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Emerging Interventions for PTSD: Future Directions for Clinical Care and Research

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Abstract

Efficacious therapeutic approaches for treating Posttraumatic Stress Disorder (PTSD) are needed given the significant psychosocial and physical impairment associated with the disorder (e.g., Hidalgo & Davidson, 2000; Jaycox & Foa, 1999; Stein, McQuaid, Pedrelli, Lenox & McCahill, 2000). Although variations of cognitive behavioral therapy (CBT) effectively treat PTSD, non-response rates and dropout rates remain relatively high (Bradley, Greene, Russ, Dutra, & Westen, 2005; Schottenbauer, Glass, Arnkoff, Tendick & Gray, 2008). Thus, treatment outcome research is needed to improve the effectiveness of existing protocols, particularly within specific populations, develop new approaches for treating individuals who cannot access or do not benefit from traditional treatments, and evaluate the types of treatment that may be effective for particular individuals. The present review provides an overview of emerging treatment approaches for PTSD that attempt to address these remaining issues in the treatment outcome literature.

1. Introduction

Posttraumatic stress disorder (PTSD) is a psychopathological response to exposure to a traumatic stressor and is characterized by recurrent re-experiencing of the event, avoidance of reminders of the trauma, emotional numbing and withdrawal, and hyper-arousal (American Psychiatric Association, 2000). Prevalence estimates suggest that about 8% of the general population is affected by PTSD (Kessler, Chiu, Demler, & Walters, 2005). The disorder typically follows a chronic course, and leads to significant work and social impairment and increased healthcare utilization (e.g., Hidalgo & Davidson, 2000; Jaycox & Foa, 1999; Stein, McQuaid, Pedrelli, Lenox & McCahill, 2000). Given the number of affected individuals and significance of functional impairment the disorder results in substantial personal and societal costs.

Empirical study from the past three decades supports the use of exposure-based therapy, such as Prolonged Exposure (PE; Foa & Rothbaum, 1998; for a review see Ponniah & Hollon; for a recent meta-analysis see Powers, Halpern, Ferenschak, Gilihan, & Foa, in press), for treatment of this condition. Cognitive therapies, including Cognitive Processing Therapy (CPT; Resick & Schnicke, 1992), also have a good evidence base suggesting that they reduce PTSD symptoms (e.g., Chard, 2005; Ehlers et al., 2003; Monson et al., 2006;

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Resick et al., 2008; Resick, Nishith, Weaver, Astin, & Feuer, 2002). Empirical support also exists for Eye Movement Desensitization Reprocessing Therapy (EMDR; Shapiro et al., 1999). Although some studies of EMDR have been questioned on methodological grounds (e.g., Taylor et al., 2003; Rothbaum, Astin, & Marsteller, 2005), a number of recent reviews (e.g., Spates, Koch, Cusack, Pagoto, & Waller, 2009; Nathan & Gorman, 2007; Ponniah & Hollon, 2009) and meta-analyses (e.g., Bisson, et al., 2007; Seidler & Wagner, 2006) conclude that EMDR is an effective treatment for PTSD. In reviewing the body of evidence for PTSD treatments to date, the recent Institutes of Medicine report (2008), however, concluded that only approaches that include an exposure component currently have sufficient evidence to be considered efficacious.

In spite of these existing approaches, there remains a need for additional PTSD treatment approaches because no single intervention is universally effective, acceptable and/or feasible (Bradley, Greene, Russ, Dutra, & Westen, 2005; Schottenbauer, Glass, Arnkoff, Tendick & Gray, 2008). Also, research surveying clinician attitudes toward exposure therapy suggests that many therapists feel uncomfortable using these treatments and infrequently use them (e.g., Becker, Zayfert, & Anderson, 2004). Thus, it is important to develop novel intervention approaches that may effectively treat PTSD in those who are unresponsive to available empirically supported approaches or who prefer a different type of intervention. Ultimately, it will be important to understand predictors of treatment response to aid clinical judgment about which treatments will be effective for which individuals.

This review provides an overview of novel treatment approaches and important modifications to established approaches based on mode of delivery or the target population. We include approaches with initial or preliminary empirical support, as well as those with a strong theoretical basis that are currently being evaluated for efficacy (see also Cukor, Spitalnick, Difede, Rizzo, & Rothbaum, 2009). Reviewed studies are constrained to psychosocial treatments for adults, organized based on the theoretical rationale for the treatment. Our objective is to review interventions that are novel and relatively understudied at this point in time, in an effort to highlight areas that will likely be addressed in future treatment outcome research.

2. Novel Interventions

We consider novel interventions to be those that have a theoretical basis that differs from those that underlie the established approaches. When sufficient data permits we report effect sizes of PTSD symptom change (Cohen's d for between-group changes when a comparison group is used or the standardized mean change g for either within-group change in the case of open trials; see Table 1).

