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Professional Education Series

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The Physiological Connections Between Diet & Sleep and Associated Risks for Developing Chronic Diseases

TODAY'S AGENDA:

- Introduction & Housekeeping
- Speaker Introduction
- Presentation
- Q&A
- Closing



WEBINAR HOST:

Acacia Wright, RD, CD
Sr. Manager of Nutrition
Communications
Orgain, LLC



WEBINAR PRESENTER:

Marie-Pierre St-Onge, Ph.D, CCSH, FAHA
Associate Professor of Nutritional Medicine,
Director of Columbia University Irving
Medical Center Sleep center of Excellence



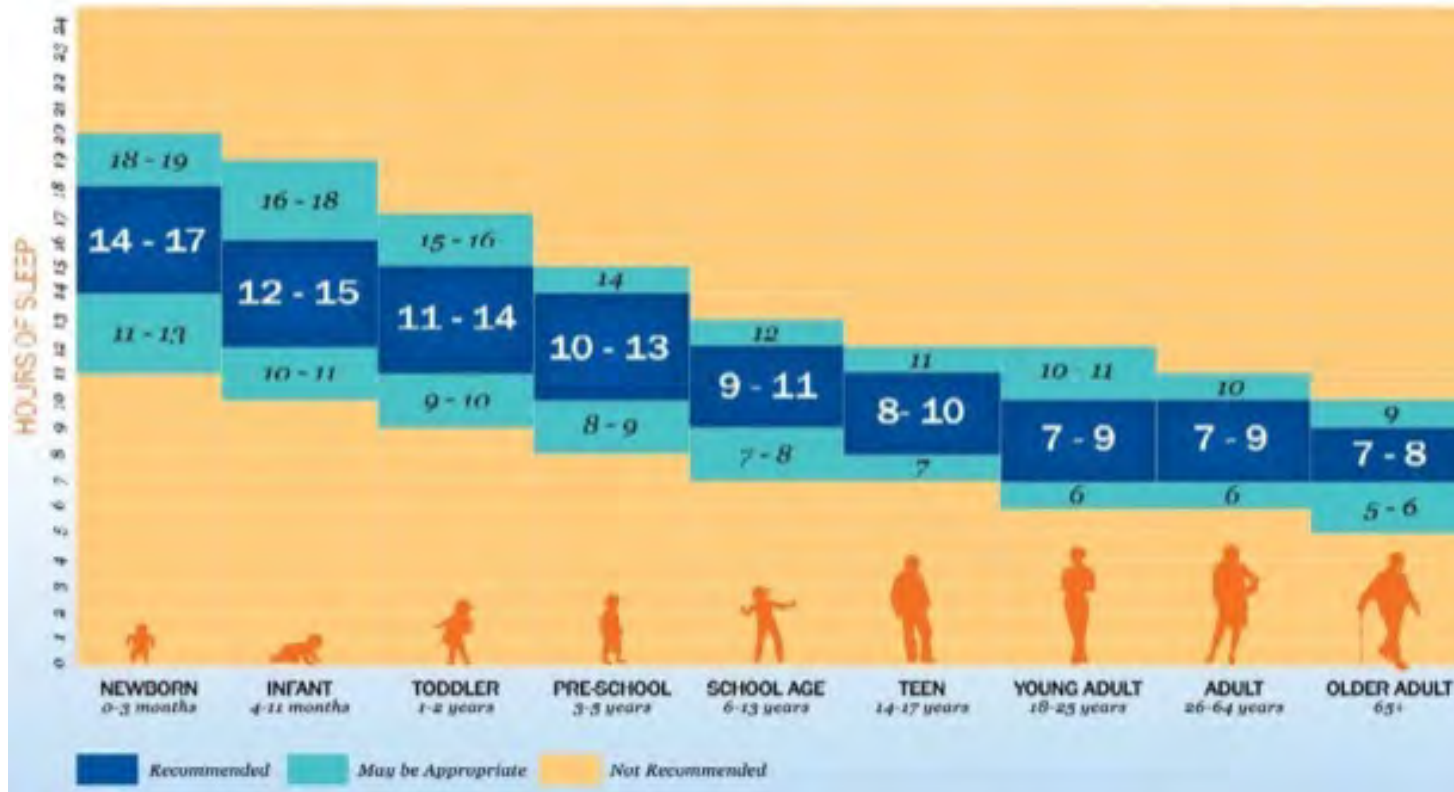
The Physiological Connections Between Diet & Sleep: Associated Risks for Developing Chronic Diseases

