after the robbery such as described in Criterion A2 of the DSM-IV PTSD diagnosis.		
	Exclusion criteria: no additional exclusion criteria.		
	Sample size: number screened not reported; 75 individuals were randomised.		
	<b>Mean age:</b> 21.8 (SD = 6.7) years		
	<b>Gender:</b> 36 (48%) men; 39 (52%) women		
	Ethnicity: not reported		
	Country: Canada		
Interventions	<b>Group 1: CISD-A:</b> n = 33		
	In the CISD-A condition (adapted from Mitchell 1995), debriefings were conducted individually and consisted of 2 × 1-hour sessions at a 1-week interval and were conducted by an experienced psychologist. The first debriefing session took place 2–22 days after the robbery and covered the following themes: goals of the session; present tense detailed description of the traumatic event; thoughts and emotions experienced by the participant during and after the event; information about normal stress reactions and the challenge of the participant's irrational beliefs; the stress management techniques; and if necessary, reference to further follow-up. The second session of debriefing followed the same structure and included a review of reactions, thoughts and emotions experienced during the week following the first session.		
	<b>Group 2: TAU:</b> n = 42		
	Participants received no interventions and participated only at the assessments.		
Outcomes	PTSD: PTSD module of the SCID; IES		
	Other: none		
	Follow-up: 1 and 3 months' postbaseline		
Notes			
Risk of bias			
Bias	Authors' judgement Support for judgement		
Random sequence generation (selection bias)	Unclear risk Not reported		



Marchand 2006 (Continued)		
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Assessments were conducted by graduate students who were blind to group assignment.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Proportion of participants unavailable for follow-up was similar for both groups of the study at 1 and 3 months (24%). Completer and ITT analyse using last observation carried forward were undertaken.
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.
Other bias	Unclear risk	There was a fairly sizeable difference between the 2 groups for IES scores at baseline. This was not statistically significant but may have contributed to differences at subsequent time points and this was not controlled for in analyses.

#### Mouthaan 2013

Methods	Design: RCT
Participants	Setting: 2 level 1 trauma centres
	<b>Type of trauma exposure:</b> injury patients transported by ambulance or helicopter. Suspected to have experienced possible severe injuries that required specialised acute medical care.
	<b>Inclusion criteria:</b> aged ≥ 18 years, proficiency in Dutch and having experienced a potential traumatic event (cf. Criterion A1 DSM-IV PTSD diagnosis).
	<b>Exclusion criteria:</b> injury resulting from deliberate self-harm; organic brain condition, psychotic disor der, bipolar disorder or depression with psychotic features; moderate-to-severe traumatic brain injury or permanent residency outside the Netherlands.
	Sample size: 1807 individuals considered; 1032 assessed for eligibility and 300 randomised
	<b>Mean age:</b> self-guided Internet-based intervention: 44.18 (SD 15.76) years; no intervention: 43.49 (SD 16.00) years
	<b>Gender:</b> 180 (60%) men; 120 (40%) women
	Ethnicity: not reported. 249 (83%) reported to be of Dutch cultural background.
	Country: Netherlands
Interventions	<b>Group 1: self-guided Internet-based intervention:</b> n = 151
	Based on established CBT techniques (Trauma TIPS, which is based on CBT techniques of psychoedu-

Based on established CBT techniques (Trauma TIPS, which is based on CBT techniques of psychoeducation, stress management/relaxation techniques and in vivo exposure). It consists of 6 steps, including introduction to the programme and basic operating instructions; assessments of acute anxiety and arousal using VAS at pre- and postintervention; video features of the trauma centre's surgical head explaining the procedures at the centre and the purpose of the programme, and of 3 patient models sharing their experiences after their injury; a short textual summary of 5 coping tips for common physical and psychological reactions after trauma; audio clips with instructions for stress management techniques; contact information for program assistance or professional help for enduring symptoms; and a Web forum for peer support.

**Group 2: TAU:** n = 149



Mouthaan 2013 (	(Continued)
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Care as usual, available to patients from both groups, consisted of incidental, non-structured talks with trauma centre staff or with a patient's general practitioner (GP), either directly following injury or during the course of the trial.

#### Outcomes

PTSD: CAPS; IES-R

**Other:** HADS; MINI-Plus, version 5.0 to diagnose major depressive disorders and other anxiety disorders; mental healthcare utilisation using the Trimbos/Institute for Medical Technology Assessment questionnaire and the Short Form Health and Labour Questionnaire.

Follow-up: 1, 3, 6 and 12 months' postinjury

Notes

The authors noted that participants were reluctant to use the intervention.

## Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Randomisation performed by a research member independent of data collection in a 1:1 ratio by a computerised program using random block sizes stratified by study centre.
Allocation concealment (selection bias)	Low risk	Randomisation performed by a research member independent of data collection in a 1:1 ratio by a computerised program using random block sizes stratified by study centre.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Patients were asked not to share information about the randomisation to the assessors, to ensure that they were blind to the allocated interventions.
Incomplete outcome data (attrition bias) All outcomes	Low risk	ITT and completer-only analysis was undertaken using mixed-model analysis. Modest loss to follow-up at 1 and 3 months. However, missing data > 50% at 6 and 12 months.
Selective reporting (reporting bias)	Low risk	Outcomes were as reported in the study protocol.
Other bias	High risk	Authors affiliated with experimental intervention.

#### **Rothbaum 2012**

Methods	Design: RCT	
Participants	Setting: participants were recruited from a hospital trauma centre.	
	<b>Type of trauma exposure:</b> patients admitted to the emergency department experiencing a DSM-IV criterion A trauma.	
	<b>Inclusion criteria:</b> adults aged 18–65 years who had experienced a traumatic event within 72 hours of presentation in the emergency department; afraid that they might be killed or seriously injured during the event; able to be contacted following discharge and to return for follow-up appointments; and alert and oriented and able to provide informed consent.	
	<b>Exclusion criteria:</b> non-English speaking; lost consciousness > 5 min during the event; history of a serious mental illness; currently suicidal or reported current substance dependence.	
	<b>Sample size:</b> 5608 individuals assessed for eligibility; 4219 did not meet inclusion criteria; 1249 refused to take part or were excluded for other reasons; 137 randomised.	



Rothbaum 2	2012	(Continued)
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**Mean age:** 31.5 (SD = 11.6) years

Gender: 48 (35.0%) men; 89 (65.0%) women

**Ethnicity:** 13.1% black; 78.8% white; 1.5% Native American; 6.6% other

Country: USA

Interventions **Group 1: modified PE:** n = 69

Patients received 3 × 1-hour sessions of a modified PE intervention, distributed 1 week apart. PE consisted of imaginal exposure, processing time, breathing retraining, psychoeducation and homework

tasks (including addressing in vivo exposure goals).

Group 2: assessment only: n = 68

Outcomes PTSD: PTSD PSS-I; PDS

Other: BDI-II, Additional Treatment Inventory; Standardized Trauma Interview

Follow-up: 4 and 12 weeks

Notes

#### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Low risk	Envelopes containing computer-generated patient random assignments (either to immediate intervention or assessment only) were given to the patient and their nurse after the initial evaluation to ensure that assessors remained blind.
Allocation concealment (selection bias)	Low risk	Envelopes containing computer-generated patient random assignments (either to immediate intervention or assessment only) were given to the patient and their nurse after the initial evaluation to ensure that assessors remained blind.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Assessors were blinded.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Missing data handled with multiple imputation with mixed-effect models being used to obtain predicted mean values for outcomes at each assessment point. There was a disproportionately higher level of missing data in the intervention group at initial outcome (31.9% compared to 19.1%) and 12 weeks' follow-up (39.1% compared to 27.9%).
Selective reporting (reporting bias)	Unclear risk	Outcomes were not reported in study protocol.
Other bias	High risk	Authors affiliated with experimental intervention.

## Ryding 1998

Methods	Design: RCT
Participants	Setting: hospital obstetrics and gynaecology department



Ryding 1998 (Continued)

Type of trauma exposure: EmCS

Inclusion criteria: Swedish-speaking women, subjected to an EmCS delivering a live infant.

**Exclusion criteria:** no additional criteria specified.

Sample size: 111 patients screened; 106 randomised and 105 agreed to take part.

Mean age: 30 (SD not reported) years

**Gender:** 99 (100%) women **Ethnicity:** not reported

Country: Sweden

Interventions

#### **Group 1: counselling:** n = 53

Intervention delivered by an obstetrician with a primary psychotherapy qualification. After informed consent, the counsellor booked the participant for a consultation at the maternity ward as soon as practicable. First, the participant was asked to tell her own story about the delivery including the Em-CS. Then her thoughts and feelings during 6 phases of the delivery experience (arrival at the unit to sight of the infant for the first time) were explored. The aim of the first consultation was to establish contact and to collect relevant information. The participant was asked what she needed help with. A second consultation took place before the woman was discharged from hospital. The participant was encouraged to talk about her worst memories of and feelings about the delivery. Questions about risks to the life or health of the woman or her baby were dealt with. Existential issues, such as the insecurity of life, were discussed if relevant. The participant was assured that her possible symptoms of post-traumatic stress reactions were normal under the circumstances. She was instructed how to 'dose' frightening memories when at home. She was encouraged in her role as a new mother. During the third consultation about 2 weeks after delivery, the participant discussed her situation at home and her contact with her baby, her partner and others. Any post-traumatic stress reactions were again considered. She was given a copy of her complete record, which was examined in detail and explained. Possible feelings of shame or guilt about her performance during or after delivery were discussed, as well as possible feelings of anger or disappointment with the staff. The consultation ended by focusing on a positive memory in relation to the delivery. The fourth consultation took place about 3 weeks after delivery. Thoughts and feelings concerning the delivery experience were again examined. The meaning of the recent EmCS for the individual woman, and what she had learnt from the experience, was discussed. The possibility of another pregnancy was discussed, as well as hopes and misgivings about a future delivery. The participant was given medical advice concerning a possible pregnancy and delivery to come. The first consultation took ≥ 1 hour. The second to fourth meetings were limited to about 45 min.

Group 2: TAU: n = 52

A member of the research team contacted participants on the maternity ward. The women completed 3 questionnaires intended to measure the cognitive appraisal of the experienced delivery, the possible presence of post-traumatic stress reactions and general mental distress. If they had questions regarding the recent delivery or expressed distress of any type, they were encouraged to contact the ward staff.

Outcomes

PTSD: IES

Other: Symptoms Check List (SCL); Wijma-Expectancy/Experience Questionnaire

Follow-up: 1 and 6 months' postpartum

Notes

Risk of bias

Bias Authors' judgement Support for judgement



Ryding 1998 (Continued)		
Random sequence generation (selection bias)	High risk	Every second EmCS patient, according to the delivery ward register, was selected for counselling, the remainder being selected for the comparison group.
Allocation concealment (selection bias)	High risk	Every second EmCS patient, according to the delivery ward register, was selected for counselling, the remainder being selected for the comparison group.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	All measures where self-report which participants completed by post.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Dropout from both groups was low and about equivalent.
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.
Other bias	Unclear risk	The study therapist was also the research leader and the study report acknowledges that participant gratitude may have contributed to participant responses. Treatment adherence was not reported.

## Ryding 2004

Methods	Design: RCT		
Participants	Setting: hospital obstetrics and gynaecology department		
	Type of trauma exposure: EmCS		
	<b>Inclusion criteria:</b> all Swedish-speaking women giving birth to a live infant by EmCS at the participating hospital		
	Exclusion criteria: non-Swedish speakers		
	Sample size: 217 women met the inclusion criteria; 162 were randomised.		
	Mean age: 32 (SD not specified) years, range 19–44 years		
	<b>Gender:</b> 162 (100%) women		
	Ethnicity: not specified		
	Country: Sweden		
	162 women who had experienced birth by EmCS. Intervention offered to all.		
Interventions	Group 1: group counselling: n = 89		
	4 or 5 women were invited to each of the EmCS groups. The group leaders were a maternity and child		

4 or 5 women were invited to each of the EmCS groups. The group leaders were a maternity and child welfare psychologist and an experienced delivery ward midwife. Consultations lasted for 2 hours, and the groups met twice at a 2- to 3-week interval. The main purpose was to arrange for participants to meet other women who had undergone EmCS so that they could share experiences. It was hoped that the women would be able to discuss both medical procedures (the midwife would have the answers) and psychological matters, such as feelings after a traumatic birth and feelings about motherhood and the baby (the psychologist would be able to provide counselling). It also hoped to identify new mothers in need of individual contact and to provide information about the possibilities of further help.

**Group 2: TAU:** n = 73



Ryc	ling	2004	(Continued)
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Women offered an individual consultation to discuss their recent delivery, if they wished, after completing outcome questionnaires at 6 months' postpartum. Standard care after an EmCS included the midwife and doctor involved in the procedure visiting the mother in the maternity ward so as to exchange information about the experience, although for practical reasons this was not always possible.

Outcomes

PTSD: IES

Other: Wijma Expectancy/Experience Questionnaire; EPDS

Follow-up: 6 months

Notes

## Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	Women who gave birth on approximately 18 predetermined days of the month were randomised to the counselling group, and the remainder to the control group.
Allocation concealment (selection bias)	High risk	Women who gave birth on approximately 18 predetermined days of the month were randomised to the counselling group, and the remainder to the control group.
Blinding of outcome assessment (detection bias) All outcomes	Low risk	All measures where self-report which participants completed by post.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Analyses were based on the number of questionnaires returned. Response rate was high for both groups. Data were reported by median and interquartile range. The reason for this was unclear but may have been because of skew.
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.
Other bias	Unclear risk	The group intervention was not manualised and was provided through semi-structured delivery. Treatment adherence was not reported.

## Taghizadeh 2008

Methods	Design: RCT
Participants	Setting: hospital
	Type of trauma exposure: traumatic birth
	<b>Inclusion criteria:</b> women who had experienced a traumatic delivery who had given birth to alive and physically normal baby; could understand and speak Persian; no history of known psychological illnesses or not using psychological drugs; no history of infertility and abortion despite the desire to preserve the foetus
	<b>Exclusion criteria:</b> mothers who themselves or their infants needed special care; mothers who used external counselling services; mothers who experienced stressful events during the study and ≤ 1 year before childbirth
	Sample size: 300 women randomised: 150 to each group



Tagh	izadel	1 2008	(Continued)
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Mean age: not reported

Gender: women

Ethnicity: Iranian or who could speak and understand Persian (Farsi)

Country: Iran

Interventions

**Group 1: consultation with midwife:** n = 150

The first face-to-face consultation session was 72 hours after giving birth. The duration of intervention

was 4-6 weeks and each session was 40-60 min.

**Group 2: TAU** 

Outcomes

PTSD: IES

Other: not reported
Follow-up: 3 months

Notes

## Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	High risk	All eligible participants in the first 3 days of week (in Iran it is Sat, Sun, Mon) were assigned to control group and all in other 4 days were assigned to intervention. The next week they exchange the sequence, which means 3 first days of week (in Iran it is Sat, Sun, Mon) were assigned to intervention group and all in other 4 days were assigned to control group.
Allocation concealment (selection bias)	High risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Low risk	The trained researcher who was gathering data for postintervention question- naires was blind about control and intervention groups.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Information on the number of participants who completed treatment and number of dropouts were not reported.
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.
Other bias	Low risk	No other biases indicated.

## Wang 2015

Methods	Design: RCT
Participants	Setting: emergency department of a hospital
	<b>Type of trauma exposure:</b> severe MVA survivors. Patients were suspected to have experienced possible severe injuries that required specialised acute medical care.
	Inclusion criteria: aged 18–65 years, and able to communicate in verbal and written Chinese.



