

## Outline



Nightmare Definition/Prevalence/Evaluation



**Nightmares in the Military** 



**Treatment** 

Non-pharmacologic Treatment

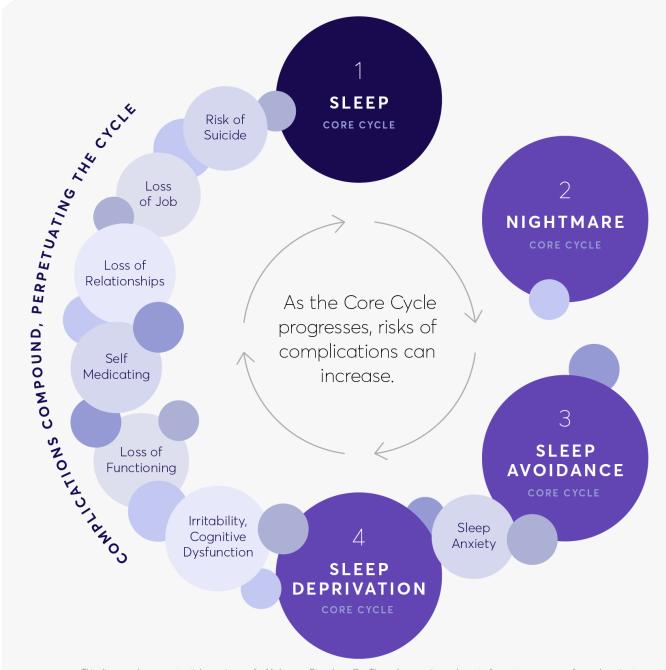
Pharmacologic Treatment

# Nightmares are the Most Prevalent Parasomnia in Pediatric and Adult Populations in the US

Parasomnia	Adult Lifetime Prevalence (%)	Children Prevalence (%)*
NREM		•
Sleepwalking	22.4	17.0
Confusional Arousal	18.5	17.3
Sleep Terror	10.4	6.5
Sleep-related eating disorder	4.5	
Sexual act during sleep	7.1	
REM		
REM behavior disorder	15.0	
Sleep related groaning	31.3	
<ul> <li>Nightmares</li> </ul>	66.2	10-50

## Nightmares

- Associated with anxiety, depression, and PTSD
- Associated with heightened risk of suicidal ideation
- Treatment can lead to improvement in comorbid anxiety, depression, and PTSD



This diagram shows a potential experience of a Nightmare Disorder suffer. The order, severity, and onset of symptoms can vary for each patient.

## Nightmare Disorder (NDO)

- Repeated occurrences of disturbing, wellremembered dreams that cause clinically significant distress or impairment
- Trauma related nightmares (TRN) follow a traumatic experience
- Clinically significant nightmares occur at least weekly
- Few studies evaluate NDO in military personnel

## Diagnostic Criteria for Nightmare Disorder (NDO)

#### Minimal Diagnostic Criteria: 3rd Edition ICSD1

**A**. Repeated occurrences of extended, extremely dysphoric, and well-remembered dreams that usually involve threats to survival, security, or physical integrity

- **B.** On awakening from the dysphoric dreams, the person rapidly becomes oriented and alert
- **C.** The dream experience, or the sleep disturbance produced by awakening from it, causes clinically significant distress or impairment in social, occupational, or other important areas of functioning as indicated by the report of at least one of the following:
  - 1. Mood disturbance (e.g., persistence of nightmare effect, anxiety, dysphoria).
  - 2. Sleep resistance (e.g., bedtime anxiety, fear of sleep/subsequent nightmares).
  - 3. Cognitive impairments (e.g., intrusive nightmare imagery, impaired concentration, or memory)
  - 4. Negative impact on caregiver or family functioning (e.g., nighttime disruption)
  - 5. Behavioral problems (e.g., bedtime avoidance, fear of the dark)
  - 6. Daytime sleepiness
  - 7. Fatigue or low energy
  - 8. Impaired occupational or educational function
  - 9. Impaired interpersonal/social function

Nightmare disorder (ND))

Per the American Academy of Sleep Medicine, NDO affects 4% of the adult population in the US<sup>2</sup>

# DSM-5: NDO Criterion

- A. Repeated occurrences of extended, extremely dysphoric, and well remembered dreams that usually involve threats to survival, security, or physical integrity and that generally occur during the second half of the major sleep episode.
- 3. On awakening from the dysphoric dreams, the person rapidly becomes oriented and alert.
- C. The sleep disturbance causes clinically significant distress or impairment in social, occupational, or other important areas of functioning.
- D. The nightmare symptoms are not attributable to the physiological effects of a substance.
- E. Coexisting mental and medical disorders do not adequately explain the predominant content of dysphoric dreams.

Descriptors								
Severity		Duration						
Mild	< 1 per week on average	Acute	≤ 1 month					
Moderate	>1 per week	Subacute	>1 month and < 6 months					
Severe	Nightly	Chronic	≥ 6 months					

**PSQI** 

Pittsburgh Sleep Quality Index

_					
Dur	na t	ha.	nact	mon	th:

1	What t	tim o	have	 mentally	, gone	to.	had	-+ ·	niah+2	

- 2. How long (in minutes) has it taken to you to fall asleep each night?
- 3. What time have you usually gotten up in the morning?
- 4. A. How many hours of actual sleep did you get at night?
- B. How many hours were you in bed?

BED TIME
NUMBER OF MINUTES
GETTING UP TIME

5. During the past month, how often have you had trouble sleeping because you:	Not during the last month (0)	Less than once a week (1)	Once or twice a week (2)	Three or more times a week (3)
A. Cannot get to sleep within 30 minutes				
B. Wake up in the middle of the night or early morning				
C. Have to get up to use the bathroom				
D. Cannot breathe comfortably				
E. Cough or snore loudly				
F. Feel too cold				
G. Peer too not				
H. Have a bad dream				
i. Have pain				
Other reason(s), please describe including how often you have had trouble sleeping because of this reason(s):				
During the past month, how often have you taken medicine (prescribed or over the counter) to help you sleep?				
7. During the past month, how often have you had trouble staying				

8.	During the past month, how much of a problem has it been for you to keep up enthusiasm to get things done?
	□ Not a problem at all (0) □ Only a very slight problem (1) □ Somewhat of a problem (2) □ A very big problem (3
٥	How would you rate your cleen quality overall?

