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CARDI•OH

Ohio Cardiovascular and Diabetes Health Collaborative



In partnership with:



Heart Health and the Science of Sleep

May 24, 2023



CARDI•OH

Ohio Cardiovascular and Diabetes Health Collaborative

Welcome

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Case Western Reserve University School of Medicine

About Cardi-OH

Founded in 2017, the mission of Cardi-OH is to improve cardiovascular and diabetes health outcomes and eliminate disparities in Ohio's Medicaid population.



CARDI•OH

Ohio Cardiovascular and Diabetes Health Collaborative

WHO WE ARE: An initiative of health care professionals across Ohio's seven medical schools.

WHAT WE DO: Identify, produce, and disseminate evidence-based cardiovascular and diabetes best practices to primary care teams.

HOW WE DO IT: Best practices resources are available via an online library at Cardi-OH.org, including monthly newsletters, podcasts, webinars, and virtual clinics using the Project ECHO® virtual training model.

Learn more at Cardi-OH.org



In partnership with:



Special Thanks



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 - Use the Q&A feature to submit questions at any point
 - Questions will be answered during the Q&A portion of the webinar
- **Post webinar evaluation survey**
 - The survey link will be shared at the end of today's webinar and also sent by email
 - Please complete by **COB Wednesday, May 31**

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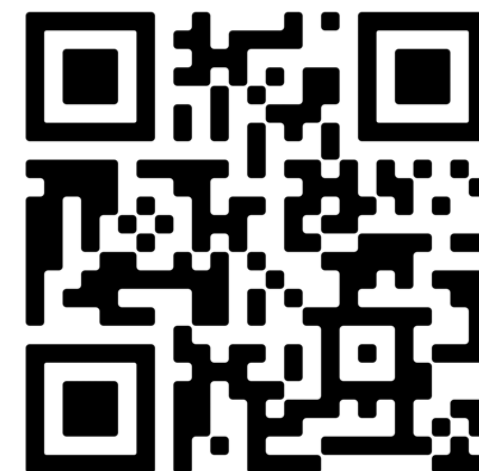


- The following speakers have no relevant financial interest or affiliation with any organizations that could be perceived as a real or apparent conflict of interest in the context of the subject of their presentation:
 - Shari Bolen, MD, MPH; Michael W. Konstan, MD; Jennifer Molano, MD; Amy Zack, MD
- The following members of the planning committee do not have any disclosures or financial relationships from any ineligible companies:
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- Complete CME Evaluation and claim credits by **Friday, June 9.**
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Agenda

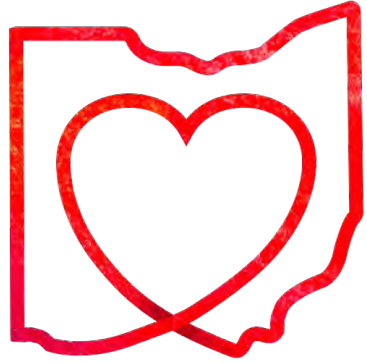
Topics	Presenter(s)	Timing
Welcome and Overview	Michael W. Konstan, MD Shari Bolen, MD, MPH	5 mins.
Heart Health and the Science of Sleep	Jennifer Molano, MD	40 mins.
Audience Question and Answer	Amy Zack, MD (Moderator) Jennifer Molano, MD	10 mins.
Next Steps and Wrap Up	Shari Bolen, MD, MPH	5 mins.



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Ohio Cardiovascular and Diabetes Health Collaborative

Heart Health and the Science of Sleep

Jennifer Molano, MD

Associate Professor

Neurology and Rehabilitation Medicine

University of Cincinnati College of Medicine

Learning Objectives



1. Identify the cardiovascular implications of sleep conditions
2. Screen patients at risk for sleep conditions
3. Counsel patients on how to optimize sleep health

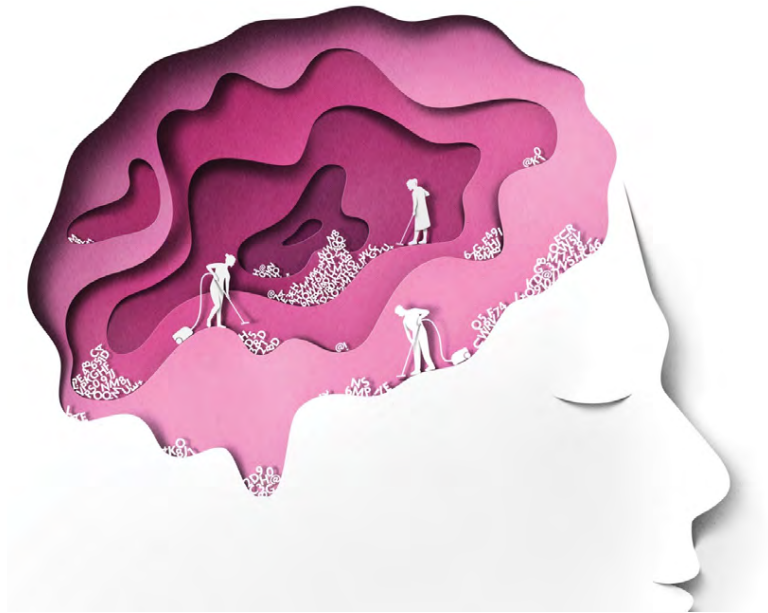
What is Sleep?

“A reversible behavioral state of perceptual disengagement from and unresponsiveness to the environment.”



Why Sleep?

- Sleep helps us perform well and feel better
- Theories include:
 - Conservation of metabolic energy
 - 15% less energy expenditure in sleep than quiet wakefulness
 - Cognition
 - Decreased intellectual performance after sleep deprivation
 - Memory consolidation
 - Regulation of temperature and other body functions



Why Do We Sleep?

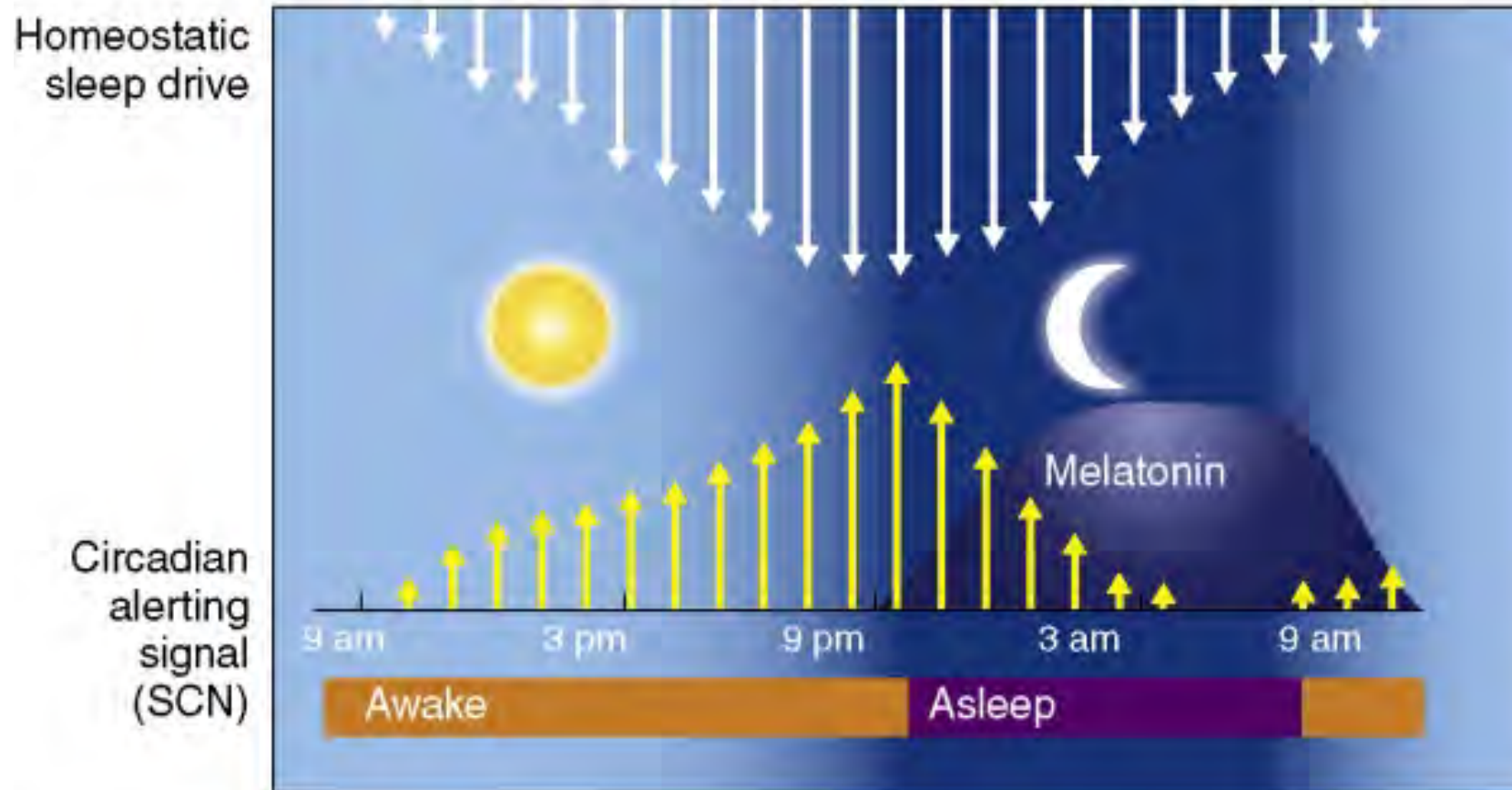
PHYSIOLOGIC REGULATION IN SLEEP		
NREMS	FUNCTIONS	REMS
ENDOCRINE		
GH early secretion PRL early secretion ACTH late secretion T late secretion TSH inhibition		
BREATHING		
Regular decrease Effective Effective Active Tone maintained	Ventilation Reflexes proprioceptive Chemoceptive Intercostal muscles Upper airways muscles	Variable amplitude Overridden Overridden Inactive Tone reduced
CIRCULATION		
Eurhythmic decrease Regular decrease Regular decrease Effective Effective	Heart rate Blood pressure Cardiac output Reflexes proprioceptive Chemoceptive	Variably arrhythmic Irregular oscillations Irregular oscillations Overridden Overridden
GASTROINTESTINAL		
Decreased function		Decreased function
RENAL		
Decreased urine flow		Decreased urine flow
SEXUAL		
		Penile erection
HOMEOSTATIC	REGULATION	POIKILOSTATIC
GH, growth hormone; PRL, prolactin; ACTH, adrenocorticotrophic hormone; T, testosterone; TSH, thyroid-stimulating hormone		



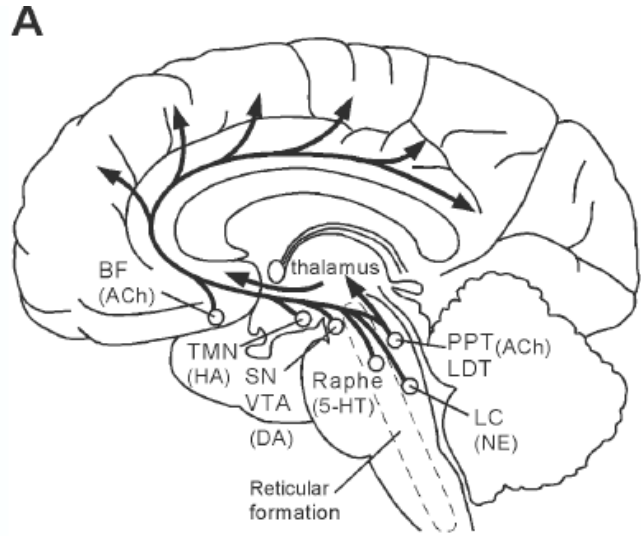
Kryger MG, Avidan AAY. Atlas of Clinical Sleep Medicine. 3rd Edition. 2010.
Lloyd-Jones DM, Allen NB, Anderson CAM, et al. Circulation. 2022;146: e18-e43.