#### Wang 2015 (Continued)

**Exclusion criteria:** history of neurological problems, brain surgery, brain damage and spinal cord injuries; current alcohol or drug abuse (or both), and current or past schizophrenic or psychotic disorders.

**Sample size:** 76 individuals assessed for eligibility; 52 randomised.

Mean age: 40.3 (SD 14.8) years, based on available data from 46 completers.

Gender: 19 (41.3%) men; 27 (58.7%) women, based on available data from 46 completers.

Ethnicity: not reported but likely to be Chinese.

Country: China

#### Interventions

#### **Group 1: creative arts:** n = 26

Intervention delivered in small groups of 4–6 participants in hospital led by a psychologically trained and supervised artist. The primary modalities utilised were creative writing and drawing. Participants were offered 8 weekly 40 min sessions. In the first 2 weeks, participants were requested to perform simple drawing techniques. Weeks 3–5 explored present emotions and issues. Final sessions focused on bringing together resources for moving forward.

**Group 2: waiting list:** n = 26

#### Outcomes

PTSD: CAPS; IES-R

Other: HADS (depression and anxiety); PTGI

Follow-up: 2, 6 and 12 months

#### Notes

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Not reported
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Participants were asked not to share information to the assessors to ensure that they were blind to interventions.
Incomplete outcome data (attrition bias) All outcomes	Low risk	Completer-only analysis, although follow-up data was available for all participants at 2-month follow-up and the majority of participants at subsequent follow-ups.
Selective reporting (reporting bias)	High risk	No study protocol identified. The study authors reported no statistically significant differences between intervention and controls, which meant that it was not possible to include this study in meta-analysis. This has the potential to lead to an overestimate of effectiveness for those outcomes.
Other bias	High risk	Intervention was not manualised so it could not be guaranteed that the therapy was delivered in the same manner to all of the participants. Sample size was small.



#### Wijesinghe 2015

Methods **Design:** RCT

Participants Setting: district general hospital

Type of trauma exposure: snakebite victims

Inclusion criteria: snakebite victims admitted to hospital identified as being envenomed and requiring

treatment with antivenom.

Exclusion criteria: aged <18 years, known mental illness, and without basic fluency in the Sinhala lan-

guage.

Sample size: 225 snakebite victims randomised.

Mean age: 42.1 (SD 12.4) years

Gender: 168 (74.7%) men; 57 (25.3%) women

Ethnicity: 100% Asian
Country: Sri Lanka

#### Interventions

#### **Group 1: PFA and psychoeducation and CBT:** n = 75

PFA and psychoeducation following hospital treatment (as described below) and a single session of CBT at 1 month. In this session, victims were initially engaged in a focused discussion on how they had functioned in their daily lives after the snakebite and whether they had any ongoing difficulties. A checklist guided the doctors providing the intervention to identify the victims' dysfunctional cognitions related to health, personal life, functional abilities and overall future expectations. If dysfunctional cognitions were elicited, they were reframed in a positive manner. The doctors also encouraged return to work and normal life. In patients who had not gone back to work to an optimal level, activity scheduling was introduced. A phased return to household and occupational activity was suggested and patients were encouraged to return to their hobbies and pleasurable activities. Other maladaptive coping methods such as substance misuse were discussed and counselling provided as needed. In situations where there was a understandable anxiety about returning to work and being bitten again by a snake, practical safety measures such as wearing boots and gloves and carrying a torch and a stick were encouraged. The duration of this intervention was typically 20 min.

## **Group 2: PFA and psychoeducation:** n = 75

Discussion with a non-specialist doctor about the patient's opinion on the causes and consequences of the snake bite. The intervention followed the normal doctor–patient interview style in which patients were initially engaged in open-ended questions and allowed to express their views. The conversation was then moved into a more structured discussion, using a structured checklist to ensure a degree of standardisation. This list included important thoughts to elicit, such as myths, negative assumptions, and future plans and expectations of the patient. If any erroneous or maladaptive ideas were identified, they were challenged in a non-confrontational manner and more plausible, evidence-based alternative views were expressed. Common misbeliefs which have a negative impact on a person's psychological adaptation and subsequent functional level were specifically addressed. Patients were encouraged to engage in a healthy lifestyle following discharge and to avoid the trap of assuming a sick role.

**Group 3: no intervention:** n = 75

Outcomes PTSD: PSS-SR

Other: HSCL-25; BDI; Sheehan Disability Inventory

Follow-up: 6 months

Notes Psychological interventions were delivered by a non-specialist doctors involved in the study were trained by a specialist psychiatrist.



### Wijesinghe 2015 (Continued)

### Risk of bias

Bias	Authors' judgement	Support for judgement
Random sequence generation (selection bias)	Unclear risk	Not reported
Allocation concealment (selection bias)	Unclear risk	Not reported
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Patients were assessed for presence of psychological morbidity and functional status 6 months following discharge from hospital by a specialist psychiatrist blind to intervention.
Incomplete outcome data (attrition bias) All outcomes	Unclear risk	Analysis was reported for those available to follow-up. Loss to follow-up for the study as a whole was low at 8.4%. However, there was proportionately more missing data for no intervention (9.3%) and PFA only (13.3%) compared to CBT (2.6%). A sensitivity analysis assuming including data for individuals who were unavailable was undertaken based on the assumption of worst clinical assumption. However, the details of these analyses were not reported.
Selective reporting (reporting bias)	High risk	A study protocol was available but the planned process of analysis was not explained. The study authors reported no statistically significant differences between intervention and controls, which meant that it was not possible to include this study in meta-analysis. This has the potential to lead to an overestimate of effectiveness for those outcomes.
Other bias	Unclear risk	Psychiatric caseness was based on the use of screening instruments.

### Zatzick 2001

Zatzick 2001				
Methods	<b>Design:</b> RCT – bias unlikely			
Participants	Setting: level 1 hospital trauma centre			
	Type of trauma exposure: injured MVA and assault victims.			
	Inclusion criteria: hospitalised MVA or assault survivors aged 14–65 years, who were English speaking.			
	Exclusion criteria: patients who sustained severe injuries.			
	Sample size: 105 patients assessed for study participation; 57 eligible to take part; 34 randomised.			
	<b>Mean age:</b> 33.8 (SD 12.1) years			
	<b>Gender:</b> 20 (58.8%) men; 14 (41.7%) women			
	Ethnicity: 61.8% white; 38.2% other			
	Country: USA			
Interventions	Group 1: collaborative care intervention: n = 16			
	On the surgical ward, each patient was assigned to a trauma support specialist who met each intervention patient at the bedside. The trauma support specialists were instructed to develop a therapeutic relationship and follow patients for 4 months through primary care outpatient appointments and community rehabilitation. To establish a basis for collaborative problem definition and shared patient–provider treatment planning, the trauma support specialists were instructed to elicit and track			



#### Zatzick 2001 (Continued)

patients' post-traumatic concerns. Patients' post-traumatic concerns were incorporated into joint problem definition and the trauma support specialists were instructed to intervene on behalf of the patients in the resolution of these concerns whenever possible. A psychotherapy module specifically targeting post-traumatic distress and substance use was also delivered as part of the multifaceted collaborative intervention. A psychoeducational component of the intervention began with a review of the traumatic event, followed by a discussion of related emotions, cognitions and possible future post-traumatic symptoms, and closed with suggested coping strategies including algorithms for contacting the support specialist. The motivational enhancement techniques focused on the evaluation of readiness to change and implementation of a motivational interview targeting post-traumatic alcohol and drug use.

**Group 2: usual care:** n = 18

Outcomes

PTSD: PCL-C

Other: CES-D; ASI; PCS of the Medical Outcomes Study 12-Item Short-Form Health Survey

Follow-up: 1 and 4 months

Notes

Patients aged 14-65 were eligible for inclusion in this study. We included it as the majority of participants were likely to have been aged 18 and over based on the mean age of participants (33.8; SD 12.1).

### Risk of bias

Bias	Authors' judgement	Support for judgement	
Random sequence generation (selection bias)	Unclear risk	Not reported	
Allocation concealment (selection bias)	Low risk	The project co-ordinator independently randomised new patients in blocks of 6 to the intervention or control group.	
Blinding of outcome assessment (detection bias) All outcomes	Low risk	Research associates conducting follow-up telephone interviews remained blinded to patient group assignments.	
Incomplete outcome data (attrition bias) All outcomes	Low risk	Outcome analyses were conducted for the both the ITT sample and for those with complete data on each outcome measure at all time points.	
Selective reporting (reporting bias)	Unclear risk	No study protocol identified.	
Other bias	High risk	Sample size was small. The authors also acknowledge that "there were some difficulties implementing the collaborative care principles of continuous case management and active sustained follow-up. For instance, patients with no insurance often required intensive service coordination efforts targeting linkage with community agencies. Because these activities required time availability and flexibility of scheduling that taxed the abilities of the highly trained, hospital-based case managers, these efforts to insure continuity of care frequently fell short." The collaborative intervention was not manualised and was likely to be implemented with marked variability across trauma support specialists.	

ASI: Addiction Severity Index; AUDIT: Alcohol Use Disorders Identification Text; BDI: Beck Depression Inventory; CAPS: Clinician-Administered PTSD Scale; CBT: cognitive behavioural therapy; CES-D: Center for Epidemiological Studies Depression Scale; CISD-A: Critical Incident Stress Debriefing – Adapted; DASS-21: Depression Anxiety and Stress Scale-21; DSM: Diagnostic and Statistical Manual of Mental Disorders; EmCS: emergency caesarean section; EPDS: Edinburgh Postnatal Depression Scale; EQ-5D: EuroQol; GAD: generalised anxiety disorder; GAD-7: Generalised Anxiety Disorder Assessment; HADS: Hospital Anxiety and Depression Scale; HRQoL: health-related



quality of life; HSCL: Hopkins Symptom Checklist depression scale; HTQ-IV: Harvard Trauma Questionnaire; ICD: implantable cardioverter defibrillator; ICU: intensive care unit; IES: Impact of Events Scale; IES-R: Impact of Events Scale – Revised; IPC: interpersonal counselling; ITT: intention-to-treat; MA: mortuary attendant; min: minute; MINI-PTSD: Mini-International Neuropsychiatric Interview – PTSD; MSI: memory structuring intervention; MSSS: Maternity Social Support Scale; MVA: motor vehicle accident; n: number; NBAS: Neonatal Behavioral Assessment Scale; NICU: neonatal intensive care unit; PCL: Post-Traumatic Stress Disorder Checklist; PCS: Physical Components Summary; PDS: Posttraumatic Diagnostic Scale; PE: prolonged exposure; PFA: psychological first aid; PHQ: Patient Health Questionnaire; PICU: paediatric intensive care unit; PPQ: Perinatal Posttraumatic Stress Disorder Questionnaire; PRIME: Promoting Resilience In Mothers' Emotions; PROMIS: Patient-Reported Outcomes Measurement Information System; PSS-PICU: Parental Stressor Scale – Paediatric Intensive Care Unit; PSS-I: PTSD Symptom Scale – Interview Version; PSS-SR: Post-traumatic Stress Symptom Scale – Self Report; PTGI: Posttraumatic Growth Inventory; PTSD: post-traumatic stress disorder; PTSS: post-traumatic stress syndrome; RCT: randomised controlled trial; SAS-SR: Social Adjustment Scale – Self-Report; SCCIP-ND: Surviving Cancer Competently Intervention Program – Newly Diagnosed; SCID: Structured Clinical Interview for DSM; SCL: Symptoms Checklist; SCS: Social Constraints Scale; SD: standard deviation; SF-36: Short-Form 36; SOFA: Sequential Organ Failure Assessment; STAI: State-Trait Anxiety Inventory; TAU: treatment as usual; VAS: visual analogue scale; WHOQOLBREF: World Health Organization Quality of Life Assessment – Brief Version.

### **Characteristics of excluded studies** [ordered by study ID]

Study	Reason for exclusion
Ben-Zion 2018	Types of participants: treatment study of symptomatic individuals
Bisson 2004	Types of participants: treatment study of symptomatic individuals
Bryant 1998	Types of participants: treatment study of symptomatic individuals
Bryant 1999	Types of participants: treatment study of symptomatic individuals
Bryant 2003	Types of participants: treatment study of symptomatic individuals
Bryant 2005	Types of participants: treatment study of symptomatic individuals
Bryant 2008	Types of participants: treatment study of symptomatic individuals
Bugg 2009	Types of participants: treatment study of symptomatic individuals
Cernvall 2015	Types of participants: treatment study of symptomatic individuals
Echeburua 1996	Types of participants: treatment study of symptomatic individuals
Ehlers 2003	Types of participants: treatment study of symptomatic individuals
Foa 2006	Types of participants: treatment study of symptomatic individuals
Freedman (in preparation)	Types of participants: treatment study of symptomatic individuals
Freedman (submitted)	Types of participants: treatment study of symptomatic individuals
Freyth 2010	Types of participants: treatment study of symptomatic individuals
Jarero 2011	Types of participants: treatment study of symptomatic individuals
Jarero 2015	Types of participants: treatment study of symptomatic individuals
Nixon 2012	Types of participants: treatment study of symptomatic individuals
Nixon 2016	Types of participants: treatment study of symptomatic individuals



Study	Reason for exclusion
O'Donnell (in preparation)	Types of participants: treatment study of symptomatic individuals
O'Donnell 2012	Types of participants: treatment study of symptomatic individuals
Resnick 2005	Types of participants: single session intervention
Rose 1999	Types of participants: single session intervention
Rothbaum (submitted)	Types of participants: single session intervention
Shalev 2012	Types of participants: treatment study of symptomatic individuals
Shapiro 2015	Types of participants: treatment study of symptomatic individuals
Shapiro 2018	Types of participants: treatment study of symptomatic individuals
Shaw 2013	Types of participants: treatment study of symptomatic individuals
Sijbrandij 2007	Types of participants: treatment study of symptomatic individuals
Skogstad 2015	Types of participants: treatment study of symptomatic individuals
Turpin 2005	Types of participants: single session intervention
van Emmerik 2008	Types of participants: treatment study of symptomatic individuals
Wagner 2007	Types of participants: treatment study of symptomatic individuals
Wu 2014	Types of participants: treatment study of symptomatic individuals
Zatzick 2004	Types of participants: treatment study of symptomatic individuals
Zatzick 2013	Types of participants: treatment study of symptomatic individuals
Zatzick 2015	Types of participants: treatment study of symptomatic individuals
Öst unpublished	Types of participants: treatment study of symptomatic individuals

## **Characteristics of studies awaiting assessment** [ordered by study ID]

### Cairns 2018

Methods	Pilot RCT
Participants	Family members of individuals admitted to ICU.
Interventions	Sensation Awareness Focused Training vs control group
Outcomes	HADS, IES, PSS
Notes	



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Methods	RCT
Participants	ICU patients treated for cardiorespiratory failure.
Interventions	A self-directed mobile app-based mindfulness programme vs a therapist-led telephone-based mindfulness programme vs a web-based crillness education programme.
Outcomes	PTSS, PHQ-9, GAD-7, EQ-5D, PHQ-15, CAMS-R, Brief COPE
Notes	

### **Guest 2018**

Methods	RCT
Participants	Adult survivor of an MVA who lodged an insurance claim within 4 months of their MVA
Interventions	E-mailed CBT with telephone support vs healthy lifestyle interventions
Outcomes	DASS and IES-R
Notes	Some participants in this study were recruited beyond 3 months. Therefore, inclusion of this study will depend on whether it will be possible to access outcome data for those recruited before 3 months only. The reported results are preliminary and the trial is ongoing.

