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Visuospatial computer game play after memory reactivation delivered three days after a traumatic film reduces the number of intrusive memories of the experimental trauma

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3

1 Abstract

Objective: The experience of intrusive memories is a core clinical symptom of posttraumatic 2 stress disorder (PTSD), and can be distressing in its own right. Notions of dual task interference 3 and reconsolidation-update mechanisms suggest novel approaches to target intrusive memories. 4 This study tested the hypothesis that a single-session cognitive intervention (memory reminder 5 6 task plus Tetris gameplay) would reduce the occurrence of experimental trauma memories even 7 when delivered 3 days post-trauma. Critically, this study tested effects against two control groups: Reminder-only, and reminder plus another computer game (a form of Quiz). 8 Methods: 86 healthy volunteers (59% female, age M = 24.35, SD = 4.59 years) watched a 9 trauma film and then recorded their intrusive memories in a diary for 3 days (pre-intervention). 10 11 They then returned to the lab. After presentation of visual reminder cues for the film plus a 10 minute wait period (memory reminder task), participants were randomized into one of three task 12 conditions (Tetris game play, Quiz game play, vs. reminder-only). They then kept the diary for a 13 further 3 days (post-intervention). 14 **Results:** As predicted, after the experimental manipulation, the reminder+*Tetris* group 15 experienced significantly fewer intrusions than the reminder-only group (d = 1.37). Further, the 16 reminder+*Tetris* group also experienced significantly fewer intrusions than the reminder+*Quiz* (d 17 = 0.65) group. Contrary to predictions, the reminder+Quiz group experienced significantly fewer 18 intrusions than the reminder-only group (d = 0.69). Prior to the experimental manipulation, there 19 was no significant difference between groups in number of intrusions. Recognition memory test 20 scores for facts of the trauma film after 6 days were comparable between groups. 21 22 Conclusions: We demonstrated that 3 days after experimental trauma (i.e. after memory consolidation) an intervention comprising a reminder task prior to a 15 minute cognitive 23 interference task (one of two computer games) led to a reduction in intrusion occurrence 24 compared to reminder only. We interpret and discuss our findings within the framework of 25 supposed reconsolidation-update mechanisms and competition for limited (visuospatial) working 26 27 memory resources. Should these effects hold true in clinical populations, this type of simple intervention approach could help contribute to reducing intrusive memories of trauma. 28 29

1 **1 Introduction**

Many people will be exposed to a traumatic event during their lifetime (WHO, 2013), a 2 significant minority of whom eventually go on to develop conditions such as posttraumatic stress 3 disorder (PTSD; Kessler et al., 2005). Although there are effective treatments for established 4 PTSD (administered at least 1 month after experiencing a traumatic event; NICE, 2005), only a 5 minority of patients receive such treatment, leading to suffering and societal costs (Kessler, 6 2000). It is possible that symptoms such as intrusive memories can develop after an even wider 7 8 range of events than those we currently classify as trauma (Mol et al., 2005). Further, evidencebased treatments to *prevent* the development of intrusive memories or future PTSD after a 9 traumatic event are lacking; new methods to ameliorate psychological distress soon after a 10 trauma are therefore needed (Rose et al., 2002; Roberts et al., 2010; Iyadurai et al., 2018). 11 A core clinical symptom (Kupfer & Regier, 2011) of PTSD and acute stress disorder (ASD) is 12 the "recurrent, involuntary and intrusive distressing memories of the traumatic event(s)" 13 (American Psychiatric Association, 2013). Intrusive memories are experienced as sensory-14 perceptual (typically visual) mental representations of the traumatic event(s) (Ehlers & Clark, 15 16 2000; Brewin, 2014). It has been argued that emotional memory is perceptual (Arntz et al., 2005) and likewise that mental imagery has a powerful impact on emotion (Holmes & Mathews, 2005). 17 Image-based intrusive memories as a symptom post-trauma may offer a translational treatment 18 19 target in their own right. Cognitive models of PTSD suggest that intrusive memories arise due to the differential 20 processing that can occur during the encoding of a traumatic event: There is a shift towards 21 enhanced perceptual processing (associated with a focus on sensory-perceptual information, 22 sometimes termed data-driven processing), and reduced 'conceptual' processing, associated with 23 24 a focus on information related to reasoning, organizing information, creating meaning from an event (Ehlers & Clark, 2000; Holmes & Bourne, 2008; Brewin, 2014). The predominance of 25 perceptual processing at encoding is thought to lead to the subsequent involuntary triggering of 26 sensory-perceptual image based memories by perceptual cues (both external and internal) in the 27 environment (Ehlers & Clark, 2000; Holmes & Bourne, 2008; Brewin, 2014). 28 We need to find novel ways of working with emotional mental imagery (Arntz et al., 2007; 29

30 Holmes et al., 2007). Recent advances in memory research, concerning memory reconsolidation,

1 open up hypotheses about new techniques that could be used to modify emotional memories.

2 Following an event, new memories undergo a time-dependent process of stabilization, involving

3 molecular and cellular processes, termed memory consolidation (Davis & Squire, 1984;

4 McGaugh, 2000). During this discrete time-window new memories are fragile and vulnerable to

interference, before becoming gradually resistant to change (Shadmehr & Holcomb, 1997; over
approximately 6 hours; McGaugh, 2000).

approximatory o nours, the Gaugh, 2000).

7 Insights into the neuroplasticity of memory suggest a consolidated memory may be rendered

8 malleable following its reactivation via a retrieval cue (Alberini, 2005; Nader & Einarsson,

9 2010), necessitating restabilization in order for the memory to persist. During memory

10 reconsolidation, a labile memory can be disrupted or enhanced via an intervention (Misanin et

al., 1968; Nader et al., 2000b). Animal studies have shown successful weakening of conditioned

12 fear memories with pharmacological agents (the protein synthesis inhibitor anisomycin) when

13 applied during reconsolidation (Nader et al., 2000a). Recent studies conducted in humans

14 provide initial evidence that conditioned fear memories (Kindt et al., 2009; Schiller et al., 2010)

and episodic memories (Kroes et al., 2014) can also be modified in line with the idea of

16 reconsolidation update mechanisms.

17 Cognitive models of working memory and dual-task studies suggest additional strategies to

reduce intrusive memories of trauma. Working memory (WM) is a limited capacity system

19 which temporarily maintains and stores information (Baddeley & Hitch, 1974; Baddeley, 2012)

20 to "...support human thought processes by providing an interface between perception, long-term

memory and action" (p. 829; Baddeley, 2003). Such WM models assume some form of limited-

22 capacity "central executive" that oversees two types of storage systems: one for visuospatial

23 information and one for auditory-verbal information.

24 Dual task experiments demonstrate reduced information processing when similar cognitive tasks

compete for shared resources. For example, performing a visuospatial task selectively interferes

with actively holding a sensory-perceptual mental image in mind. Research has shown that

visuospatial dual-task interference can result in a visual image being held in WM becoming less

vivid and, consequently, less emotional (e.g. Andrade et al., 1997; Baddeley & Andrade, 2000;

Van den Hout et al., 2001; Engelhard et al., 2010; Engelhard et al., 2011).