2.1 Interpersonal Psychotherapy (IPT)

Interpersonal therapy (IPT; Weissman, Markowitz, Klerman, 2007; Markowitz, Milrod, Bleiberg, & Marhsall, 2009) is an empirically supported treatment originally developed to modify the social context of individuals with depression (e.g., de Mello, Mari, Bacaltchuk, Verdelli, & Neugebauer, 2005). Although the treatment was developed over thirty years ago, only recently has IPT been empirically evaluated for other disorders, including PTSD. IPT as applied to PTSD emphasizes the impact of the traumatic event on interpersonal behavior, with a focus on the individual's role disputes, role transitions, and loss, in an effort to both improve social functioning and decrease PTSD symptoms (Markowitz, 2010). There is reason to believe that IPT may be useful for patients with PTSD. PTSD is associated with significant social impairment (Schnurr, Lunney, Bovin, & Marx, 2009), and many symptoms of PTSD manifest in the context of social relations, including feelings of detachment, social withdrawal and isolation, irritability and angry outbursts, and lack of

trust of others (APA, 2000). Further, deterioration in social functioning is one of the strongest predictors of return for additional care after initial treatment, suggesting the importance of addressing this domain in treatment (Fontana & Rosenheck, 2010).

Few studies have evaluated IPT for PTSD and the estimated effect sizes vary widely (see Table 1), but preliminary results appear promising. In a pilot study of 14 individuals who completed IPT, 12 individuals no longer met diagnostic criteria for PTSD at post-test, and overall participants experienced significant reductions in PTSD symptoms, depression, anger, and social functioning (Bleiberg & Markowitz, 2005). In a second open trial, Robertson, Rushton, Batrim, Moore and Morris (2007) found that an eight-week IPT program led to decreased symptoms of PTSD and improved mood and general health ratings in 13 individuals. PTSD and depression symptom improvements have also been documented in Veteran samples completing IPT (Ray & Webster, 2010). Moreover, in a study that randomly assigned women to IPT or a waitlist control, PTSD and depression symptom reduction was relatively greater in the IPT group than those in the waitlist group (Krupnick et al., 2008). Given its promise and different theorized mechanism from other PTSD treatment approaches, the effects of IPT have also been evaluated for individuals who are non-responsive to other treatment. Campanini et al. (2010) provided IPT to 40 individuals who continued to have PTSD after an adequate pharmacology trial. Results indicated that after IPT, these individuals on average had lower PTSD, depression, and general anxiety and distress symptoms.

2.2. Acceptance and Commitment Therapy

The “third wave” psychotherapies, such as Acceptance and Commitment Therapy (ACT), Dialectical Behavior Therapy (DBT) and Mindfulness Based Cognitive Therapy, aim to modify the relationship between the individual and psychological experiences, rather than focusing on reduction of symptomatology (Hayes, Luoma, Bond, Masuda, & Lillis, 2006). In recent years, interest in these treatments has rapidly increased (e.g., Hayes, 2008), but only ACT has been empirically evaluated in PTSD.

These third wave therapies share the use of mindfulness to emphasize one's ability to allocate attention and awareness to the current moment, including internal experiences and environmental stimuli. In addition, these interventions emphasize acceptance of such experiences rather than passing judgment or attempting to control these experiences (Baer, Smith, Hopkins, Krietemeyer, & Toney, 2006; Bishop et al., 2004). Empirical data suggest that mindfulness qualities are positively associated with better psychological health and negatively associated with general distress (e.g., depression, anxiety; Vujanovic, Niles, Pietrefesa, Schmertz, & Potter, 2011). ACT includes mindfulness with the aim of decreasing experiential avoidance (i.e., efforts to blunt distressing inner experiences) and increasing the ability to be present in the current moment (Hayes, Strosahl, & Wilson, 1999). Additionally, ACT helps the individual to articulate his/her values and make behavioral changes that are consistent with these ideals. Published trials indicate that ACT is effective relative to waitlist control, TAU, and psychological placebo for disorders such as depression (e.g., Powers, Zum Vörde Sive Vörding, & Emmelkamp, 2009) and is comparable to cognitive therapy for depression and anxiety (Forman, Herbert, Moitra, Yeomans, & Geller, 2007).

PTSD is characterized by ineffective attempts to control and regulate internal experiences, including attempts to avoid memories and feelings associated with trauma. This experiential avoidance is associated with the maintenance or exacerbation of PTSD over time (Tull, Gratz, Salter, & Roemer, 2004), suggesting that ACT may be a useful intervention for this population. Data from open trials with inpatient veterans with PTSD indicate that ACT was associated with decreased PTSD symptoms and depression from pre to post treatment (Walser et al., 2005). ACT was also associated with increased self-reported acceptance and

decreased thought suppression (Ulmer et al., 2005). Varra and colleagues (2009) also evaluated ACT for veterans with PTSD. Results from that open trial similarly showed clinically significant decreases in PTSD and depression. Future research is needed to fully evaluate the effectiveness of this treatment for PTSD. In an effort to better understand the potential efficacy of ACT for Veterans, we are currently conducting a multi-site RCT of ACT. With a single trial reporting sufficient data to calculate an effect size, it is premature to comment on the potential efficacy of the approach. ACT merits further evaluation, however, based on its strong theoretical rationale and evidence base in other conditions (see Ost, 2008 for a meta analysis).