## Kilpatrick 1984

Methods	Not stated
Participants	Victims of rape
Interventions	Repeated assessment vs delayed assessment vs therapy – Brief Behavioral Intervention
Outcomes	Not stated
Notes	To date, it has not been possible to access this publication

### **Kredentser 2018**

Methods	Pilot RCT
Participants	ICU survivors
Interventions	Usual care, ICU diary, psychoeducation, or ICU diary and psychoeducation
Outcomes	IES-R, HADS
Notes	



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Methods	RCT
Participants	Parents of children admitted to PICU, parental anxiety, depression, global health, post-traumatic stress, care satisfaction, decision regret and team collaboration
Interventions	PICU supports, a navigator-based communication intervention vs informational brochure
Outcomes	Family satisfaction
Notes	Information based on conference abstract only.

## Navidian 2017

Methods	RCT
Participants	Mothers after still birth
Interventions	Grief counselling vs usual care
Outcomes	PPQ
Notes	

### Nielsen 2019

Methods	RCT
Participants	ICU patients and their relatives
Interventions	Family authored diaries vs usual care
Outcomes	PTSS-14
Notes	

### **Rodin 2019**

Methods	Phase II RCT
Participants	Adults with acute leukaemia
Interventions	EASE plus usual care vs usual care
Outcomes	SASRQ, BDI, Physical symptom burden as measured by the MSAS, Brief Pain Inventory, FACITSp
Notes	



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Methods	RCT
Participants	Pregnant women with foetal abnormalities requiring pregnancy termination
Interventions	Family-support programme vs routine care
Outcomes	Family APGAR, IES-R, Edinburgh Postnatal Depression Scale
Notes	

#### Wade 2019

Methods	RCT
Participants	Newly diagnosed or recently relapsed adults with acute myeloid leukaemia or acute lymphocytic leukaemia within 1 month of inpatient admission to hospital.
Interventions	EASE for acute leukaemia plus usual care vs usual care alone
Outcomes	SASRQ
Notes	SASRQ evaluates symptoms of acute stress disorder but can reasonably be consider a proxy measure for PTSD.

#### Wendlandt 2019

Methods	RCT
Participants	Surrogate decision makers of patients with chronic critical illness
Interventions	Structured family informational and emotional support meetings led by palliative care specialists vs usual care.
Outcomes	IES-R
Notes	

BDI: Beck Depression Inventory; Brief COPE: Brief Coping Orientation to Problems Experienced Inventory; CAMS-R: Cognitive and Affective Mindfulness Scale - Revised; CBT: cognitive behavioural therapy; DASS: Depression Anxiety Stress Scales; EASE: Emotion And Symptom-focused Engagement; FACITSp: Functional Assessment of Chronic Illness Therapy-Spiritual Well-Being Scale; Family APGAR: Family Adaptation Partnership Growth Affection and Resolve index; GAD-7: Generalised Anxiety Disorder Assessment; HADS: Hospital Anxiety and Depression Scale; ICU: intensive care unit; IES: Impact of Events Scale; IES-R: Impact of Events Scale - Revised; MSAS: Memorial Symptom Assessment Scale; MVA: motor vehicle accident; PHQ-9: 9-item Patient Health Questionnaire; PICU: paediatric intensive care unit; PPQ: Prenatal Posttraumatic Stress Questionnaire; PSS: Perceived Stress Scale; PTSD: post-traumatic stress disorder; PTSS: post-traumatic stress syndrome; RCT: randomised controlled trial; SASRQ: Stanford Acute Stress Reaction Questionnaire.

**Characteristics of ongoing studies** [ordered by study ID]



SRCTN39318241	
Trial name or title	Randomized controlled study of Internet-based cognitive behavioural therapy for women having post-traumatic stress after childbirth
Methods	RCT
Participants	Women, aged ≥ 18 years with a negative birth experience or an immediate caesarean section or a major postpartum haemorrhage (or a combination of these)
Interventions	iCBT plus TAU vs TAU
Outcomes	Traumatic Event Scale, Edinburgh Postnatal Depression Scale, Satisfaction with Life Scale, Ways of Coping Questionnaire, Communication sub scale from the Evaluation and Nurturing Relationship Issues, Communication and Happiness, Revised Dyadic Adjustment Scale, Postpartum Bonding Questionnaire, EQ-5D 5D, 36-item Short Form Health Survey, Hospital Anxiety and Depression Scale
Starting date	28 April 2016
Contact information	Prof Agneta Skoog Svanberg: agneta.skoog_svanberg@kbh.uu.se
Notes	Recruitment begins around 8 weeks' postpartum.

### NCT02085512

Trial name or title	Prevention of PTSD III: neurocognitive training of emotional regulation
Methods	RCT
Participants	Adult survivors of traumatic events consecutively admitted to a general hospital emergency department
Interventions	Neurobehavioural training group vs control for specific neurocognitive tasks vs control for eventual beneficial effect of performing active computer games
Outcomes	CAPS-IV, PCL-5, SCID, PDI, BDI, neurocognitive measures
Starting date	13 March 2014
Contact information	Naomi Fine: nomsfine@gmail.com
Notes	

### NCT02591472

Trial name or title	An integrated-delivery-of-care approach to improve patient outcomes, safety, well-being after orthopaedic trauma
Methods	RCT
Participants	Patients aged 18–85 years; admitted with severe or multiple orthopaedic trauma (any major bone fractures that impairs mobility or participation in activities of daily living and self-care, or both); and have received or will receive ≥ 1 surgical procedure for their orthopaedic injuries.



NCT02591472 (Continued)	
Interventions	Integrated care, involving acute care therapies, postacute rehabilitation and follow-up clinic visits after discharge plus simultaneous psychosocial support vs usual care
Outcomes	Physical functioning through various means, PCL, BDI, STAI, Tampa Scale of Kinesiophobia-11
Starting date	29 October 2015
Contact information	Heather Vincent: vincehk@ortho.ufl.edu
Notes	

## NCT03438175

Trial name or title	Intensiva 2.0: improve the communication towards families of critically ill patients (Intensiva2)
Methods	Cluster RCT
Participants	Family members of individuals admitted to an intensive care unit
Interventions	Enhanced communication by brochure, website and posters plus TAU vs TAU
Outcomes	Comprehension of medical information, Hospital Anxiety and Depression Scale, Short Screening Scale for Symptoms of Post-traumatic Stress Disorder, PCL-5, Jefferson Scale for Physician Empathy, Maslach Burnout Inventory
Starting date	1 March 2018
Contact information	Giovanni Mistraletti: giovanni.mistraletti@unimi.it
Notes	

### NCT03496714

Trial name or title	Online psychoeducation for the prevention of PTSD
Methods	RCT
Participants	Adults who have experienced a Criteria A trauma within the past 30 days, as assessed by the Life Events Checklist for DSM-5
Interventions	Psychoeducation on safety behaviours and how to fade them vs psychoeducation only vs monitoring only
Outcomes	PSS-SR-5, PSSI-5, PCL-5, PTSD Safety Behavior Inventory, Posttraumatic Cognitions Inventory, Trauma Coping Self-Efficacy scale, Anxiety Sensitivity Index-3, BDI, AUDIT-C, PHQ-9, GAD-7
Starting date	12 April 2018
Contact information	Anna Foulser: afoulser@utexas.edu; Michael Telch: telch@austin.utexas.edu
Notes	



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Trial name or title	Effects of a neuroscience-based technique on post-traumatic stress disorder symptoms, inflammation, and survival in cancer patients announced of a palliative disease progression and their partners (NeuroPrevPTSD)
Methods	RCT
Participants	Adults who received in the last 7 days the diagnosis of metastatic incurable bladder, prostate, kidney, colorectal or sarcoma cancer
Interventions	Memory structuring intervention plus vagal breathing vs support and attention (usual care)
Outcomes	PCL-5, EQ-5D, Brief Approach/Avoidance Coping Questionnaire
Starting date	25 July 2018
Contact information	Yori Gidron: yori.gidron2@univ-lille3.fr; Georges-Michel Reich: M-Reich@o-lambret.fr
Notes	

#### Wells 2018

Trial name or title	Improving the effectiveness of psychological interventions for depression and anxiety in the cardiac rehabilitation pathway using group-based metacognitive therapy (PATHWAY Group MCT): study protocol for a randomised controlled trial
Methods	Multicentre RCT
Participants	Heart disease patients
Interventions	Group-based metacognitive therapy plus usual cardiac rehabilitation vs usual cardiac rehabilitation alone
Outcomes	HADS, IES-R, MCQ-30, EQ-5D-5L
Starting date	2015
Contact information	Adrian Wells: adrian.wells@manchester.ac.uk
Notes	

AUDIT-C: Alcohol Use Disorders Identification Test – C; BDI: Beck Depression Inventory; CAPS-IV: Clinician-Administered PTSD Scale; DSM-5: Diagnostic and Statistical Manual of Mental Disorders – 5th edition; EQ-5D; EuroQol; GAD-7: Generalised Anxiety Disorder Assessment; HADS: Hospital Anxiety and Depression Scale; iCBT: Internet-based cognitive behavioural therapy; IES-R: Impact of Events Scale – Revised; MCQ-30: Metacognitions Questionnaire 30; PCL-5: Post-Traumatic Stress Disorder Checklist; PDI: Peritraumatic Distress Inventory; PHQ-9: Patient Health Questionnaire; PSS-SR-5: Post-traumatic Stress Symptom Scale – Self Report; PSSI-5: Posttraumatic Stress Disorder Symptom Scale Interview for DSM-5; PTSD: post-traumatic stress disorder; RCT: randomised controlled trial; STAI: State-Trait Anxiety Inventory; TAU: treatment as usual.

### DATA AND ANALYSES



## Comparison 1. Any early psychological intervention versus waiting list/usual care

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 PTSD diagnosis post-treatment	5	556	Risk Ratio (M-H, Random, 95% CI)	1.06 [0.85, 1.32]
1.1 Brief individual trauma processing therapy vs treatment as usual (TAU)	3	262	Risk Ratio (M-H, Random, 95% CI)	1.10 [0.87, 1.40]
1.2 Self-guided Internet-based intervention vs TAU	1	228	Risk Ratio (M-H, Random, 95% CI)	0.91 [0.40, 2.06]
1.3 Brief dyadic cognitive-behavioural intervention vs TAU	1	66	Risk Ratio (M-H, Random, 95% CI)	0.59 [0.21, 1.61]
2 PTSD diagnosis 3–6 months	5	758	Risk Ratio (M-H, Random, 95% CI)	0.62 [0.41, 0.93]
2.1 Brief individual trauma processing therapy vs TAU	3	251	Risk Ratio (M-H, Random, 95% CI)	0.73 [0.44, 1.22]
2.2 Self-guided Internet-based intervention vs TAU	1	185	Risk Ratio (M-H, Random, 95% CI)	0.50 [0.18, 1.45]
2.3 Intensive care diaries vs delayed intensive care diaries	1	322	Risk Ratio (M-H, Random, 95% CI)	0.38 [0.17, 0.82]
3 PTSD diagnosis 7–12 months	1	132	Risk Ratio (M-H, Random, 95% CI)	0.94 [0.20, 4.49]
3.1 Self-guided Internet-based intervention vs TAU	1	132	Risk Ratio (M-H, Random, 95% CI)	0.94 [0.20, 4.49]
4 Dropouts from treatment	11	1154	Risk Ratio (M-H, Random, 95% CI)	1.34 [0.91, 1.95]
4.1 Brief individual trauma processing therapy vs TAU	5	571	Risk Ratio (M-H, Random, 95% CI)	1.06 [0.67, 1.68]
4.2 Brief dyadic therapy vs TAU	2	112	Risk Ratio (M-H, Random, 95% CI)	2.09 [0.76, 5.75]
4.3 Group therapy vs TAU	1	162	Risk Ratio (M-H, Random, 95% CI)	0.72 [0.27, 1.89]
4.4 Collaborative care vs TAU	1	34	Risk Ratio (M-H, Random, 95% CI)	0.75 [0.14, 3.94]
4.5 Brief IPT vs TAU	1	90	Risk Ratio (M-H, Random, 95% CI)	3.06 [1.39, 6.75]
4.6 Telephone-based CBT vs TAU	1	185	Risk Ratio (M-H, Random, 95% CI)	2.42 [0.79, 7.44]
5 Severity of PTSD symptoms post- treatment	9	1326	Std. Mean Difference (IV, Random, 95% CI)	-0.09 [-0.29, 0.12]
5.1 Brief individual trauma process- ing therapy vs TAU	4	465	Std. Mean Difference (IV, Random, 95% CI)	0.04 [-0.34, 0.42]
5.2 Self-guided Internet-based intervention vs TAU	1	300	Std. Mean Difference (IV, Random, 95% CI)	-0.38 [-0.61, -0.15]



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
5.3 Group counselling vs TAU	1	147	Std. Mean Difference (IV, Random, 95% CI)	-0.09 [-0.41, 0.24]
5.4 Collaborative care vs TAU	1	29	Std. Mean Difference (IV, Random, 95% CI)	-0.50 [-1.24, 0.25]
5.5 Intensive care diaries vs delayed access to intensive care diaries	1	330	Std. Mean Difference (IV, Random, 95% CI)	-0.22 [-0.44, -0.01]
5.6 Three-step early intervention vs TAU	1	55	Std. Mean Difference (IV, Random, 95% CI)	0.33 [-0.20, 0.86]
6 Severity of PTSD symptoms: 3–6 months	15	1921	Std. Mean Difference (IV, Random, 95% CI)	-0.10 [-0.22, 0.02]
6.1 Brief individual trauma process- ing therapy vs TAU	4	466	Std. Mean Difference (IV, Random, 95% CI)	-0.05 [-0.41, 0.30]
6.2 Self-guided Internet-based intervention vs TAU	1	300	Std. Mean Difference (IV, Random, 95% CI)	-0.27 [-0.50, -0.04]
6.3 Brief dyadic therapy vs TAU	2	103	Std. Mean Difference (IV, Random, 95% CI)	-0.41 [-0.81, -0.02]
6.4 Collaborative care vs TAU	1	26	Std. Mean Difference (IV, Random, 95% CI)	0.41 [-0.37, 1.19]
6.5 Brief interpersonal therapy (IPT) vs TAU	1	58	Std. Mean Difference (IV, Random, 95% CI)	0.10 [-0.42, 0.61]
6.6 Intensive care diaries vs delayed access to intensive care diaries	1	322	Std. Mean Difference (IV, Random, 95% CI)	0.0 [-0.22, 0.22]
6.7 Three-step early intervention vs TAU	1	55	Std. Mean Difference (IV, Random, 95% CI)	-0.08 [-0.61, 0.45]
6.8 Telephone-based cognitive behavioural therapy (CBT) vs TAU	1	185	Std. Mean Difference (IV, Random, 95% CI)	-0.20 [-0.49, 0.09]
6.9 Supported psychoeducation vs TAU	1	23	Std. Mean Difference (IV, Random, 95% CI)	-0.65 [-1.61, 0.30]
6.10 Communication facilitator in an intensive care setting vs usual care	1	168	Std. Mean Difference (IV, Random, 95% CI)	0.02 [-0.29, 0.32]
6.11 Nurse-led intensive care recovery programme vs TAU	1	215	Std. Mean Difference (IV, Random, 95% CI)	-0.02 [-0.29, 0.25]
7 Severity of PTSD symptoms: 7–12 months	4	765	Std. Mean Difference (IV, Random, 95% CI)	-0.09 [-0.32, 0.14]
7.1 Self-guided Internet-based intervention vs wait list/TAU	1	300	Std. Mean Difference (IV, Random, 95% CI)	0.0 [-0.23, 0.23]
7.2 Three-step early intervention vs TAU	1	55	Std. Mean Difference (IV, Random, 95% CI)	-0.18 [-0.71, 0.35]
7.3 Telephone-based CBT	1	185	Std. Mean Difference (IV, Random, 95% CI)	-0.38 [-0.67, -0.09]