30 Within experimental psychopathology, the trauma film paradigm (Holmes & Bourne, 2008;

James et al., 2016), has long been used to investigate the generation of emotional intrusive

mental images after trauma, and their modification – for example with imagery rescripting
(Hagenaars & Arntz, 2012). Arntz and colleagues have argued that stimuli used in traditional
experimental psychopathology approaches such as fear conditioning are too simple to model the
learning / unlearning of complex emotional memories, and thus alternative approaches are
needed to evolve novel therapeutic interventions concerning trauma memories (Kunze et al.,
2015).

Using the trauma film paradigm has allowed the investigation of the impact of engaging in
visuospatial cognitive tasks on subsequent image-based intrusive memory development.

9 Volunteers view a film with distressing content in the laboratory (an experimental trauma) and

10 then over the following days keep a diary in which they record any intrusive memories of the

11 film. Visuospatial tasks administered either during (e.g. undertaking a complex sequence tapping

12 task; Holmes et al., 2004) or soon after (e.g. playing the computer game 'Tetris'; Holmes et al.,

13 2009; Holmes et al., 2010a) an experimental trauma have led to significantly fewer intrusions

14 during the following week compared to no-task (control). Further, a study by Holmes et al.

15 (2010a) suggested that these effects may be modality specific: A task (*Tetris*) supposedly taxing

visuospatial WM (Lau-Zhu et al., 2017) reduced intrusive memory frequency compared to a no-

17 task control, whereas a verbal-based task (the general knowledge computer game *Quiz*) did not.

18 While a review of the literature by Brewin (2014) suggests that, on balance, visuopatial tasks

19 more consistently lead to a reduction in intrusions than do so-called verbal ones, it is important

20 to note that in some studies verbal tasks have also been shown to reduce intrusions (Krans et al.,

21 2009; Hagenaars et al., 2017).

22 Thus far, most studies investigating cognitive task interference on intrusive memory have

focused on a time window of 24 hours or less between experimental trauma exposure and

24 intervention. For example, studies examined the effects of cognitive interference during or

shortly after (e.g. up to four hours) watching the trauma film (e.g. Holmes et al., 2009; 2010b). A

26 recent study showed that a visuospatial interference task (reminder cue plus Tetris game play)

27 reduced subsequent intrusive memory frequency after a longer time delay of 24 hours (i.e. the

experimental manipulation was administered 24 hours after experimental trauma; James et al.,

29 2015). However, from a clinical perspective there are still challenges associated with reaching

30 people within 24 hours of a traumatic event. Hence, an extension of the time frame of

intervention delivery beyond 24 hours post trauma within a secondary prevention approach could
 benefit greater numbers of individuals who have recently experienced a trauma.

3 In the current research we built upon the above-mentioned study (James et al., 2015), which had

4 used a similar cognitive interference procedure 24 hours after experimental trauma. James et al.

5 (2015) compared four between-subject conditions, and found that only a memory reminder task

6 in combination with *Tetris* game play (but neither task in isolation) led to a reduction in intrusive

7 memories of a trauma film. Results appear to be in line with reconsolidation update mechanisms

8 in that both memory reactivation (reminder cue) and a blockade (*Tetris* game play) were

9 required. However, this study did not test modality-specific task effects during this time frame

10 since no non-visuospatial control task was used (James et al., 2015). Here we extended the

11 approach taken by this previous study and had three aims:

12 i) Can a visuospatial interference intervention (involving reminder cue and *Tetris*) reduce

13 intrusive memories of a trauma film even if delivered 72 hours (3 days) after the film? Were

such an effect obtained, this might enhance the range of applications for this kind of secondary

15 prevention intervention for people who seek help later than 24 hours after a traumatic event.

ii) Is such an effect of the "reminder cue and *Tetris*" condition reducing intrusions also apparent

17 when compared not only to a reminder cue alone condition – as used in the James et al (2015)

18 study – but also to an *active* control condition (i.e. with an alternative task to Tetris game play,

19 akin to a placebo group)?

20 iii) Is any effect on reducing intrusions modality specific, i.e. would the non-visuospatial control

21 intervention (reminder cue plus a *Quiz* game) also reduce intrusions when compared to the

22 reminder cue condition?

23 Thus, the current study tested the hypotheses that a single session cognitive intervention

24 (memory reminder task followed by 15 min *Tetris* gameplay) would reduce later intrusive

trauma memories when delivered 3 days after experimental trauma (a film) compared to two

26 control groups: (1) reminder only: a memory reminder task followed by no cognitive interference

task (sitting quietly in the laboratory for an equivalent period of time to the computer gameplay);

and critically (2) an *active control group*: the memory reminder task followed by 15 min *Quiz*

29 gameplay (an active control for the computer game play component). Intrusions of the trauma

30 film were recorded in a diary, both pre-intervention (days 1-3 after film viewing) and again post-

31 intervention (days 4-6). We assessed recognition memory (voluntary recall) for the trauma film

on day 7. We predicted recognition memory test scores would be comparable in all three groups
in line with findings showing an interference task involving *Tetris* selectively impedes the
intrusive nature of the memory and not the content of recall per se (Lau-Zhu et al., manuscript
under review).

5 2 Materials and methods

6 2.1 Participants

Ninety participants were recruited from two university campuses and a nursing school. Hence, 7 8 all participants were students. Participants were screened and judged ineligible to participate in the study if they reported previously experiencing 6 or more traumatic events on the Essen 9 Trauma – Inventory (ETI; Talgay et al., 2007). Further, participants were excluded from analysis 10 if they failed to participate in the second session of the study, or did not send in their completed 11 intrusion diary on Day 7 (n = 2 reminder+*Tetris* group, n = 2 reminder-only group). The final 12 sample included 86 participants (51 females, age range = 18 - 42; Supplementary Table 1). 13 Participants were reimbursed for their participation (cinema tickets). The study was approved by 14 the ethics committee of the medical faculty of the Ruhr-University of Bochum, Germany (Ref. 15 Nr. 4902-14). 16

17 **2.2 Procedure**

18 Pre-laboratory baseline questionnaires (ETI, SCL-90-R, ERQ, and STAI) were completed via

email prior to attending the laboratory on Day 1. On the first laboratory session (Day 1, see

20 Figure 1) participants provided written informed consent. Demographic information (age,

21 gender) was collected before viewing the trauma film. Participants completed mood ratings

22 immediately prior to watching the film. Immediately afterwards they repeated the mood ratings

and completed further ratings of attention and emotional response to the film.

24 Participants were then given instruction on how to keep the Intrusion Diary to record any

25 intrusive memories of the film they experienced over the subsequent 3 days (Holmes & Bourne,

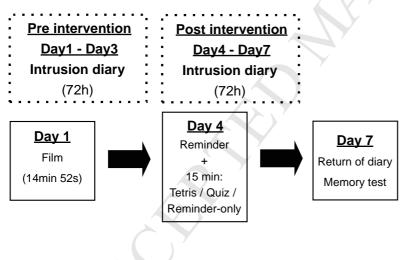
26 2008; Holmes et al., 2010a; James et al., 2015; James et al., 2016).