☐ Very good (0) ☐ Fairly good (1) ☐ Fairly bad (2) ☐ Very bad (3)



#### Psychiatry Research

Volume 28, Issue 2, May 1989, Pages 193-213



The Pittsburgh sleep quality index: A new instrument for psychiatric practice and research

## PSQI-A

Pittsburgh Sleep Quality Index Addendum for PTSD

					~	
During the past month, how often (can be based on report of roommate or bed	Never	Not during the past	Less than once a week	Once or twice a week	Three or more times a	I do not know
partner)		month	Office & Week	twice a week	week	KIIOW
do you wake with a headache?						
do you wake with a dry mouth?						
do you stop breathing while sleeping?						
do you wake gasping or choking?						
do you snore loudly enough to be noticed?						
do you snore loudly enough that you were						
told you bothered some else's sleep?						
do you have leg twitching or jerking while you sleep?						
are you BOTHERED by the urge to move your legs for comfort as you fall asleep or sit?						
do you grind/clench your teeth at night?						
do you walk in your sleep?						
do you experience very vivid dreams while falling asleep or waking up?						
are you unable to move while falling asleep or waking up?						
are you unable to move arms or legs when						
laugning or reeling other strong emotions?						
have you had memories or nightmares of a traumatic experience?						
had episodes of terror or screaming during						
sleep without fully awakening?						
had episodes of "acting out" your dreams such as kicking, punching, running, or screaming						



#### Journal of Anxiety Disorders

Volume 19, Issue 2, 2005, Pages 233-244



A brief Sleep Scale for Posttraumatic Stress Disorder: Pittsburgh Sleep Quality Index Addendum for PTSD

Anne Germain <sup>a</sup> 🙎 🖂 , Martica Hall <sup>a</sup>, Barry Krakow <sup>b c</sup>, M. Katherine Shear <sup>a</sup>, Daniel J. Buysse <sup>a</sup>



SLEEPJ, 2021, 1-11

doi: 10.1093/sleep/zsaa254

Advance Access Publication Date: 27 November 2020

Original Article

### ORIGINAL ARTICLE

## The Nightmare Disorder Index: development and initial validation in a sample of nurses

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Sophie Wardle-Pinkston<sup>2,†</sup>, Danica C. Slavish<sup>4,o</sup>, Brett Messman<sup>4</sup>, Rosemary Estevez<sup>5</sup>,
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Tucson, AZ <sup>3</sup>Department of Psychiatry and Behavioral Sciences, University of Texas Health Sciences Center at San Antonio, San Antonio, TX
<sup>4</sup>Department of Psychology, University of North Texas, Denton, TX <sup>5</sup>Mental Health Flight, Maxwell Air Force Base, AL

## Nightmare Disorder Index

### Thinking about a typical night in the last month ...

1.	<ol> <li>How many nights a week did you have nightmares (i.e., disturbing, extended, well- remembered dreams)?</li> </ol>	0 nights per week	< 1 nights per week	1-3 nights per week	4-6 nights per week	7 nights per week
		(0)	(1)	(2)	(3)	(4)

### Skip to next questionnaire if answered "0" above.

2.	How often do you wake up from your nightmares AND quickly become alert?	Never (0)	Rarely (1)	Sometimes (2)	Often (3)	Always (4)
3.	To what extent have nightmares troubled/distressed you in general?	Not at all (0)	A little (1)	Somewhat (2)	Much (3)	Very Much (4)
4.	To what extent have nightmares caused difficulties in social, work, or other areas of life?	Not at all (0)	A little (1)	Somewhat (2)	Much (3)	Very Much (4)
5.	How long have you been bothered by nightmares?	Less than 1 week (0)	Less than 1 month (1)	1-6 months (2)	6-12 months (3)	>12 months (4)

#### Categorical Scoring Instructions:

Items 1-4 on the NDI correspond to the DSM-5 criteria for nightmare disorder and thus are used to obtain diagnostic categories. Item 5 is not used in the assessment of categorical scores but can be used to denote the acuity specifier. Please note that a diagnosis cannot be made without a clinical interview; the NDI is meant to serve as a screening tool and should not be used to assign a definitive diagnosis.

No Nightmare Disorder: A respondent is assigned this category if they answer "0" for item 1.

Subthreshold/Partial Nightmare Disorder Symptoms: A respondent is assigned this category if they respond 1-4 (i.e., <1 night per week to 7 nights per week) on item 1 AND respond 0 or 1 for any item 2-4. This indicates that while the respondent has experienced nightmares on a consistent basis in the past month, they do not endorse all symptoms at a level severe enough to be included in the category of probable nightmare disorder.

Probable Nightmare Disorder: A respondent is assigned this category if they respond 1-4 on item 1 AND ≥2 on all items 2-4.

A <u>severity</u> specifier can be assigned to respondents falling in this category based on the response to item 1 as follows: 1 - Mild; 2-3 - Moderate; 4 - Severe.

An <u>acuity</u> specifier can be assigned to respondents falling in this category based on the response to item 5 as follows: 1 – Acute; 2 – Subacute; 3-4 – Persistent



## Degree of Trauma Exposure

- Dream recall/nightmares after Loma Prieta earthquake
  - Assessed college students
  - Witnessed direct impa
- School shooting night
  - 63% in school yard
  - 56% of those in the school
  - 43% of those at home
  - 33% of children geographically away

## Proximity effects dream content

# Trauma and Dream Recall Frequency

## Changes in dreaming

- Severely traumatize dreams
- Palestinian childre dreams compared to children living in peaceful Galilee
  - •Increased trauma

## Temporal changes in dream recall

- Recall increases with stress/trauma
- Tends to decrease months to years after trauma

## Nightmares: Prevalence among the Finnish General Adult Population and War Veterans during 1972-2007

Nils Sandman, MSc<sup>1,2</sup>; Katja Valli, PhD<sup>2,3</sup>; Erkki Kronholm, PhD<sup>4</sup>; Hanna M. Ollila, PhD<sup>1</sup>; Antti Revonsuo, PhD<sup>2,3</sup>; Tiina Laatikainen, PhD<sup>4,5</sup>; Tiina Paunio, MD, PhD<sup>1,6</sup>