What Drives Our Sleep Wake Cycle?

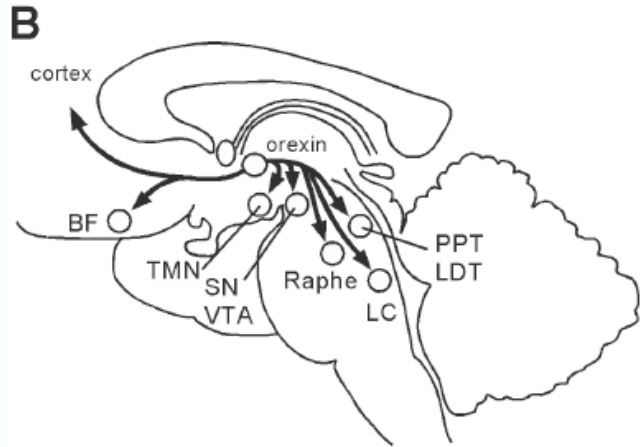
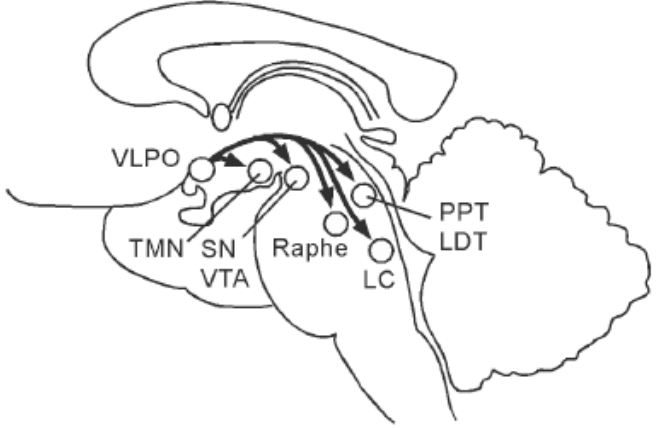
Sleep-wake cycle: Two-process model



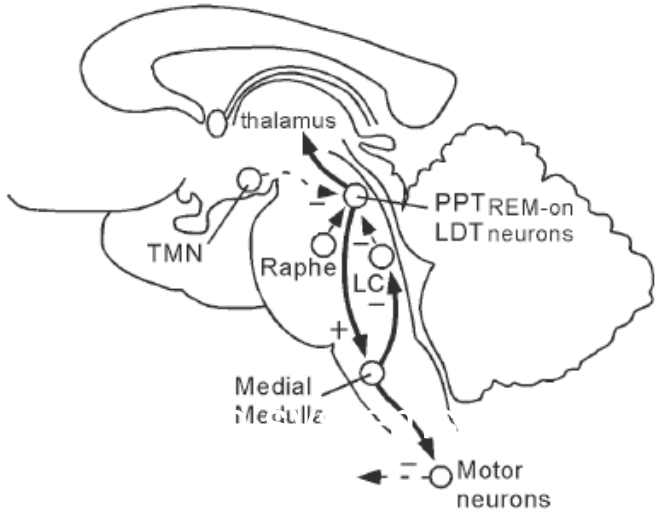
What Drives Our Sleep Wake Cycle?



NREM Sleep



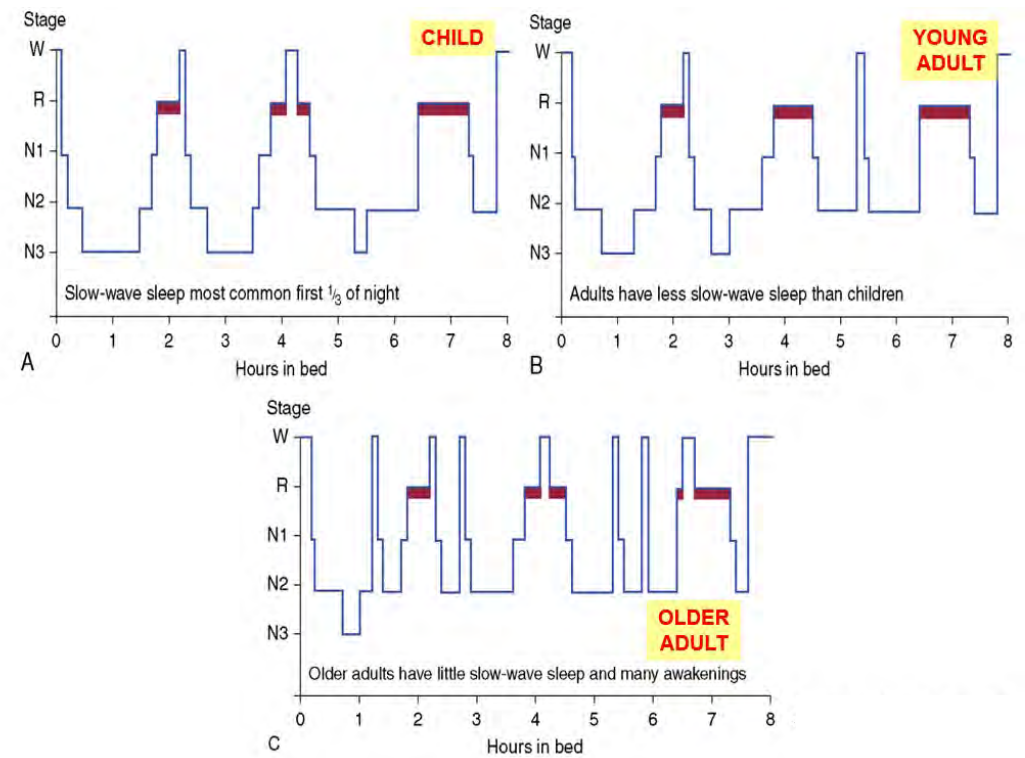
REM sleep



España RA, Scammell TE. Sleep. 2004; 27(4): 811-820.

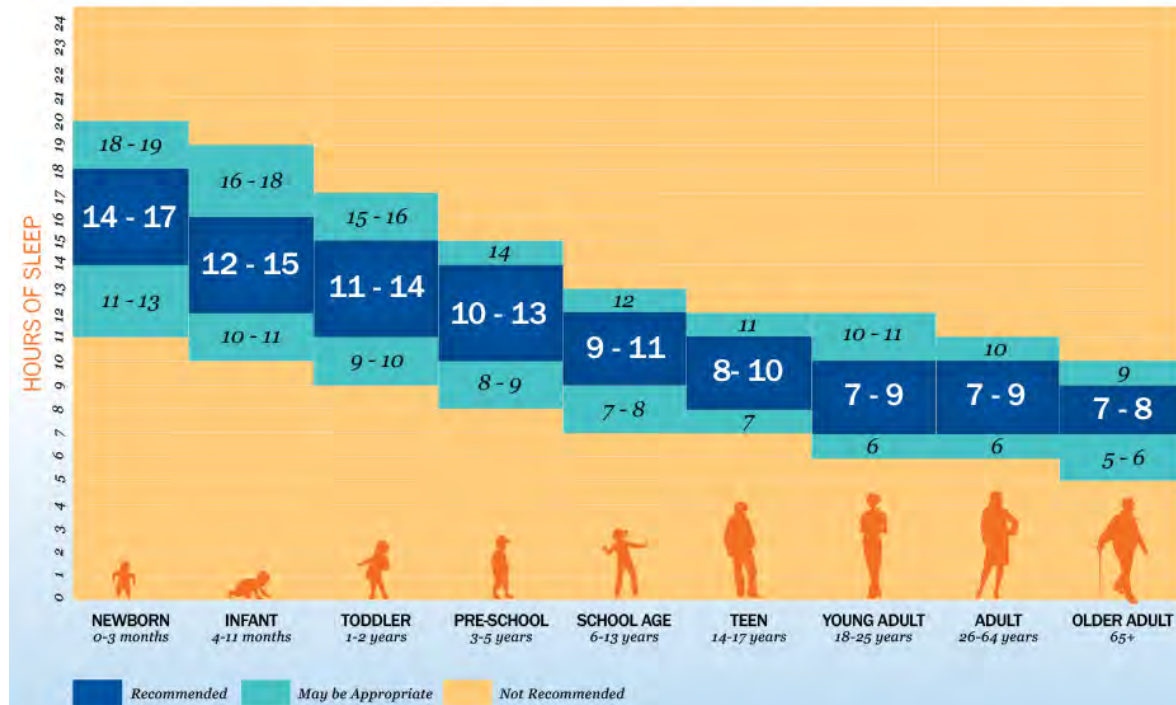
How is Sleep Organized?

- Non-REM (N1,N2,N3) and REM Sleep
- Sleep cycle is about 90-110 minutes
- Repeated 4-6 times per night
- Stage N3 decreases, REM increases as night progresses
- Young adults spend:
 - 50-60% of the night in stage N2
 - REM is ~20-25% of the night
 - Stage N3 is 15-20% of the night



How Much Sleep Do We Need?

SLEEP DURATION RECOMMENDATIONS



SLEEPFOUNDATION.ORG | SLEEP.ORG

Hirshkowitz M, The National Sleep Foundation's sleep time duration recommendations: methodology and results summary, Sleep Health (2015), <http://dx.doi.org/10.1016/j.sleh.2014.12.010>

What Happens When Someone Does Not Sleep Well?