27 Participants returned to the laboratory 72 h after the experimental trauma. They were randomly

allocated to group (reminder-only, reminder+*Tetris*, or reminder+*Quiz*) using a minimization

- 1 scheme (see below). All participants practiced playing *both* computer games (*Tetris* and *Quiz*)
- 2 for 3 min. This was to ensure that participants would know how to play their allocated game (if
- 3 applicable) after the reminder task, and to keep all experimental procedures prior to the reminder
- 4 task matched across groups. After this practice, participants in all groups completed the memory
- 5 reminder task. They then played *Tetris* or *Quiz*, or sat quietly for 15 min, according to their
- 6 group allocation. After this period, participants allocated to *Tetris* or *Quiz* provided ratings of
- 7 concentration, enjoyment, difficulty and distraction.
- 8 All participants were then reminded of the instructions for the diary and were asked to record
- 9 intrusions in the diary for a further 3 days, after which (on Day 7) participants were asked to
- send back their completed diary by post. On Day 7 participants also completed three last
- 11 questionnaires as a final assessment via email (IES-R, verbal recognition memory test, and
- 12 finally game-related impact rating).

13 Figure 1. General study design.



14 2.3 Tasks

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15 2.3.1 Memory Reminder Task

16 The memory reminder task procedure consisted of two parts, i) film reminder cue presentation

17 followed by ii) a music filler task for 10 minutes:

18 2.3.1.1 Film reminder cue presentation

Two static visual images were presented side by side (using PowerPoint) as reminder cues for 1 the trauma film (viewed 3 days previously). Altogether there were 32 images, two for each of the 2 16 scenes within the film. All images were taken from a moment briefly before the traumatic 3 content, i.e. they were *not* of the worst moments themselves. Participants were instructed to view 4 each pair of images and to press a key when they identified which film clip the pair of images 5 6 related to. A subsequent slide prompted participants to close their eyes and recall that film clip as vividly as possible. When they had finished recalling the clip they were instructed to press the 7 key again in order to have a short self-paced break before moving on to the next pair of images. 8 The time participants took to recognize the still images as being from the film, time spent 9 recalling the film clip with closed eyes, and time taken for the subsequent break were all 10 11 recorded.

12 2.3.1.2 Music filler task

13 Following the presentation of the reminder cues, to allow time for the potential memory

14 reconsolidation processes to be initiated, there was a short time interval before the intervention.

15 This was based on the time interval used in both some animal (Nader et al., 2000a) and human

studies (Schiller et al., 2010; Agren et al., 2012). We used a 10 min interval containing a music

- 17 filler task, in which participants listened to and rated the pleasantness of music clips (as used in
- 18 James et al., 2015).

19 2.3.1.3 Computer game (cognitive interference) tasks

20 2.3.1.3.1 Tetris computer game

In Tetris, 7 differently shaped geometric blocks fall one at a time from the top to the bottom of 21 the screen in a random sequence. As they fall, the blocks can be moved (left, right, rotated 90° 22 23 clockwise or accelerated) using the keyboard arrow keys. The aim is to fit the blocks together to 24 create complete horizontal lines across the playing area. Each time a full horizontal line is created it disappears, and the participant is awarded points. In the current experiment participants 25 26 were instructed to focus on the three blocks due to fall after the one that they were currently manipulating (these blocks were displayed in a preview to the right of the screen). Participants 27 were asked to work out in their 'mind's eye' where best to place these blocks in order to create 28

1 the horizontal lines. The computer game *Tetris* (set to "Marathon" mode, *Tetris* Zone version

2 1.2.1) was played for 15 min on a 15 inch laptop (n = 16) or a 22 inch (n = 12) colour monitor

3 with no sound. The participants with the highest score in *Tetris* and in *Quiz* (see below) were told

4 that they would each win a prize (two cinema tickets).

5 2.3.1.3.2 Quiz computer game

- 6 In the *Quiz* game a series of questions, each with four possible multiple-choice answers, are
- 7 displayed one at a time on the screen. The objective of the game is to gain points by selecting the
- 8 correct answer using the mouse. Participants were instructed to answer as many questions
- 9 correctly as possible within the 15 minutes of game play, and were told that the participant with
- 10 most correct answers would win a prize, to ensure that both the verbal and visuospatial computer
- 11 games had similar temporal constraints. Questions were from the general knowledge domain
- 12 (e.g., with what item of clothing would you associate the word Panama? A= scarf, B=Gloves,
- 13 *C=hat, D=coat*). The computer game itself is called QUIZPro IV (<u>https://litschi.de/edv-</u>
- 14 <u>service/software-2/quizpro</u>) and was played on a 15 inch laptop (n = 16) or a 22 inch (n = 14)
- 15 color monitor with no sound. Note this also a commercially available computer game (freeware),
- but differs from the one used by Holmes et al (2010), called "PubQuiz", mainly in that it is in

17 German.

18 **2.3.1.3.3 Reminder-only**

Participants in the reminder-only group sat quietly in the laboratory for 15 min and were told they could think about anything they like unrestricted. However they were not permitted to undertake any other activity, such as looking at their mobile phone.

22 **2.4 Trauma Film**

The trauma film lasted 14 min 52 sec and consisted of 16 different scenes depicting traumatic events to people, i.e. events involving exposure to actual or threatened death, or serious injury to others (American Psychiatric Association, 2013), as well as aversive scenes of trauma involving non-human animals. Each scene of the film contained footage with different content, e.g. a young girl hit by a car with blood dripping out of her ear; a boy gets hit by a van while texting on his mobile phone and falls to the road; beheading of an animal. The film contained some new

- 1 scenes compared to earlier studies, and details about the scenes and their source can be found in
- 2 Supplementary Table 3. The film was presented in a darkened room using a 15 inch laptop
- 3 computer (n = 45) or a 22 inch computer monitor (n = 41), both with a viewing distance of
- 4 approximately 12 inches. Participants were instructed not to view the film as they might do
- 5 normally, but to pay close attention and to imagine they were there as a bystander at the scene.
- 6 The experimenter waited outside the room while participants viewed the film.

7 2.5 Measures

8 2.5.1 Pre-laboratory baseline questionnaire

9 Prior to laboratory testing participants completed questionnaire measures administered via a

- 10 standardized email for inclusion eligibility and baseline measures:
- 11 **2.5.1.1 Essen Trauma Inventory** (ETI; Talgay et al., 2007)

12 Prior trauma history was reported using the ETI trauma list (not the complete ETI questionnaire).

- 13 Participants were provided with a list of 15 different traumatic events and indicated which they
- 14 had experienced during their lifetime. A cut-off of 6 events was used to exclude participants
- 15 from the study (developed from the experience with other studies with healthy participants in our16 lab).

17 **2.5.1.2 Symptom Checklist-90-Revised** (SCL-90-R; Franke, 1995)

- Psychological symptoms and distress were measured using the SCL-90-R. This self-report
 symptom inventory has 90 items spanning nine symptom dimensions. Distress symptoms can be
 measured using individual symptom dimensions or by 3 summary global scores referred to as the
 Global Severity Index (GSI), the Positive Symptom Distress Index (PSDI) and the Positive
 Symptom Total (PST). The GSI global score was used to assess participants' distress levels.
 Internal consistency of the GSI (as well as of PSDI and PST) is reported to be between 0.96 and
- 0.98. Test-retest reliability for GSI is r = 0.90 (Franke, 1995).