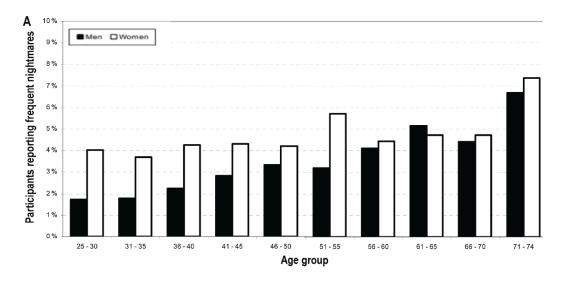
## **Frequent nightmares**

• Men: 3.5%

Women: 4.8% (p<0.0001)</li>

## Prevalence affected by

- Sex
- Age
- Year of survey



**Table 5**—Nightmares and symptoms of insomnia, depression, and anxiety among the war generation and the general population in the combined sample of participants of the years 1972, 1977, 1982, 1987, and 1997

## War experiences

•	n	İ	g	h	t	m	a	r
•	İ	n	S	0	m	n	i	a

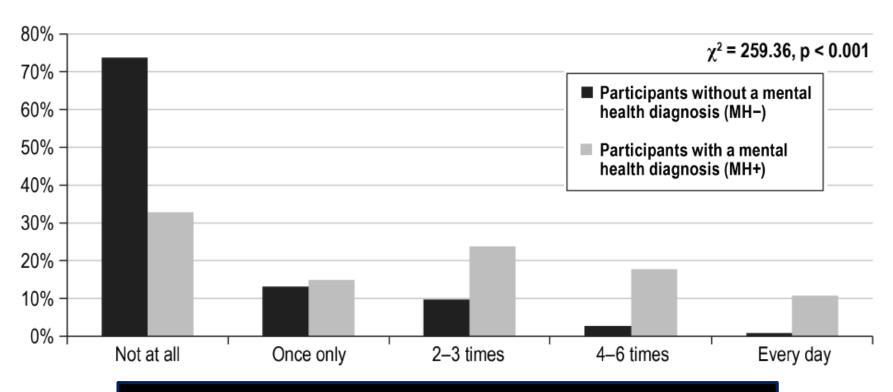
	General	oopulation	War generation			
e s	Men	Women	Men	Women		
n	17,705	17,754	5,300	5,915		
Nightmares often (%)	2.8	4.6	7.2	7.0		
Insomnia often (%)	5.4	6.1	10.9	13.9		
Depressed often (%)	3.7	6.5	7.1	9.6		
Anxious often (%)	7.3	9.8	11.0	12.0		

SLEEP 2013;36(7):1041-1050.

## A Comparison of Sleep Difficulties among Iraq/Afghanistan Theater Veterans with and without Mental Health Diagnoses

Christi S. Ulmer, PhD<sup>1,2</sup>; Elizabeth Van Voorhees, PhD<sup>1,2</sup>; Anne E. Germain, PhD<sup>3</sup>; Corrine I. Voils, PhD<sup>1,4</sup>; Jean C. Beckham, PhD<sup>1,2,5</sup>; the VA Mid-Atlantic Mental Illness Research Education and Clinical Center Registry Workgroup<sup>5</sup>

<sup>1</sup>Durham Veterans Affairs Medical Center, Durham, NC; <sup>2</sup>Department of Psychiatry and Behavioral Sciences, Duke University Medical Center, Durham, NC; <sup>3</sup>University of Pittsburgh, Departments of Psychiatry and Psychology, Pittsburgh, PA, <sup>4</sup>Department of General Internal Medicine, Duke University Medical Center, Durham, NC; <sup>5</sup>VISN 6 Mental Illness Research, Education, and Clinical Center, Durham, NC J Clin Sleep Med 2015;11(9):995–1005.

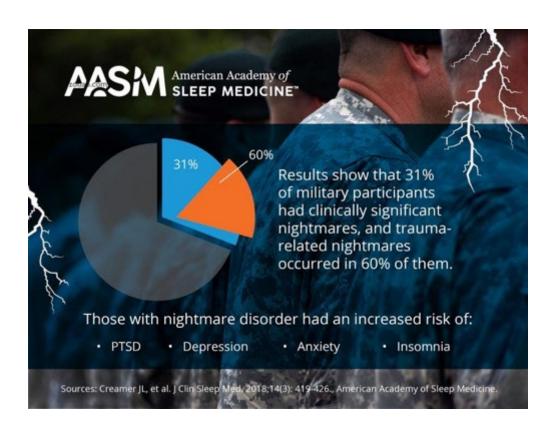


Distressing dreams of a traumatic event (frequency)

### Nightmares in United States Military Personnel With Sleep Disturbances

Jennifer L. Creamer, MD1; Matthew S. Brock, MD2; Panagiotis Matsangas, PhD3; Vida Motamedi, BA4; Vincent Mysliwiec, MD2

<sup>1</sup>Martin Army Community Hospital, Sleep Center, Fort Benning, Georgia; <sup>2</sup>San Antonio Military Medical Center, Department of Sleep Medicine, JBSA-Lackland, Texas; <sup>3</sup>Operations Research Department, Naval Postgraduate School, Monterey, California; <sup>4</sup>National Institutes of Health, National Institutes of Nursing Research, Bethesda, Maryland



- Despite 31% reporting at least weekly nightmares, only 3.9% reported nightmares as a reason for sleep evaluation
- A majority of those with NDO, reported having TRN (60%)
- NDO/TRN were more common in patients with comorbid anxiety, depression, PTSD, and TBI
  - TRN appeared to be the driving force behind this association
- 86% of those with NDO had insomnia
  - Insomnia symptoms worse in TRN
  - Sleep quality improved on PSG in TRN

### 0669

#### PREVALENCE AND CORRELATES OF NIGHTMARES IN ACTIVE DUTY SERVICE MEMBERS

Kristi Ensor Pruiksma, PhD<sup>1</sup>, Danica Slavish, PhD<sup>2</sup>,
Sophie Wardle, BS<sup>2</sup>, Alyssa Ojeda, BS<sup>3</sup>, Daniel Taylor, PhD<sup>4</sup>,
Alan Peterson, PhD<sup>1</sup>, Kevin Kelly, MD<sup>5</sup>, Douglas Maur<sup>5</sup>,
Jim Mintz, PhD<sup>1</sup>, Brett Litz, PhD<sup>6</sup>, Elisa Borah, PhD<sup>7</sup>,
Antoinette Brundige<sup>1</sup>, Stacey Young-McCaughan, PhD<sup>1</sup>,
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<sup>1</sup>UT Health Science Center at San Antonio, San Antonio, TX, USA, <sup>2</sup>University of North Texas, Denton, TX, USA, <sup>3</sup>UT Health Science Center at San Antonio, UT Health San Antonio, TX, USA, <sup>4</sup>UT Health Science Center at San Antonio, Denton, TX, USA, <sup>5</sup>Carl R Darnall Army Medical Center, Fort Hood, TX, USA, <sup>6</sup>Boston University School of Medicine, Boston, MA, USA, <sup>7</sup>University of Texas at Austin, Austin, TX, USA, <sup>8</sup>Duke University Medical Center, Durham, NC, USA.