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
7.4 Nurse-led intensive care recovery programme vs TAU	1	225	Std. Mean Difference (IV, Random, 95% CI)	0.12 [-0.14, 0.38]
8 Severity of depressive symptoms post-treatment	5	671	Std. Mean Difference (IV, Random, 95% CI)	-0.19 [-0.40, 0.01]
8.1 Brief individual trauma processing therapy vs TAU	1	137	Std. Mean Difference (IV, Random, 95% CI)	-0.46 [-0.80, -0.12]
8.2 Self-guided Internet-based intervention vs TAU	1	300	Std. Mean Difference (IV, Random, 95% CI)	-0.17 [-0.39, 0.06]
8.3 Collaborative care vs TAU	1	29	Std. Mean Difference (IV, Random, 95% CI)	-0.26 [-0.99, 0.48]
8.4 Group therapy vs TAU	1	147	Std. Mean Difference (IV, Random, 95% CI)	-0.22 [-0.55, 0.10]
8.5 Brief IPT vs TAU	1	58	Std. Mean Difference (IV, Random, 95% CI)	0.32 [-0.20, 0.84]
9 Severity of depressive symptoms at 3–6 months	7	1009	Std. Mean Difference (IV, Random, 95% CI)	-0.04 [-0.19, 0.10]
9.1 Brief IPT vs TAU	1	58	Std. Mean Difference (IV, Random, 95% CI)	0.0 [-0.52, 0.52]
9.2 Self-guided Internet-based intervention vs TAU	1	300	Std. Mean Difference (IV, Random, 95% CI)	-0.12 [-0.35, 0.10]
9.3 Collaborative care vs TAU	1	26	Std. Mean Difference (IV, Random, 95% CI)	0.86 [0.05, 1.68]
9.4 Supported psychoeducational intervention vs TAU	1	23	Std. Mean Difference (IV, Random, 95% CI)	-0.07 [1.00, 0.87]
9.5 Telephone-based CBT	1	185	Std. Mean Difference (IV, Random, 95% CI)	0.0 [0.0, 0.0]
9.6 Communication facilitator in an intensive care setting vs TAU	1	171	Std. Mean Difference (IV, Random, 95% CI)	-0.07 [-0.37, 0.23]
9.7 Nurse-led intensive care recovery programme vs TAU	1	246	Std. Mean Difference (IV, Random, 95% CI)	-0.04 [-0.29, 0.21]
10 Severity of depressive symptoms at 7–12 months	3	745	Std. Mean Difference (IV, Random, 95% CI)	0.01 [-0.14, 0.15]
10.1 Self-guided Internet-based intervention vs TAU	1	300	Std. Mean Difference (IV, Random, 95% CI)	0.06 [-0.17, 0.28]
10.2 Telephone-based CBT	1	185	Std. Mean Difference (IV, Random, 95% CI)	-0.16 [-0.45, 0.13]
10.3 Nurse-led intensive care recovery programme vs TAU	1	260	Std. Mean Difference (IV, Random, 95% CI)	0.06 [-0.18, 0.30]
11 Severity of anxiety symptoms post-treatment	3	358	Std. Mean Difference (IV, Random, 95% CI)	-0.41 [-0.98, 0.16]

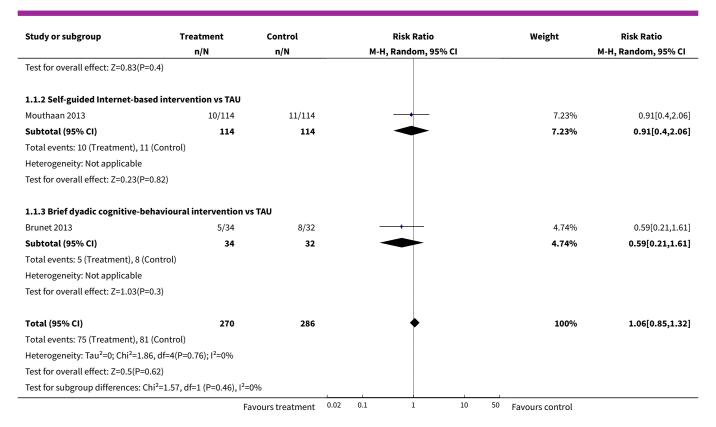


Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
11.1 Self-guided Internet-based intervention vs TAU	1	300	Std. Mean Difference (IV, Random, 95% CI)	-0.03 [-0.25, 0.20]
11.2 Brief dyadic therapy vs TAU	1	29	Std. Mean Difference (IV, Random, 95% CI)	-0.76 [-1.54, 0.02]
11.3 Brief IPT vs TAU	1	29	Std. Mean Difference (IV, Random, 95% CI)	-0.76 [-1.54, 0.02]
12 Severity of anxiety symptoms at 3–6 months	6	945	Std. Mean Difference (IV, Random, 95% CI)	-0.05 [-0.19, 0.10]
12.1 Supported psychoeducational intervention vs TAU	1	23	Std. Mean Difference (IV, Random, 95% CI)	-0.15 [-1.08, 0.78]
12.2 Self-guided Internet-based intervention vs TAU	1	300	Std. Mean Difference (IV, Random, 95% CI)	-0.09 [-0.32, 0.14]
12.3 Brief dyadic therapy vs TAU	1	29	Std. Mean Difference (IV, Random, 95% CI)	-0.76 [-1.54, 0.02]
12.4 Telephone-based CBT	1	185	Std. Mean Difference (IV, Random, 95% CI)	0.0 [0.0, 0.0]
12.5 Communication facilitator in an intensive care setting vs usual care	1	162	Std. Mean Difference (IV, Random, 95% CI)	0.04 [-0.27, 0.35]
12.6 Nurse-led intensive care recovery programme vs TAU	1	246	Std. Mean Difference (IV, Random, 95% CI)	0.03 [-0.22, 0.28]
13 Severity of anxiety symptoms at 7–12 months	3	746	Std. Mean Difference (IV, Random, 95% CI)	-0.04 [-0.27, 0.18]
13.1 Self-guided Internet-based intervention vs TAU	1	300	Std. Mean Difference (IV, Random, 95% CI)	0.09 [-0.13, 0.32]
13.2 Telephone-based CBT	1	185	Std. Mean Difference (IV, Random, 95% CI)	-0.30 [-0.59, -0.01]
13.3 Nurse-led intensive care recovery programme vs TAU	1	261	Std. Mean Difference (IV, Random, 95% CI)	0.04 [-0.21, 0.28]

Analysis 1.1. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 1 PTSD diagnosis post-treatment.

Study or subgroup	Treatment	Control			Risk Ratio			Weight	Risk Ratio
	n/N	n/N		M-	H, Random, 95% CI	l			M-H, Random, 95% CI
1.1.1 Brief individual trauma (TAU)	processing therapy vs tre	eatment as usual						-	
Gamble 2005	17/50	16/53			<del>-</del>			15.19%	1.13[0.64,1.98]
Marchand 2006	11/25	11/32			+-			11.33%	1.28[0.67,2.46]
Rothbaum 2012	32/47	35/55			<del></del>			61.52%	1.07[0.81,1.42]
Subtotal (95% CI)	122	140			<b>•</b>			88.04%	1.1[0.87,1.4]
Total events: 60 (Treatment),	62 (Control)								
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0	.27, df=2(P=0.88); I <sup>2</sup> =0%								
	F	avours treatment	0.02	0.1	1	10	50	Favours control	

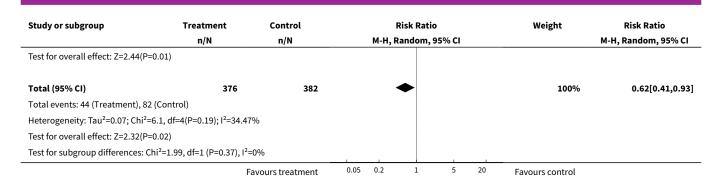




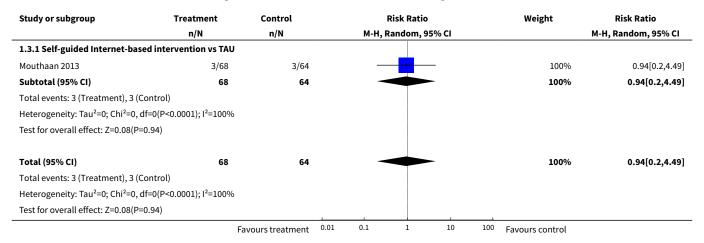
Analysis 1.2. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 2 PTSD diagnosis 3-6 months.

Study or subgroup	Treatment	Control	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Random, 95% CI		M-H, Random, 95% CI
1.2.1 Brief individual trauma proc	essing therapy vs TA	U			
Gamble 2005	3/50	9/53	<del></del>	9.11%	0.35[0.1,1.23]
Marchand 2006	10/25	11/32		22.6%	1.16[0.59,2.29]
Rothbaum 2012	18/42	32/49	-	37.59%	0.66[0.44,0.98]
Subtotal (95% CI)	117	134	<b>*</b>	69.29%	0.73[0.44,1.22]
Total events: 31 (Treatment), 52 (Co	ontrol)				
Heterogeneity: Tau <sup>2</sup> =0.09; Chi <sup>2</sup> =3.4,	df=2(P=0.18); I <sup>2</sup> =41.18	%			
Test for overall effect: Z=1.21(P=0.23	3)				
1.2.2 Self-guided Internet-based i	ntervention vs TAU				
Mouthaan 2013	5/97	9/88	<del></del>	12.03%	0.5[0.18,1.45]
Subtotal (95% CI)	97	88		12.03%	0.5[0.18,1.45]
Total events: 5 (Treatment), 9 (Cont	rol)				
Heterogeneity: Not applicable					
Test for overall effect: Z=1.27(P=0.2)	)				
1.2.3 Intensive care diaries vs dela	ayed intensive care d	iaries			
Jones 2010	8/162	21/160	<del></del>	18.68%	0.38[0.17,0.82]
Subtotal (95% CI)	162	160		18.68%	0.38[0.17,0.82]
Total events: 8 (Treatment), 21 (Cor	ntrol)				
Heterogeneity: Not applicable					





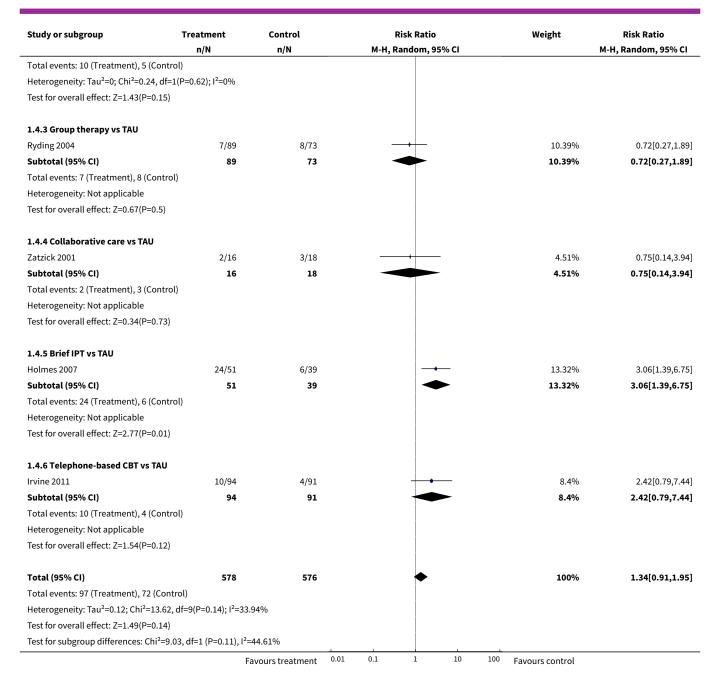
## Analysis 1.3. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 3 PTSD diagnosis 7-12 months.



Analysis 1.4. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 4 Dropouts from treatment.

Study or subgroup	Treatment	Control	Risk Ratio	Weight	Risk Ratio
	n/N	n/N	M-H, Random, 95% CI		M-H, Random, 95% CI
1.4.1 Brief individual trauma p	processing therapy vs TAI	J			
Brom 1993	11/68	20/83	<b>-</b> •	16.12%	0.67[0.35,1.3]
Gamble 2005	0/50	0/53			Not estimable
Marchand 2006	8/33	10/42	<del></del>	12.97%	1.02[0.45,2.29]
Rothbaum 2012	22/69	13/68	<del>  • -</del>	17.71%	1.67[0.92,3.03]
Ryding 1998	3/53	3/52		5.03%	0.98[0.21,4.64]
Subtotal (95% CI)	273	298	<b>*</b>	51.82%	1.06[0.67,1.68]
Total events: 44 (Treatment), 46	(Control)				
Heterogeneity: Tau <sup>2</sup> =0.06; Chi <sup>2</sup> =	4.05, df=3(P=0.26); I <sup>2</sup> =25.8	4%			
Test for overall effect: Z=0.26(P=	:0.79)				
1.4.2 Brief dyadic therapy vs T	AU				
Brunet 2013	5/37	3/37	<del>-   •</del>	6.29%	1.67[0.43,6.47]
Kazak 2005	5/18	2/20	+	5.27%	2.78[0.61,12.59]
Subtotal (95% CI)	55	57		11.56%	2.09[0.76,5.75]
	Fa	avours treatment 0.0	1 0.1 1 10 10	00 Favours control	

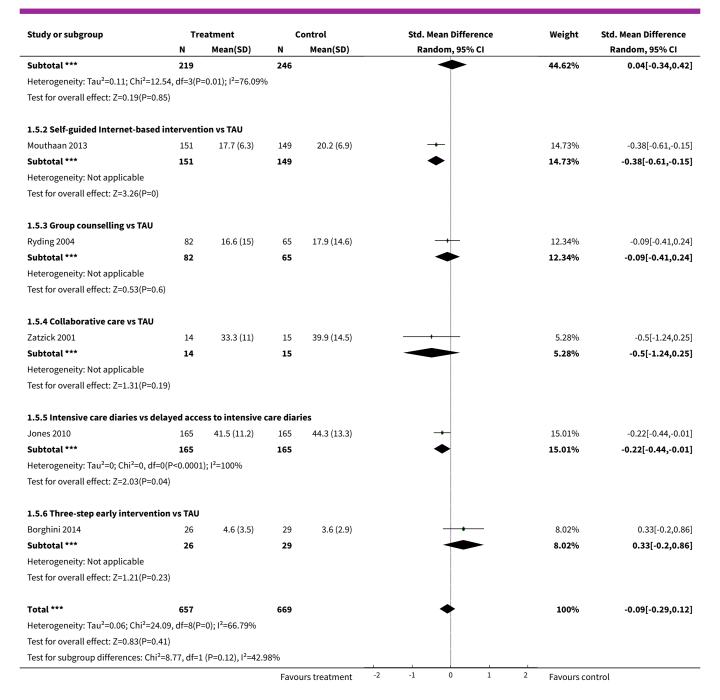




Analysis 1.5. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 5 Severity of PTSD symptoms post-treatment.