Marie-Pierre St-Onge, PhD, CCSH, FAHA
Associate Professor, Division of General Medicine
Director, Center of Excellence for Sleep & Circadian Research
Department of Medicine, Columbia University Irving Medical Center

Learning Objectives

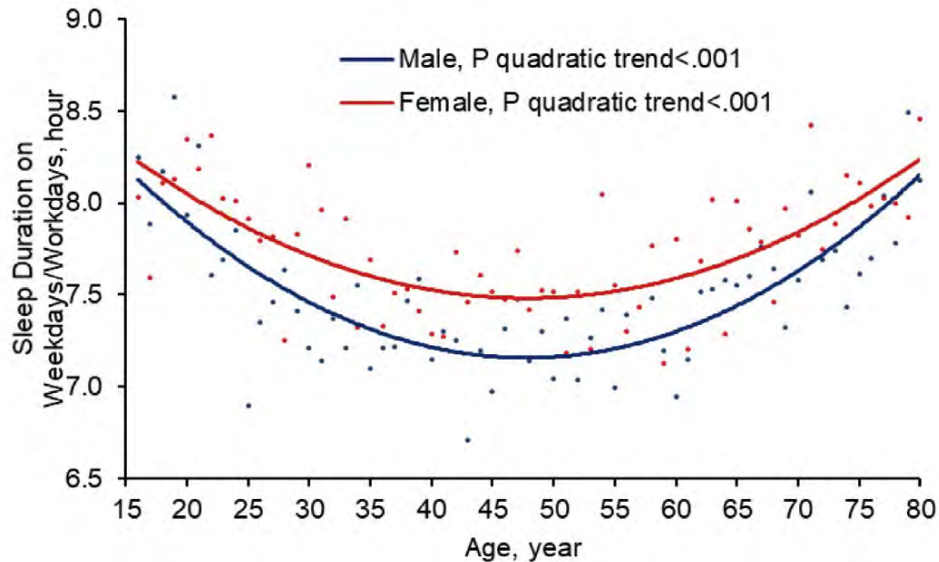
- Explain the relationships between sleep, obesity, and chronic diseases
 - Epidemiological evidence
 - Clinical intervention studies
- Describe mechanisms by which sleep influences obesity and chronic disease risk
 - Short & long-term effects of sleep restriction
 - Patterns of sleep
- Discuss influence of diet on sleep