- Pre-deployment processing
- 2010-2011
- 39.9% experienced nightmares

## Nightmare disorder in active-duty US military personnel



Brian A. Moore, PhDa\*, Allison Brager, PhDb, Jason Judkins, DSc, PhDc, Vincent Mysliwiec, MDd



- Data taken from Defense Medical Epidemiology Database
- NDO incidence (per 10,000 service members) between 2016-2019
  - High 2016: 11.3
  - Low 2018: 6.9
- Higher rates in older senior service members, female, and Black service members

<sup>\*</sup> Department of Psychological Science, Kennesaw State University, Kennesaw, Georgia, USA

b Marketing & Engagement Brigade, United States Army Recruiting Command, Fort Knox, Kentucky

<sup>6</sup> Military Performance Division, United States Army Research Institute of Environmental Medicine, Natick, Massachusetts

<sup>&</sup>lt;sup>d</sup> Department of Psychiatry and Behavioral Health, University of Texas Health Science Center at San Antonio, San Antonio, Texas

## Unexplained Clinical Findings in Combat Veterans

## Constellation of:

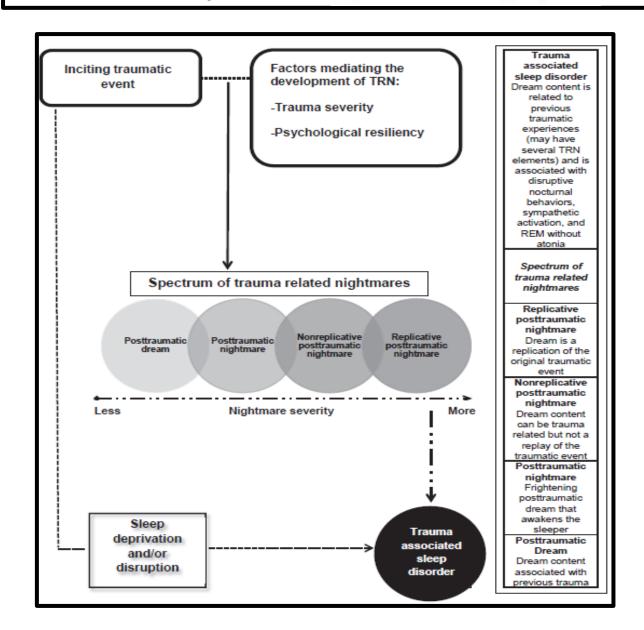
- 1. Nightmares
  - a. With and without dream recall
- 2. Disruptive nocturnal behaviors
  - a. Combative behaviors
  - b. Vocalizations
  - c. Somnambulism
- 3. Autonomic hyperactivity
  - a. Night sweats
  - b. Racing heart



C.Machado

#### Trauma associated sleep disorder: A parasomnia induced by trauma

Vincent Mysliwiec <sup>a, \*</sup>, Matthew S. Brock <sup>a</sup>, Jennifer L. Creamer <sup>b</sup>, Brian M. O'Reilly <sup>b</sup>, Anne Germain <sup>c, d</sup>, Bernard J. Roth <sup>b</sup> Sleep Medicine Reviews 37 (2018) 94–104



		Normal dreaming			
	]	Normal dysphoric dreaming			
	Increasing affect load	Bad dreaming			
		Idiopathic nightmares - low distress			
		Idiopathic nightmares - high distress		in I	Disruptive N
		Posttraumatic nightmares – trauma-related	Increasing affect distress  Increasing  Increasing  Increasing  trauma severity	reasing affe	
		Posttraumatic nightmares - replicative		wakening ct distress	Nocturnal
		Trauma associated Sleep Disorder	<b>V</b>	٧	behaviors

## Diagnostic Criteria for TSD

- A. Onset of symptoms after combat or other traumatic experience
- **B.** A history of altered dream mentation that is related to prior traumatic experience
- C. Self or witnessed reports of disruptive nocturnal behaviors to include at least one of the following:
  - 1. Abnormal vocalizations
    - a. Moaning, screaming, or yelling
  - 2. Abnormal motor behaviors in sleep
    - b. Tossing, turning, or thrashing
    - c. Combative behaviors such as striking bedpartner
- **D.** Symptoms of autonomic hyperarousal or PSG monitoring demonstrates one or more of the following associated with dream mentation:
  - 1. Tachycardia
  - 2. Tachypnea
  - 3. Diaphoresis
- E. There is an absence of EEG epileptiform activity on PSG and the disturbance is not better explained by another sleep disorder, mental disorder, medical disorder, medication, or substance use

#### Notes:

- 1. PSG may demonstrate:
  - a. Variable amounts of REM sleep without atonia
  - b. Dream enactment behavior in REM sleep
- 2. Onset is typically in close temporal proximity to trauma exposure, often in the setting of sleep deprivation/disruption
- 3. Patients with TSD frequently have comorbid insomnia and/or obstructive sleep apnea

REVIEW Open Access

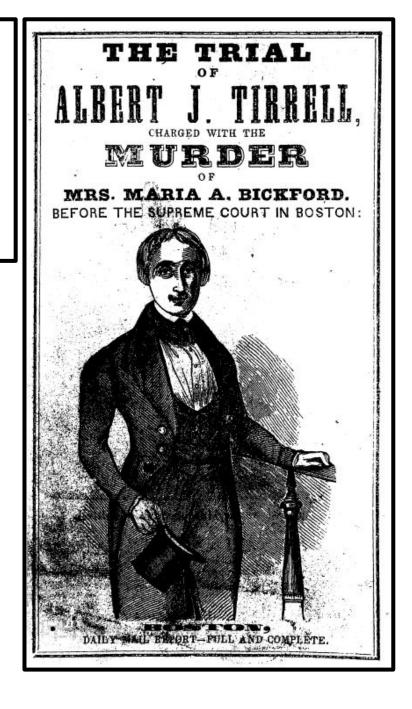
## The spectrum of disorders causing violence during sleep