Study or subgroup	Tre	Treatment		Control		Std. Mean Difference		Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI				Random, 95% CI
1.5.1 Brief individual traum	a processing th	erapy vs TAU							
Brom 1993	68	21.9 (15.9)	83	17.4 (15)		<del> </del>		12.42%	0.29[-0.03,0.61]
Gamble 2005	49	4.8 (3.7)	53	5.5 (3)		<del>-+</del>		10.85%	-0.19[-0.58,0.2]
Marchand 2006	33	19.4 (17.5)	42	12 (13.9)		<del>  • </del>		9.31%	0.47[0,0.93]
Rothbaum 2012	69	19.1 (15.2)	68	24.5 (14)		<u>→</u>		12.04%	-0.37[-0.71,-0.03]
			Favo	urs treatment	-2	-1 0 1	2	Favours cont	rol





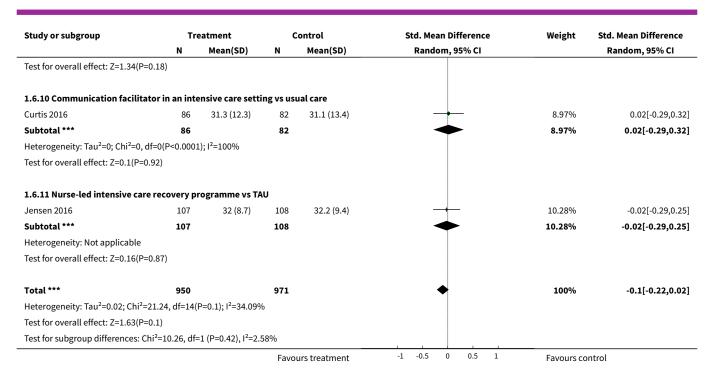
Analysis 1.6. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 6 Severity of PTSD symptoms: 3-6 months.

Study or subgroup	Tre	atment	c	ontrol	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI		Random, 95% CI
1.6.1 Brief individual trauma	a processing th	erapy vs TAU					
Brom 1993	68	9.3 (11)	83	7.4 (11.3)	+-	8.35%	0.17[-0.15,0.49]
Gamble 2005	50	2.5 (2.4)	53	3.8 (3.6)	<del></del>	6.45%	-0.41[-0.81,-0.02]
Marchand 2006	33	15.5 (15.7)	42	9.7 (13.8)		5.07%	0.39[-0.07,0.85]
			Favo	urs treatment	-1 -0.5 0 0.5 1	Favours co	ontrol



Study or subgroup		eatment		ontrol	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI		Random, 95% CI
Rothbaum 2012	69	15.5 (16.5)	68	20.3 (14.8)	<del></del>	7.86%	-0.31[-0.65,0.03
Subtotal ***	220		246			27.74%	-0.05[-0.41,0.3
Heterogeneity: Tau <sup>2</sup> =0.09; Chi <sup>2</sup> =10.8	7, df=3(P	=0.01); I <sup>2</sup> =72.4%					
Test for overall effect: Z=0.29(P=0.77)	)						
1.6.2 Self-guided Internet-based in	tervent	ion vs TAU					
Mouthaan 2013	151	14.3 (7.2)	149	16.8 (10.9)		12.02%	-0.27[-0.5,-0.04
Subtotal ***	151		149		•	12.02%	-0.27[-0.5,-0.04
Heterogeneity: Not applicable							
Test for overall effect: Z=2.33(P=0.02)	)						
1.6.3 Brief dyadic therapy vs TAU							
Brunet 2013	37	21.7 (18.9)	37	29.9 (18.4)	+	5.06%	-0.43[-0.89,0.03
Kazak 2005	11	28.3 (13.3)	18	35.4 (22.4)	<del></del>	2.21%	-0.35[-1.11,0.4
Subtotal ***	48		55			7.26%	-0.41[-0.81,-0.02
Heterogeneity: Tau²=0; Chi²=0.03, df	=1(P=0.8	6); I <sup>2</sup> =0%					
Test for overall effect: Z=2.05(P=0.04)	)						
1.6.4 Collaborative care vs TAU							
Zatzick 2001	12	42.5 (18.2)	14	36.1 (11.7)	<del>-  </del>	2.09%	0.41[-0.37,1.19
Subtotal ***	12		14			2.09%	0.41[-0.37,1.19
Heterogeneity: Tau $^2$ =0; Chi $^2$ =0, df=0(	P<0.0001	L); I <sup>2</sup> =100%					
Test for overall effect: Z=1.04(P=0.3)							
1.6.5 Brief interpersonal therapy (	PT) vs T	AU					
Holmes 2007	27	38 (19.1)	31	36.2 (17)		4.24%	0.1[-0.42,0.61
Subtotal ***	27		31			4.24%	0.1[-0.42,0.61
Heterogeneity: Not applicable							
Test for overall effect: Z=0.37(P=0.71)	)						
1.6.6 Intensive care diaries vs dela	yed acce	ess to intensive	care diar	ies			
Jones 2010	162	24 (12.2)	160	24 (11.6)	+	12.44%	0[-0.22,0.22
Subtotal ***	162		160		<b>*</b>	12.44%	0[-0.22,0.22
Heterogeneity: Not applicable							
Test for overall effect: Not applicable							
1.6.7 Three-step early intervention	vs TAU						
Borghini 2014	26	3.3 (3.4)	29	3.6 (3.3)		4.07%	-0.08[-0.61,0.45
Subtotal ***	26		29			4.07%	-0.08[-0.61,0.45
Heterogeneity: Not applicable							
Test for overall effect: Z=0.31(P=0.76)	)						
1.6.8 Telephone-based cognitive b	ehaviou	ral therapy (CB	Γ) vs TAU				
Irvine 2011	94	0.6 (0.7)	91	0.7 (0.5)	-+-	9.45%	-0.2[-0.49,0.09
Subtotal ***	94		91			9.45%	-0.2[-0.49,0.09
Heterogeneity: Not applicable							
Test for overall effect: Z=1.33(P=0.18	)						
1.6.9 Supported psychoeducation	vs TAU						
Als 2015	17	19.5 (7.5)	6	25.8 (13.8)	+	1.44%	-0.65[-1.61,0.3
Subtotal ***	17	. ,	6	· <i>'</i>		1.44%	-0.65[-1.61,0.3
					-		,

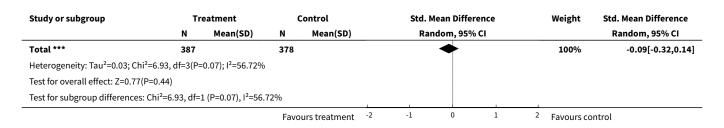




Analysis 1.7. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 7 Severity of PTSD symptoms: 7–12 months.

Study or subgroup	Treatment		Control		Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI		Random, 95% CI
1.7.1 Self-guided Internet-based in	terventi	on vs wait list/	ΓΑυ				
Mouthaan 2013	151	13 (11.3)	149	13 (10)	-	31.69%	0[-0.23,0.23]
Subtotal ***	151		149		<b>*</b>	31.69%	0[-0.23,0.23]
Heterogeneity: Not applicable							
Test for overall effect: Not applicable							
1.7.2 Three-step early intervention	vs TAU						
Borghini 2014	26	2.7 (2.6)	29	3.2 (2.7)		13.25%	-0.18[-0.71,0.35]
Subtotal ***	26		29			13.25%	-0.18[-0.71,0.35]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.67(P=0.51)							
1.7.3 Telephone-based CBT							
Irvine 2011	94	0.5 (0.5)	91	0.7 (0.7)		26.37%	-0.38[-0.67,-0.09]
Subtotal ***	94		91		•	26.37%	-0.38[-0.67,-0.09]
Heterogeneity: Not applicable							
Test for overall effect: Z=2.57(P=0.01)							
1.7.4 Nurse-led intensive care reco	very pro	gramme vs TAl	J				
Jensen 2016	116	31.3 (9.5)	109	30.1 (9.7)	-	28.7%	0.12[-0.14,0.38]
Subtotal ***	116		109		<b>*</b>	28.7%	0.12[-0.14,0.38]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.89(P=0.37)							
			Favoi	ırs treatment -2	-1 0 1	<sup>2</sup> Favours co	ontrol





Analysis 1.8. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 8 Severity of depressive symptoms post-treatment.

Study or subgroup	Tre	eatment	C	Control	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI		Random, 95% CI
1.8.1 Brief individual trauma proce	ssing th	erapy vs TAU					
Rothbaum 2012	69	15 (14.1)	68	21.4 (13.4)		22.62%	-0.46[-0.8,-0.12]
Subtotal ***	69		68			22.62%	-0.46[-0.8,-0.12]
Heterogeneity: Not applicable							
Test for overall effect: Z=2.64(P=0.01)							
1.8.2 Self-guided Internet-based in	terventi	ion vs TAU					
Mouthaan 2013	151	3.6 (2.5)	149	4.1 (3.4)	-	34.33%	-0.17[-0.39,0.06]
Subtotal ***	151		149			34.33%	-0.17[-0.39,0.06]
Heterogeneity: Not applicable							
Test for overall effect: Z=1.44(P=0.15)							
1.8.3 Collaborative care vs TAU							
Zatzick 2001	14	21.1 (12.2)	15	24.6 (14.3)	+	6.96%	-0.26[-0.99,0.48]
Subtotal ***	14		15			6.96%	-0.26[-0.99,0.48]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.68(P=0.49)							
1.8.4 Group therapy vs TAU							
Ryding 2004	82	6.4 (4.9)	65	7.5 (4.9)	<del></del>	23.71%	-0.22[-0.55,0.1]
Subtotal ***	82		65			23.71%	-0.22[-0.55,0.1]
Heterogeneity: Not applicable							
Test for overall effect: Z=1.34(P=0.18)							
1.8.5 Brief IPT vs TAU							
Holmes 2007	27	13.1 (12.4)	31	9.8 (7.8)	+	12.38%	0.32[-0.2,0.84]
Subtotal ***	27		31			12.38%	0.32[-0.2,0.84]
Heterogeneity: Not applicable							
Test for overall effect: Z=1.2(P=0.23)					İ		
Total ***	343		328		•	100%	-0.19[-0.4,0.01]
Heterogeneity: Tau <sup>2</sup> =0.02; Chi <sup>2</sup> =6.16,	df=4(P=	0.19); I <sup>2</sup> =35.08%					
Test for overall effect: Z=1.83(P=0.07)							
Test for subgroup differences: Chi <sup>2</sup> =6	.16, df=1	(P=0.19), I <sup>2</sup> =35.	08%				

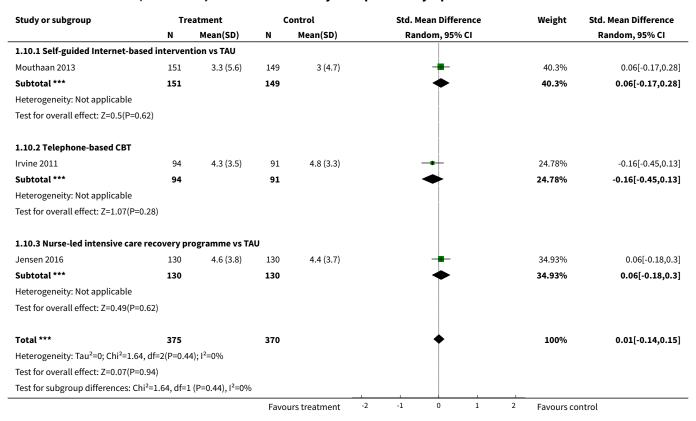


Analysis 1.9. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 9 Severity of depressive symptoms at 3-6 months.

Study or subgroup	Tre	eatment	С	ontrol	Std. Mean Difference	Weight	Std. Mean Difference
, , ,	N	Mean(SD)	N	Mean(SD)	Random, 95% CI	J	Random, 95% CI
1.9.1 Brief IPT vs TAU							·
Holmes 2007	27	12.3 (11.2)	31	12.3 (11.5)		7.66%	0[-0.52,0.52]
Subtotal ***	27		31		•	7.66%	0[-0.52,0.52]
Heterogeneity: Not applicable							
Test for overall effect: Not applicable							
1.9.2 Self-guided Internet-based in	terventi	ion vs TAU					
Mouthaan 2013	151	3.5 (3.1)	149	3.9 (3.4)	-	35.48%	-0.12[-0.35,0.1]
Subtotal ***	151		149		•	35.48%	-0.12[-0.35,0.1]
Heterogeneity: Not applicable							
Test for overall effect: Z=1.05(P=0.29)							
1.9.3 Collaborative care vs TAU							
Zatzick 2001	12	29.6 (17.1)	14	16.7 (11.8)	<del></del>	3.15%	0.86[0.05,1.68]
Subtotal ***	12		14			3.15%	0.86[0.05,1.68]
Heterogeneity: Not applicable							
Test for overall effect: Z=2.08(P=0.04)							
1.9.4 Supported psychoeducationa	l interve	ention vs TAU					
Als 2015	17	2.8 (3.2)	6	3 (4.4)	<del></del>	2.4%	-0.07[-1,0.87]
Subtotal ***	17		6			2.4%	-0.07[-1,0.87]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.14(P=0.89)							
1.9.5 Telephone-based CBT							
Irvine 2011	94	0 (0)	91	0 (0)			Not estimable
Subtotal ***	94		91				Not estimable
Heterogeneity: Not applicable							
Test for overall effect: Not applicable							
1.9.6 Communication facilitator in	an inten	sive care settin	g vs TAU				
Curtis 2016	87	5.1 (5.7)	84	5.5 (5.8)	-	21.47%	-0.07[-0.37,0.23]
Subtotal ***	87		84		<b>*</b>	21.47%	-0.07[-0.37,0.23]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.45(P=0.65)							
1.9.7 Nurse-led intensive care recov	ery pro	gramme vs TAL	ı				
Jensen 2016	123	5 (3.9)	123	5.2 (4)	+	29.85%	-0.04[-0.29,0.21]
Subtotal ***	123		123		<b>*</b>	29.85%	-0.04[-0.29,0.21]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.3(P=0.76)							
Total ***	511		498		•	100%	-0.04[-0.19,0.1]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =5.3, df=5	(P=0.38)	); I <sup>2</sup> =5.7%					
Test for overall effect: Z=0.59(P=0.55)							
Test for subgroup differences: Chi <sup>2</sup> =5.	2 df-1/	D-0 20) 12-E 704					



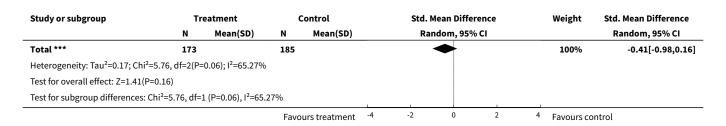
Analysis 1.10. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 10 Severity of depressive symptoms at 7–12 months.



Analysis 1.11. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 11 Severity of anxiety symptoms post-treatment.

Study or subgroup	Tre	eatment	c	ontrol	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI		Random, 95% CI
1.11.1 Self-guided Internet-based in	iterven	tion vs TAU					
Mouthaan 2013	151	4.6 (7.1)	149	4.8 (7.3)	<b>#</b>	47.43%	-0.03[-0.25,0.2]
Subtotal ***	151		149		<b>*</b>	47.43%	-0.03[-0.25,0.2]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.24(P=0.81)							
1.11.2 Brief dyadic therapy vs TAU							
Kazak 2005	11	36.4 (10.5)	18	46.8 (14.9)	-	26.29%	-0.76[-1.54,0.02]
Subtotal ***	11		18		•	26.29%	-0.76[-1.54,0.02]
Heterogeneity: Not applicable							
Test for overall effect: Z=1.9(P=0.06)							
1.11.3 Brief IPT vs TAU							
Holmes 2007	11	36.3 (10.5)	18	46.8 (14.9)	<del></del>	26.29%	-0.76[-1.54,0.02]
Subtotal ***	11		18		•	26.29%	-0.76[-1.54,0.02]
Heterogeneity: Not applicable							
Test for overall effect: Z=1.91(P=0.06)							
				L		I	
			Favo	urs treatment -4	-2 0 2	<sup>4</sup> Favours co	ontrol





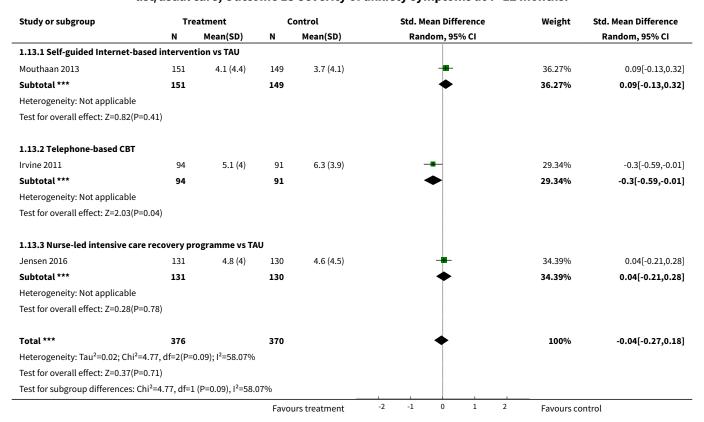
Analysis 1.12. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 12 Severity of anxiety symptoms at 3-6 months.