Hypersomnia



Insomnia

A General Approach to Sleep Issues

- Sleep quantity
 - Number of hours in bed
- Sleep quality
 - Perception of sleeping the entire time while in bed
- What time do you go to bed?
- What time do you wake up?
- Do you have trouble falling asleep or staying asleep?
 - Number of minutes or hours awake
 - Sleep onset
 - Sleep maintenance
 - Number of awakenings
- Do you nap in the day?

A General Approach to Sleep Issues

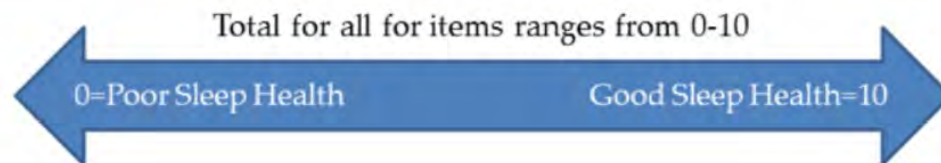
Other history to obtain:

- Caffeine, alcohol, drug use
- Past medical and surgical history
- Medications
- Changes in weight
- Social or life stressors
 - Especially if history of insomnia
- Review of systems
 - Snoring
 - Dry mouth, morning headache, nasal congestion
 - Nocturia
 - Pain
 - Restless legs



Screening Tools for Sleep Issues

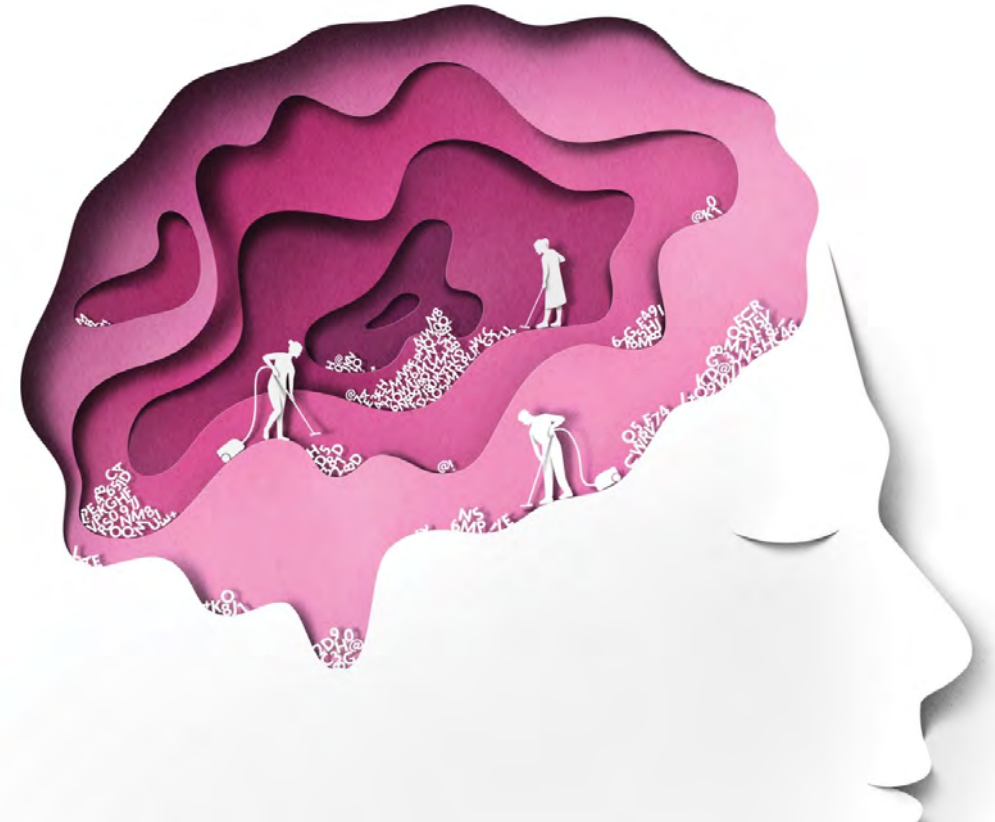
		Rarely/ Never (0)	Sometimes (1)	Usually/ Always (2)
<u>S</u> atisfaction	Are you satisfied with your sleep?			
<u>A</u> lertness	Do you stay awake all day without dozing?			
<u>T</u> iming	Are you asleep (or trying to sleep) between 2:00 a.m. and 4:00 a.m.?			
<u>E</u> fficiency	Do you spend less than 30 minutes awake at night? (This includes the time it takes to fall asleep and awakenings from sleep.)			
<u>D</u> uration	Do you sleep between 6 and 8 hours per day?			



- **RU-SATED**
 - Focuses on sleep health
- **Epworth Sleepiness Scale**
 - Obstructive Sleep Apnea (OSA)
- **OSA-Specific Screening Tools**
 - Berlin Questionnaire
 - STOP-BANG Questionnaire

Why is it Important to Address Sleep Issues?

- Fatigue
- Slower response time
- Increased errors
- Reduced learning
- Reduced flexible thinking
- Increased risk taking
- Higher emotional dysregulation
- Decreased resilience



Why is it Important to Address Sleep Issues?

Sleep and Cardiovascular Effects

- Sleep duration
- Insomnia
- Sleep and blood pressure
- Obstructive sleep apnea (OSA)
- Restless legs syndrome (RLS)
- Shift work disorder

Sleep Duration

Health Risks

- Sleep duration affects survival
- U-shaped effect, based on meta-analyses of 137 prospective cohort studies comprised of ~5 million participants

Short duration

- 12% increased risk for all-cause mortality
- 37-38% increased risk for incident diabetes mellitus and obesity
- 26% increased risk for coronary heart diseases
- 16-17% increased risk for cardiovascular disease and hypertension

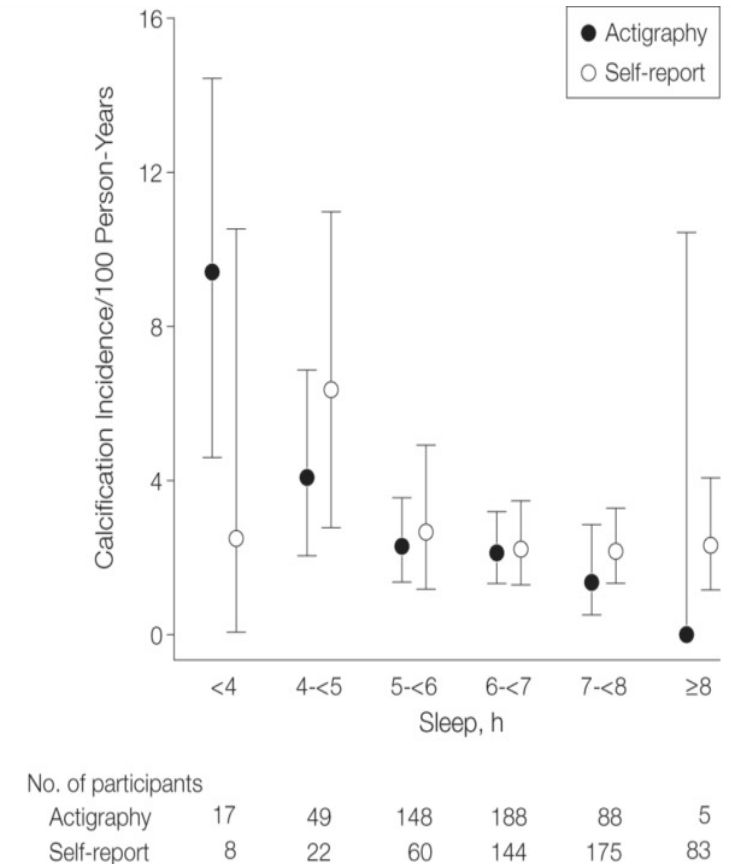
Long duration

- 39% increased risk for all-cause mortality
- ~25% increased risk for incident diabetes mellitus, cardiovascular disease, and coronary artery disease
- 46% increased risk for stroke

Sleep Duration

Coronary Artery Calcification Risk

- CARDIA cohort ages 18 to 30 in 1985-86
- Years 15 and 20, a subset of 495 subjects studied by sleep questionnaire and actigraphy in addition to the standard chest CT
- Odds of incident coronary calcification inversely related to hours of sleep
- An extra hour of sleep decreased estimated odds of calcifications by 33%
- Modeled effect of 1 extra hour of sleep nightly was equal to the modeled effect of a 16.5 mmHg drop in systolic blood pressure



Insomnia

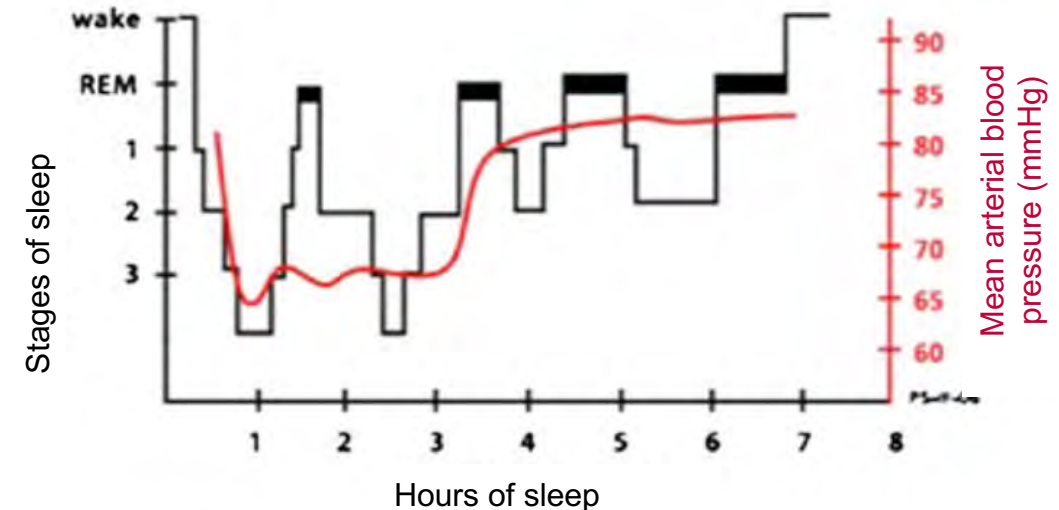
- **Definition:** Difficulty falling asleep, staying asleep, or early morning awakenings
- Health Professionals Follow-Up Study
 - Prospective cohort of 23,447 male health professionals
- Insomnia complaints associated with less physical activity, higher BMI, increased prevalence of depression, hypertension, cholesterol, triglycerides, diabetes, myocardial infarction, and stroke.
- Frequent sleep onset insomnia had a 55% (HR 1.55;95% CI, 1.19-2.04; p -trended=0.01) increased risk of cardiovascular disease mortality, particularly myocardial infarction
- No increased risk for difficulty maintaining sleep or early morning awakenings