SLEEP DURATION RECOMMENDATIONS

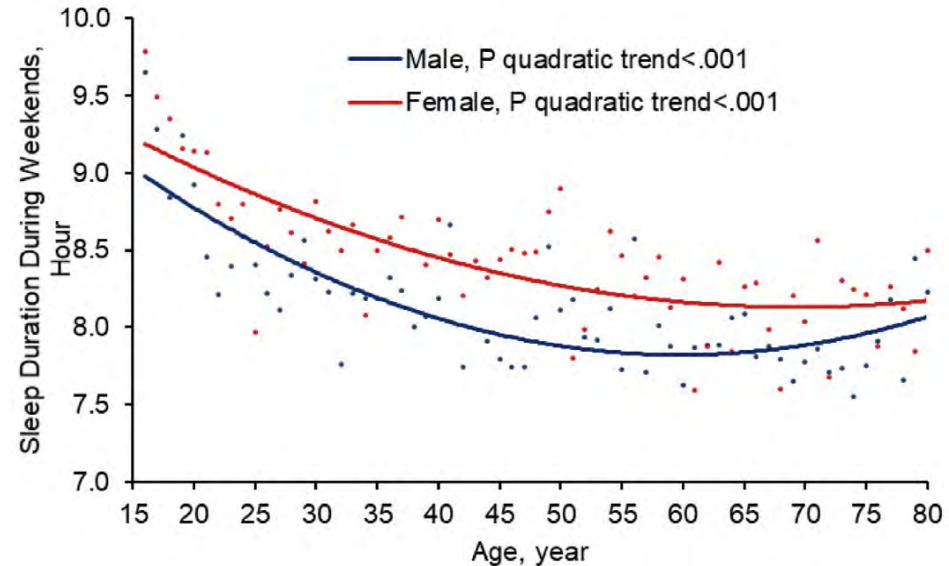


Sleep Duration in Males vs Females: Weekdays vs Weekends

Weekdays



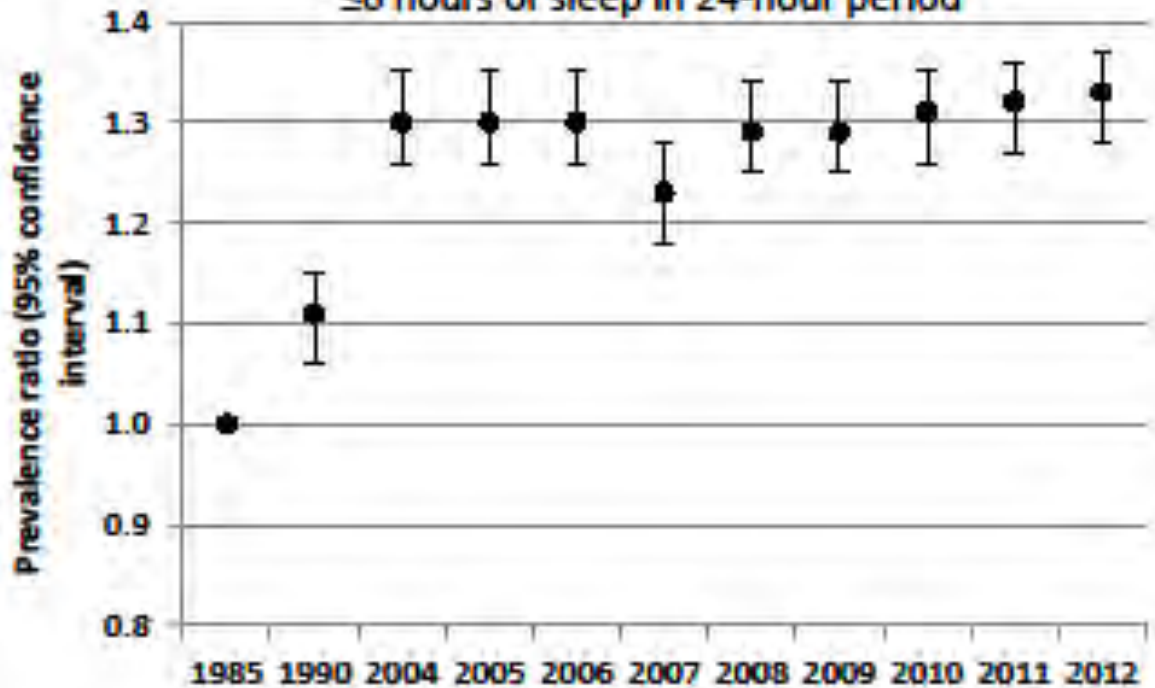
Weekends



In general, females report 20 minutes longer sleep than males across the lifespan & fewer report short sleep duration