Study or subgroup	Tre	eatment	C	ontrol	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI		Random, 95% CI
1.12.1 Supported psychoeducation	nal inter	vention vs TAU					
Als 2015	17	6.5 (4.3)	6	7.2 (4.8)		2.43%	-0.15[-1.08,0.78]
Subtotal ***	17		6			2.43%	-0.15[-1.08,0.78]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.32(P=0.7	5)						
1.12.2 Self-guided Internet-based	l interven	tion vs TAU					
Mouthaan 2013	151	4 (3.1)	149	4.3 (3.4)	-	39.59%	-0.09[-0.32,0.14]
Subtotal ***	151		149		•	39.59%	-0.09[-0.32,0.14]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.79(P=0.4	3)						
1.12.3 Brief dyadic therapy vs TAI	J						
Kazak 2005	11	36.4 (10.5)	18	46.8 (14.9)	<del></del>	3.48%	-0.76[-1.54,0.02]
Subtotal ***	11		18			3.48%	-0.76[-1.54,0.02]
Heterogeneity: Not applicable							
Test for overall effect: Z=1.9(P=0.06	)						
1.12.4 Telephone-based CBT							
Irvine 2011	94	0 (0)	91	0 (0)			Not estimable
Subtotal ***	94		91				Not estimable
Heterogeneity: Not applicable							
Test for overall effect: Not applicab	le						
1.12.5 Communication facilitator	in an inte	ensive care sett	ing vs usi	ual care			
Curtis 2016	85	4.6 (5.6)	77	4.4 (4.9)	+	21.76%	0.04[-0.27,0.35]
Subtotal ***	85		77		<b>*</b>	21.76%	0.04[-0.27,0.35]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.24(P=0.8	1)						
1.12.6 Nurse-led intensive care re	covery pi	ogramme vs TA	<b>U</b>				
Jensen 2016	123	5.4 (3.7)	123	5.2 (4.5)	-	32.74%	0.03[-0.22,0.28]
Subtotal ***	123		123		<b>*</b>	32.74%	0.03[-0.22,0.28]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.27(P=0.7	9)				į		
Total ***	481		464		•	100%	-0.05[-0.19,0.1]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =4.07, d	f=4(P=0.4	)· I <sup>2</sup> =1 84%			İ		



Study or subgroup	Tr	Treatment		Control	Std. Mean Difference					Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)		Ran	dom, 95	% CI			Random, 95% CI
Test for overall effect: Z=0.63(P=0	).53)										
Test for subgroup differences: Ch	i <sup>2</sup> =4.07, df=	1 (P=0.4), I <sup>2</sup> =1.84 <sup>0</sup>	%								
			Fav	ours treatment	-2	-1	0	1	2	Favours cor	ntrol

# Analysis 1.13. Comparison 1 Any early psychological intervention versus waiting list/usual care, Outcome 13 Severity of anxiety symptoms at 7–12 months.



### Comparison 2. Any early psychological intervention versus active control condition

Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
1 Dropouts from treatment	2	425	Risk Ratio (M-H, Random, 95% CI)	1.61 [1.11, 2.34]
1.1 Brief individual trauma processing therapy vs parenting support	1	262	Risk Ratio (M-H, Random, 95% CI)	1.78 [1.12, 2.84]
1.2 Guided self-help vs physical educational intervention	1	163	Risk Ratio (M-H, Random, 95% CI)	1.33 [0.71, 2.50]



Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
2 Severity of PTSD symptoms post-treatment	2	392	Std. Mean Difference (IV, Random, 95% CI)	0.13 [-0.06, 0.33]
2.1 Brief individual trauma processing therapy vs parenting support	1	217	Std. Mean Difference (IV, Random, 95% CI)	0.14 [-0.13, 0.41]
2.2 Guided self-help vs physical educational intervention	1	175	Std. Mean Difference (IV, Random, 95% CI)	0.13 [-0.17, 0.42]
3 Severity of PTSD symptoms at 3–6 months	4	465	Std. Mean Difference (IV, Random, 95% CI)	-0.02 [-0.31, 0.26]
3.1 Brief individual trauma processing therapy vs supportive listening	2	51	Std. Mean Difference (IV, Random, 95% CI)	-0.54 [-1.42, 0.34]
3.2 Brief individual trauma processing therapy vs parenting support	1	239	Std. Mean Difference (IV, Random, 95% CI)	0.06 [-0.19, 0.31]
3.3 Guided self-help vs physical educational intervention	1	175	Std. Mean Difference (IV, Random, 95% CI)	0.13 [-0.16, 0.43]
4 Severity of PTSD symptoms at 7–12 months	1	200	Mean Difference (IV, Random, 95% CI)	1.27 [-0.60, 3.14]
4.1 Brief individual trauma processing therapy vs parenting support	1	200	Mean Difference (IV, Random, 95% CI)	1.27 [-0.60, 3.14]
5 Severity of depression symptoms post- treatment	2	392	Std. Mean Difference (IV, Random, 95% CI)	0.12 [-0.08, 0.32]
5.1 Brief individual trauma processing therapy vs parenting support	1	217	Std. Mean Difference (IV, Random, 95% CI)	0.06 [-0.21, 0.33]
5.2 Guided self-help vs physical educational intervention	1	175	Std. Mean Difference (IV, Random, 95% CI)	0.19 [-0.11, 0.49]
6 Severity of depressive symptoms at 3–6 months	2	409	Std. Mean Difference (IV, Random, 95% CI)	0.04 [-0.16, 0.23]
6.1 Brief individual trauma processing therapy vs parenting support	1	234	Std. Mean Difference (IV, Random, 95% CI)	0.09 [-0.17, 0.34]
6.2 Guided self-help vs physical educational intervention	1	175	Std. Mean Difference (IV, Random, 95% CI)	-0.04 [-0.33, 0.26]
7 Severity of depressive symptoms at 7–12 months	1	198	Mean Difference (IV, Random, 95% CI)	0.79 [-0.66, 2.24]
7.1 Brief individual trauma processing therapy vs parenting support	1	198	Mean Difference (IV, Random, 95% CI)	0.79 [-0.66, 2.24]
8 Severity of anxiety symptoms post-treatment	2	392	Std. Mean Difference (IV, Random, 95% CI)	0.08 [-0.12, 0.28]

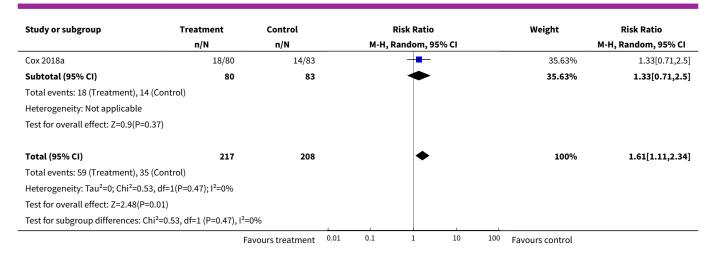


Outcome or subgroup title	No. of studies	No. of partici- pants	Statistical method	Effect size
8.1 Brief individual trauma processing therapy vs parenting support	1	217	Std. Mean Difference (IV, Random, 95% CI)	0.10 [-0.17, 0.37]
8.2 Guided self-help vs physical educational intervention	1	175	Std. Mean Difference (IV, Random, 95% CI)	0.05 [-0.24, 0.35]
9 Severity of anxiety symptoms at 3–6 months	2	414	Std. Mean Difference (IV, Random, 95% CI)	0.00 [-0.19, 0.19]
9.1 Brief individual trauma processing therapy vs parenting support	1	239	Std. Mean Difference (IV, Random, 95% CI)	0.03 [-0.22, 0.28]
9.2 Guided self-help vs physical educational intervention	1	175	Std. Mean Difference (IV, Random, 95% CI)	-0.04 [-0.33, 0.26]
10 Severity of anxiety symptoms at 7–12 months	1	199	Mean Difference (IV, Random, 95% CI)	-0.07 [-1.58, 1.44]
10.1 Brief individual trauma processing therapy vs parenting support	1	199	Mean Difference (IV, Random, 95% CI)	-0.07 [-1.58, 1.44]
11 General functioning post-treatment	1	218	Mean Difference (IV, Random, 95% CI)	-0.02 [-0.05, 0.01]
11.1 Brief individual trauma processing therapy vs parenting support	1	218	Mean Difference (IV, Random, 95% CI)	-0.02 [-0.05, 0.01]
12 General functioning at 3–6 months	1	239	Mean Difference (IV, Random, 95% CI)	-0.03 [-0.06, 0.00]
12.1 Brief individual trauma processing therapy vs parenting support	1	239	Mean Difference (IV, Random, 95% CI)	-0.03 [-0.06, 0.00]
13 General functioning at 7–12 months	1	199	Mean Difference (IV, Random, 95% CI)	0.01 [-0.03, 0.05]
13.1 Brief individual trauma processing therapy vs parenting support	1	199	Mean Difference (IV, Random, 95% CI)	0.01 [-0.03, 0.05]

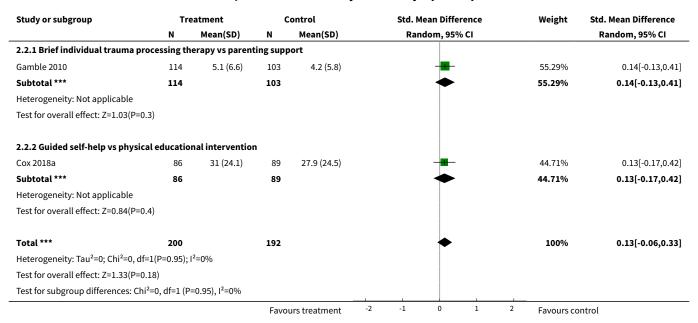
# Analysis 2.1. Comparison 2 Any early psychological intervention versus active control condition, Outcome 1 Dropouts from treatment.

Study or subgroup	Treatment	Control			Risk Ratio			Weight	Risk Ratio
	n/N	n/N		М-Н,	Random, 95%	6 CI			M-H, Random, 95% CI
2.1.1 Brief individual trauma	processing therapy vs par	enting support							
Gamble 2010	41/137	21/125			-			64.37%	1.78[1.12,2.84]
Subtotal (95% CI)	137	125			•			64.37%	1.78[1.12,2.84]
Total events: 41 (Treatment), 2	1 (Control)								
Heterogeneity: Not applicable									
Test for overall effect: Z=2.42(P	2=0.02)								
2.1.2 Guided self-help vs phy	sical educational interven	tion							
	Fa	avours treatment	0.01	0.1	1	10	100	Favours control	





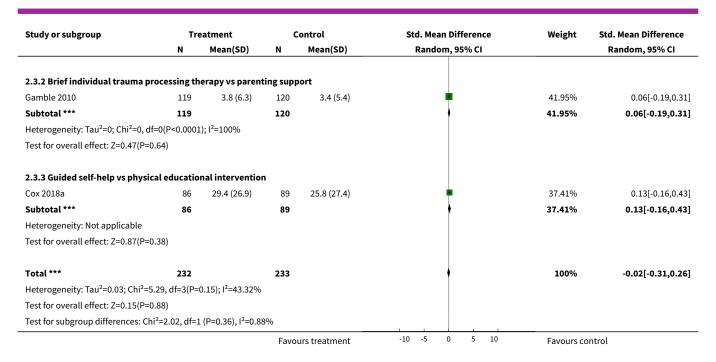
Analysis 2.2. Comparison 2 Any early psychological intervention versus active control condition, Outcome 2 Severity of PTSD symptoms post-treatment.



Analysis 2.3. Comparison 2 Any early psychological intervention versus active control condition, Outcome 3 Severity of PTSD symptoms at 3–6 months.

Study or subgroup	Tre	atment	c	ontrol	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI		Random, 95% CI
2.3.1 Brief individual trauma	a processing th	erapy vs suppo	rtive list	ening			
Gidron 2001	8	8.1 (9.1)	9	18.5 (9.1)	+	6.77%	-1.09[-2.13,-0.05]
Gidron 2007	19	9.5 (3.4)	15	10.1 (3.4)	+	13.87%	-0.17[-0.85,0.51]
Subtotal ***	27		24		•	20.64%	-0.54[-1.42,0.34]
Heterogeneity: Tau <sup>2</sup> =0.22; Chi	i <sup>2</sup> =2.09, df=1(P=0	0.15); I <sup>2</sup> =52.23%					
Test for overall effect: Z=1.2(P	P=0.23)						
			Favo	urs treatment	-10 -5 0 5 10	Favours co	ontrol





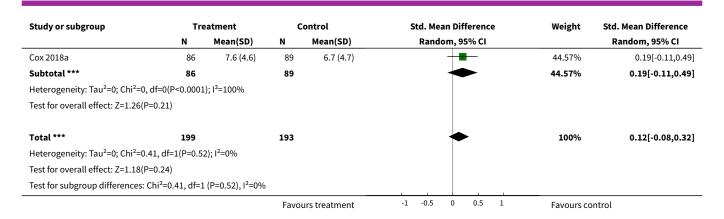
Analysis 2.4. Comparison 2 Any early psychological intervention versus active control condition, Outcome 4 Severity of PTSD symptoms at 7–12 months.

Study or subgroup	Tre	eatment	С	ontrol	Mean	Difference	Weight	Mean Difference
	N	Mean(SD)	N	Mean(SD)	Rand	om, 95% CI		Random, 95% CI
2.4.1 Brief individual trauma proce	ssing th	erapy vs paren	ting supp	ort				
Gamble 2010	96	5.4 (6.8)	104	4.1 (6.7)		-	100%	1.27[-0.6,3.14]
Subtotal ***	96		104			•	100%	1.27[-0.6,3.14]
Heterogeneity: Not applicable								
Test for overall effect: Z=1.33(P=0.18)	)							
Total ***	96		104			•	100%	1.27[-0.6,3.14]
Heterogeneity: Not applicable								
Test for overall effect: Z=1.33(P=0.18)	)							
			Favoi	urs treatment	-10 -5	0 5 10	Favours contr	ol

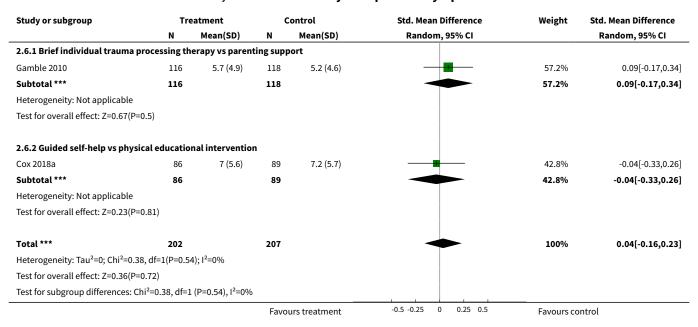
Analysis 2.5. Comparison 2 Any early psychological intervention versus active control condition, Outcome 5 Severity of depression symptoms post-treatment.