2.3. Mantram Repetition

After a traumatic event, individuals may feel a foreshortened sense of future in conjunction with changes in the strength or nature of their spiritual or religious beliefs (Fontana & Rosenheck, 2004). Given the effect of trauma on these types of beliefs, psycho-spiritual interventions have been considered as a complimentary therapy for the treatment of PTSD. One such intervention, mantram repetition involves selecting a mantram or meaningful phrase (e.g., Shalom), to be repeated during times of stress. The mantram is used to create a state of relaxation when autonomic arousal is high, effectively interrupting the fight or flight response (Bormann, Thorp, Wetherell, & Golshan, 2008). Other components of the treatment involve using mantram repetition to decrease reactivity to the environment and increase mindful, focused attention.

Two studies of mantram repetition suggest it may be a useful adjunct to more traditional treatments for PTSD. In the first, mantram repetition compared to TAU was conducted in a group of combat Veterans to determine feasibility and gather preliminary data on its effectiveness (Bormann et al., 2008). Comparison of the groups indicated that mantram repetition led to lower scores on measures of PTSD and quality of life (Quality of Life Enjoyment and Satisfaction scale: $d = .70$), and mindfulness (Mindfulness Attention Awareness Scale: $d = .72$). In the second study comparing mantram repetition therapy to TAU (Borman, Thorp, Wetherall, Golshan, & Lang, in press), results indicated that the mantram repetition group demonstrated significant reduction in self report and interviewer-rated PTSD as well as spiritual wellbeing (Functional Assessment of Chronic Illness Therapy Spiritual Well-being scale: $d = .61$). Thirty percent of individuals in the mantram group compared to thirteen percent in the TAU group no longer met criteria for PTSD at the end of the intervention. Moreover, drop out was low (6%) and treatment satisfaction was moderate to high in 97% of the mantram sample. Thus, this intervention is well tolerated in this population and appears to impact aspects of spiritual well-being and mindfulness qualities as well as reducing PTSD symptoms. All studies of the approach have been by a single investigative team, however, so other groups should be encouraged to attempt to replicate their results.

2.4. Psychodynamic Therapy

The psychodynamic tradition emphasizes the importance of change in internal dynamics over the course of therapy in an effort to create symptom reduction. Within this framework, transference interpretations based on the therapeutic relationship are used to resolve internal conflict and understand how the patient relates to the world and others. Focusing on the relationship, it is thought, increases insight and as a result improves the patient's functioning (e.g., Johansson et al., 2010). Although psychodynamic therapy is relatively understudied, time-limited psychodynamic treatments have received empirical support. These treatments, based on conceptualizations of trauma-focused psychotherapy outlined by Horowitz et al. (1997), include components designed to increase the patient's awareness of unconscious internal conflicts and develop insight into the patient's difficulties based on transference and

counter transference in the therapeutic relationship (e.g., Krupnick, 2002; Schnyder, 2005). The therapist works with the patient to understand and process underlying beliefs and attitudes that have prevented integration of the traumatic event and to understand the personal meaning of the event.

A number of published case studies outline the conceptual rationale and implementation of psychodynamic approaches with PTSD patients (e.g., Krupnick, 2002; Moss, 2009), but few published randomized trials of psychodynamic treatments exist. Brom, Kleber, and Defares (1989) randomized patients to psychodynamic treatment, hypnotherapy, systemic desensitization, or waitlist. Results revealed the active conditions were equally effective at treating PTSD symptoms and that the psychodynamic treatment was more effective than the waitlist condition on symptoms related to trauma. Thus, although these treatments may appear to be promising based on this single study, future research will be necessary before any conclusions about efficacy can be drawn.

2.5. Modification of Cognitive Processing

Extant empirical data indicate that anxiety disorders are characterized by specific neuropsychological and information processing abnormalities relative to healthy individuals, including biases in attention (e.g., Bar-Haim et al., 2007), interpretation (Amir, Beard, Przeworski, 2005), memory (Mitte, 2008) and executive functioning (Eysenck, Derakshan, Santos, & Calvo, 2007). Theoretical accounts suggest these abnormalities maintain anxiety by biasing cognition in favor of negative, threatening information. In support of this proposition, a number of studies indicate that information processing characteristics, such as attention and interpretation bias, may be causally implicated in anxiety (e.g., Mathews & MacLeod, 2002; Wilson, MacLeod, Mathews, & Rutherford, 2006). Evidence for a causal relationship between cognitive processes and anxiety symptoms suggests that experimental manipulation of these processes may be a promising treatment target for disorders such as PTSD. PTSD is characterized by at least two forms of aberrant cognitive processing that may be appropriate targets for this type of intervention, including information processing biases and executive functioning deficits.