Rising prevalence of short sleep in US adults

≤6 hours of sleep in 24-hour period



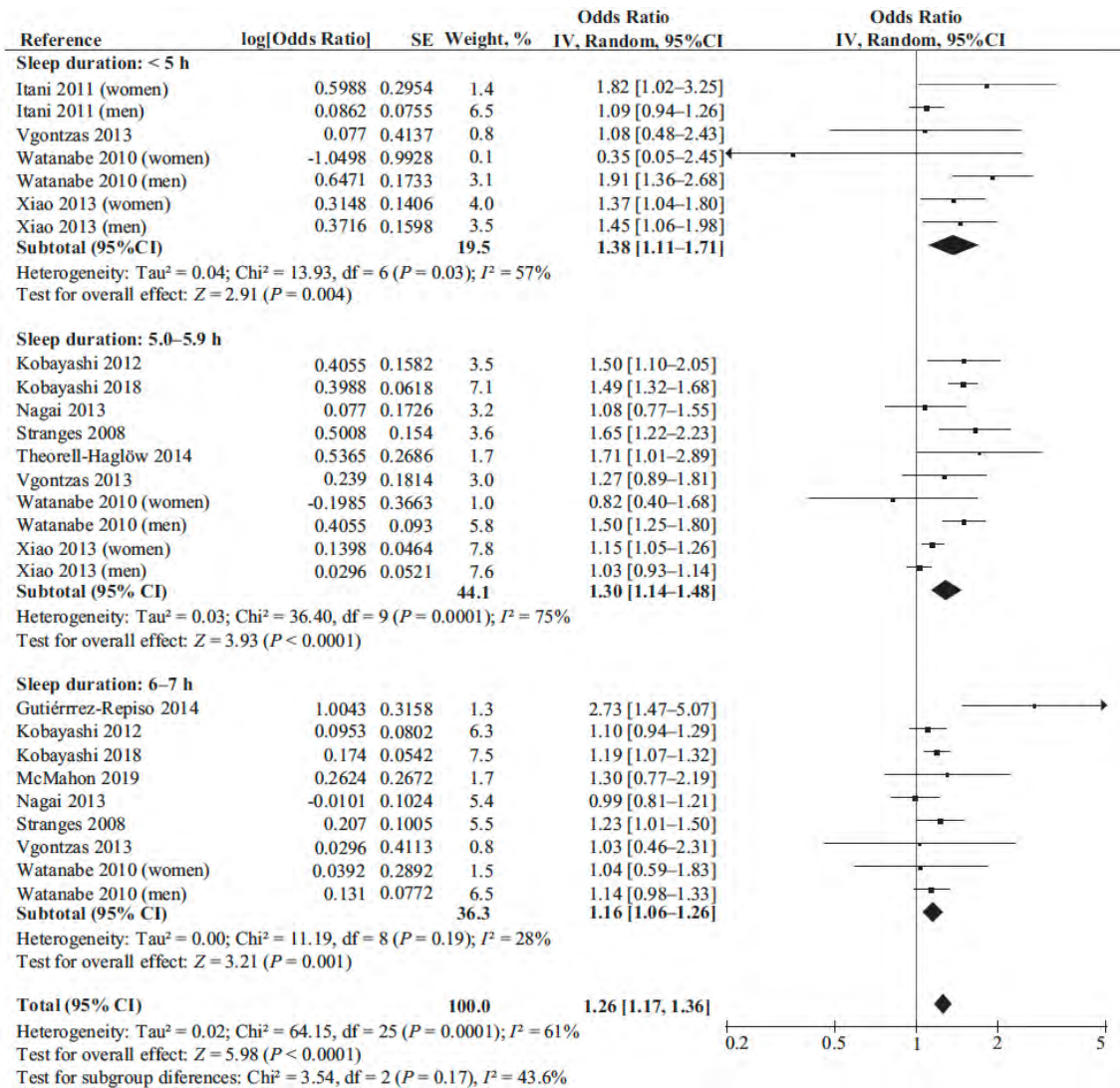
Age-adjusted prevalence of sleeping ≤6 h/night:

- 1985: 22.3%
- 2012: 29.2%

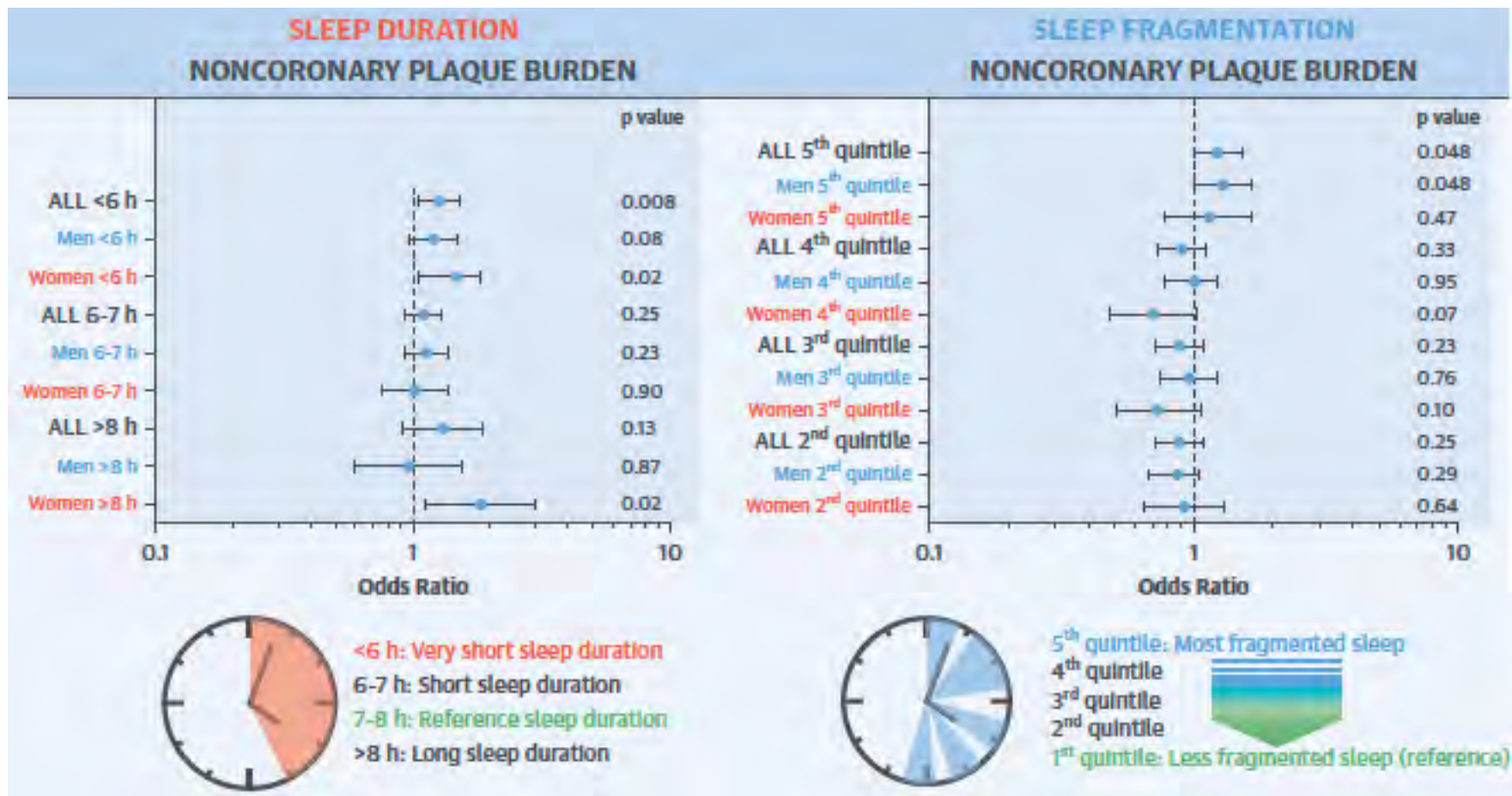
Age-adjusted prevalence of sleeping 7-8 h/night:

- 1985: 65.9%
- 2012: 62.8%

Increased odds of developing obesity in short sleepers

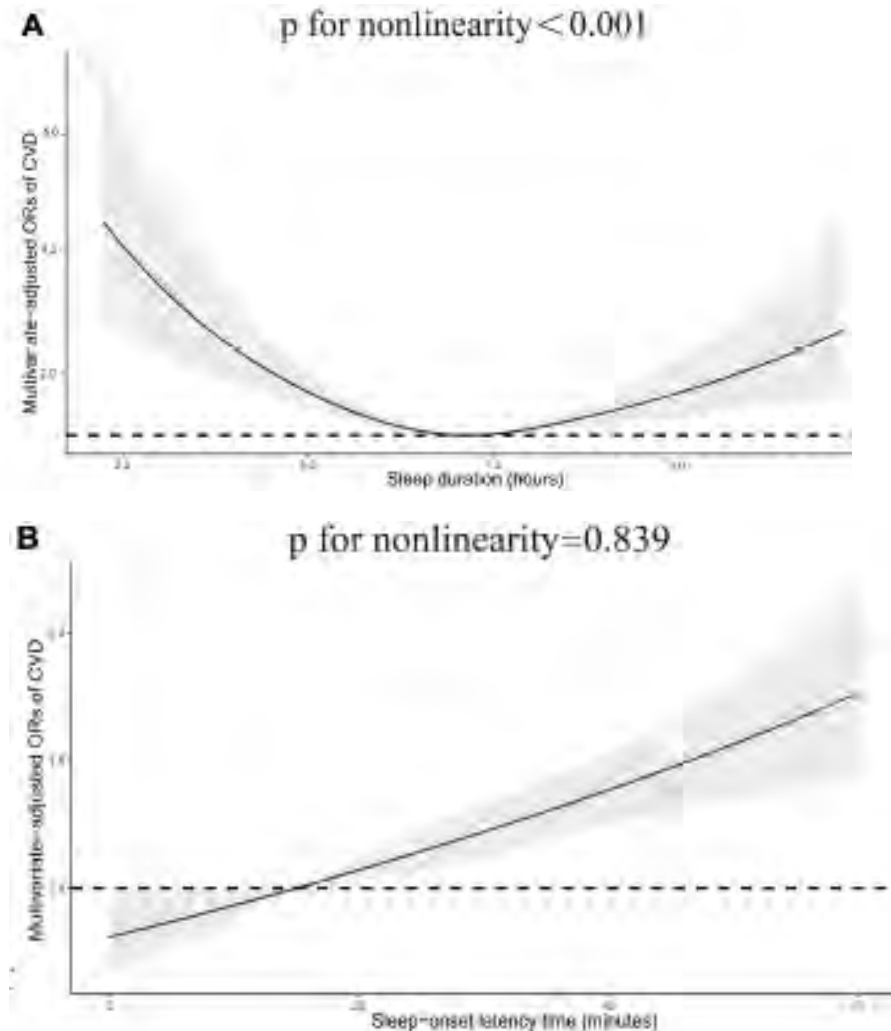