Study or subgroup	Tre	eatment	c	ontrol	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI		Random, 95% CI
2.5.1 Brief individual trauma	processing th	erapy vs paren	ting supp	oort			
Gamble 2010	113	5.4 (4.7)	104	5.1 (4.1)	_	55.43%	0.06[-0.21,0.33]
Subtotal ***	113		104		<b>*</b>	55.43%	0.06[-0.21,0.33]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.45(P	=0.65)						
2.5.2 Guided self-help vs phys	sical educatio	nal interventio	n				
			Favo	urs treatment	-1 -0.5 0 0.5 1	Favours co	ontrol





Analysis 2.6. Comparison 2 Any early psychological intervention versus active control condition, Outcome 6 Severity of depressive symptoms at 3–6 months.



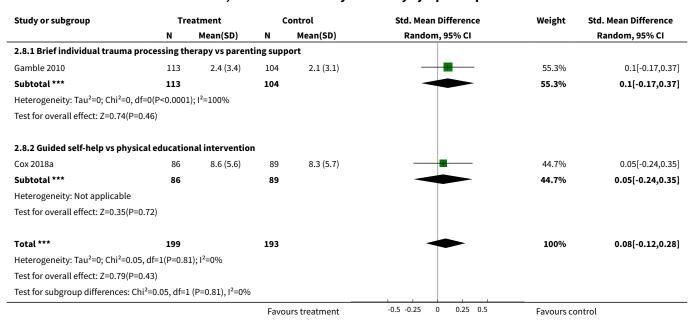
Analysis 2.7. Comparison 2 Any early psychological intervention versus active control condition, Outcome 7 Severity of depressive symptoms at 7–12 months.

Study or subgroup	Tre	eatment	С	ontrol		Mea	an Difference	1	Weight	Mean Difference
	N	Mean(SD)	N	Mean(SD)		Rar	idom, 95% CI			Random, 95% CI
2.7.1 Brief individual trauma pro	cessing th	erapy vs paren	ting supp	ort						
Gamble 2010	96	6.6 (4.9)	102	5.8 (5.5)					100%	0.79[-0.66,2.24]
Subtotal ***	96		102						100%	0.79[-0.66,2.24]
Heterogeneity: Not applicable										
Test for overall effect: Z=1.07(P=0.2	29)									
Total ***	96		102						100%	0.79[-0.66,2.24]
Heterogeneity: Not applicable								1		
			Favo	ırs treatment	-4	-2	0 2	4 F	avours control	



Study or subgroup	Treatment Con		Control	Mean Difference				Weight	Mean Difference		
	N	Mean(SD)	N	Mean(SD)		Ran	dom, 95%	% CI			Random, 95% CI
Test for overall effect: Z=1.07(P=0.29)										,	
			Favo	ours treatment	-4	-2	0	2	4	Favours contr	ol

## Analysis 2.8. Comparison 2 Any early psychological intervention versus active control condition, Outcome 8 Severity of anxiety symptoms post-treatment.



Analysis 2.9. Comparison 2 Any early psychological intervention versus active control condition, Outcome 9 Severity of anxiety symptoms at 3–6 months.

Study or subgroup	Tre	eatment	С	ontrol	Std. Mean Difference	Weight	Std. Mean Difference
	N	Mean(SD)	N	Mean(SD)	Random, 95% CI		Random, 95% CI
2.9.1 Brief individual trauma ¡	processing th	erapy vs paren	ting supp	ort			
Gamble 2010	119	2.6 (4.1)	120	2.5 (4.2)		57.74%	0.03[-0.22,0.28]
Subtotal ***	119		120			57.74%	0.03[-0.22,0.28]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.22(P=	=0.82)						
2.9.2 Guided self-help vs phys	ical educatio	nal interventio	n				
Cox 2018a	86	8.3 (5.6)	89	8.5 (5.7)		42.26%	-0.04[-0.33,0.26]
Subtotal ***	86		89			42.26%	-0.04[-0.33,0.26]
Heterogeneity: Not applicable							
Test for overall effect: Z=0.23(P=	=0.81)						
Total ***	205		209		•	100%	0[-0.19,0.19]
Heterogeneity: Tau <sup>2</sup> =0; Chi <sup>2</sup> =0.1	l, df=1(P=0.75	); I <sup>2</sup> =0%					
Test for overall effect: Z=0.02(P=	=0.99)						
Test for subgroup differences: C	Chi <sup>2</sup> =0.1, df=1	(P=0.75), I <sup>2</sup> =0%					
			Favoi	urs treatment	-0.5 -0.25 0 0.25 0.5	Favours co	ontrol



# Analysis 2.10. Comparison 2 Any early psychological intervention versus active control condition, Outcome 10 Severity of anxiety symptoms at 7–12 months.

Study or subgroup	Tre	eatment	c	ontrol		Mea	n Difference	Weight	Mean Difference
	N	Mean(SD)	N	Mean(SD)		Ran	dom, 95% CI		Random, 95% CI
2.10.1 Brief individual trauma proce	ssing t	herapy vs parei	nting sup	port					
Gamble 2010	95	3.3 (4.7)	104	3.4 (6.1)				100%	-0.07[-1.58,1.44]
Subtotal ***	95		104			~		100%	-0.07[-1.58,1.44]
Heterogeneity: Not applicable									
Test for overall effect: Z=0.09(P=0.93)									
Total ***	95		104			•		100%	-0.07[-1.58,1.44]
Heterogeneity: Not applicable									
Test for overall effect: Z=0.09(P=0.93)									
			Favoi	ırs treatment	-4	-2	0 2	4 Favours cont	rol

Analysis 2.11. Comparison 2 Any early psychological intervention versus active control condition, Outcome 11 General functioning post-treatment.

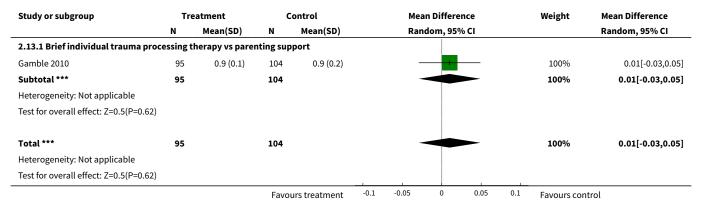
Study or subgroup	Tre	eatment	С	ontrol		Me	an Difference		Weight	Mean Difference
	N	Mean(SD)	N	Mean(SD)		Rar	ndom, 95% CI			Random, 95% CI
2.11.1 Brief individual trauma proce	essing t	herapy vs parei	nting sup	port						
Gamble 2010	114	0.9 (0.1)	104	0.9 (0.1)					100%	-0.02[-0.05,0.01]
Subtotal ***	114		104				•		100%	-0.02[-0.05,0.01]
Heterogeneity: Not applicable										
Test for overall effect: Z=1.23(P=0.22)										
Total ***	114		104				•		100%	-0.02[-0.05,0.01]
Heterogeneity: Not applicable										
Test for overall effect: Z=1.23(P=0.22)										
			Favoi	urs treatment	-0.2	-0.1	0 0.1	0.2	Favours contro	

Analysis 2.12. Comparison 2 Any early psychological intervention versus active control condition, Outcome 12 General functioning at 3–6 months.

Study or subgroup	Tre	eatment	C	Control		Mean	Differ	ence		Weight	Mean Difference
	N	Mean(SD)	N	Mean(SD)		Rande	om, 95	% CI			Random, 95% CI
2.12.1 Brief individual trauma proc	essing t	herapy vs pare	nting sup	port							
Gamble 2010	119	0.9 (0.1)	120	0.9 (0.1)			+			100%	-0.03[-0.06,0]
Subtotal ***	119		120				1			100%	-0.03[-0.06,0]
Heterogeneity: Not applicable											
Test for overall effect: Z=1.84(P=0.07)											
Total ***	119		120							100%	-0.03[-0.06,0]
Heterogeneity: Not applicable											
Test for overall effect: Z=1.84(P=0.07)											
			Favo	urs treatment	-2	-1	0	1	2	Favours contro	l



## Analysis 2.13. Comparison 2 Any early psychological intervention versus active control condition, Outcome 13 General functioning at 7–12 months.



#### **APPENDICES**

### **Appendix 1. CCMDCTR core MEDLINE search**

The search strategy listed below is the weekly Ovid Medline search which was used to inform the Group's specialised register to June 2016. It is based on a list of terms for all conditions within the scope of the Cochrane Common Mental Disorders Group plus a sensitive RCT filter.

## Ovid MEDLINE search strategy, used to inform the Cochrane Common Mental Disorders Group's Specialised Register

A weekly search alert based on condition + RCT filter only

1. [MeSH Headings]:

eating disorders/ or anorexia nervosa/ or binge-eating disorder/ or bulimia nervosa/ or female athlete triad syndrome/ or pica/ or hyperphagia/ or bulimia/ or self-injurious behavior/ or self mutilation/ or suicide/ or suicidal ideation/ or suicide, attempted/ or mood disorders/ or affective disorders, psychotic/ or bipolar disorder/ or cyclothymic disorder/ or depressive disorder/ or depressive, postpartum/ or depressive disorder, major/ or depressive disorder, treatment-resistant/ or dysthymic disorder/ or seasonal affective disorder/ or neurotic disorders/ or depression/ or adjustment disorders/ or exp antidepressive agents/ or anxiety disorder/ or agoraphobia/ or neurocirculatory asthenia/ or obsessive-compulsive disorder/ or obsessive hoarding/ or panic disorder/ or phobic disorders/ or stress disorders, traumatic/ or combat disorders/ or stress disorders, post-traumatic/ or stress disorders, traumatic, acute/ or anxiety/ or anxiety, castration/ or koro/ or anxiety, separation/ or panic/ or exp anti-anxiety agents/ or somatoform disorders/ or body dysmorphic disorders/ or conversion disorder/ or hypochondriasis/ or neurasthenia/ or hysteria/ or munchausen syndrome by proxy/ or munchausen syndrome/ or fatigue syndrome, chronic/ or obsessive behavior/ or compulsive behavior/ or behavior, addictive/ or impulse control disorders/ or firesetting behavior/ or gambling/ or trichotillomania/ or stress, psychological/ or burnout, professional/ or sexual dysfunctions, psychological/ or vaginismus/ or Anhedonia/ or Affective Symptoms/ or \*Mental Disorders/

### 2. [Title/ Author Keywords]:

(eating disorder\* or anorexia nervosa or bulimi\* or binge eat\* or (self adj (injur\* or mutilat\*)) or suicide\* or suicidal or parasuicid\* or mood disorder\* or affective disorder\* or bipolar i or bipolar ii or (bipolar and (affective or disorder\*)) or mania or manic or cyclothymic\* or depression or depressive or dysthymi\* or neurotic or neurosis or adjustment disorder\* or antidepress\* or anxiety disorder\* or agoraphobia or obsess\* or compulsi\* or panic or phobi\* or ptsd or posttrauma\* or post trauma\* or combat or somatiform or somatifation or medical\* unexplained or body dysmorphi\* or conversion disorder or hypochondria\* or neurastheni\* or hysteria or munchausen or chronic fatigue\* or gambling or trichotillomania or vaginismus or anhedoni\* or affective symptoms or mental disorder\* or mental health).ti,kf.

3. [RCT filter]:

(controlled clinical trial.pt. or randomized controlled trial.pt. or (randomi#ed or randomi#ation).ab,ti. or randomly.ab. or (random\* adj3 (administ\* or allocat\* or assign\* or class\* or control\* or determine\* or divide\* or distribut\* or expose\* or fashion or number\* or place\* or recruit\* or subsitut\* or treat\*)).ab. or placebo\*.ab,ti. or drug therapy.fs. or trial.ab,ti. or groups.ab. or (control\* adj3 (trial\* or study or studies)).ab,ti. or ((singl\* or doubl\* or tripl\* or trebl\*) adj3 (blind\* or mask\* or dummy\*)).mp. or clinical trial, phase ii/ or clinical trial, phase ii/ or randomized controlled trial/ or pragmatic clinical trial/ or (quasi adj (experimental or random\*)).ti,ab. or ((waitlist\* or wait\* list\* or treatment as usual or TAU) adj3 (control or group)).ab.)

4. (1 and 2 and 3)



Records were screened for reports of RCTs within the scope of the Cochrane Common Mental Disorders Group. Secondary reports of RCTs were tagged to the appropriate study record.

Similar weekly search alerts were also conducted on OVID Embase and PsycINFO, using relevant subject headings (controlled vocabularies) and search syntax, appropriate to each resource.

### Appendix 2. Database searches (March 2018/2019)

Date of search: 3 March 2018 Date limits: 2014 onwards

Database hits:

- MEDLINE (1742)
- Embase (3319)
- CENTRAL (2028)
- PsycINFO (1449)
- PILOTS (879)

Total=9417

Duplicates removed=4620 Studies screened for RCTs=4797

Records excluded=3632

RCT records identified=1165

### **Databases: CENTRAL**

Host: Wiley interface

Data Parameters: Cochrane Central Register of Controlled Trials: Issue 2 of 12, February 2018

Date Searched: Monday, March 3rd 2018

Searched by: Chris Cooper

Hits: 2028 IDSearchHits

#1 MeSH descriptor: [Stress Disorders, Post-Traumatic] this term only (1492)

#2 (PTSD or ((posttrauma\* or post-trauma\* or post trauma\*) near/3 (stress\* or disorder\* or psych\* or symptom\*)) or acute stress disorder\* or combat disorder\* or war neuros\*) (5065)

#3 (((acute or traumatic) near/1 stress\*) and (expos\* or psyc\*)) (1525)

#4 (traumatised near/1 (victim\* or survivor\*)) 2

#5 (traumatized near/1 (victim\* or survivor\*)) 4

#6 (trauma\* near/2 (event\* or memor\* or flashback\* or nightmare\*)) 553

#7 ((trauma\* or posttrauma\* or post-trauma\* or victim\* or survivor\*) and (exposure near/3 (therap\* or psychotherap\* or training or counsel\*))) 417

#8 MeSH descriptor: [Crisis Intervention] this term only 166

#9 (critical incident near/1 (stress or debrief\* or de-brief\*)) 24

#10 (debriefing or de-briefing) 328

#11 (crisis intervention\* or CISD) 1003

#12 ((stress or group\* or psychological or crisis) near/3 (debrief\* or de-brief\*)) 107

#13 (trauma\* near/2 (event\* or memor\* or flashback\* or nightmare\*)) 553

#14 (EMDR or (eye movement desensitization and reprocessing)) 225

#15 (EMDR or (eye movement desensitisation and reprocessing)) 197

#16 (#1 or #2 or #3 or #4 or #5 or #6 or #7 or #8 or #9 or #10 or #11 or #12 or #13 or #14 or #15)

Publication Year from 2014 to 2018 (2893)

File: VO1 CENTRAL n2028.txt

## Databases: Ovid MEDLINE(R) Epub Ahead of Print, In-Process & Other Non-Indexed Citations, Ovid MEDLINE(R) Daily and Ovid MEDLINE(R) 1946 to Present

Host: OVID

Data Parameters: 1946-Current

Date Searched: Monday, March 3rd 2018

Searched by: Chris

Hits: 1742 #SearchesResults

1 Stress Disorders, Post-Traumatic/ 27503

2 (PTSD or ((posttrauma\* or post-trauma\* or post trauma\*) adj3 (stress\* or disorder\* or psych\* or symptom?)) or acute stress disorder\* or combat disorder\* or war neuros\*).ti,ab,kf. 31111