25 2.5.1.3 Emotional Regulation Questionnaire (ERQ; Gross & John, 2003; Abler & Kessler, 26 2009)

- 1 Emotional experience and emotional expression was measured using the EQR. The self-report
- 2 questionnaire asked participants to rate 10 statements such as '*I control my emotions by not*
- 3 expressing them' (expressive suppression) and 'when I want to feel less negative emotion, I
- 4 change the way I'm thinking about the situation' (cognitive reappraisal) on a scale from 1
- 5 'strongly disagree' to 7 'strongly agree'. For each scale a sum score was calculated. Internal
- 6 consistency of the German version is 0.74 for the expressive suppression subscale, and 0.76 for
- 7 the reappraisal subscale (Abler & Kessler, 2009). Test-retest reliability over 3 months is r = 0.69
- 8 for both scales (Gross & John, 2003).

9 **2.5.1.4 State-Trait Anxiety Inventory – Trait** (STAI-T; Spielberger et al., 1983)

The STAI-T measured levels of anxiety and is a 20 item self-report measure. Each item is rated on a four point scale, with scores ranging from a minimum of 20 to a maximum of 80, where higher scores represent greater levels of trait anxiety. Internal consistency is 0.90. Test-retest reliability after 63 days is between r = 0.77 and r = 0.90 (Laux et al., 1981).

14 2.5.2 Memory Measures

15 **2.5.2.1 Intrusion diary**

Participants were asked to keep a pen and paper diary to record any intrusive image-based 16 memories of the experimental trauma (film) they experienced during their daily lives (e.g. 17 18 Holmes et al., 2004; James et al., 2015). The diary was kept over the course of 6 day; days 1 to 3 (pre-intervention) and then days 4 to 6 (post-intervention) and returned to the experimenter on 19 day 7. Each day in the diary was represented by a separate printed box, each further split into 20 three sections (morning, afternoon and evening). Participants were asked to tick a box in the 21 appropriate section when they experienced an intrusive memory (or to indicate that they had 22 23 experienced none by drawing a line though that section of that particular day). They were then asked to briefly describe the content of each of their intrusions marked in the diary (e.g. a silver 24 car crashing on the freeway) so that the experimenter could later use these descriptions to 25 26 confirm whether the intrusion related to the film. Participants were asked to record all intrusions immediately after they had occurred, or as soon as possible thereafter, and to set aside regular 27 time slots each day to check that their diary was up-to-date. Verbal and written instructions were 28

provided with the diary. Intrusive memories were defined as any memory of scenes of the film 1 that appeared spontaneously in their mind (e.g. "spontaneously pop into your mind when you are 2 not deliberately thinking about the film"). Participants were instructed not to include memories 3 that they deliberately recalled and were also given instructions about the form of intrusive 4 memories, that is 'mental images' (e.g. "in the form of pictures in your mind's eye") rather than 5 verbal thoughts in the form of words/ phrases. Experimenters checked that the intrusion 6 descriptions in the diary were matched to scenes in the film (rather than for example of different 7 films, or everyday life). In this study 100% were matched to the films and were included in 8 subsequent data analyses. 9 Importantly, days 1-6 were defined by 24 h intervals and not by calendar days (as in other studies 10 using similar set-ups, e.g. James et al., 2015; Hagenaars et al., 2017). That is, day 1 was the first 11 24 h period after the first lab session (e.g. if this session ended at 4pm, it would run from 4pm 12 that day to 4pm the next day), day 2 the second 24 h period, and so on. Day 4 was the 24 h 13 period immediately following the second lab session, and the beginning of day 4 as a post-14 intervention time period was marked clearly (visually) in each diary to ensure that days 1 to 3 15 and days 4 to 6 were clearly separated as pre-intervention and post-intervention for all 16

17 participants.

18 2.5.2.2 Intrusion diary compliance rating

Participants rated their compliance with the diary at 2 time points, in the second laboratory
session for the first 3 days (pre-intervention), and on day 7 via email for the subsequent 3 days
(post-intervention). Ratings were made on a scale anchored from 1 *'not compliant'* to 10 *'very compliant'*.

23 **2.5.2.3 Impact of Event Scale-Revised** (IES-R; Weiss & Marmar, 1997)

The IES-R was administered on final day of the experiment (Day 7) using the German version (Maercker & Schützwohl, 1998). It was used to assess participants' distress related to the experimental trauma during the past 7 days (i.e. items were anchored to the trauma film). The IES-R is a 22-item self-report measure. Each item is rated on a scale from 0 'not at all' to 4 'extremely'. The IES-R Total score ranges from a minimum of 0 to a maximum of 88, where 1 "Intrusion", "Avoidance" and "Hyperarousal". Internal consistency (Cronbach's α) for the

2 German version is: 0.90 (intrusion subscale), 0.90 (hyperarousal subscale), and 0.79 (avoidance

3 subscale). Test-retest reliability is 0.80 for the intrusion subscale, 0.79 for the hyperarousal

4 subscale, and 0.66 for the avoidance subscale (Maercker & Schützwohl, 1998). Although used

5 most commonly to measure levels of distress on a continuum, a cut-off of 33 has been proposed

6 (Creamer et al., 2003) for a likely diagnosis of PTSD, with a sensitivity of 0.91.

7 2.5.2.4 Verbal recognition memory test

8 The recognition memory test comprised 48 'true/false' written statements relating to each of the

9 16 scenes that made up the trauma film (3 per scene). Examples included 'Scene 1: *The sun was*10 *shining in the cemetery*' and 'Scene 3: *A white car was involved with the accident*'. Participants

indicated 'yes' or 'no' (on paper) as to whether or not they recognized the written statement as

12 belonging to the film viewed 7 days before.

13 2.5.3 Film Measures

14 **2.5.3.1 Pre- to post-film mood ratings**

Participants rated the emotions sad, scared, calm, disgusted, hopeless, happy, aghast, frightened, and helpless on 9 visual analogue scales (VAS) given pre- and post-film. Participants rated how they felt '*right at this very moment*' on scales anchored from 0 '*not at all*' to 10 '*extremely*'. A composite mood score was calculated by summing the 9 emotions, with happy and calm reverse-

19 scored.

20 2.5.3.2 Film attention and affect rating

Following film viewing participants also rated how much attention they had paid to the film, and how much they were emotionally affected by the film on scales anchored from 0 '*not at all*' to 10 '*extremely*'.

24 2.5.4 Computer Game Play Ratings

- 25 Following computer game play in both active conditions, ratings for levels of game
- 26 concentration [How well did you concentrate during game play?], enjoyment [Did you enjoy the

- 1 game you just played?], difficulty [How difficult did you find the game you just played?], and
- 2 distraction [How distracted were you when you played the game?] were made on separate scales
- anchored from 0 'not at all' to 10 -'very well' / 10 -'very much' / 10 -'extremely' / 10 -
- 4 'maximally';
- 5 Participants were asked to rate how much they believed playing the game Tetris or Quiz 3 days
- 6 after the emotional film would increase or decrease intrusive images of the film (of the type
- 7 recorded in the diary) from -10 'extreme decrease' to 0 'no effect' through to 10 'extreme
- 8 *increase*' to check for demand characteristics at the end of the experiment (impact rating).
- 9 2.6 Data Analysis and Statistics

10 **2.6.1 Sample Size**

- 11 Sample size was calculated based on findings of prior studies (Holmes et al., 2009; Holmes et al.,
- 12 2010a) albeit at an earlier intervention time point, i.e. we expected a medium to large effect size
- 13 (d = 0.70, corresponding to f = 0.42). Given this effect size, we calculated a total sample size of
- 14 n = 90 with power set to 0.95 and $\alpha = 0.05$ (G*Power 3.1.7).