Carlos H. Schenck

## **Table 1** Differential diagnosis of sleep-related injury and violence

- 1. REM Sleep Behavior Disorder (RBD)
- 2. NREM Sleep Parasomnias (Sleepwalking, Sleep Terrors)
- 3. Parasomnia Overlap Disorder (RBD + NREM Sleep Parasomnias)
- 4. Obstructive Sleep Apnea
- Sexsomnia (Sleep Related Abnormal Sexual Behaviors)
- 6. Sleep Related Dissociative Disorder
- 7. Trauma-Associated Sleep Disorder/Post-Traumatic Stress Disorder
- 8. Periodic Limb Movement Disorder
- 9. Rhythmic Movement Disorder
- 10. Noctumal Scratching Disorder
- 11. Sleep Related Eating Disorder
- 12. Nocturnal Seizures
- 13. Miscellaneous/Mixed Disorders



## Only a Small Percentage of Patients Experiencing Nightmares Seek Professional Help

Seeking professional help for nightmares: A representative study	"Every eighth person with frequent nightmares, defined as every other week or more often, sought at one time of his/her life for professional help for coping with nightmares." - Schredel 2013	
Eur. J. Psychiat. Vol. 27, N.° 4, (259-264) 2013  Michael Schredl  Central Institute of Mental Health, Medical Faculty Mannheim/Heidelberg University	<ul> <li>"Nightmare frequency correlated with seeking professional help" - Schredel 2013</li> </ul>	
Interest in Information about Nightmares in Patients with Sleep Disorders  Michael Schredl, PhD'; Lara Dehmlow, cand. psych.'; Judith Schmitt, MD²  'Sleep Laboratory, Central Institute of Mental Health, Medical Faculty Mannheim / Heidelberg University, Mannheim, Germany; 'Abteilung für Schlafmedizin, Theresienkrankenha und St. Hedwig-Klinik GmbH, Mannheim, Germany	□ "Overall, <b>12.24</b> % of 972 participants with valid answers for this item <b>reported that they had sought help for their nightmares</b> . The figure for the patients with frequent nightmares (once a week or more often) was higher (33.00%)" - Schredl 2016	
Nightmares: Under-Reported, Undetected, and Therefore Untreated  Michael R. Nadorff, PhD <sup>1,2</sup> ; Danielle K. Nadorff, PhD <sup>1</sup> ; Anne Germain, PhD <sup>3</sup>	"Our results indicate that <b>most</b> participants with clinically significant nightmare symptoms <b>are unlikely to report nightmares to a healthcare provider</b> " - Nadorff 2015	
Nightmares in United States Military Personnel With Sleep Disturbances  Jennifer L. Creamer, MD¹; Matthew S. Brock, MD²; Panagiotis Matsangas, PhD³; Vida Motamedi, BA⁴; Vincent Mysliwiec, MD²	□ "Only 3.9% of military receiving a sleep study reported nightmares as the reason for evaluation" Creamer 2018	
Barry Krakow, MD  Michael Hollifield, MD  Lisa Johnston, MA, MPH  Mary Koss, PhD  Ron Schrader, PhD  Roll D. W. C. D. D. C.	"Only <b>14 [of 142]</b> women [with a history of chronic nightmares and a history of sexual or physical assault] ever <b>sought help for the problem of nightmares</b> " Krakow 2002	

<sup>1.</sup> Schredl M. Eur. J. Psychiat 2013;27(4):259-264. 2. Schredl M. et al. J Clin Sleep Med 2016;12(7):973-977. 3. Nadorff MR. et al. J Clin Sleep Med 2015;11(7):747-750. 4. Creamer JL, et al. J Clin Sleep Med. 2018;14(3):419-426. 5. Krakow B, et al. JAMA 2001; 286(5):537-545

Teddy D. Warner, PhD

# Psychologist/Psychotherapist are More Likely to Hear About Nightmares than other Providers

Barry Krakow, MD	
Michael Hollifield, MD	
Lisa Johnston, MA, MPH	
Mary Koss, PhD	
Ron Schrader, PhD	
Teddy D. Warner, PhD	

Imagery Rehearsal Therapy for Chronic Nightmares in Sexual Assault Survivors With Posttraumatic Stress Disorder A Randomized Controlled Trial

"Women who sought help for nightmares **sought help in the context of psychotherapy** for other psychiatric symptoms" Krakow 2002

#### SCIENTIFIC INVESTIGATIONS

#### Interest in Information about Nightmares in Patients with Sleep Disorders

Michael Schredl, PhD1; Lara Dehmlow, cand. psych.1; Judith Schmitt, MD2

Sleep Laboratory, Central Institute of Mental Health, Medical Faculty Mannheim / Heidelberg University, Mannheim, Germany; <sup>1</sup>Abteilung für Schlafmedizin, Theresienkrankenhau: und St. Hedwig-Klinik GmbH, Mannheim, Germany

□ "The largest group of healthcare professionals that were asked for help were **psychologists/psychotherapists** (about 70%), psychiatrists (about 10%), general practitioners (about 6%), sleep specialists (about 5%)" - Schredl 2016

Prevalence, correlates and treatment of nightmares in secondary mental healthcare

Annette van Schagen

- "Nightmare disorder might be ignored in mental health care because according to DSM-IV-TR criteria nightmares can only be classified as a nightmare disorder if they do not occur exclusively during the course of another mental disorder" -van Schagen 2016
- □ "Nightmares are highly prevalent in psychiatric practice and our data suggest they are often missed in psychiatric assessment" van Schagen 2016

## Medications Associated with Nightmares

Medication Category	Examples		
Acetylcholinesterase inhibitors	Donepezil, galantamine, rivastigmine		
Antidepressants	Bupropion, MAOIs, mirtazapine, SNRIs, SSRIs, TCAs		
Antihistamines	Chlorpheniramine		
Antimicrobials	Ciprofloxacin, efavirenz, erythromycin, ganciclovir, mefloquine		
Atypical antipsychotic medications	Clozapine, olanzapine, risperidone		
Beta blockers	Labetalol metoprolol, propranolol		
Calcium channel blockers	Verapamil		
Dopaminergic drugs	Amantadine, amphetamine, levodopa, methylphenidate, pramipexole, ropinirole, selegiline		
Sedative hypnotics	Alcohol, barbiturates, temazepam, triazolam		
Statins	Atorvastatin		
OTC	Melatonin		



