# Stronger Together



# Integrative Approaches to Improve Sleep During Cancer Treatment and Survivorship

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# **Objectives:**

- Define Integrative Medicine and describe our oncology care model
- Screen patients for sleep concerns in the context of cancer care and survivorship
- Apply evidence-based, Integrative methods to improve sleep quality and outcomes





# **Complementary Versus Alternative**

"Complementary" generally refers to using a non-mainstream approach together with conventional medicine.

 "Alternative" refers to using a non-mainstream approach in place of conventional medicine.





# **Integrative Medicine**

 Unfortunately, some patients gravitate to the use of widely promoted disproved or unproven "alternative" modalities to achieve their goals.

 No less than mainstream cancer therapies in common use, complementary therapies must be evidence-based or, lacking firm evidence, must at least have a rational basis.





# What is Integrative Medicine?

- "Integrative medicine"—increasingly has replaced CAM as a preferred term
- Integrative oncology is a synthesis of mainstream treatment and complementary therapies in cancer care.
  - Noninvasive, non-pharmacologic adjuncts to mainstream treatment that improve patients' strength and control the physical and emotional symptoms associated with cancer and cancer treatment.
  - Provide patients with a sense of control and self-empowerment at a time when many feel vulnerable and life seems out of control.

Deng, G, Frenkel M, Cohen L, et al: Evidence-based clinical practice guidelines for integrative oncology: Complementary therapies and <u>J Soc Integr Oncol 7:85-120, 2009</u>







Conventional and non conventional modalities

Patient centered

**Partnership** 

Integrative Medicine

Natural, effective, non-invasive interventions

Prevention to treatment





# **Integrative Medicine**

Engages mind, body, spirit and community

Encourages providers to model healthy lifestyles for their patients

- Focuses attention on lifestyle choices for prevention & maintenance of health
- Maintains that healing is always possible even when cure is not





# **Symptom Clusters**

- Fatigue
- Weight loss/gain
   Pain
- Poor sleep quality
   Physical
- Depression
- Anxiety
- Neuropathy

- Cognitive slowing
- - deconditioning
- Sexual dysfunction
- Lymphedema







# **Integrative Medicine in Cancer Care**

Nutrition

Acupuncture

Exercise

Massage

Mindfulness

Yoga

Sleep









Sleep is a natural and innate process that is inherent to our physiology.





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## **Integrative Sleep Medicine**

- Sleep is a multidimensional process
  - "A four movement symphony"
  - Stage 1, 2, 3, and REM
- Brain and body cooling
- Multidisciplinary field
- Growing science
- Behavior-based, not an organ based science



"Sleep medicine focuses on the clinical assessment, physiologic testing, diagnosis, management and prevention of sleep and circadian rhythm disorders. Sleep specialists treat patients of any age and use multidisciplinary approaches"

— American Academy of Sleep Medicine





## Importance of restful sleep

- Reverse damage from wakefulness
  - Reverse oxidative stress
  - Replete energy stores
- Brain and body cooling
- Stimulate brain development (REM sleep)
- Regulate ion channels
- Learning and memory
- Optimize regulation of other physiological functions





## **Biochemistry of sleep**

- Substance P, glutamate (pronociceptive arousal provoking)
- GABA (inhibitory) alpha 2 delta receptors of Ca channels
- Serotonin Dorsal raphe nucleus
- Almost every medication for depression (SSRI, MAO-i) will decrease REM sleep

Marshall NS et al. Sleep Medicine Reviews, 2008.





## **Metabolic Changes Associated with Poor Sleep**

- Sleep loss adversely affects metabolic hormones
- 1 Evening cortisol
- Sympathetic activation
- Thyrotropin
- Glucose tolerance
- ↑ Ghrelin ↓ Leptin
- These endocrine changes contribute to altered signaling of hunger and appetite which may promote weight gain and obesity

Leproult, R et al; Endoct Dev 2010.





## Physiologic effects of sleep loss

- Sleep loss also alters immune function
  - Reduced natural killer cell activity
  - Changes in circulating levels of leukocytes and cytokines (increased TNF, IL-6)
  - Decreased antibody titers influenza vaccination
  - Inflammatory markers increased (CRP)
  - What is the most common sleep condition?

Banks S, et al. J Clin Sleep Med, 2007.