- 3 (((acute or traumatic) adj stress\*) and (expos\* or psyc\*)).ti,ab,kf. 10567
- 4 (traumati#ed adj (victim? or survivor?)).ti,ab,kf. 34
- 5 (trauma\* adj2 (event? or memor\* or flashback\* or nightmare?)).ti,ab,kf. 8174
- 6 ((trauma\* or post-trauma\* or victim\* or survivor?) and (exposure adj3 (therap\* or psychotherap\* or training or counsel\*))).ti,ab,kf,hw. 901
- 7 Crisis Intervention/ 5457
- 8 (critical incident adj (stress or debrief\* or de-brief\*)).ti,ab,kf. 223
- 9 (debriefing or de-briefing).ti,kf. 577
- 10 (crisis intervention? or CISD).ti,ab,kf.1744
- 11 ((stress or group? or psychological or crisis) adj3 (debrief\* or de-brief\*)).ti,ab,kf. 406
- 12 (trauma\* adj2 (event? or memor\* or flashback\* or nightmare?)).ti,kf. 1150
- 13 (EMDR or (eye movement desensiti#ation and reprocessing)).ti,ab,kf,sh. 510
- 14 (1 or 2 or 3 or 4 or 5 or 6 or 7 or 8 or 9 or 10 or 11 or 12 or 13) 52168
- 15 randomized controlled trial.pt. 454849
- 16 controlled clinical trial.pt. 92204
- 17 randomized.ab. 404382
- 18 placebo.ab. 186843
- 19 clinical trials as topic.sh. 182777
- 20 randomly.ab. 285994
- 21 trial.ti. 178689
- 22 (15 or 16 or 17 or 18 or 19 or 20 or 21) 1136215
- 23 (14 and 22) 4000
- 24 (2014\* or 2015\* or 2016\* or 2017\* or 2018\*).yr,dt,ed,ep. 5444042
- 25 (23 and 24) 1742

### **Databases: Embase**

Host: OVID

Data Parameters: 1974 to 2018 March 02 Date Searched: Monday, March 3rd 2018

Searched by: Chris Cooper

Hits: 3319

**#Searches Results** 

- 1 posttraumatic stress disorder/ 48854
- 2 "trauma and stressor related disorders" / 34962
- 3 combat disorders/ 26663
- 4 psychological trauma/ 5351
- 5 stress disorders, post-traumatic/ 16743
- 6 stress disorders, traumatic, acute/751
- 7 (PTSD or ((posttrauma\* or post-trauma\* or post trauma\*) adj3 (stress\* or disorder\* or psych\* or symptom?)) or acute stress disorder\* or combat disorder\* or war neuros\*).ti,ab,kw. 39945
- 8 (((acute or traumatic) adj stress\*) and (expos\* or psyc\*)).ti,ab,kw. 15122
- 9 (traumati#ed adj (victim? or survivor?)).ti,ab,kw. 51
- 10 (trauma\* adj2 (event? or memor\* or flashback\* or nightmare?)).ti,ab,kw. 10514
- 11 (EMDR or (eye movement desensiti#ation and reprocessing)).ti,kw. 527
- 12 ((trauma\* or posttrauma\* or victim\* or survivor?) and (exposure adj3 (therap\* or psychotherap\* or training or counsel\*))).ti,ab,kw. 1096
- 13 (critical incident adj (stress or debrief\* or de-brief\*)).ti,ab,kw. 275
- 14 (debriefing or de-briefing).ti,ab,kw. 4133
- 15 (crisis intervention? or CISD).ti,ab,kw. 2273
- 16 ((stress or group? or psychological or crisis) adj3 (debrief\* or de-brief\*)).ti,ab,kw. 602
- 17 (trauma\* adj2 (event? or memor\* or flashback\* or nightmare?)).ti,ab,kw. 10514
- $18 \ (1 \ or \ 2 \ or \ 3 \ or \ 4 \ or \ 5 \ or \ 6 \ or \ 7 \ or \ 8 \ or \ 9 \ or \ 10 \ or \ 11 \ or \ 12 \ or \ 13 \ or \ 14 \ or \ 15 \ or \ 16 \ or \ 17) \ 74063$
- 19 crossover-procedure/ or double-blind procedure/ or randomized controlled trial/ or single-blind procedure/ or (random\* or factorial\* or crossover\* or cross over\* or placebo\* or (doubl\* adj blind\*) or (singl\* adj blind\*) or assign\* or allocat\* or volunteer\*).tw. 1970074 20 (18 and 19) 7601
- 21 (2014\* or 2015\* or 2016\* or 2017\* or 2018\*).yr,dc. 7084132
- 22 (20 and 21) 3319

### **Databases: PsycINFO**

Host: OVID

Data Parameters: 1806 to February Week 4 2018 Date Searched: Monday, March 3rd 2018



Searched by: Chris

Hits: 1449

**#Searches Results** 

1 posttraumatic stress disorder/ or complex ptsd/ or desnos/ or acute stress disorder/ or combat experience/ or "debriefing (psychological)"/ or emotional trauma/ or post-traumatic stress/ or exp stress reactions/ or traumatic neurosis/ 50806

2 exp disasters/8186

3 (PTSD or ((posttrauma\* or post-trauma\* or post trauma\*) adj3 (stress\* or disorder\* or psych\* or symptom?)) or acute stress disorder\* or combat disorder\* or war neuros\*).ti,ab. 38985

4 (((acute or traumatic) adj stress\*) and (expos\* or psyc\*)).ti,ab. 16755

5 (traumati#ed adj (victim? or survivor?)).ti,ab. 68

6 (trauma\* adj2 (event? or memor\* or flashback\* or nightmare?)).ti,ab. 11819

7 (EMDR or (eye movement desensiti#ation and reprocessing)).ti,ab. 1640

8 ((trauma\* or post-trauma\* or victim\* or survivor?) and (exposure adj3 (therap\* or psychotherap\* or training or counsel\*))).ti,ab. 1086

9 crisis intervention/3314

10 (critical incident adj (stress or debrief\* or de-brief\*)).ti,ab. 443

11 (debriefing or de-briefing).ti,ab. 2186

12 (crisis intervention? or CISD).ti,ab. 3505

13 ((stress or group? or psychological or crisis) adj3 (debrief\* or de-brief\*)).ti,ab. 596

14 (trauma\* adj2 (event? or memor\* or flashback\* or nightmare?)).ti,ab. 11819

15 (1 or 2 or 3 or 4 or 5 or 6 or 7 or 8 or 9 or 10 or 11 or 12 or 13 or 14) 80813

16 clinical trials.sh. 10820

17 (randomi#ed or randomi#ation or randomi#ing).ti,ab,id. 72509

18 (RCT or at random or (random\* adj3 (assign\* or allocat\* or control\* or crossover or cross-over or design\* or divide\* or division or number))).ti,ab,id. 82020

19 (control\* and (trial or study or group) and (placebo or waitlist\* or wait\* list\* or ((treatment or care) adj2 usual))).ti,ab,id,hw.25590

20 ((single or double or triple or treble) adj2 (blind\* or mask\* or dummy)).ti,ab,id. 24054

21 trial.ti. 25583

22 placebo.ti,ab,id,hw. 37267

23 treatment outcome.md. 18762

24 treatment effectiveness evaluation.sh. 21858

25 mental health program evaluation.sh. 2028

26 (16 or 17 or 18 or 19 or 20 or 21 or 22 or 23 or 24 or 25) 169119

27 (15 and 26) 4124

28 (2014\* or 2015\* or 2016\* or 2017\* or 2018\*).yr,dc,mo. 782907

29 (27 and 28) 1449

### **Database: PILOTS: Published International Literature On Traumatic Stress**

Host: Pro Quest

Data Parameters: 1871-Current

Date Searched: Monday, March 3rd 2018

Searched by: Chris

Hits: 879 Search Strategy

Set#: S1 Searched for: ti((posttrauma\* near/4 (stress\* or disorder\* or psych\* or symptom\*))) OR ab((posttrauma\* near/4 (stress\* or disorder\* or psych\* or symptom\*))) Results: 16999\*

Set#: S2 Searched for: ti((post-trauma\* near/4 (stress\* or disorder\* or psych\* or symptom\*))) OR ab((post-trauma\* near/4 (stress\* or disorder\* or psych\* or symptom\*))) Results: 6647°

Set#: S3 Searched for: ti((post trauma\* near/4 (stress\* or disorder\* or psych\* or symptom\*))) OR ab((post trauma\* near/4 (stress\* or disorder\* or psych\* or symptom\*))) Results: 7214°

Set#: S4 Searched for: ti((PTSD or acute stress disorder\* or combat disorder\* or war neuros\*) ) OR ab((PTSD or acute stress disorder\* or combat disorder\* or war neuros\*)) Results: 30435\*

Set#: S5 Searched for: ti((((acute or traumatic) near/2 stress\*) and (expos\* or psyc\*)) ) OR ab((((acute or traumatic) near/2 stress\*) and (expos\* or psyc\*)) ) Results: 2341°

Set#: S6 Searched for: ti((traumatised near/2 (victim\* or survivor\*))) OR ab((traumatised near/2 (victim\* or survivor\*))) Results: 84°

Set#: S7 Searched for: ti((trauma\* near/3 (event\* or memor\* or flashback\* or nightmare\*))) OR ab((trauma\* near/3 (event\* or memor\* or flashback\* or nightmare\*))) Results: 6974°

Set#: S8 Searched for: ti(((trauma\* or posttrauma\* or post-trauma\* or victim\* or survivor\*) and (exposure near/4 (therap\* or psychotherap\* or training or counsel\*)))) OR ab(((trauma\* or posttrauma\* or post-trauma\* or victim\* or survivor\*) and (exposure near/4 (therap\* or psychotherap\* or training or counsel\*)))) Results: 787°

Sei#: S9 Searched for: ti((critical incident near/2 (stress or debrief\* or de-brief\*)) ) OR ab((critical incident near/2 (stress or debrief\* or de-brief\*)) ) Results: 385°



Set#: S10 Searched for: ti((debriefing or de-briefing)) OR ab((debriefing or de-briefing)) Results: 685°

Set#: S11 Searched for: ti((crisis intervention\* or CISD)) OR ab((crisis intervention\* or CISD)) Results: 784°

Set#: S12 Searched for: ti(((stress or group\* or psychological or crisis) near/4 (debrief\* or de-brief\*)) ) OR ab(((stress or group\* or psychological or crisis) near/4 (debrief\* or de-brief\*)) ) Results: 464°

Set#: S13 Searched for: ti((trauma\* near/3 (event\* or memor\* or flashback\* or nightmare\*))) OR ab((trauma\* near/3 (event\* or memor\* or flashback\* or nightmare\*))) Results: 6974°

Set#: S14 Searched for: ti((EMDR or (eye movement desensitisation and reprocessing))) OR ab((EMDR or (eye movement desensitisation and reprocessing))) Results: 888°

Set#: S15 Searched for: ti((EMDR or (eye movement desensitiZation and reprocessing))) OR ab((EMDR or (eye movement desensitiZation and reprocessing))) Results: 888°

Set#: S16 Searched for: (s1 or s2 or s3 or s4 or s5 or s6 or s7 or s8 or s9 or s10 or s11 or s12 or s13 or s14 or s15)

Results: 36840\*

Set#: S17 Searched for: MAINSUBJECT.EXACT("Randomized Clinical Trial") Results: 1210° Set#: S18 Searched for: ab((randomized or randomised or placebo or randomly)) Results: 2931°

Set#: S19 Searched for: ti(trial) Results: 784°

Set#: S20 Searched for: (S17 or S18 or S19) Results: 3226° Set#: S21 Searched for: S16 and s20 Results: 2654°

Set#: S22 Searched for: (S16 and s20) AND pd(20140101-20180301) Results: 879° \* Duplicates are removed from your search, but included in your result count.

### PTSD update search (15 March 2019):

- CLib:CENTRAL (Issue 3 of 12, March 2019, date limited 2018 onwards), n=514 (116 of these are from ClinicalTrials.gov)
- Ovid MEDLINE (2018 to 15-Mar-2019), n=599
- Ovid Embase (2018 to 15-Mar-2019), n=1035
- Ovid PsycINFO (2018 to 15-Mar-2019), n=445
- Proquest PTSDpubs, (2018-03-01 to 2019-03-15) n=197

Total=2790
Duplicates removed, n=1178
Records to screen, n=1612
RCTs, n=781
Reviews, n=157
Irrelevant, n=674

### WHAT'S NEW

Date	Event	Description
7 August 2019	New citation required and conclusions have changed	Review updated. New studies added.
7 August 2019	New search has been performed	We updated the searches on 3 March 2018; we identified 16 new studies.

### HISTORY

Protocol first published: Issue 1, 2008 Review first published: Issue 3, 2009

<sup>°</sup> Duplicates are removed from your search and from your result count.



Date	Event	Description
24 February 2010	Amended	Minor changes to contact details of three authors (including contact author)
17 February 2010	Amended	Search strategy amended; review link updated; contact author's email address updated; co-author's name and title corrected
1 February 2009	Amended	Converted to new review format.
1 October 2008	Amended	The decision was taken to split the research question proposed in the original protocol 'Multiple session early psychological interventions for prevention and treatment of post-traumatic stress disorder' into two reviews (one on prevention, and one on treatment). The title of this review changed at that time. The original protocol remains on the Cochrane Library until such time as the review on treatment is published

### CONTRIBUTIONS OF AUTHORS

NPR: writing of the protocol and review. Undertook quality assessment and data entry.

NJK: commentary on the protocol and review. Undertook quality assessment and recording of data.

JK: commentary on the protocol and review. Undertook quality assessment and recording of data.

LR: writing of the review.

CL: screening of titles and abstracts.

JIB: offered supervision of the protocol development and commentary on the protocol and review. Writing of the 'Discussion' section of the review.

### **DECLARATIONS OF INTEREST**

NPR: none.

NJK: none.

JK: none.

LR: none.

CL: none.

JIB: none.

### SOURCES OF SUPPORT

## Internal sources

• No sources of support provided, UK.

### **External sources**

• National Institute for Health Research (NIHR), UK.

LR is funded by Cochrane Infrastructure funding to the Common Mental Disorders Cochrane Review Group

### DIFFERENCES BETWEEN PROTOCOL AND REVIEW

In the previous version of this review, we reported severity of PTSD symptoms as measured by a clinician-administered measure or a self-report measure separately. Although clinician-administered measures are considered to be the 'gold standard' in the traumatic stress field



many studies only report outcomes for self-report measures. Therefore, we used both clinician-administered and self-report measures as an index of PTSD severity. When a study used both types of outcome, we gave priority to clinician-administered measures.

Previously we evaluated risk of bias using a 19-item checklist that we developed for the review. Since publication of this review, the 'Risk of bias' tool has become the established means by which to evaluate study quality and we, therefore, used this in this review.

Previously we reported findings by the follow-up periods seven to nine months and 10 to 12 months separately. For this update, we decided to combine outcomes for these periods to seven to 12 months. Previously we did not specify the primary outcome period. For this review, we identified this as three to six months' post-trauma.

In the previous review, we planned to undertake separate ITT and completer-only analyses. In this review, we endeavoured to use ITT data wherever possible, but given the heterogeneity of interventions and the limited number of studies, we decided not to separate ITT and completer analyses in this version.

### INDEX TERMS

### **Medical Subject Headings (MeSH)**

\*Cognitive Behavioral Therapy [methods]; Desensitization, Psychologic; Psychotherapy [methods]; Quality of Life; Randomized Controlled Trials as Topic; Stress Disorders, Post-Traumatic [\*prevention & control]; Time Factors; Waiting Lists

#### MeSH check words

Humans