Experimental psychopathology studies indicate that individuals with PTSD are characterized by information processing biases including attention bias toward threatening cues (e.g., Buckley, Blanchard, & Neill, 2000; Constans, 2005; Pineles, Shipherd, Mostoufi, Abramovitz, & Yovel, 2009; Pineles, Shipherd, Welch, & Yovel, 2007). This bias is thought to contribute to heightened sense of threat and increased distress and, potentially by increasing behavioral avoidance, to prevent the distress associated with encountering threat (e.g., Pineles et al., 2009). Consequently, programs designed to alter this type of “mental habit” by training attentional allocation *away from* threat at a basic information processing level may provide an effective way to both test theoretical models of causality and develop novel interventions. Researchers have utilized a computer-based program to facilitate attention disengagement from threat (relative to neutral stimuli) in individuals with generalized social phobia (Amir et al., 2009; Schmidt, Richey, Buckner, & Timpano, 2009) and generalized anxiety disorder (Amir, Beard, Burns, & Bomyea, 2009). These interventions utilize brief, computer-administered programs completed over relatively short duration (e.g., 6 weeks) to alter these “mental habits”. In spite of their brevity and lack of therapist contact, results from these studies indicate that individuals in these programs demonstrate decreased anxiety symptoms relative to those in control groups.

To date no published studies have evaluated the effect of attention modification in individuals with PTSD. However, pilot data from ongoing trials of attention modification programs in PTSD samples is promising (Amir et al., personal communication). For example, a preliminary feasibility study of 29 active duty military participants given

attention modification in addition to treatment as usual demonstrated that participants experienced a significant decrease in symptoms [assessed using the Posttraumatic Stress Disorder checklist (PCL-M) and Beck Depression Inventory –II (BDI-II)]. In addition, data from a pilot study of 37 PTSD in-patients randomized to TAU, TAU plus attention modification, or TAU plus the attention control condition indicate that individuals in the combined treatment condition demonstrated relatively greater decreases in symptoms on the PCL-M and BDI-II relative to those in the TAU plus attention control condition. In summary, these preliminary data and the success of these programs in treating other anxiety disorders in a relatively brief and cost efficient manner points to their potential utility for PTSD. Trials evaluating this type of intervention are currently being conducted (N. Amir, B. Olatunji, personal communication).

Executive functioning deficits may be another form of cognitive processing suitable to a computer-based intervention. A number of studies demonstrated that specific cognitive deficits related to executive control ability differentiate individuals with and without PTSD (for reviews see Buckley et al., 2000; Constans, 2005; Mathews & MacLeod, 2005). Corroboration for these findings comes from neuroimaging studies indicating that neural activation in areas implicated in executive control appears to be relatively diminished in individuals with PTSD compared to healthy individuals, while areas regulating emotional reactivity are over-active (for reviews see Etkin & Wagner, 2007; Francati, Vermetten, & Bremner, 2007). Recent theoretical accounts of this phenomenon posit that the persistence of these thoughts in clinical disorders such as PTSD stems from deficits in basic cognitive systems that regulate the inhibition of information (e.g., Anderson & Levy, 2009; Joormann, Yoon, & Siemer, 2010; Verwoerd, de Jong, & Wessel, 2008). Consistent with these theories, research suggests that individuals with poor executive functioning performance – particularly on tasks relying on inhibitory control - have difficulty suppressing and regulating intrusive cognitions (e.g., Brewin & Smart, 2005; Verwoerd, Wessel, & de Jong, 2009; Wessel, Overwijk, Verwoerd, & de Vrieze, 2008). Moreover, disorders characterized by intrusive symptoms, such as PTSD, are characterized by difficulty with executive functioning tasks, including those that rely on inhibition (e.g., Constans, 2005). By this theoretical account, doing activities that improve components of executive functioning and thus strengthen prefrontal cortex activation may ameliorate PTSD symptoms (McNally, 2007).

Although no published empirical data on executive functioning training program exists in anxiety, cognitive research suggests that such training can potentially improve executive functioning (e.g., Persson & Reuter-Lorenz, 2007). Recent data also indicates that working memory capacity improvements as a result of such training improve individuals' ability to suppress negative, personally relevant memories (Bomyea & Amir, 2010). Our laboratory is currently conducting a larger-scale clinical trial using this executive functioning training program in a sample of individuals with PTSD to determine the effect of such training on cognitive functioning and symptoms. Clearly this work is in its nascence, but it has the advantage of potentially appealing to those who do not want to engage in traditional psychotherapy.

3. Modified Mode of Delivery: Telemedicine and Computer delivered CBT

The interventions described in this section generally employ exposure-based therapy or cognitive therapy. Thus, they are not departures in terms of the nature of the intervention. They do, however, strive to reduce barriers to accessing effective care for PTSD. There are a number of advantages to delivering CBT by telephone, videoconferencing, computer or Internet. Many people do not live near a provider that is trained in empirically supported approaches or cannot access care for logistical reasons or because of avoidance of leaving

home or traveling. The option to contact a mental health provider remotely may enhance access to good care. Telemedicine and computer-delivered therapy also lends privacy that may appeal to populations concerned about stigma attached to seeking mental health care (e.g., military personnel; Hoge et al., 2004). Meta-analyses suggest this type of intervention is effective for anxiety more generally relative to waitlist and placebo control groups and is equivalent to treatment as usual (TAU) conditions (telehealth: Cuijper et al., 2009; computer-delivered CBT: Reger & Gahm, 2009). A concern about such interventions, however, is that the therapeutic alliance may be compromised thereby rendering the interventions less effective.