Increased subclinical atherosclerotic burden associated with poor sleep

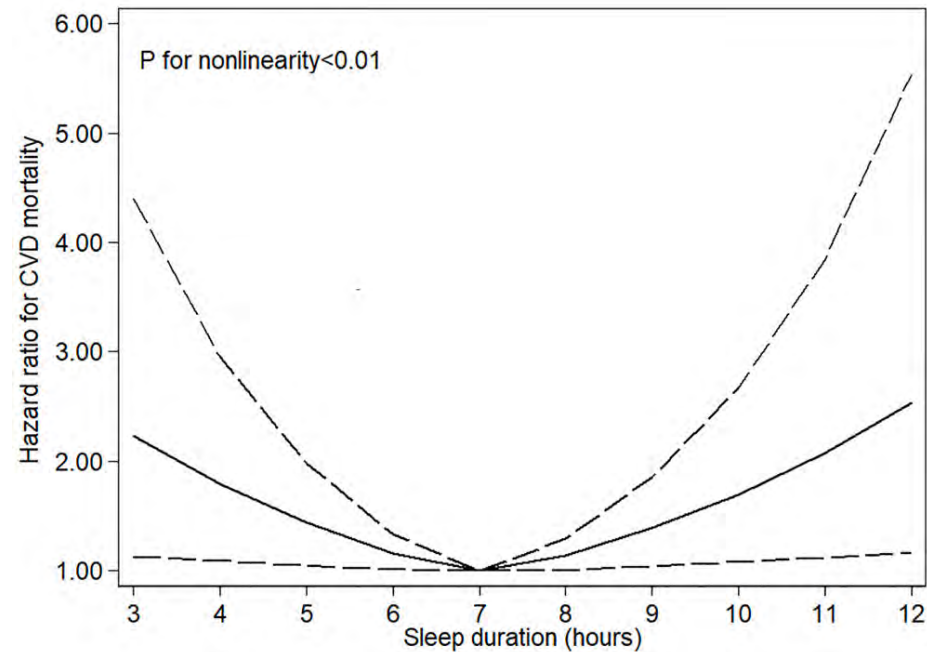
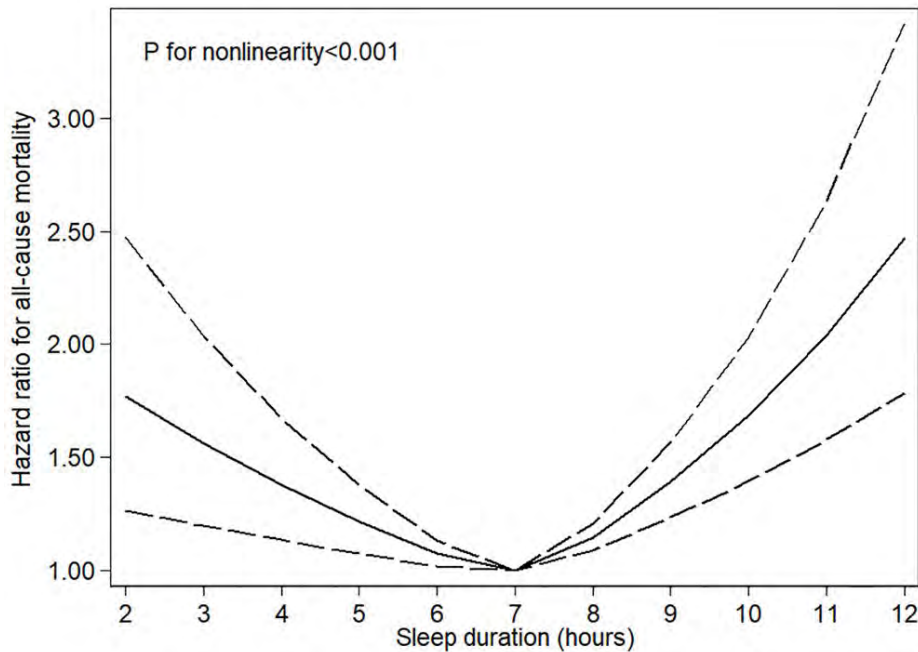