## јсѕм Journal of Clinical Sleep Medicine

## Best Practice Guide for the Treatment of Nightmare Disorder in Adults

Standards of Practice Committee:

R. Nisha Aurora, M.D.<sup>1</sup>; Rochelle S. Zak, M.D.<sup>2</sup>; Sanford H. Auerbach, M.D.<sup>3</sup>; Kenneth R. Casey, M.D.<sup>4</sup>; Susmita Chowdhuri, M.D.<sup>5</sup>; Anoop Karippot, M.D.<sup>6</sup>; Rama K. Maganti, M.D.<sup>7</sup>; Kannan Ramar, M.D.<sup>8</sup>; David A. Kristo, M.D.<sup>9</sup>; Sabin R. Bista, M.D.<sup>10</sup>;

- Level A (both PTSD and idiopathic nightmares):
  - Prazosin
  - Image Rehearsal Therapy
- Level B

 Systematic desensitization and progressive deep muscle relaxation training for idiopathic nightmares

- Level C
  - Clonidine for PTSD-associated nightmares

Term	Level	Evidence Levels	Explanation
Recommended / Not recommended	Α	1 or 2	Assessment supported by a substantial amount of high quality (Level I or II) evidence and/or based on a consensus of clinical judgment
Suggested / Not Suggested	В	1 or 2—few studies 3 or 4—many studies and expert consensus	Assessment supported by sparse high grade (Level I or II) data or a substantial amount of low-grade (Level III or IV) data and/or clinical consensus by the task force
May be considered / Probably should not be considered	С	3 or 4	Assessment supported by low grade data without the volume to recommend more highly and likely subject to revision with further studies

# Prazosin summary

- 12 studies
  - 5 placebo controlled trials (
  - 4 open label
  - 4 retrospective chart reviews
- Prazosin effective, but doses up to 10-15mg are often needed.
- Consider daytime prazosin if BP can tolerate

#### Position Paper for the Treatment of Nightmare Disorder in Adults: An American Academy of Sleep Medicine Position Paper

Timothy I. Morgenthaler, MD<sup>1</sup>; Sanford Auerbach, MD<sup>2</sup>; Kenneth R. Casey, MD, MPH<sup>3</sup>; David Kristo, MD<sup>4</sup>; Rama Maganti, MD<sup>5</sup>; Kannan Ramar, MD<sup>1</sup>; Rochelle Zak, MD<sup>6</sup>; Rebecca Kartie, MD, MSHI, MBA<sup>7</sup>



## **Behavioral therapy**

- Image Rehearsal therapy
- PTSD Nightmares: CBT, CBT-I, eye desensitization and reprocessing, and exposure, relaxation, and rescripting therapy
- NDO: CBT, hypnosis, lucid dreaming therapy, progressive deep muscle relaxation, sleep dynamic therapy, self-exposure therapy, systemic desensitization, and testimony method

### Pharmacologic therapy

- PTSD Nightmares: prazosin, atypical antipsychotics, clonidine, cyproheptadine, fluvoxamine, gabapentin, nabilone, phenalzine, topiramate, trazodone, and tricyclic antidepressants
- NDO: prazosin, nitrazepam, and triazolam





# VA/DOD CLINICAL PRACTICE GUIDELINE FOR THE MANAGEMENT OF POSTTRAUMATIC STRESS DISORDER AND ACUTE STRESS DISORDER

Version 3.0 – 2017

e. Prazosin					
28a	For global symptoms of PTSD, we suggest against the use of prazosin as mono-or augmentation therapy.	Weak Against	Reviewed, New-replaced		
28b	For nightmares associated with PTSD, there is insufficient evidence to recommend for or against the use of prazosin as mono- or augmentation therapy.	N/A	Reviewed, New-replaced		

#### Discussion

Four small, published trials of variable quality met the threshold for review. [185-188] These trials contained a total of 167 subjects, all of whom were Veterans or active duty Service Members. Most of these trials had promising results, particularly for nightmares. However, in a much larger, well-designed VA Cooperative multi-site trial with 304 subjects, prazosin failed to separate from placebo in the treatment of both global symptoms of PTSD and nightmares. [189] Interestingly, this study had not been published at the time of our review, three years after its completion. Nonetheless, we believed it was important to include in our analysis due to its significance and availability in the public domain (www.clinicaltrials.gov, identifier NCT00532493).

## The NEW ENGLAND JOURNAL of MEDICINE

ESTABLISHED IN 1812

FEBRUARY 8, 2018

VOL. 378 NO. 6

### Trial of Prazosin for Post-Traumatic Stress Disorder in Military Veterans

M.A. Raskind, E.R. Peskind, B. Chow, C. Harris,\* A. Davis-Karim, H.A. Holmes, K.L. Hart, M. McFall, T.A. Mellman, C. Reist, J. Romesser, R. Rosenheck, M.-C. Shih, M.B. Stein, R. Swift, T. Gleason, Y. Lu, and G.D. Huang

concern that the condition of potential participants would deteriorate clinically during 6 months of receiving placebo could have motivated providers to use available open-label prazosin rather than refer patients for trial participation. Further-

more, the exclusion of participants who were unwilling or unable to discontinue trazodone, an antidepressant with  $\alpha_1$ -adrenoreceptor antagonist activity, may have eliminated potential participants who would have had a response to

less likely to be ameliorated with antiadrenergic treatment. Concern about the increasing incidence of suicide and of violent behavior among veterans led the planning committee to make psychosocial instability an exclusion criterion for participation in the trial. None of the previous smaller randomized trials of prazosin for PTSD<sup>10-15</sup>

## Active Duty Service Members Prazosin Use

- Army, Navy, Marines, Air Force
- MTF Encounters
- 2004 to 2021
- Prescribed at least a 90 day supply of prazosin

Calendar Year	Nbr with Sleep Nightmares diagnosis	Nbr with Meds in year of diagnosis
2004	1094	8
2005	1124	15
2006	1383	31
2007	1824	98
2008	2609	252
2009	4827	382
2010	4811	549
2011	5968	974
2012	6573	1346
2013	5999	1557
2014	5612	1330
2015	5915	1385
2016	7226	1611
2017	6459	1392
2018	6141	1068
2019	6577	1144
2020	5974	1027
2021	7472	1291