Technology-facilitated methods for delivering traditional care remotely typically involve some form of live clinician contact, such as videoconferencing with a therapist. A number of pilot studies have used this type of remote therapy (e.g., Frueh et al., 2007; Morland, Pierce, & Wong, 2004; Tuerk, Yoder, Ruggiero, Gros, & Acierno, 2010; Germain, Marchand, Bouchard, Drouin, & Guay, 2009). In general results suggest that remote therapies show promise as effective treatments for reducing PTSD and general distress. Examination of effect sizes indicates that the magnitude of treatment effects for telehealth delivery versus in-person contact is roughly equivalent (e.g., Germain et al., 2009). Moreover, recent research examining therapist variables indicate that ratings of therapist adherence, rapport building, and expressive empathy are similar across these delivery forms (Frueh et al., 2007).

In addition to CBT delivered remotely through the use of technology, the possibility of using computer-administered CBT (either via written therapist instructions or computer-guided modules) has also been evaluated. Pilot work suggests that programs delivering therapy contents directly via the computer are effective in reducing PTSD symptoms (e.g., Klein et al., 2009). In addition, initial RCTs using computer-assisted interventions indicate this modality effectively decreases PTSD symptoms. Knaevelsrud and Maerker (2007) report that relative to a waitlist control group, individuals ($N = 96$) randomized to 10 sessions of therapist-assisted internet-based CBT showed decreases in PTSD symptoms. Later analysis indicated that these effects were maintained over an 18-month follow-up period (Knaevelsrud & Maerker, 2010). In a sample of 45 active duty personnel, Litz and colleagues (2007) also demonstrated that relative to internet-based supportive therapy individuals completing the active DESTRESS program demonstrated relatively greater improvement in PTSD symptoms. Ratings from these programs indicate high treatment satisfaction and, in cases where therapist contact is completed (e.g., via email), high ratings of therapeutic alliance (Klein et al., 2009; Knaevelsrud et al., 2007).

Difficulties inherent in the comparison of two active treatments (e.g., large samples required to detect relatively small effects) limit conclusions that can be drawn presently regarding the effects of telehealth CBT relative to traditional in-person approaches. Nonetheless, in trials comparing remote-delivery methods and in-person treatments the effect on PTSD and general distress symptoms appear to be comparable (e.g., Germaine et al., 2009). In addition, the combined data on effectiveness, therapeutic adherence and satisfaction with the therapeutic alliance suggest that interpersonal rapport building may be attainable even when interactions are not “in vivo”. Head-to-head comparison is currently being evaluated in larger scale trials to further evaluate efficacy relative to in-person approaches (e.g., S. Thorp, personal communication). The completion of these non-inferiority trials will facilitate conclusions about whether telehealth CBT can match gold-standard in-person interventions for those who are unable or unwilling to obtain treatment in traditional clinical settings.

4. Adaptations to Target Specific Populations

In this final section, the interventions generally are built upon cognitive behavioral principles, but they have been modified to better meet the needs of specific populations. Up to this point, there has been little differentiation in the treatment of PTSD based on trauma type. These interventions reflect the developing understanding that the nature of the event may affect the challenges faced by the traumatized person and that interventions should be tailored accordingly.

4.1. Cognitive Trauma Therapy for Battered Women

Cognitive Trauma Therapy for Battered Women (CTT-BW; Kubany & Watson, 2002) relies on cognitive behavioral techniques with particular emphasis on guilt. The rationale for this is that guilt-related cognitions that are common to this population (e.g., beliefs about preventability of events, failure to consider retrospective justification for actions and decisions). Also included are components addressing assertiveness skills and prevention of revictimization by the individual's spouse or other future partners.

Two published studies evaluate CTT-BW for this population. In the first, 37 female domestic violence survivors with PTSD were randomly assigned to complete CTT-BW either immediately or after a delay of approximately six weeks (Kubany, Hill, & Owens, 2003). Results indicated that after participation in the active CTT-BW component, women reported lower symptoms of PTSD, depression and guilt, and higher self-esteem scores. Overall, 94% of the women completing the therapy no longer met diagnostic criteria for PTSD at post-assessment, and treatment gains were maintained at 3-month follow up. In the second study, 125 women were assigned to either immediate or delayed CTT-BW (Kubany et al., 2004). PTSD symptoms, depression, self esteem, and guilt were assessed before treatment, at post treatment, and at 3 and 6 month follow up. Consistent with Kubany et al. (2003), women receiving treatment demonstrated relatively greater symptom improvements on PTSD, depression, and guilt scores from pre to post assessment relative to those in the waitlist condition, and gains were maintained at follow up. Given these results, CTT-BW holds promise for this population and, perhaps in a modified form, other traumatized groups characterized by excessive guilt (Kubany & Watson, 2002). These types of interventions are currently being evaluated in female survivors of domestic violence (C. Allard, personal communication) as well as combat veterans (S. Norman, personal communication). It remains to be seen, however, if these modifications improve on gold-standard interventions.