Sleep and Blood Pressure

Normal Dipping

- Slow wave sleep (SWS)
 - Characterized by increased vagal tone and decreased sympathetic activity with subsequent lower heart rate and blood pressure
 - Decreased sympathetic vasoconstriction in the muscular vascular bed concomitantly
- REM sleep
 - Increased and variable sympathetic activity
- Overall blood pressure decreases by ≥ 10 mmHg during sleep, known as “dipping”

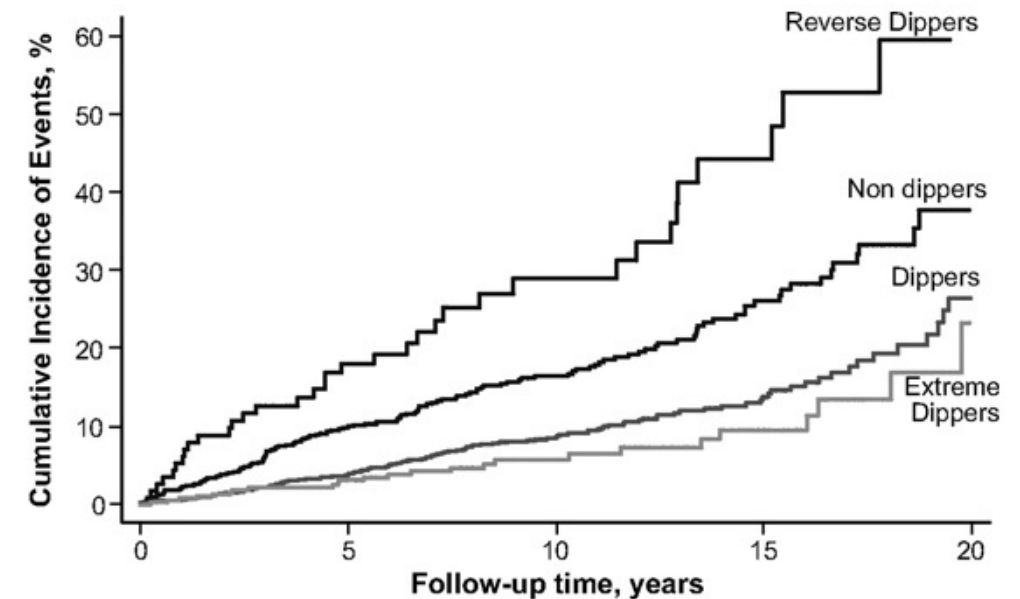
Normal Dipping Pattern in a Healthy Individual During a Typical Night of Sleep



Hypertension and Sleep

- Absence of dipping predicts poorer health outcomes and increased mortality
- Nocturnal hypertension or “reverse dipping” is more predictive of cardiovascular morbidity than daytime readings

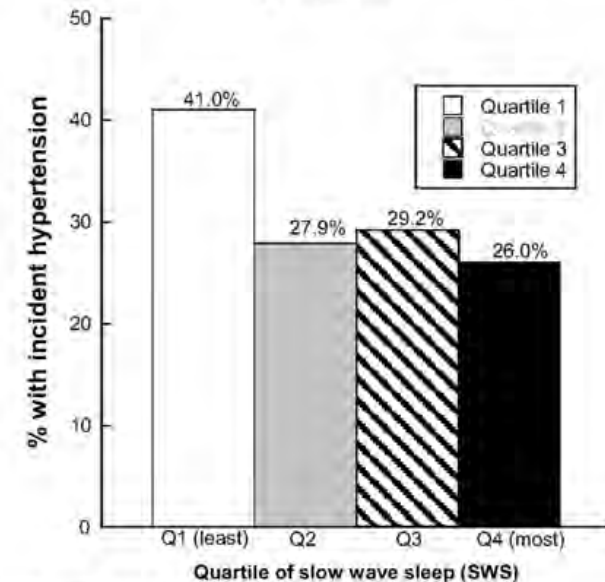
Kaplan-Meier curves reporting the cumulative incidence of cardiovascular disease in the 4 categories of dipping pattern.



Hypertension and Sleep

- MrOs Sleep Cohort, a longitudinal study of males >67 years old
- 784 normotensive males at baseline developed incident hypertension over the 3.5 years of the study
- 243 of them had significantly less slow wave sleep (SWS) at baseline
- Those with the lowest amount of SWS had 70% increased odds of developing hypertension vs those with higher amounts

Percentage of participants in each quartile of slow wave sleep (SWS) who developed incident hypertension.



Fung M M et al. Hypertension. 2011;58:596-603



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Hypertension and Obstructive Sleep Apnea (OSA)



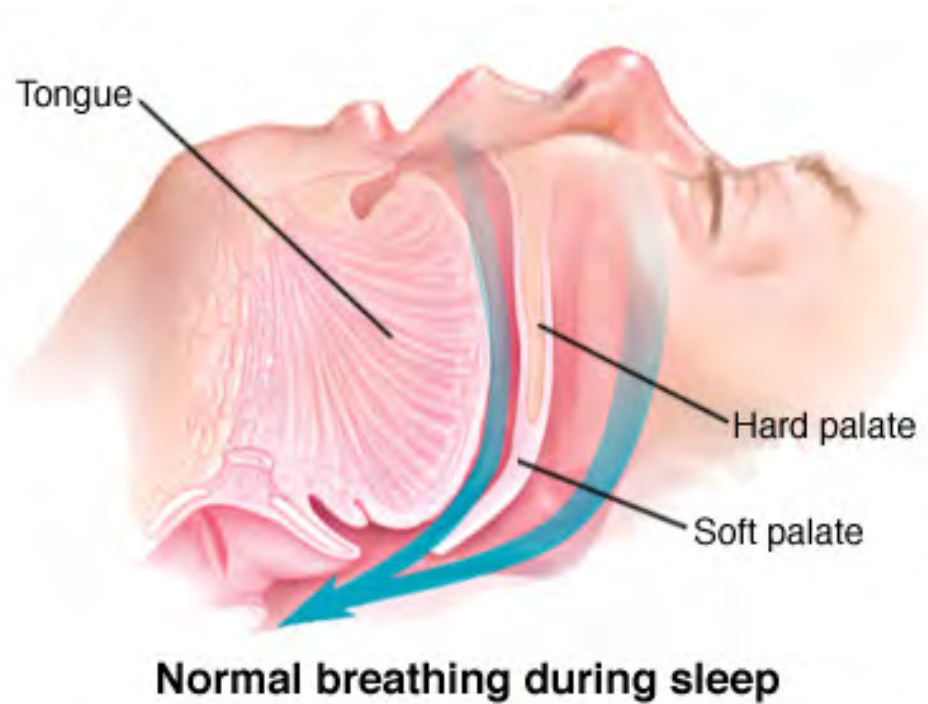
- OSA is common in those with hypertension
 - 30% in hypertensive adults and 80% in those with drug-resistant hypertension
- Untreated OSA is associated with increased hypertension
 - 18% increased risk in mild OSA
 - 56% increased risk in severe OSA
- OSA is associated with elevated nocturnal hypertension
- Severe OSA is associated with 8-fold risk of poorly controlled blood pressure in Black patients
- Increased severity of OSA is associated with a non-dipping pattern of blood pressure
- Non-dippers have an increased risk for stroke, cardiovascular disease-related and all-cause mortality
- Treatment for OSA may decrease blood pressure

What is Obstructive Sleep Apnea (OSA)?

- Seen in 5% of the population
 - But can be seen in up to 60% in those over 65 years old
- Higher incidence
 - Obesity, cardiac disease, pulmonary disease
- More common in males
- Snoring is most common associated symptom (90%)
 - Other symptoms
 - Daytime sleepiness
 - Morning headaches
 - Dry mouth

What Happens in Obstructive Sleep Apnea (OSA)?

Open Airway

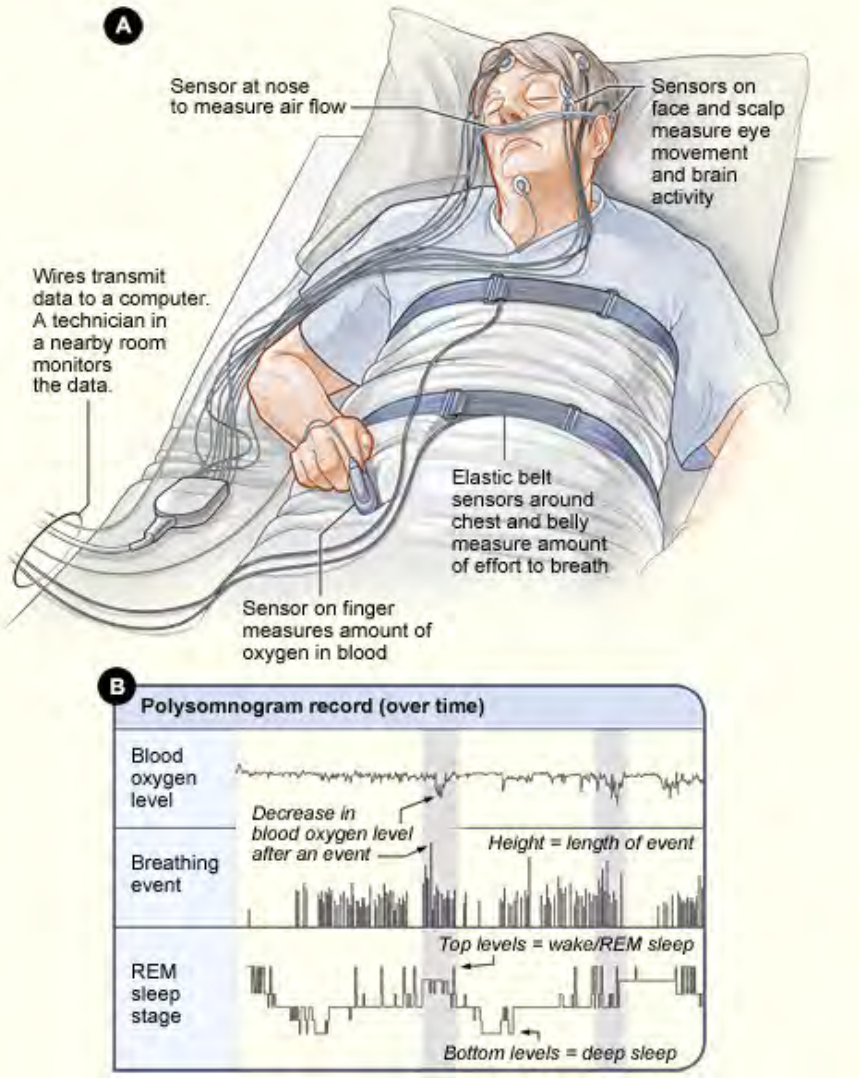


Blocked Airway (OSA)