## **Sleep Medicine: Insomnia and Cancer**

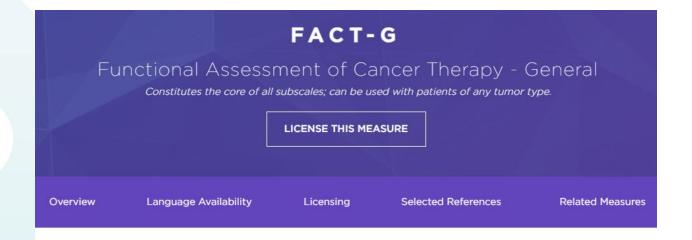
- Natural course of insomnia comorbid with cancer: an 18-month longitudinal study
  - Purpose: assess prevalence and natural course of comorbid insomnia with cancer over an 18-month period
  - Insomnia is a frequent and enduring problem in patients with cancer
  - Early intervention strategies could prevent the problem from becoming more severe and chronic
    - Cognitive-behavioral therapy (i)
    - Mindfulness techniques

Savard J, et al. J Clin Oncol. 2011 Sep 10; 29(26):3580-6.





# **Care Design: Screening**



#### Overview

The Functional Assessment of Cancer Therapy - General (FACT-G) is a 27-item questionnaire designed to measure four domains of HRQOL in cancer patients: Physical, social, emotional, and functional well-being.





## **Sleep Quality Related to Cancer Symptoms**



- Sleep dysfunction and psychosocial adaptation among women undergoing treatment for non-metastatic breast cancer
  - O Purpose:
    - Determine frequency of sleep disturbances in women prior to adjuvant therapy for breast cancer
    - Determine whether greater sleep dysfunction predicts poorer functional outcomes
  - Results of the Study
    - Consistent associations between sleep quality and psychosocial adaptation
    - Associations independent of anxiety and depression
    - Important: Comprehensive psychosocial interventions that consider sleep problems





# **Integrative Sleep Clinical Pearl**

- 3 hours no food
- 2 hours limit fluids
- 1 hour avoid bright-screened electronic devices





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#### Prolonged Nightly Fasting and Breast Cancer Prognosis.

Marinac CR1, Nelson SH2, Breen Cl3, Hartman SJ4, Natarajan L4, Pierce JP4, Flatt SW3, Sears DD5, Patterson RE4.

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2413 women with breast cancer but without DM, aged 27 to 70 years at diagnosis and participated in the prospective Women's Healthy Eating and Living study between March 1, 1995, and May 3, 2007

Clinical outcomes were invasive breast cancer recurrence and new primary breast tumors during a mean of 7.3 years of study follow-up as well as death from breast cancer or any cause during a mean of 11.4 years of surveillance

Nightly fasting duration was estimated from 24-hour dietary recalls collected at baseline, year 1, and year 4 – sleep duration self reported, archived blood samples used for HgbA1c and CRP





- Fasting < 13 hours per night (lower 2 tertiles of nightly fasting distribution) was assoc with an increase in the risk of breast cancer recurrence compared with fasting 13 or more hours per night (hazard ratio, 1.36; 95% CI, 1.05-1.76)
- Not assoc with higher risk of breast ca mortality or all cause
- Prolonging the length of the nightly fasting interval may be a simple, non-pharmacologic strategy for reducing the risk of breast cancer recurrence. Improvements in glucoregulation and sleep may be mechanisms linking nightly fasting with breast cancer prognosis.





June 10, 2019

# Association of Exposure to Artificial Light at Night While Sleeping With Risk of Obesity in Women

Yong-Moon Mark Park, MD, PhD<sup>1</sup>; Alexandra J. White, PhD<sup>1</sup>; Chandra L. Jackson, PhD, MS<sup>1</sup>; et al

≫ Author Affiliations | Article Information

JAMA Intern Med. 2019;179(8):1061-1071. doi:10.1001/jamainternmed.2019.0571

Instead of using our phones or watching TV prior to bedtime, what shall we recommend?





## Mindfulness Improves Sleep and Quality of Life



- Effectiveness of Mindfulness-Based Stress Reduction in Mood, Breast- and Endocrine-Related Quality of Life, and Well-Being in Stage 0 to III Breast Cancer: A Randomized, Controlled Trial
  - Mindfulness was more effective than standard care for improving:
    - Mood
    - Breast- and endocrine-related quality of life
    - Well-being
  - Improvements maintained at three months
  - Mindfulness can help alleviate long-term emotional and physical adverse effects of medical treatments, including endocrine treatments





## **Sleep: The Mind-Body Connection**

- Mind-body treatments for the pain-fatigue-sleep disturbance symptom cluster in persons with cancer
  - Co-occurring pain, fatigue, and sleep disturbance comprise a common symptom cluster in patients with cancer
  - Identify treatment approaches that target the cluster of symptoms
    - Mind-body interventions could help patients manage all symptoms in the cluster with a single treatment strategy
  - Studies needed to examine the impact of:
    - Relaxation
    - Imagery/hypnosis
    - > CBT/CST
    - Meditation
    - Music interventions