Sleep Medicine Reviews 50 (2020) 101272



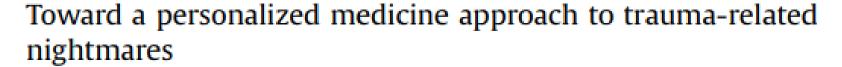
Contents lists available at ScienceDirect

## Sleep Medicine Reviews

journal homepage: www.elsevier.com/locate/smrv



**GUEST EDITORIAL** 





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20 January 2020 Available online 25 January 2020

#### Prazosin Treatment for Nightmares

#### Prazosin Dosing:

- 1mg 30 minutes prior to sleep for 3 days.
- 2mg 30 minutes prior to sleep for 3 days.
- 4mg 30 minutes prior to sleep for 5 days.
- 4. 5mg 30 minutes prior to sleep until you are seen back in clinic.
- The usual dose to result in nightmare improvement is 6-8 mg. Some patients require a higher dose (i.e. 12-16mg), some patients require a lower dose (i.e. 2-4 mg). This depends on how you respond to the medication.

#### Prazosin Side effects:

- 1. Light headedness/dizziness. To minimize this side effect, get out of bed slowly.
- Passing out, rarely.
- 3. Never take with Viagra.

### PTSD and OSA in veterans

#### PAP ADHERENCE IN VETERANS WITH PTSD

## Positive Airway Pressure Adherence in Veterans with Posttraumatic Stress Disorder

Ali A. El-Solh, MD, MPH<sup>1,2,3</sup>; Lakshmy Ayyar, MD<sup>2</sup>; Morohonfolu Akinnusi, MD<sup>2</sup>; Sachin Relia, MD<sup>4</sup>; Opeoluwa Akinnusi, MD<sup>4</sup>

<sup>1</sup>The Veterans Affairs Western New York Healthcare System, Buffalo, NY; <sup>2</sup>Division of Pulmonary, Critical Care, and Sleep Medicine, Department of Medicine, <sup>3</sup>Department of Social and Preventive Medicine, and <sup>4</sup>Department of Psychiatry, State University of New York at Buffalo School of Medicine and Biomedical Sciences and School of Public Health and Health Professions, Buffalo, NY

**Study Objectives**: To determine the short-term positive airway pressure (PAP) adherence rates and to identify non–mask-related risk factors associated with 30-day nonadherence to PAP in a population of veterans with obstructive sleep apnea (OSA) and posttraumatic stress disorder (PTSD).

Design: A retrospective study.

Settings: A Veterans Affairs hospital.

Patients: One hundred forty-eight PTSD veterans newly diagnosed with OSA and a control group of OSA without PTSD matched for age, gender, BMI, and severity of OSA.

- PTSD associated with poor CPAP adherence
- Increased EDS associated with improved PAP adherence
- Increased PAP adherence associated with decreased nightmare frequency

El-Solh et al 2010



# Journal of Clinical Sleep Medicine

http://dx.doi.org/10.5664/jcsm.2260

## The Impact of Posttraumatic Stress Disorder on CPAP Adherence in Patients with Obstructive Sleep Apnea

Jacob F. Collen, M.D.; Christopher J. Lettieri, M.D., F.A.A.S.M.; Monica Hoffman, M.D. Pulmonary, Critical Care and Sleep Medicine, Walter Reed National Military Medical Center, Bethesda, MD

45 Veterans (PTSD+OSA) vs. 45 age-matched control+OSA

More comorbid insomnia w/PTSD (25.8% vs 11.1%)

PTSD = significantly less CPAP use

Nights used: 61.4% vs. 76.8%, p = 0.001

Mean nightly: 3.4 vs 4.7, p < 0.001

Regular use: 25.2% vs. 58.3%, p = 0.01

58% relative reduction in PAP use J Clin Sleep Med 2012;8(6):667-672.

## Co-Morbid PTSD, OSA & Insomnia

Insomnia, nightmares, and sleep avoidance common Insomnia reported in 64-100%

Likely due to a hypervigilant state

Portends worse outcomes
Increased risk for Major depressive disorder
Independent risk factor for suicidality
Less response to anti-depressant therapy
Increased risk of relapse

Negative impact on PAP use/adherence

## Low Arousal Threshold in Patients with PTSD+OSA

Explored prevalence and impact of low arousal threshold in patients with Co-morbid PTSD+OSA with and without Insomnia

Insomnia common in PTSD, with increased prevalence in those with comorbid OSA

Hyperaroused state in PTSD may reflect a LAT

Found 55% of those with PTSD+OSA have LAT

LAT in 76% of those w/ PTSD+OSA+Insomnia

Represents a significant barrier to PAP use

### Journal of Clinical Sleep Medicine

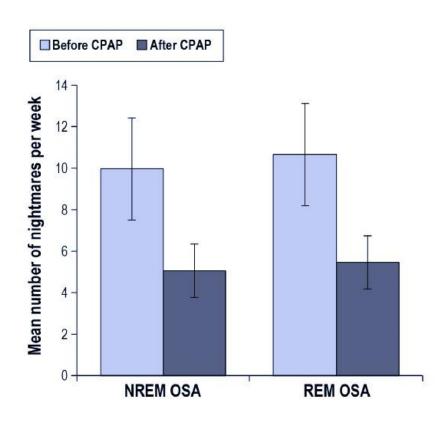
#### SCIENTIFIC INVESTIGATIONS

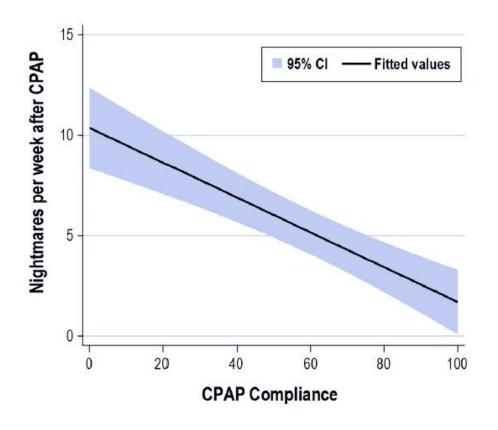
#### Treatment of OSA with CPAP Is Associated with Improvement in PTSD Symptoms among Veterans

Jeremy E. Orr, MD<sup>1</sup>; Carolina Smales, BS<sup>2</sup>; Thomas H. Alexander, MD, MHSc<sup>2,3</sup>; Carl Stepnowsky, PhD<sup>2</sup>; Giora Pillar, MD<sup>4</sup>; Atul Malhotra, MD<sup>1</sup>; Kathleen F. Sarmiento, MD<sup>1,2</sup>