4.2. Adaptive Disclosure

Many models of PTSD focus on the experience of fear during and after trauma. There is growing recognition that combat is not always experienced as fear-provoking but is associated with a spectrum of emotional reactions, including disgust, loss or grief (e.g., Neria & Litz, 2004) and reactions (e.g., guilt and betrayal) to perpetrating or witnessing acts that violate one's moral values – also referred to as “moral injury” (e.g., Litz et al., 2009). Accordingly, Litz and colleagues (see Steenkamp et al., in press) developed a manualized intervention for active duty populations called Adaptive Disclosure (AD) to specifically address grief and moral injury. AD uses a trauma narrative to increase an individual's awareness of strongly held beliefs about the event and to disconfirm maladaptive beliefs about disclosure of the event (cf. PE wherein the narrative is used to foster habituation to the trauma memory). Once beliefs are brought to one's awareness, a variety of techniques, including Socratic questioning, in vivo exposure and the “empty chair” technique are used to modify beliefs. AD also was carefully designed to be respectful of and congruent with military culture. Data from a nonrandomized program evaluation involving 56 Marines showed a positive impact of AD on PTSD, depression, trauma-related cognitions and coping

(Litz, unpublished data). Our group is testing this intervention in a large RCT at the Marine base at Camp Pendelton, California to evaluate efficacy as compared to CPT.

4.3. Narrative Exposure Therapy

Narrative Exposure Therapy (NET) was developed for civilian individuals who have experienced war-related traumas, including torture, terrorism or being a refugee or displaced person. Socioeconomic and political factors, such as extreme poverty and relocation, in conjunction with the severe trauma histories (often including chronic trauma exposure) and cultural/linguistic differences suggest that specifically tailored treatment interventions may be helpful for these individuals (e.g., Neuner, Schauer, Klaschik, Karunakara, & Elbert, 2004).

Drawing on techniques from PE and Testimony Therapy (Cienfuegos & Monelli, 1983), NET (Schauer, Neuner, & Elbert, 2005) attempts integrate and organize the individual's autobiographical memory of the traumatic event and to bring meaning to the experience by allowing the individual to “bear witness” to what they have experienced (Neuner, Schauer, Roth, & Ebert, 2002). This is accomplished by the patient and therapist collaboratively constructing a full biography of the patient's life, including the traumatic events. This process helps the patient to access and process cognitive and emotional reactions to the trauma, while simultaneously creating a written account of the trauma that becomes a history or can contribute to legal proceedings. Similar to exposure-based therapy, NET requires the individual to recount traumatic events in rich emotional detail and, therefore, may help to integrate emotional memories with autobiographical memory (Robjant & Fazel, 2010). NET differs from traditional exposure-based therapy, however, in that the narrative is of the individual's *entire* life (rather than a single “worst” traumatic event) and is not repeated over and over in therapy. NET is also specifically tailoring to be administered in settings such as refugee camps. In addition, NET greatly emphasizes making meaning of and bearing witness to the events.

A growing body of research indicates that NET is a promising treatment for refugee populations. Bichescu, Neuner, Schauer & Elbert (2007) randomly assigned individuals who had been political detainees to NET or psychoeducation treatment. Six months after treatment, the NET group, as compared to the psychoeducation group, experienced relatively fewer symptoms of PTSD and depression. Similarly, Neuner et al. (2004) found that African refugees who were randomly assigned to NET had lower PTSD symptoms at one-year follow-up than those in a psychoeducation group or a supportive counseling group. Neuner et al. (2010) also found that asylum-seekers randomly assigned to NET had relatively better PTSD symptom improvement at six month follow-up relative to those in a TAU condition (although all but one participant in the study continued to meet diagnostic criteria for PTSD). However, in a different sample of African refugees, Neuner and colleagues (2008) found that NET was superior to a no-treatment group, but not a trauma counseling group, on symptoms of PTSD and depression. In a trial of NET versus interpersonal therapy, these two active treatments appear to fare equally well (Schaal, Elbert, & Neuner, 2009). See Robjant and Fazel (2010) for a currently review of published and unpublished trials of NET in children and adults. NET remains to be compared to the well established approaches.

Although NET trials typically conduct assessments later than traditional treatment trials and contain variability in the number and setting of treatments (e.g., in rural communities, hospital settings) they may also possess greater ecological validity and be perceived as very acceptable by these populations. Anecdotal reports from these studies indicate that participants felt the writing of the narrative was both personally meaningful and significant, because although the majority of the individuals could not read, they felt there was value in

making their story accessible to future generations. It is important to note that these trials are typically conducted in individuals who are still experiencing unsafe conditions and persecution, threat of repatriotization (Neuner et al., 2010) and take place in settings with minimal resources, and yet still indicate that NET may possess therapeutic potential for these types of populations. Currently at least one randomized controlled study is underway in the Netherlands to better understand the effect of NET relative to treatment as usual for individuals with Borderline PD and PTSD (Clinicaltrials.gov).