15 **2.6.2 Random allocation to groups**

A minimization scheme was used to allocate participants to group (Scott et al., 2002; Altman & Bland, 2005). Initially, participants were randomly allocated to one of the three groups. Then, after an interim check, to reduce differences in baseline measures between the groups occurring by chance, the final 45 participants were allocated to group based upon three selected baseline scores (reported number of intrusions over the first three days, age, and STAI). This allowed us to minimize possible baseline differences between the three groups on these three variables, which were deemed likely to affect intrusion frequency.

23 2.6.3 Data analysis

Number of intrusions, baseline variables (Age, SCL-90-R, STAI-T, ERQ), trauma questionnaires
(ETI, IES-R), diary compliance, film ratings, recognition memory test, ratings of computer game
play and the reminder cue presentation measures were skewed, thus between-group comparisons
(reminder+*Tetris* vs. reminder+*Quiz* vs. reminder-only) were conducted using the Kruskal-

Wallis-test for independent samples, and planned comparisons were conducted using the MannWhitney-test. For comparisons of mood pre- and post-film we first conducted a Friedman test for
dependent variables to analyze mood deterioration and then compared difference scores of mood
pre- to post-film by a Kruskall-Wallis test. Gender was analyzed between groups using a Chisquared test.

Primary outcome. A Kruskal-Wallis test was used for between-group comparisons in the 6 mean number of intrusions at baseline (pre-intervention, days 1 to 3). Baseline imbalances were 7 subsequently controlled for using a Quade test (non-parametric ANCOVA; Quade, 1967) for 8 9 between-group comparisons in the mean number of intrusions post-intervention (days 4-6) followed by planned comparisons (see Figure 2). The Quade test involves a ranking of both the 10 dependent (intrusions post-intervention) and the covariate variable (intrusions pre-intervention) 11 ignoring the grouping factor. In a second step a regression analysis of the dependent variable on 12 13 the covariate is conducted saving the unstandardized residuals as a new dependent variable. In a 14 final step these residuals are tested between groups by running a one-way analysis of variance. Additionally, day-to-day differences in the number of intrusions were calculated, and entered 15 into a two-way mixed ANOVA with time as within-subjects factor and group as between-16 subjects factor. Post-hoc one-way ANOVAs were performed to detect for which day-to-day 17 changes there were differences between the groups, followed by independent sample t-tests. 18 19 Two-tailed tests and an alpha level of 0.05 were used for all statistical comparisons.

3 Results

20 3.1 Pre-Laboratory Baseline Questionnaires

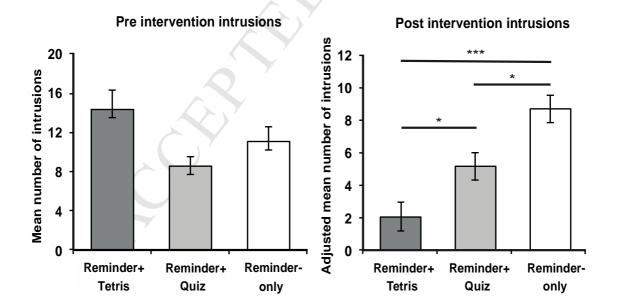
There were no statistically significant differences between groups on prior trauma history (ETI; $X^2(2, N = 86) = 3.37$, p = 0.185, d = 0.404), the GSI of the SCL-90-R $X^2(2, N = 85) = 0.602$, p = 0.740, d = 0.169, ERQ cognitive reappraisal $X^2(2, N = 86) = 0.339$, p = 0.844, d = 0.126, ERQ expressive suppression $X^2(2, N = 86) = 0.756$, p = 0.685, d = 0.188, STAI-T $X^2(2, N = 86) =$ 0.933, p = 0.627, d = 0.210. Further there were no group differences for gender $X^2(2, N = 86) =$ 0.31, p = 0.86, d = 0.120, or age $X^2(2, N = 85) = 1.96$, p = 0.376, d = 0.307 see Supplementary Table 1.

1 **3.2** Primary outcome measure: diary intrusions post-intervention

A Kruskal-Wallis test on the distribution of number of intrusions per group pre-intervention (day 2 1 to 3) revealed no significant differences between groups at baseline $X^2(2, N = 86) = 4.798$, p = 3 0.091, $\eta^2 = 0.067$ (Figure 2 and Table 1). As the distribution of intrusions pre-intervention was 4 skewed ($M_{Tetris} = 14.25 (10.67), M_{Ouiz} = 8.50 (4.97), M_{reminder-only} = 11.00 (8.21)$) and contained 5 6 outliers, a Quade test controlling for pre-intervention intrusions was calculated to analyze 7 differences between groups in post-intervention intrusions (see methods section 2.6.3). As a part of the Quade test, the one-way ANOVA of unstandardized residuals of a regression analysis 8 (pre-intervention intrusions on post-intervention intrusions) showed a significant effect of 9 intervention type (F_{2.83} = 12.89, p < 0.001, $\eta_p^2 = 0.237$). 10 As predicted, planned comparisons (Games-Howell corrected) between all groups showed that 11 the reminder+Tetris group experienced significantly fewer intrusions post-intervention (days 4-12 6) compared to the reminder-only group ($M_{Tetris} = 3.18$ (2.83), $M_{reminder-only} = 8.61$ (8.10); Mean 13 difference_{Residuals} = -25.29, SE = 4.90, p < 0.001, d = 1.37). Critically, the reminder+*Tetris* group 14 also experienced significantly fewer intrusions post-intervention than did the reminder+Quiz 15 group ($M_{Quiz} = 4.13$ (4.06); Mean difference_{Residuals} = -11.22, SE = 4.48, p = 0.040, d = 0.653). 16 Contrary to predictions, we found that the reminder+*Quiz* group experienced significantly fewer 17 intrusions post-intervention (days 4-6) compared to the reminder-only group (Mean difference = 18 -13.97, SE = 5.32, p = 0.030, d = 0.693), see Figure 2. 19 Figure 3 depicts the trajectory of the mean number of intrusions over the 6 days of continuous 20 recording split by intervention group allowing visual inspection of the within-groups 21 22 comparison. Interestingly, only in the reminder-only group do intrusions increase significantly from day 3 to day 4 ($M_{day 3} = 2.64$ (3.12), $M_{day 4} = 5.68$ (7.63), z = -2.24, p = 0.025, d = 0.935). 23 In a more detailed analysis, to test whether changes in the number of intrusions from one day to 24 the next differed between groups, a two-way mixed ANOVA was performed, with time as a 25 within-subject factor, and group as a between-subjects factor. This analysis revealed a significant 26 main effect of time (F_{2.29, 190.11} = 12.92; p < 0.001; $\eta_p^2 = 0.135$; Greenhouse-Geisser-Correction 27 used), a significant main effect of group (F_{2, 83} = 5.80; p = 0.004; $\eta_p^2 = 0.123$), and a time x group 28 interaction (F_{4.58, 190,11} = 3.89; p = 0.003; $\eta_p^2 = 0.086$). Post-hoc one-way ANOVAs showed 29 group differences in the change in number of intrusions from day 3 to day 4 ($F_{2,83} = 6.51$; p = 30