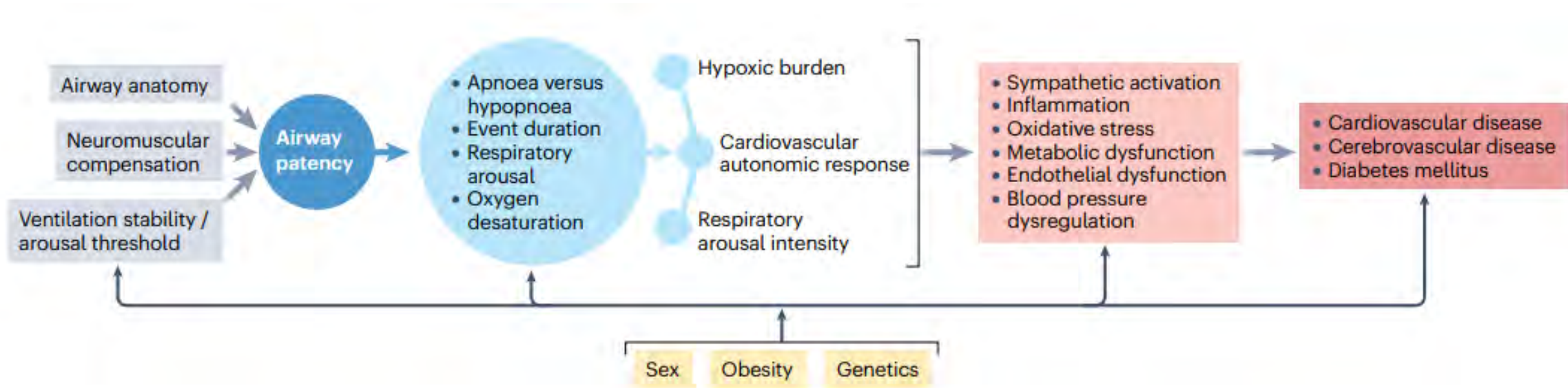


How Do We Evaluate for Obstructive Sleep Apnea (OSA)?

Polysomnography (Sleep Study)



Obstructive Sleep Apnea (OSA) and Cardiovascular Risk



Obstructive Sleep Apnea (OSA) and Diabetes Mellitus

- OSA is common in those with diabetes mellitus
 - Seen in 15-30% of patients with diabetes
- OSA is associated with an increased risk for diabetes mellitus
 - 35% increased risk in a meta-analysis of nine prospective cohort studies
- Untreated OSA is associated with increased diabetic complications
 - Peripheral neuropathy
 - Retinopathy
 - Nephropathy
- CPAP (continuous positive airway pressure) may improve glycemic control
 - Especially if used >6 hours per night

Obstructive Sleep Apnea (OSA) and Coronary Artery Disease

- OSA associated with increased coronary artery calcium scores (>400)
 - Seen in 15-30% of patients with diabetes
- OSA increases the risk for coronary artery disease
 - 3-fold risk in moderate-severe OSA in females and severe OSA in males
- Mixed results on the effects of CPAP on decreasing OSA risk
 - CPAP (continuous positive airway pressure) compliance is a factor

Obstructive Sleep Apnea (OSA) and Stroke

- OSA is associated with an increased risk for stroke
 - Risk may increase due to the degree of OSA
 - Males with severe OSA had a 3-fold increased risk for stroke
- Treatment for OSA may decrease stroke risk
- Treatment for OSA may improve neurological outcomes in those with stroke

Other Sleep Conditions and Cardiovascular Risk

Restless Legs Syndrome (RLS)



Diagnostic Criteria

- **Urg**e to move limbs, usually accompanied or caused by uncomfortable and unpleasant feelings in the limbs
- **Rest** worsens or inactivity precipitates symptoms
- **Getting** up or moving improves the urge to move
- **Evening** worsening or nighttime appearance of symptoms

Note: Symptoms should be associated with functional difficulties.

Restless Legs Syndrome (RLS) and Cardiovascular Disease

- Epidemiological data is mixed
- Increased sympathetic nervous system activation may contribute
 - Similar to Obstructive Sleep Apnea (OSA)

Cohort	N (RLS%)	Mean age, duration of follow-up	RLS information (duration, frequency, 1°/2°)	CVD outcomes
Study of Health in Pomerania [9]	4 308 (10.1%)	50.3 5 years	None; no change w/out DM cases	- MI -CVA
Women's Health Study [7]	29 756 (11.7%)	63.4 6 years	None; no change w/out ESRD/PVD	- first CVD event -MI -CVA
Physician's Health Study [6]	19 182 (7.2%)	66.6 7.3 years	None; no w/out ESRD/PVD	- first CVD event -MI -CVA
Nurse's Health Study [10]	70 977 (2.1%)	67 5.6 years	Duration > 3 years; no change w/out DM/ESRD	+(fatal) CHD +first MI event
Kaiser Permanente [11]	~12 000 with RLS	1°=58; 2°=65 3.9 years	Physician diagnosis; 2°=many comorbidities	1°:-CVD/MI/CVA 2°:+CVD/MI/CVD
Veterans Administration [12]	7 392 (0.1%)	59.8 8.1 years	"Incident RLS"	+CHD/CVA

Abbreviations: "-", no relationship; "+", positive relationship with RLS; DM, diabetes mellitus; ESRD, end-stage renal disease; CHD, chronic heart disease; MI, myocardial infarction; PVD, peripheral vascular disease; CVA, cerebrovascular accident; 1°, primary RLS; 2°, secondary RLS.

Restless Legs Syndrome (RLS) and Cardiovascular Disease

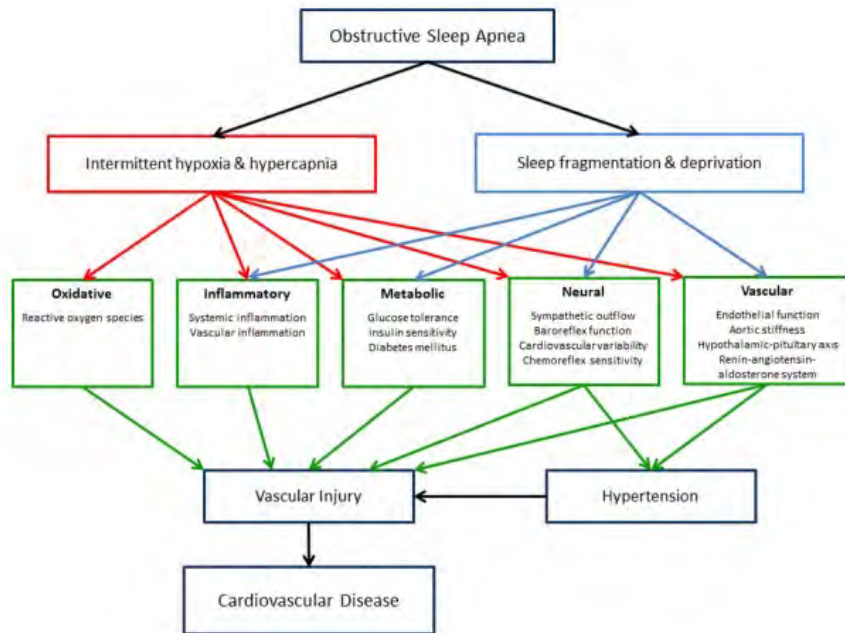


Figure 1. Proposed mechanisms linking obstructive sleep apnea to cardiovascular disease
While intermittent hypoxemia is unlikely to be relevant to RLS, there is evidence that sleep fragmentation and sleep deprivation may contribute to neural, metabolic, vascular, and inflammatory mechanisms. These pathways may therefore inform research in to the possible cardiovascular consequences of RLS.