- 32 Veterans with PTSD+OSA, treated w/ CPAP
- Significant reduction in PTSD symptoms, measured by PCL-S score
  - (60.6 ± 2.7 versus 52.3 ± 3.2 points; p < 0.001)</li>
- Less reported sleepiness
- Improved subjective sleep quality
- · Increased daytime functioning and quality of life
- Less depression

# PAP Effect on PTSD Nightmares







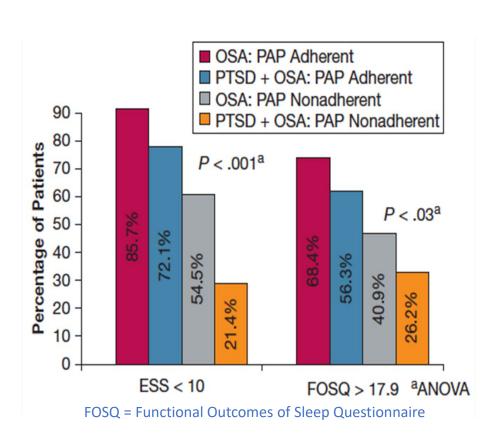
#### OSA Syndrome and Posttraumatic Stress Disorder



Clinical Outcomes and Impact of Positive Airway Pressure Therapy

Christopher J. Lettieri, MD; Scott G. Williams, MD; and Jacob F. Collen, MD

CHEST 2016; 149(2):483-490



- Sleepiness, function, and QoL worse in those with OSA+PTSD compared to either condition alone
- Patients with PTSD+OSA
  - Diminished response to PAP
  - Less resolution of symptoms

## Oral Appliances in OSA

OAs similar to PAP with improvements in neurocognitive function somnolence, depressive symptoms

Adherence with OA consistently greater than PAP Benefit in PTSD?

### A Randomized Crossover Trial Evaluating Continuous Positive Airway Pressure Versus Mandibular Advancement Device on Health Outcomes in Veterans With Posttraumatic Stress Disorder

Ali A. El-Solh, MD, MPH<sup>1,2,3</sup>; Gregory G. Homish, PhD<sup>3,4</sup>; Guy Ditursi, DDS, MBA<sup>1</sup>; John Lazarus, DDS<sup>1</sup>; Nithin Rao, DDS<sup>1</sup>; David Adamo, RPSGT<sup>1</sup>; Thomas Kufel, MD<sup>1,2</sup>

<sup>1</sup>VA Western New York Healthcare System, Buffalo, New York; <sup>2</sup>Division of Pulmonary, Critical Care, and Sleep Medicine, Department of Medicine, School of Medicine and Biomedical Sciences, University at Buffalo, Buffalo, New York; <sup>3</sup>Department of Epidemiology and Environmental Health, School of Public Health and Health Professions, University at Buffalo, New York; <sup>4</sup>Department of Community and Health Behavior, School of Public Health and Health Professions, University at Buffalo, New York

Randomized Cross Over Trial in PTSD newly diagnosed with OSA 12 weeks each of CPAP and MAD

CPAP was more efficacious in reducing AHI and improving nocturnal oxygenation than MAD (P < .001 and P = .04, respectively)

Both treatments – similar reductions in PTSD severity scores and PSQI

Adherence to MAD significantly higher than CPAP (P < .001)

58% preferred MAD to CPAP

# A randomized sham-controlled clinical trial of a novel wearable intervention for trauma-related nightmares in military veterans

Nicholas D. Davenport, PhD<sup>1,2</sup>; J. Kent Werner, MD, PhD<sup>3,4,5</sup>

Research, Minneapolis VA Health Care System, Minneapolis, MN

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<sup>4</sup>Department of Neurology, Uniformed Services University, Bethesda, MD

<sup>5</sup>Sleep Disorders Center, Walter Reed National Military Medical Center, Bethesda MD

Number Nine article on JCSM Top 10 Articles of 2022
 Via Altmetric Scores

# A randomized sham-controlled clinical trial of a novel wearable intervention for trauma-related nightmares in military veterans

- 65 Veterans with impaired sleep d/t to TRN
- Sleep quality, PTSD/depression symptoms, and QOL were looked at over 30 days
- Both groups demonstrated statistically significant improvements on all measures
  - Post hoc analysis with significant improvement in perceived sleep quality with Active Device vs. Sham





# NightWare is a therapeutic platform that treats nightmares

- Digital prescription therapeutic
- Software on an Apple Watch and iPhone
  - Hardware has almost no other functions
  - Not downloadable to a patient's own devices
- Interrupts nightmares without disrupting sleep
- NightWare improves sleep in patients with nightmares



### Machine Learning In Action

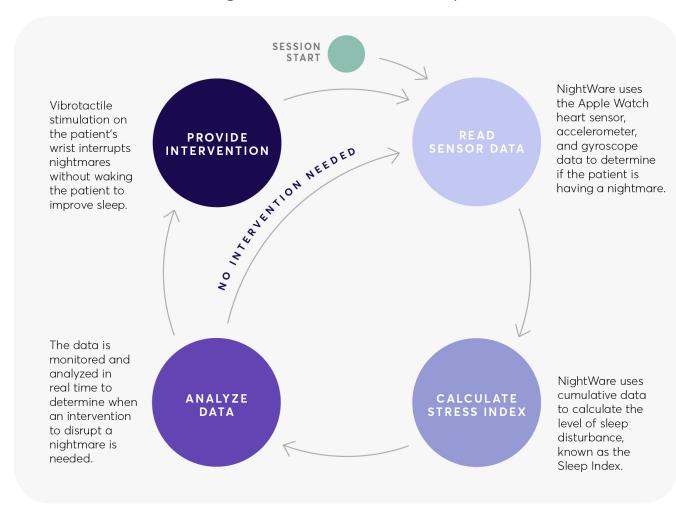
Initial Learning Period of 500 minutes

Recalibration with each session

NightWare creates a unique intervention specific to each patient's individual sleep patterns.

NightWare continuously adjusts to the patient's movement and heart rate

#### NightWare Intervention Cycle



11/1/2023 N I G H T W A R E

### Under-recognized

## Nightmare Disorder

Evaluation can lead to earlier treatment and improved outcomes

Treatment is an individualized approach