1	0.002; $\eta^2 = 0.136$), and from day 4 to day 5 (F _{2,83} = 4.48; p = 0.014; $\eta^2 = 0.097$), but not between
2	the other days. From day 3 to day 4 (i.e., from the 24 hour period prior to the intervention period
3	to the 24 hour period immediately afterwards), the number of intrusions increased in the
4	reminder-only group (M = 3.04, SD = 7.41), remained stable in the reminder+ $Quiz$ group (M =
5	0.13, SD = 2.21), but decreased in the reminder+ <i>Tetris</i> group (M = -1.46, SD = 2.98). T-tests
6	indicated significant differences between all 3 groups (reminder+ <i>Tetris</i> vs. reminder-only: $t_{54} =$
7	2.98, p = 0.004, d = 0.81; reminder+ <i>Tetris</i> vs. reminder+ <i>Quiz</i> : t_{56} = 2.33, p = 0.023, d = 0.62;
8	reminder+ $Quiz$ vs. reminder-only: $t_{56} = 2.05$, $p = 0.045$, $d = 0.55$). From day 4 to day 5, number
9	of intrusions tended to decrease, with the most pronounced decrease in the reminder-only group
10	(M = -4.11, SD = 7.79; reminder + Tetris: M = -0.43, SD = 1.53; reminder + Quiz: M = -1.63, SD = -1.
11	2.08), with significant differences between the reminder+ <i>Tetris</i> and the reminder-only groups (t_{54}
12	= 2.45; p = 0.018, d = 0.67), the reminder+ <i>Tetris</i> and the reminder+ <i>Quiz</i> groups (t_{56} = 2.50; p =
13	0.015, d = 0.67), but not between the reminder+ $Quiz$ and reminder-only groups (t ₅₆ = 1.68; p =
14	0.099, d = 0.45).

- 16 Figure 2. Intrusive memories of experimental trauma post intervention controlling for pre-
- **intervention intrusive memories.** Error bars depict s.e.m. * p < 0.05, *** p < 0.001

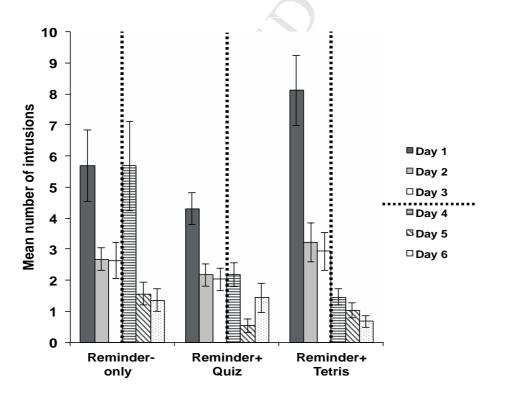


- 1 Table 1. Number of intrusive memories of experimental trauma pre- and post-intervention,
- 2 and verbal recognition memory test scores, in each of the three conditions.

	Reminder-only $(n = 28)$		Reminder $+Tetris$ $(n = 28)$		Reminder + $Quiz$ ($n = 30$)	
Measure	М	SD	М	SD	М	SD
Pre-intervention number of	11.00	8.21	14.25	10.67	8.50	4.97
intrusions in Diary (days 1-3)						
Post-intervention number of	8.61	8.10	3.18	2.83	4.13	4.06
intrusions in Diary (days 4-6)			2			
Verbal recognition memory test	36.25	5.01	34.07	5.79	36.27	5.23
scores			\mathcal{N}			

3 4

> Figure 3. Trajectory of mean number of intrusive memories of experimental trauma over all 6 diary days per condition. Days 1, 2 and 3 are pre- reminder and intervention, while Days 4, 5 and 6 are post- reminder and intervention. Error bars depict s.e.m.



1 **3.3 Intrusion diary compliance**

2 For the 3 days pre-intervention, there was no significant difference between groups on ratings of

3 diary compliance $X^2(2, N = 86) = 0.884$, p = 0.643, $\eta^2 = 0.010$. This was also the case for the 3

4 days post-intervention $X^2(2, N = 72) = 1.07$, p = 0.585, $\eta^2 = 0.015$, see Supplementary Table 2.

5 3.4 Impact of Event Scale-Revised Intrusion Subscale

Analysis of the IES-R showed a significant difference post-intervention between groups for the "Intrusion" subscale $X^2(2, N = 86) = 6.64$, p = 0.036, $\eta^2 = 0.077$. However, planned comparisons Dunn-Bonferroni-corrected showed no significant differences between the reminder+*Tetris* group and the reminder-only group (z = -2.12, p = 0.103, d = -0.470), nor between the reminder+*Quiz* group and the reminder-only group (z = -0.191, p = 1.00, d = -0.041). There was no significant difference between the reminder+*Tetris* group and the reminder+*Quiz* group (z = -2.34, p = 0.057, d = -0.522), see Supplementary Table 1.

13 **3.5 Verbal recognition memory test**

Scores on the written recognition memory test for the film did not differ significantly between groups ($X^2(2, N = 86) = 2.53$, p = 0.282, $\eta^2 = 0.029$, Table 1, Supplementary Figure 1).

16 **3.6 Film Measures**

17 **3.6.1 Mood pre- to post-film**

18 The Friedman test revealed a significant difference between pre-film and post-film mood ($X^2(2,$

19 N = 86 = 82.05, p < 0.001, $\eta^2 = 0.954$). Comparison of the difference scores showed no

- significant difference of mood deterioration between groups ($X^2(2, N = 86) = 1.06, p = 0.588, \eta^2$
- 21 = 0.012), see Supplementary Table 2. This verified that mood scores deteriorated after viewing
- the trauma film, but that this deterioration was not different between groups,

23 **3.6.2 Film attention and affect**

1 There was no significant difference between groups for ratings of how much attention

participants paid to the film, $X^2(2, N = 86) = 0.099$, p = 0.952, $\eta^2 = 0.001$, and how much they were affected by the film, $X^2(2, N = 86) = 5.042$, p = 0.067, $\eta^2 = 0.059$; see Supplementary Table 2.

5 3.7 Computer Game Play Ratings

- 6 Participants in the reminder+*Tetris* and reminder+*Quiz* groups showed no significant difference
- 7 between rating scores for concentration, enjoyment, difficulty and distraction: Concentration, z =
- 8 0.103, p = 0.918, d = 0.027; Enjoyment, z = 0.024, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, p = 0.981, d = 0.006; Difficulty, z = 0.581, d = 0.006; Difficulty,
- 9 = 0.561, d = 0.153; Distraction, z = 1.26, p = 0.209, d = 0.336, see Supplementary Table 2.
- 10 Participants in the reminder+*Tetris* and reminder+*Quiz* groups did not differ in their impact
- 11 (demand) ratings for their allocated intervention (i.e. *Tetris* in the reminder+*Tetris* group and
- 12 *Quiz* in the reminder+*Quiz* group), z = 1.19, p = 0.234, d = 0.319, or their unallocated
- intervention, z = -0.360, p = 0.719, d = -0.097, see Supplementary Table 2.