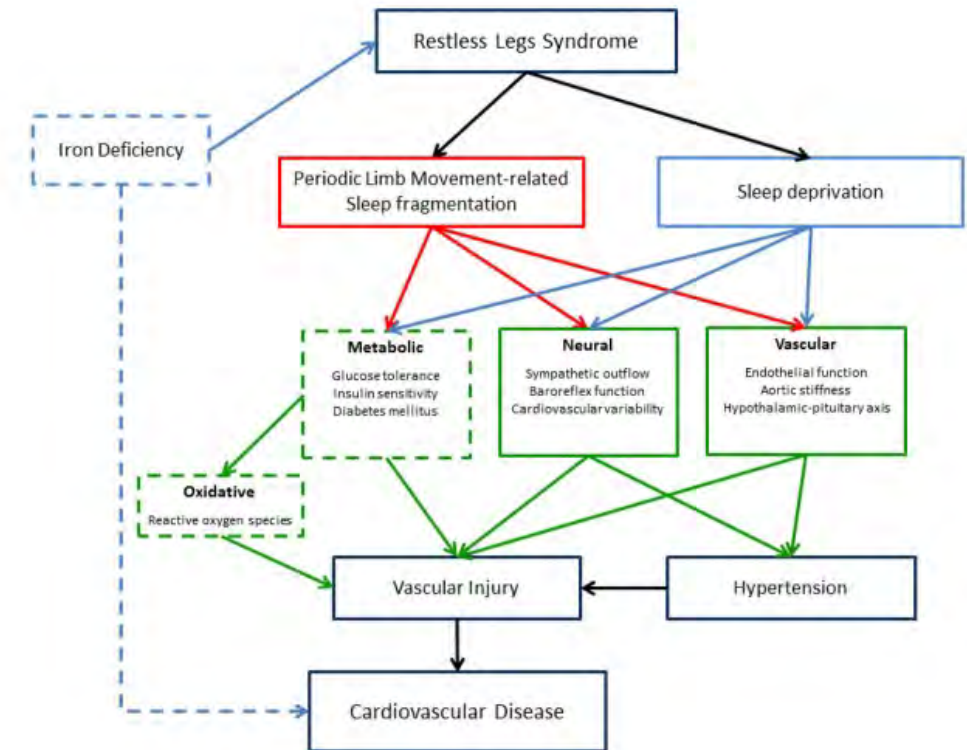
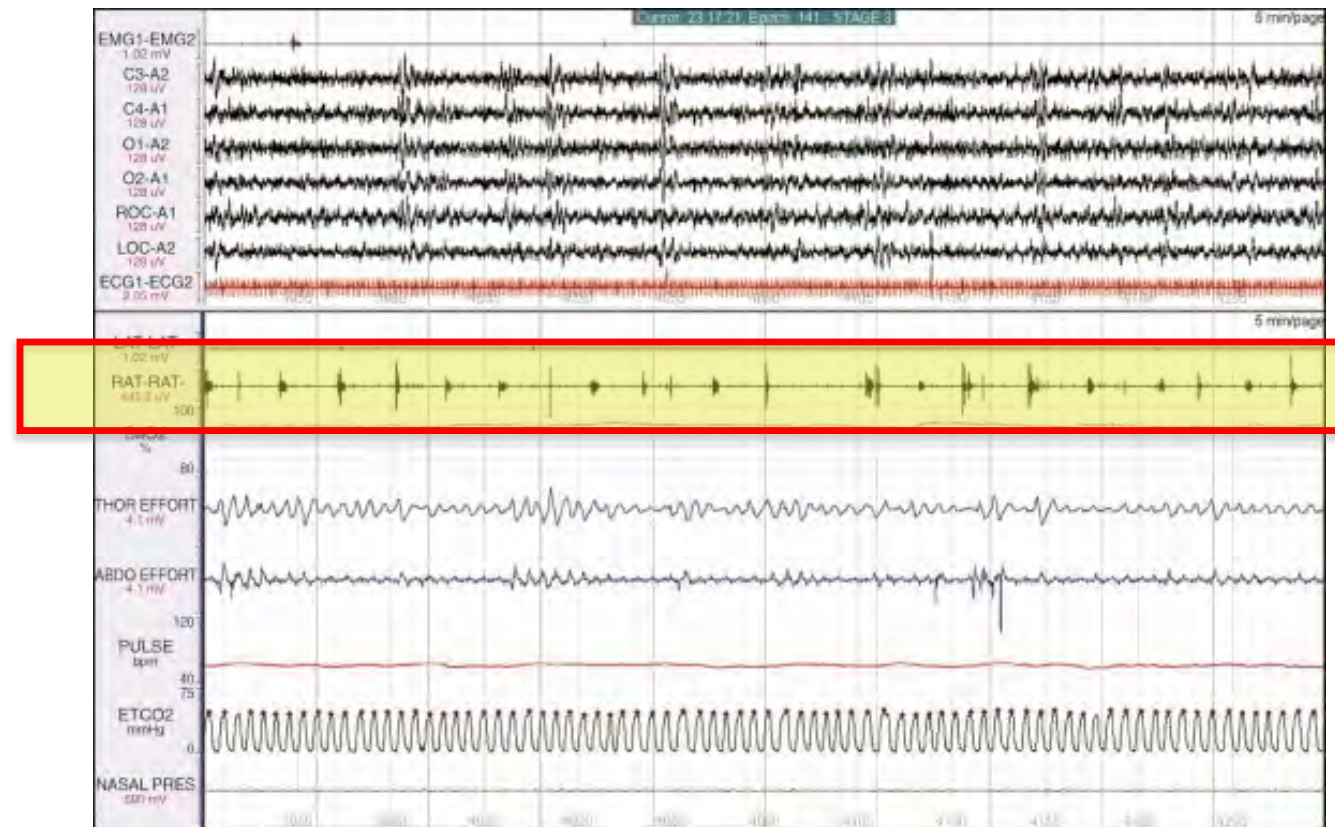


Figure 2. Plausible mechanisms linking restless legs syndrome to cardiovascular disease
Those mechanisms for which there is some experimental evidence of an association with RLS or periodic limb movements are shown in solid boxes.

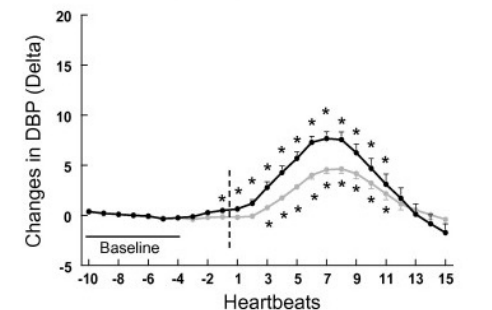
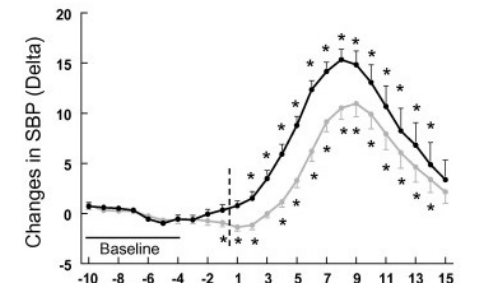
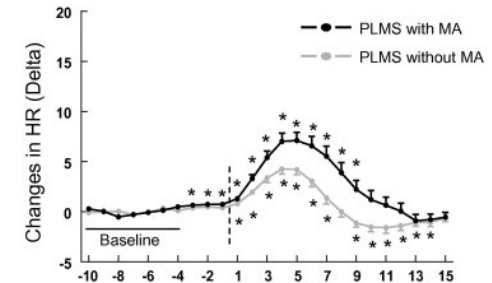
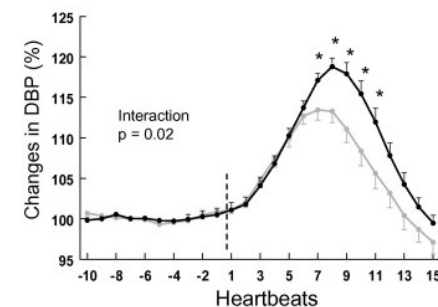
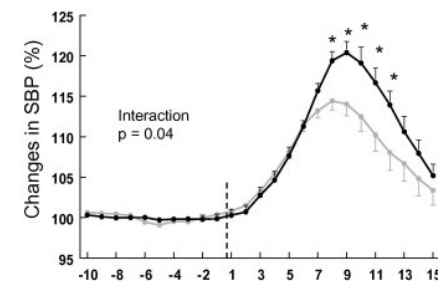
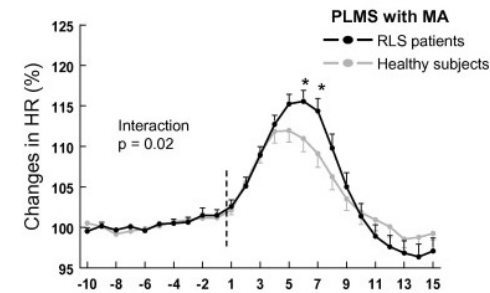
Periodic Limb Movements of Sleep

- Commonly seen in those with restless legs syndrome (RLS)
- Also, can be observed independently of RLS



Periodic Limb Movements (PLMs) of Sleep *Cardiovascular Effects*

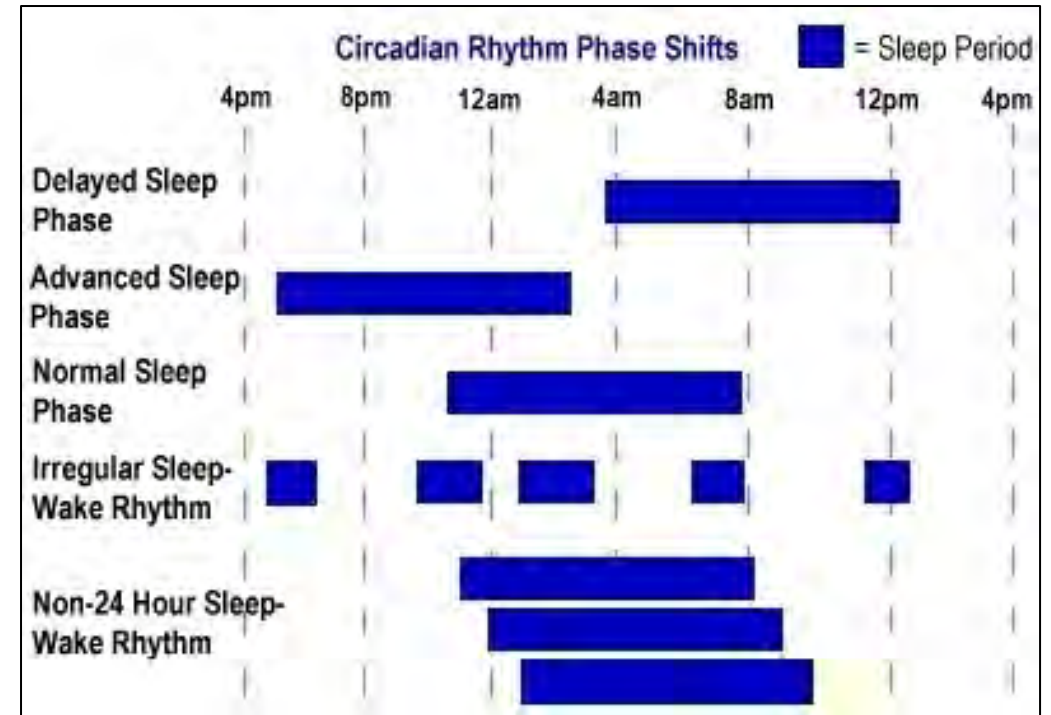
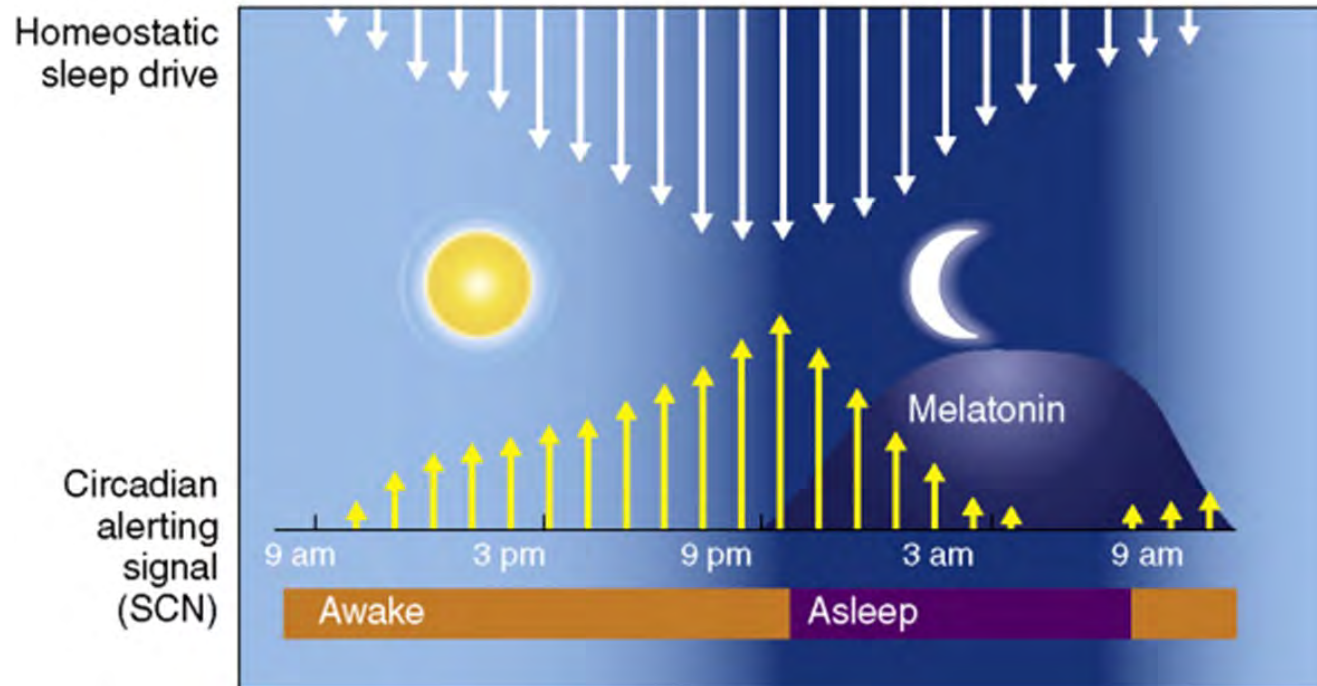
- Study with 14 restless legs syndrome (RLS) participants and 14 healthy controls
- PLMs associated with increased heart rate and blood pressure
 - More prominent in those with microarousals (MA)
- Cardiovascular effects more increased in those with RLS