4.4. Imagery Modification

Focusing on anxiety-relevant images has a long history in CBT (for a review see Holmes, Arntz, & Smucker, 2007) and has recently been applied specifically to change images in the context of sleep disturbance. Based on the premise that triggering traumatic imagery activates is distress (e.g., Brett & Orstroff, 1985), the aim of these treatments is to replace trauma-based images with more positive ones to reduce the associated negative emotion (for reviews of theoretical mechanisms see Moore & Krakow, 2010; Long & Quevillon, 2009). Such treatments include Imagery Rehearsal Therapy (IRT; e.g., Krakow et al., 2000) and Exposure, Relaxation, and Rescripting Therapy (ERRT; Davis, 2003). In contrast to exposure, which entails specific review of and habituation to the traumatic memory, imagery based treatments typically minimize recounting of the trauma. Instead, participants are guided to substitute images and nightmares related to trauma with alternate mental images.

Given the high prevalence of sleep disturbances in PTSD, imagery rescripting techniques for nightmares have been specifically examined in this population. Although the exact mechanism of imagery rescripting techniques is unknown, it is theorized that repeated practice “over-writing” the traumatic image with a non-distressing image may increase mastery over nightmare content (Nappi et al., 2010). Support for the approach comes from preliminary randomized trials and open trials (e.g., Krakow et al., 2000; Forbes, Phelps & McHugh, 2001; Lu, Wagner, Van Male, Whitehead, & Boehnlein, 2009; Nappi, Drummond, Thorp, & McQuaid, 2010). To date, three larger RCTs of imagery rescripting techniques for PTSD-related nightmares have been published. Krakow et al. (2001) randomized 168 women with PTSD related to sexual assault to a three-session group IRT protocol or a waitlist control group. At the conclusion of treatment, women in the IRT condition experienced significantly fewer nightmares per week, improved sleep quality, and lower PTSD symptoms than the control group. These treatment gains were maintained at three and six-month follow-up. Davis and Wright (2007) randomly assigned 43 individuals to ERRT or a waitlist. Participants in the ERRT group experienced decreased symptoms of depression, PTSD, and sleep problems. Cook et al. (2010) randomly assigned 124 Vietnam War veterans with PTSD to weekly group IRT or a sleep and nightmare management control group that included psychoeducation and components of CBT for insomnia. Results indicated that the IRT group had not improved significantly more than the control group on nightmare frequency, sleep quality or PTSD. Based on these mixed results, imagery rescripting procedures need further evaluation before implementation (for a review of recent sleep treatments in PTSD see Nappi et al., 2010; this issue).

5. Summary and Discussion

Recent advancements using methodologically sound designs increase confidence that PTSD can be effectively ameliorated using manual-based treatments including exposure, cognitive restructuring, or a combination of the two, as well as using EMDR. The availability of such evidence-based treatments paints a promising view for individuals suffering with PTSD. Nonetheless, a significant portion of individuals remain symptomatic after treatment, drop out of treatment, or never access empirically-supported treatments (Hoge et al., 2004;

Schottenbauer et al., 2008). Thus continued research on PTSD interventions and delivery models is certainly warranted.

Emerging interventions covered in the present review begin to address remaining questions in the PTSD treatment literature. First, there are a set of developing interventions that have different conceptual bases for treating PTSD. These interventions are derived from different theoretical models and emphasize different etiologic and maintenance factors in PTSD. Thus, they may be better suited to patients who are not helped by the well established approaches. For example, IPT may be particularly well suited to individuals whose symptoms are maintained or exacerbated by interpersonal dysfunction. Third wave approaches may be a particularly good fit for those who are compelled by living in accordance with one's values. Cognitive process modification programs provide an option for individuals who are unwilling to participate in traditional psychosocial interventions (e.g., those with high levels of avoidance behavior). A review of the effect sizes of these novel interventions yields considerable variability. In trials examining IPT, effect sizes for this type of intervention appear similar to the range of cognitive behavioral approaches in prior reviews of pre-post change within cognitive behavioral approaches (range 1.4–1.7) and between group change (1.2–1.5; Bradley et al., 2005). These findings, although preliminary, are consistent with studies demonstrating the efficacy of IPT for depression. Other studies, such as those of mantram repetition, appear to have relatively smaller effects on PTSD but do impact outcomes related to mindfulness and spirituality. These findings both highlight the need for future work both establish efficacy and point to the importance of evaluating secondary related variables that may be directly targeted in these specific treatment modalities.

A second set of interventions has attempted to increase accessibility, engagement in, or acceptability of CBT. Remotely delivered treatments capitalize on the accessibility of the World Wide Web and telehealth to disseminate treatments to individuals who might not otherwise access empirically based interventions. These interventions appear beneficial for symptom reduction. In addition, trials presented do not substantiate concerns about difficulty creating or maintaining therapeutic alliance or therapist adherence without face-to-face contact.

Finally, we have reviewed treatments such as CTT, AD, and NET, which tailor CBT for specific populations (i.e., active duty military, refugees). These efforts aim to build upon extant literature regarding the effectiveness of trauma-focused CBT in such a way that treatment outcomes are improved or bolstered for individuals who might otherwise benefit less from these treatments. These approaches appear well-tolerated in their specific populations. However, ultimately, all of these interventions will need to be evaluated against the well established treatments in terms of acceptability and effectiveness.