14 **3.8 Reminder Cue Presentation Measures**

The duration of the cue presentation (presentation of images on power point), recall for each clip following the cue, and break was not statistically different between groups: cue presentation duration, $X^2(2, N = 86) = 3.50$, p = 0.174, $\eta^2 = 0.041$; recall duration, $X^2(2, N = 86) = 2.41$, p = 0.299, $\eta^2 = 0.028$; break, $X^2(2, N = 86) = 0.121$, p = 0.941, $\eta^2 = 0.001$, see Supplementary Table 2.

20 **4 Discussion**

Results showed that the frequency of intrusive memories (reported in a daily diary) of an 21 22 experimental trauma (film) was lower for participants who underwent an intervention procedure - a memory reminder task followed by *Tetris* game play - delivered 3 days after the experimental 23 trauma, compared to two control conditions (1) participants who underwent a memory reminder 24 task followed by Quiz (a verbal computer game), and (2) participants who completed only the 25 memory reminder task only, both in line with predictions. Contrary to predictions, results 26 showed that a reminder task followed by Quiz game play led to a significant reduction of 27 intrusive memories compared to the reminder-only condition, which is of interest for a variety of 28

reasons. Declarative memory (recognition memory test) for the film appeared to be left intact 1 across conditions. Hence our study provides results relating to all three aims formulated in the 2 introduction: i) The visuospatial intervention (memory reminder task followed by Tetris game 3 play) was effective even if administered 3 days after the film; ii) this effect held not only against 4 a simple control task (reminder only) but also against an *active* control condition with another 5 computer game (memory reminder task followed by Quiz); iii) The effect of the active control 6 condition (memory reminder task followed by Quiz) in reducing intrusive memories (albeit 7 weaker than that for the memory reminder task followed by Tetris) questions the postulated 8 modality specificity of the intervention rationale. 9 Participants in the three groups did not differ in terms of baseline variables (e.g. trauma history, 10 emotion regulation), with similar ratings for diary compliance and mood deterioration from pre-11 to post-film. Crucially, participants in the two gameplay conditions also reported comparable 12 13 levels of enjoyment, concentration, difficulty, distraction and expected impact regarding the two computer games. Differences in pre-intervention intrusions were controlled for in the analyses. 14 Hence, these variables are unlikely to have accounted for the effect of the active intervention 15 conditions. 16 A detailed view of the day-to-day changes confirms the main results. Crucially, between days 3 17 and 4, i.e. from immediately before to immediately after the intervention, the three groups differ 18 19 significantly in their change in number of intrusions: Overall, there is a similar pattern of change in intrusions, with a greater decrease in the number of intrusions from pre- to post-intervention in 20 the reminder+*Tetris* group compared to each of the other two groups, in line with the hypotheses. 21 22 Further, there is a greater decrease in the reminder+Quiz group compared to the reminder-only group. The only other point where groups differ significantly is the change between days 4 and 5, 23 with greater decreases in number of intrusions in the reminder-only and reminder+Quiz groups 24 25 than the reminder+*Tetris* group. This could in part simply reflect the more substantial reduction

26 in number of intrusions that occurred the previous day in the reminder+*Tetris* group, potentially

27 precluding the possibility of observing further large decreases. However, if the pattern of results

28 is interpreted as indicating that reminder+*Tetris* led to an immediate reduction in intrusion

29 frequency, but no greater rate of decay thereafter, this could raise questions about the longevity

30 of effects over time. A limitation of the study is that it cannot answer questions about longevity

of effects, as there is no data beyond day 6, and the rapid natural decay in the number of

1 intrusions resulting from the film stimuli used means that we would not expect to be able to

2 observe longer-term effects in such an experimental study. However, within clinical studies it

would be useful to investigate the longevity of any effects on intrusions and other clinical
outcomes.

Interestingly, our current data indicates that the memory reminder task delivered in isolation in the intervention session (72 h after the experimental trauma), without a further interference task led to a significant *increase* in intrusions over the next day (Figure 3 from day 3 to day 4). Such effects remain to be better understood, but one hypothesis is that by coming back for the second visit the memory trace was reactivated in the same context where the film was shown (laboratory) and the memory for the film may have been strengthened leading to more intrusions. Further work should examine this possibility.

A driving interest in the current study was the time frame of the delivery of the 12 experimental intervention. Thus, a key finding of our study is that the intervention involving 13 memory reminder plus Tetris was effective 3 days after an experimental trauma. Previously, the 14 study on which the current design was based had tested the intervention only at 24 hours post 15 experimental trauma (James et al 2015). Prior to this the time interval between the event and the 16 intervention had been even shorter and within 6 hours post-event (e.g. Holmes et al, 2009, 2010). 17 These early time window studies inspired clinical translation to a related intervention delivered 18 19 within 6 hours of real trauma (Iyadurai et al, 2018; Horsch et al, 2017). However, there is both theoretical interest and clinical relevance of pushing the time window back even further. Thus 20 the James et al. (2015) experimental study was the first to our knowledge to test the experimental 21 22 intervention outside the so-called memory consolidation time window, i.e. once the memory was assumed to have consolidated. Subsequently, another study has also extended the intervention 23 time window to several days post analogue trauma (Hagenaars et al., 2017). This study also 24 25 showed effects of competing task-based interventions on older intrusive memories, in line with reconsolidation-based approaches (Monfils & Holmes, 2018). Further extending the time 26 27 window for intervention is of interest both as a conceptual test of replication at longer time intervals, and because demonstrating effects at longer time intervals post-event would enhance 28 the range of applications in real life. Many people experiencing a traumatic event may not seek 29 30 professional help until later than 24 hours post-trauma. Thus, if we seek to develop a secondary

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prevention intervention to reduce intrusive memories in the aftermath of trauma, a time-window
 of effectiveness of more than 24 hours will be useful.