Other Sleep Conditions and Cardiovascular Risk

Circadian Rhythm Disorders

Sleep-wake cycle: Two-process model



Shift Work and Cardiovascular Risk



- Shift work has been associated with an increased risk of myocardial infarction, stroke, and coronary events
- May have dose-response relationship to increased years of shift work
- May be more likely to have a non-dipping blood pressure pattern
- Increased risk for higher systolic and diastolic blood pressures

How Do We Optimize Sleep?



Optimizing Sleep Duration



- Educate patients about the importance of adequate sleep time and good sleep habits
- Cognitive behavioral treatment for insomnia (CBT-i) has good, long-term results in both primary insomnias and insomnia associated with disease states, e.g., cancer, depression, rheumatologic conditions, among others.

Optimizing Sleep Duration

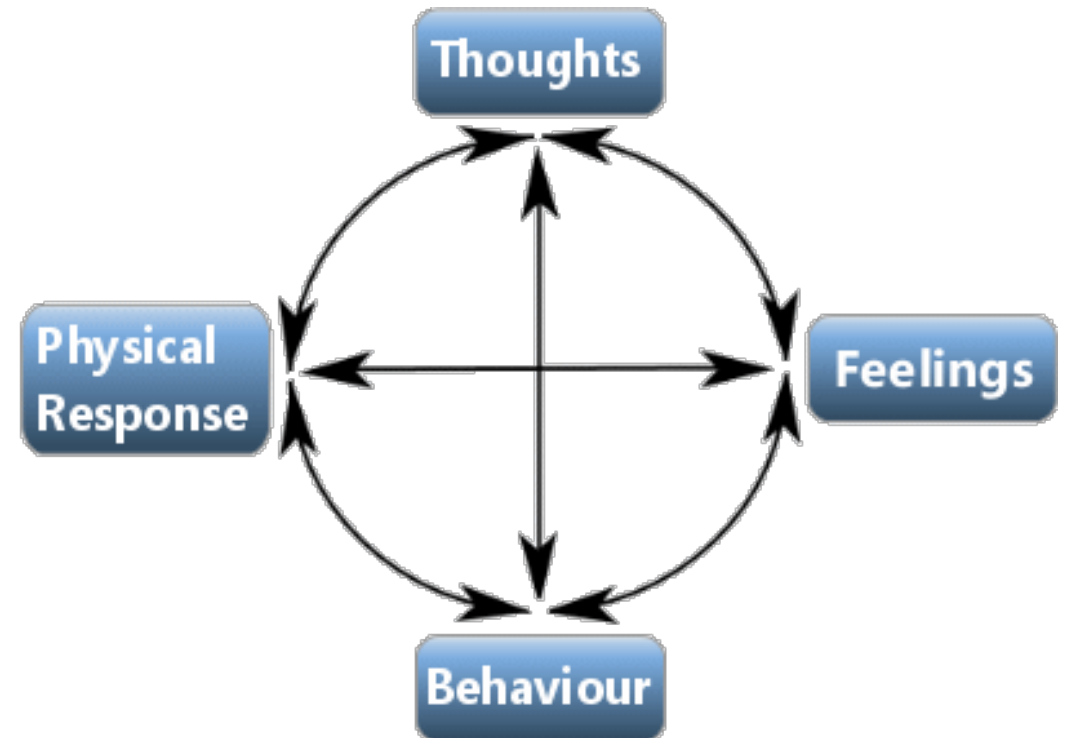
Pharmacotherapy for Insomnia

- Ideally used for acute insomnia
- Chronic, intermittent use may be beneficial
 - Anti-depressants or anti-epileptics with sedating side effects
- Studies show that behavioral treatments result in more sustained benefits over time

Optimizing Sleep Duration

Cognitive-Behavioral Therapy

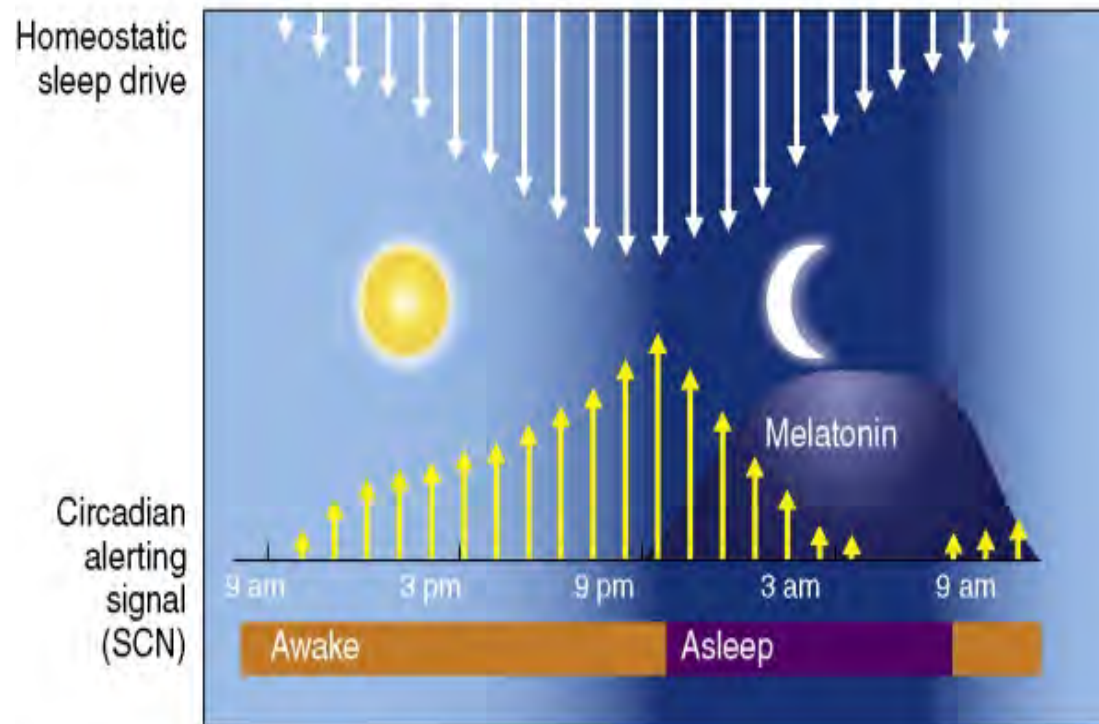
- Cognitive
 - Identify and correct negative thinking associated with sleep
- Behavioral
 - Practice good sleep habits
 - Alter schedule to promote sleep



Optimizing Sleep Duration

Why Practice Good Sleep Habits?

Sleep-wake cycle: Two-process model



Homeostatic drive for sleep

- Behaviors

Circadian rhythm

- Internal clock

It's All About The Sleep-Wake Cycle

Optimizing Sleep Duration

What Behaviors Can Hinder Our Ability to Fall Asleep?



Caffeine



Napping

Optimizing Sleep Duration

Non-Pharmacological Strategies for Insomnia



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- Eliminate naps
- No caffeine after 2 pm
- Keep a consistent sleep schedule
- Stimulus control - Keep the bed for sleep only!
 - Trains the brain to keep the room for sleep by advising the patient to go to bed only when sleepy
 - If unable to fall asleep or stay asleep after 20 minutes, the patient should be instructed to go into another room and perform a relaxing activity
- Relaxation training
 - Tensing and relaxing various muscles in the body, with the goal of calming a patient's state of arousal

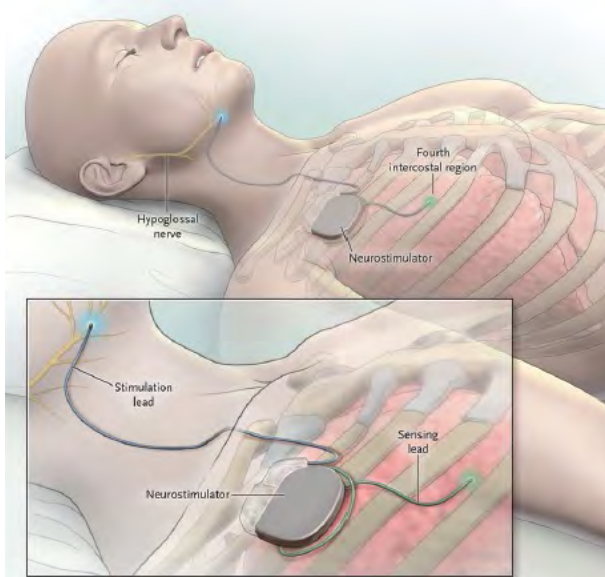
How Do We Treat Obstructive Sleep Apnea (OSA)?



Continuous Positive
Airway Pressure
(CPAP)

How Do We Treat Obstructive Sleep Apnea (OSA)?