Reviewing future directions for PTSD treatment research highlights critical questions in the literature that transcend the specific interventions studied. A review of extant treatment outcome literature in PTSD reveals that studies directly examining mechanisms of treatment response are relatively rare (although see for example Smith et al., 2007). It is likely that treatments tapping different targets than CBT do not have entirely distinct mechanisms. For example, techniques requiring the patient to complete activities (e.g., committed action in ACT, in-vivo exposure, interpersonal psychotherapy) inherently contain behavioral activation components. All psychosocial treatments (with the exception of computer-based techniques such as cognitive process modification or Internet based self-help) contain a supportive interpersonal relationship that likely drives a portion of therapeutic change. Thus, relatively little is known about what makes empirically supported interventions effective or how treatments tapping “different” mechanisms might be therapeutic.

Moreover, it has been proposed that interventions might be matched to individuals. Although empirical examination of predictors of outcome is beginning (e.g., Evans, Cowlshaw, & Hopwood, 2009; Iverson, Resick, Suvak, Walling, & Taft, in press; Tarrier, Sommerfield, Pilgrim, & Faragher, 2000), the lack of research on individual-level treatment moderators in RCTs leaves this proposal relatively understudied (although see for example Thrasher, Power, Morant, Marks, & Dalgleish, 2010). To date, no published study has examined the potential for matching treatment protocol based on symptom profile. Knowledge regarding the efficacy of more personalized interventions is thus warranted and will likely emerge in future work. Knowledge derived from these studies will be critical for understanding both who is an ideal candidate for what treatment and may also inform theoretical accounts of PTSD regarding the mechanisms of specific treatments.

In summary, the intervention approaches reviewed show promise for improving PTSD treatment. Because these interventions are in relatively early stages of treatment development, many suffer from methodological limitations such as small samples or lack of randomization. Thus, future work is needed to evaluate efficacy of these treatments using increasingly stringent research methodology. The treatment approaches highlighted here point to future directions for research, including increasing accessibility and acceptability of treatment for specific populations and testing alternative treatment models. In addition, studies designed to better understand the mechanisms of different treatments and inform clinical decision-making about treatment selection have the potential to inform our clinical practice and future research. Ultimately, this knowledge will translate into a better understanding of not only what works, but what works for whom and under what circumstances.

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Highlights

- Novel PTSD interventions have been proposed, but they are inadequately studied and effect size estimates vary considerably.
- Tailoring of empirically supported treatments to remote delivery formats appears to be generally well tolerated and efficacious.
- Modification of current treatments to specific populations shows promise but merits additional study.

Table 1

Effect sizes of novel interventions of PTSD symptoms

Therapy Type	Study	Sample	Design	Measure	Effect Size
Interpersonal Psychotherapy	Bleiberg & Markowitz, 2005	14 men and women Mixed trauma type	Open trial	CAPS	$g = 2.8$
				PSS	$g = 2.4$
	Robertson et al. 2007	13 men and women Mixed trauma type	Open trial	IES – R Avoidance	$g = .57$
				IES – R Hyperarousal	$g = .64$
	Campanini et al. 2010	40 men and women Treatment refractory sample	Open trial	CAPS	$g = 1.29$
Krupnick et al. 2007	48 women Low income sample	Random assignment: IPT (n = 32) Waitlist (n = 16)	CAPS	$d = 1.17$	
Acceptance and Commitment Therapy	Ray & Webster, 2010	9 veterans	Open trial	IES - R	ES not available
	Varra et al., 2009			CAPS	0.33
Mantram Repetition	Walser et al., 2005	161 inpatient veterans	Open trial	PCL – M	ES not available
	Bormann et al. 2008	29 combat veterans	Random assignment: Mantram repetition (n = 14) Delayed mantram repetition (n = 15)	CAPS	$d = 0.33$
Psychodynamic Therapy	Borman et al. in press	136 veterans	Random assignment: Mantram repetition and TAU (n = 66) TAU only (n = 70)	PCL	$d = 0.72$
				CAPS	$d = 0.39$
				PCL	$d = 0.34$
Attention Modification Program	Amir et al. (unpublished data)	29 Active duty personnel	Open trial	IES-R	$d = 0.67$
		37 Active duty personnel	Randomized to TAU, TAU+AMP, TAU+ACC	SCL-90 Trauma scale	$d = 0.83$
				PCL-M	ES not available

Note: CAPS = Clinician Administered PTSD scale; PSS = Posttraumatic Stress Scale; HRSD IES-R = Impact of Events Scale-Revised; PCL = Posttraumatic Symptoms Check List; SCL-90 = Symptom checklist-90 subscale with trauma-related items; TAU = Treatment as Usual; AMP = Attention Modification Program; ACC = Attention Control Condition. Given the heterogeneity in effect size calculations presented in the open trial studies reviewed, a standard mean change effect size was calculated for each study as per Becker (1998): $g = (x_1 - x_2)/s_{pre-test}$