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Special Articles

### Integrative Oncology Care of Symptoms of Anxiety and Depression in Adults With Cancer: SIO-ASCO Guideline Summary and Q&A

Linda E. Carlson, PhD, RPsych 🕞; Nofisat Ismaila, MD 🕞; Elizabeth L. Addington, PhD 🕞; Gary N. Asher, MD, MPH ; Joke Bradt, PhD, MT-BC 5; Ashwin Mehta, MD6; and Julia H. Rowland, PhD7 (D); on behalf of the SIO-ASCO Expert Panel

DOI https://doi.org/10.1200/0P.23.00358

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- Overview of recommendations
- How can people access these therapies?
- Are there disparities in accessing integrative therapies?

PRESENTED BY: Depression in Adults With Cancer

Questions regarding dietary supplements?











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## **Yoga for Cancer Survivors**

Integr Cancer Ther. 2015 Nov 29. pii: 1534735415617021. [Epub ahead of print]

YOCAS©® Yoga Reduces Self-reported Memory Difficulty in Cancer Survivors in a Nationwide Randomized Clinical Trial: Investigating Relationships Between Memory and Sleep.

 $\underline{Janelsins\ MC}^1, \underline{Peppone\ LJ}^2, \underline{Heckler\ CE}^2, \underline{Kesler\ SR}^3, \underline{Sprod\ LK}^4, \underline{Atkins\ J}^5, \underline{Melnik\ M}^6, \underline{Kamen\ C}^2, \underline{Giquere\ J}^7, \underline{Messino\ MJ}^5, \underline{Mohile\ SG}^2, \underline{Mustian\ KM}^2.$ 

- Previously showed in a phase III randomized clinical trial that yoga a program that consists of breathing exercises, postures, and meditation -significantly improved sleep quality in cancer survivors
- 328 participants from 12 centers who provided data on the memory difficulty item of the MD Anderson symptom inventory included – 8 sessions of yoga for 75 minutes
- Sleep quality measured using the Pittsburgh Sleep Quality Index
- Yoga significantly reduced patient-reported memory difficulty in cancer survivors

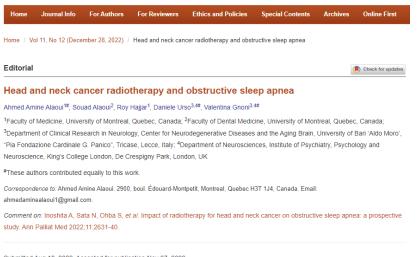




# **Obstructive Sleep Apnea**



- Screening questions
  - Berlin Index (Stop BANG)
  - Epworth Scale



Submitted Aug 18, 2022. Accepted for publication Nov 07, 2022





## **OSA** in Head and Neck Cancer

- 81.3% of patients diagnosed with HNC presented with OSA.
- Mean AHI 20.8±19.0 events/hr.
- OSA was found in 72% of patients with nasopharyngeal cancer.





Journal of Clinical Sleep Medicine

#### SCIENTIFIC INVESTIGATIONS

### Impact of Arterial Stiffness on WatchPAT Variables in Patients With Obstructive Sleep Apnea

Taku Kinoshita, MD¹; Misuzu Yahaba, MD, PhD¹; Jiro Terada, MD, PhD¹; Takuma Matsumura, MD¹; Yoriko Sakurai, MD, PhD¹; Kengo Nagashima, PhD²; Seiichiro Sakao, MD, PhD¹; Koichiro Tatsumi, MD, PhD¹

<sup>1</sup>Department of Respirology, Graduate School of Medicine, Chiba University, Chiba, Japan; <sup>2</sup>Department of Global Clinical Research, Graduate School of Medicine, Chiba

University, Chiba, Japan







## Nap time

- Sleep loss recovery
  - REM rebound
- A restful space
  - After lunch, rest a while. After dinner, walk a mile.





## **Caffeine**

- Often used to address fatigue
- Associated with disrupted sleep architecture





# What about Ashwagandha and Cannabinoids?

- Be careful about drug-herb, drug-supplement interactions
- Magnesium, Vitamin D 25-OH, B12, Folate



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### **Summary**

- Sleep disturbances are common in cancer survivors
- Screening for and treatment of sleep concerns can help improve quality of life in people with cancer
- Integrative medicine modalities can be used to address sleep complaints in people living beyond cancer
- Lifestyle modification is at the core of addressing sleep concerns
- More research needs to be done on sleep and how to use integrative medicine treatments effectively in cancer
- Contact: asmehta@mhs.net