Alternative Treatments



Surgical options



Oral appliances



Positional therapy

How Do We Treat Restless Legs Syndrome (RLS)?

Non-Pharmacological Management



- Check iron status
- Recommend mentally alerting activities
 - Decrease times of boredom
- Wean or abstain from caffeine
- Review the medication list
- Moderate (not intense) exercise may also be helpful


How Do We Treat Restless Legs Syndrome (RLS)?

Pharmacological Management



- Dopamine agonists
 - Ropinirole or pramipexole
 - Caveat: Impulse control disorders and augmentation
- Alpha-2-delta calcium channel ligands
 - Gabapentin or Pregabalin
- In refractory cases, low-dose opioids

How Can We Optimize Sleep in Shift Workers?

thebmj Visual summary 

Optimising sleep for night shifts

The below diagram describes a sleep strategy, based on the evidence included in the article by Helen McKenna and Matt Wilkes. It is supported by their wider reading and discussions with experts in the field, although they note that quality of evidence is low. They offer this as a starting point from which to develop your own sleep strategy.



How Can We Optimize Sleep in Shift Workers?



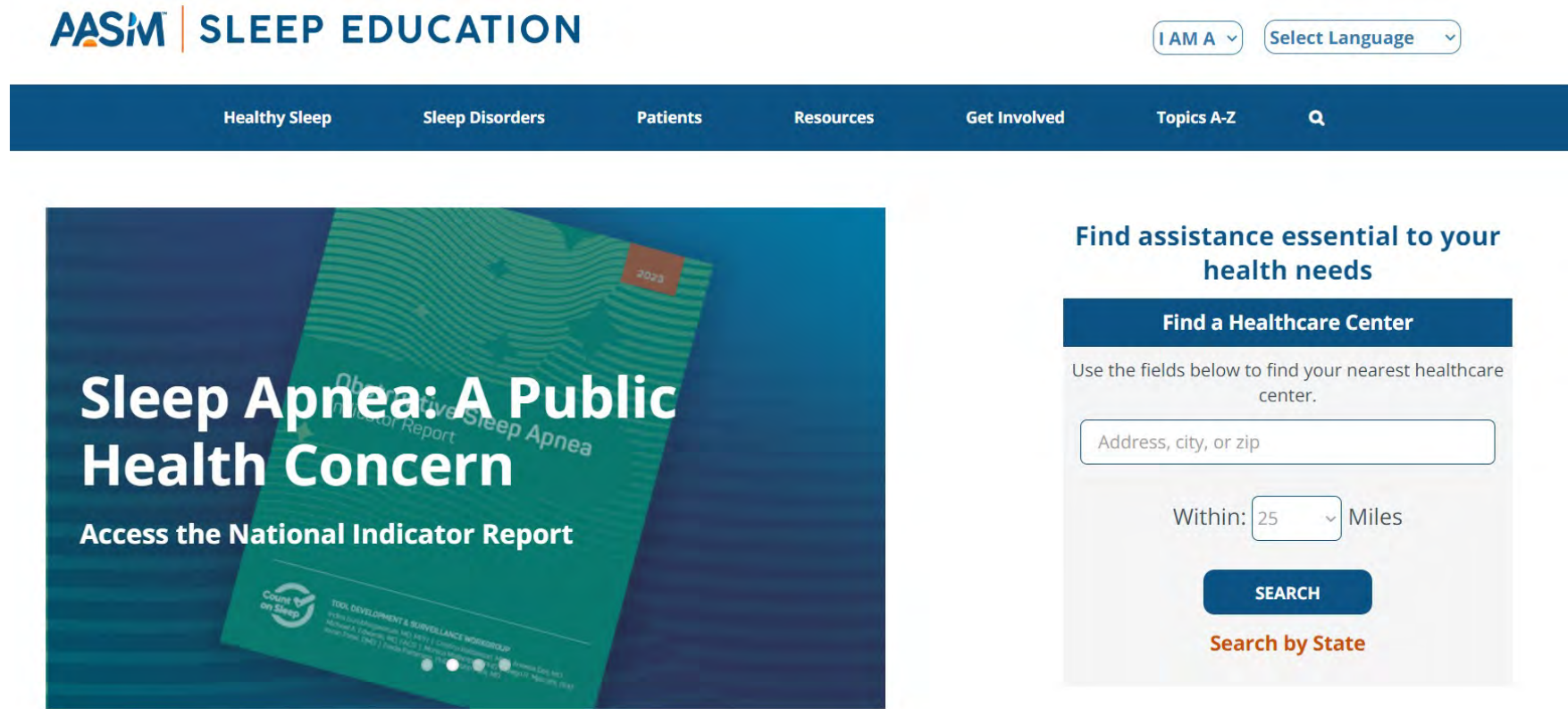
General Tips to Improve Sleep

- Consistent sleep schedule
 - Regular sleep time and bedtime routine
 - Seven to nine hours of sleep a night (18-65 years old)
- Optimal treatment of medical and psychiatric conditions
- Review of medication lists
- No naps
 - Limit to less than 30 minutes a day if necessary
- No caffeine use after 2 pm
- Avoid nicotine and alcohol before bedtime
- Maintain bedroom for nighttime activities only
- Cool, comfortable environment

Learning Objectives

1. Identify the cardiovascular implications of sleep conditions
2. Screen patients at risk for sleep conditions
3. Counsel patients on how to optimize sleep health

Resources



AASM | SLEEP EDUCATION I AM A Select Language

Healthy Sleep Sleep Disorders Patients Resources Get Involved Topics A-Z

Sleep Apnea: A Public Health Concern

2023

Access the National Indicator Report

Count on Sleep

TOOL DEVELOPMENT & SURVEILLANCE WORKGROUP

Find assistance essential to your health needs

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Use the fields below to find your nearest healthcare center.

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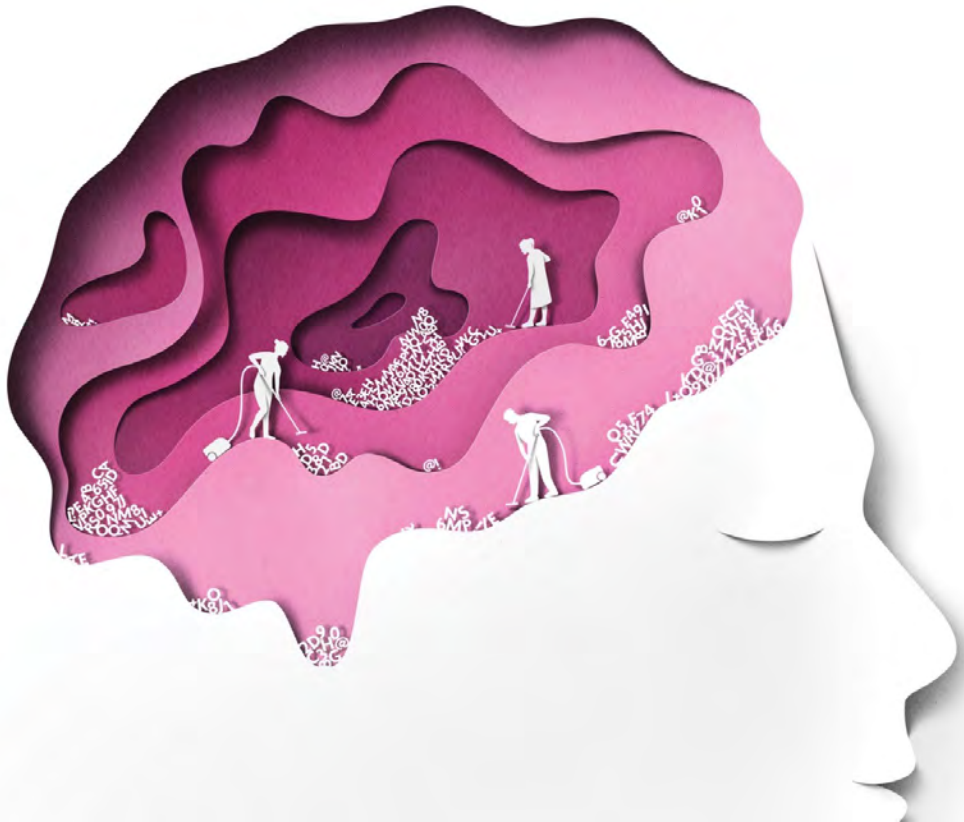
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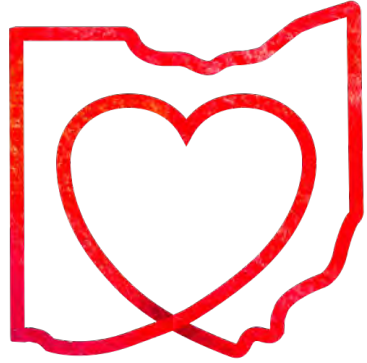
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sleepeducation.org

Thank You



Konnikova M. NY Times Sunday Opinion. 2014.
Lloyd-Jones DM, Allen NB, Anderson CAM, et al. Circulation. 2022;146:e18-e43.



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Ohio Cardiovascular and Diabetes Health Collaborative

Audience Question and Answer

Amy Zack, MD

Case Western Reserve University School of Medicine

Speakers

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Jennifer Molano, MD
University of Cincinnati College of Medicine



Amy Zack, MD (Moderator)
Case Western Reserve University School of Medicine



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Ohio Cardiovascular and Diabetes Health Collaborative

Next Steps and Wrap Up

Shari Bolen, MD, MPH

Case Western Reserve University School of Medicine

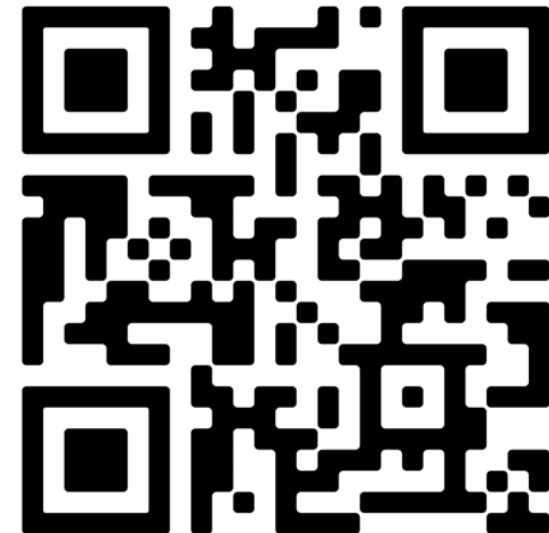
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The Ohio Cardiovascular and Diabetes Health Collaborative is funded by the Ohio Department of Medicaid and administered by the Ohio Colleges of Medicine Government Resource Center.

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