A critical limitation of the James et al. (2015) study was that it lacked a non-visuospatial 3 control task (James et al., 2015). We addressed this here by the addition of the reminder+Ouiz 4 group. We do not know why the *Quiz* game also led to a reduction in intrusions compared to the 5 control condition (reminder-only). It may be that working memory tasks may help reduce 6 intrusions by a general taxation that is not modality specific (e.g. Engelhard et al., 2011; Van den 7 Hout & Engelhard, 2012), an important point that remains to be further explored (see later). 8 Previous studies that have tested Quiz (or other verbal tasks, e.g. backward counting) in the 9 memory consolidation (rather than reconsolidation) time window have not typically found a 10 significant reduction in intrusions between such verbal tasks compared to no-task control groups. 11 On the contrary, in some studies verbal tasks even led to an increase in intrusions (Holmes et al., 12 2004; Bourne et al., 2010; Holmes et al., 2010a). However, a recent study, like the current one, 13 also indicates a beneficial effect of a word game task at a longer time interval post experimental 14 trauma, also when compared to a reactivation-only condition (Hagenaars et al., 2017). In this 15 study, the reactivation plus word games condition appeared to result in fewer intrusions than did 16 the reactivation plus Tetris condition. Another intriguing possibility in the current experiment is 17 that the impact of the verbal task compared to control is driven not simply by the verbal task, but 18 19 rather by the *increase* in intrusions seen in the reminder-only condition. This makes the comparison between the two computer games (which both include the reminder) particularly 20 important. Critically, when comparing the memory reminder task followed by *Tetris* game play 21 22 with the reminder task followed by *Quiz* game play, there is a significantly lower frequency of intrusions in the procedure including Tetris. On balance, our results indicate that the procedure 23 using Tetris may have a more powerful effect on reducing intrusions than did the other computer 24 25 game. However, given the findings by Hagenaars et al., 2017 this should be explored further. Nevertheless, results from our study thus call into question the modality specificity of the effect 26 of cognitive task interference on subsequent intrusion frequency. Task modality is an important 27 area of future enquiry given the mixed findings in the field (Gunter & Bodner, 2008; Krans et al., 28 2009; Engelhard et al., 2010; Krans et al., 2010; Logan & O'Kearney, 2012; Brewin, 2014). 29 Future work should continue to contrast alternative games hypothesized to share visuospatial or 30 verbal working memory resources, to test the role of modality-specificity in reducing intrusions 31

and ascertain which tasks are mostly likely to be helpful. Objective tests that could tease apart 1 both how much load, and of which modalities, would help in this endeavor. For future studies, it 2 might therefore be fruitful to also assess verbal intrusions in the diary in order to potentially 3 discover differential effects of task modality on intrusion modality. Tied to this, future studies 4 could also implement a *Quiz* control condition in a less visuospatial way, providing *Quiz*. 5 questions orally rather than on a screen. From a translational perspective we are interested in all 6 tasks that might be helpful in reducing intrusions (and should take care not to use ones that might 7 increase intrusions). 8

9 Returning to the translational interest in helping ameliorate post-traumatic stress symptoms, early

10 stage trials with patients have provided positive results suggesting that emotional memories can

be influenced by the noradrenergic beta-blocker propranolol (Kindt & van Emmerik, 2016).

12 There have also been negative findings, i.e. traumatic memories in PTSD were *not* influenced by

13 propranolol or the glucocorticoid antagonist mifepristone across three studies (Wood et al.,

14 2015). However, drawing on concepts from reconsolidation-update mechanisms may open up a

new way to encourage treatment innovation for PTSD and other disorders (Debiec, 2012;

16 Monfils & Holmes, 2018), here within a cognitive behavioral rather than pharmacological

approach. To date, early steps at clinical translation of this approach have predominantly looked

18 at the memory consolidation window (within 6 hours of the traumatic events; Horsch et al.,

19 2017; Iyadurai et al., 2017) rather than older trauma memories (i.e. a reconsolidation time

20 window; > 24 hr). Future clinical translation studies at longer time intervals post-trauma are

21 warranted and are underway (Kessler et al., 2018).

22 One clinically important aspect of our results is the fact that both the *Tetris* and *Quiz* intervention

23 left recognition memory for aspects of the trauma film intact while reducing diary intrusions.

24 This finding is in keeping with previous research (e.g. Brewin & Saunders, 2001; Holmes et al.,

25 2009; Holmes et al., 2010a; Krans et al., 2010; Deeprose et al., 2012). Findings may suggest a

26 dissociation between perceptual memory intrusions versus episodic memory for traumatic events

27 (e.g. Brewin, 2014), see also Lau-Zhu et al. (manuscript under review). Future work is required

to understand the precise mechanisms of action. From a legal and subjective perspective, it is

29 often important for patients not to forget the events per se but to be able to discuss them without

30 intense reliving in the form of intrusive memories (Holmes et al., 2010b).

There are several limitations of this work. Although the idea of "reconsolidation-update 1 mechanisms" guided our hypotheses, study design and explanation of results, our study itself 2 does not provide evidence for reconsolidation processes per se. Due to the lack of 'no-3 reminder+*Tetris/Quiz*' control groups, we could not demonstrate that reactivation is a necessary 4 prerequisite for an interference task (*Tetris* or *Quiz*) to be effective. Another limitation relates to 5 6 the use of film footage as an experimental analogue of traumatic events. One could argue that this does not mirror real-life trauma and hence conclusions are hard to draw regarding clinical 7 applications (James et al., 2016). Although this remains a valid criticism, it should be noted that 8 the DSM-5 (American Psychiatric Association, 2013) includes (work-related) viewing of 9 aversive film footage as a possible source of traumatization (A criterion). Another limitation 10 concerns the fact, that participants rated their compliance with the diary on a self-report scale. 11 This is potentially biased and does not necessarily capture true variations in how diligent 12 participants were in recording their intrusions. 13 14 In sum, in the current study we demonstrated that 3 days after experimental trauma (after the time frame for memory consolidation to have occurred), an intervention comprising a memory 15 reminder task and short delay prior to a 15 minute cognitive interference task (Tetris) led to a 16 reduction in intrusion occurrence, compared to both an active control (reminder+Quiz), and 17 compared to a reminder-only condition. The fact that both computer games showed an effect on 18 19 reducing subsequent intrusion frequency compared to reminder-only questions the assumed modality-specificity of the intervention, and also calls for further examination of the reminder-20 only condition in possibly increasing intrusions. Of the two games, the Tetris procedure showed 21 22 a significantly greater reduction in intrusions, and therefore it is possible that visuospatial tasks may have additional benefits as a choice of task in translating such work to clinical populations. 23 Our findings open the possibility of developing new interventions to reduce the impact of 24 25 intrusions after trauma. Should related effects hold true for older memories in clinical populations (as an initial case series study suggests; Kessler et al., 2018), then such a brief 26 27 intervention approach (memory reminder task plus cognitive interference) might provide a nonverbal, cost-effective and scalable specific treatment technique to help people reduce intrusive 28 memories of trauma that happened more than 24 hours ago (Laux et al., 1981; Creamer et al., 29 30 2003; Monfils & Holmes, 2018). This could significantly enhance the range of possible

- 1 applications as a secondary prevention intervention, as many people may seek help only more
- 2 than 24 hours after experiencing a traumatic event.
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1 Conflict of interest

- 2 The authors declare that the research was conducted in the absence of any commercial or
- 3 financial relationships that could be construed as a potential conflict of interest.

4 **Author Contributions**

- 5 This information has been removed to ensure double-blindness of the review process, and will be
- 6 included in the final manuscript.

7 Funding

- 8 This information has been removed to ensure double-blindness of the review process, and will be
- 9 included in the final manuscript.

10 Data availability

- 11 The full dataset (raw and processed) is stored on local systems and can be made available on
- 12 reasonable request.

1 Highlights

2	•	Intrusive visuospatial memories can be induced in healthy participants by watching a
3		film containing visually disturbing material (trauma film)
4	•	72 hours later, all participants were reminded of the film
5	•	A visuospatial task (computer game Tetris) administered after the reminder reduces
6		intrusive visuospatial memories
7	•	This effect is significantly superior compared to a verbal control task (Quiz game) and a
8		control group without a specific task after the reminder
9	•	The Quiz game is more effective than the reminder-only group

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