Increased odds of CVD in individuals with sleep problems: NHANES 2005-2008

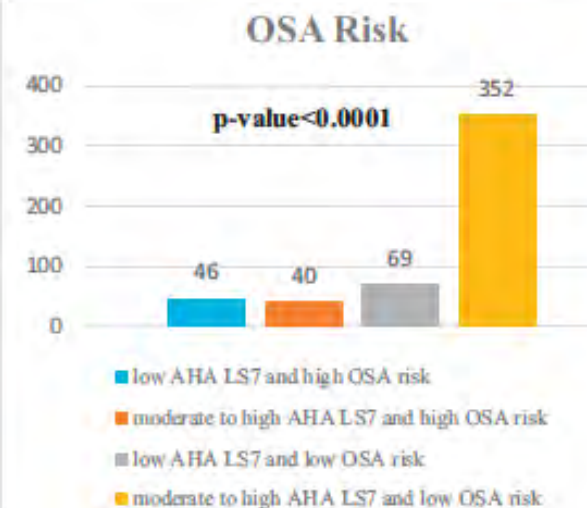
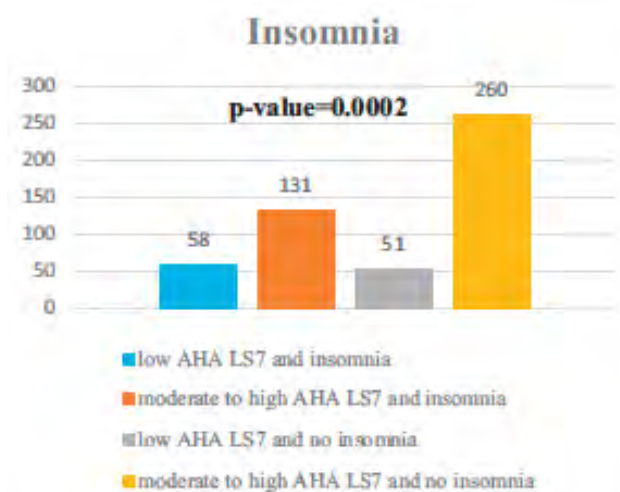
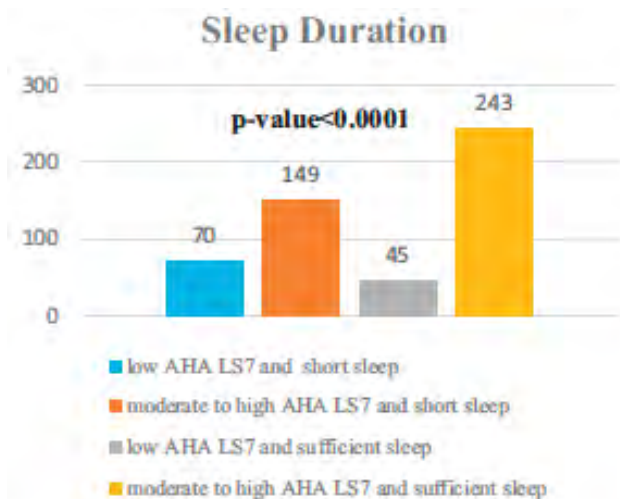
	Model 1 OR (95% CI) <i>P</i>	Model 2 OR (95% CI) <i>P</i>	Model 3 OR (95% CI) <i>P</i>
Sleep duration			
<7 vs. 7-8 h	1.57 (1.26, 1.97) *** <i>P</i> < 0.001	1.44 (1.16, 1.80) ** <i>P</i> = 0.008	1.42 (1.13, 1.78) * <i>P</i> = 0.025
>8 vs. 7-8 h	1.54 (1.03, 2.31) * <i>P</i> = 0.047	1.51 (0.97, 2.33) <i>P</i> = 0.092	1.43 (0.92, 2.22) <i>P</i> = 0.163
Sleep-onset latency time			
<5 vs. 5-30 min	0.79 (0.59, 1.06) <i>P</i> = 0.130	0.77 (0.58, 1.03) <i>P</i> = 0.108	0.77 (0.57, 1.02) <i>P</i> = 0.121
>30 vs. 5-30 min	1.77 (1.35, 2.32) *** <i>P</i> < 0.001	1.57 (1.17, 2.11) * <i>P</i> = 0.012	1.59 (1.17, 2.15) * <i>P</i> = 0.025
Sleep problems			
No	Reference	Reference	Reference
Yes	1.96 (1.62, 2.38) *** <i>P</i> < 0.001	1.74 (1.42, 2.13) *** <i>P</i> < 0.001	1.75 (1.41, 2.16) ** <i>P</i> = 0.001
OSA symptoms			
No	Reference	Reference	Reference
Yes	1.32 (1.08, 1.61) * <i>P</i> = 0.011	1.13 (0.91, 1.40) <i>P</i> = 0.303	1.12 (0.89, 1.40) <i>P</i> = 0.367
Daytime sleepiness			
No	Reference	Reference	Reference
Yes	1.75 (1.44, 2.13) *** <i>P</i> < 0.001	1.52 (1.25, 1.85) ** <i>P</i> = 0.001	1.54 (1.25, 1.89) ** <i>P</i> = 0.004



Increased risk of all-cause/CVD mortality in short & long sleepers, NHANES 2005-2014



Associations between Life's Simple 7 and sleep quality in women



Women who slept ≥ 7 h/night have higher LS7 scores and are more likely to meet >4 metrics

Poor sleep is associated with poor dietary intakes in women

Predictor	Outcome	B (SE)	p-value
Sleep quality (PSQI >5 vs. ≤5)	Food weight	79.6 (49.1)	0.106
	Added sugars	3.41 (1.57)	0.031
	% Unsaturated fats	-1.41 (0.50)	0.005
	Energy intake	108 (82)	0.184
Sleep onset latency (≤ 15 m vs. > 60 m)	Food weight	235.2 (79.6)	0.003
	Added sugars	2.97 (2.59)	0.252
	% Unsaturated fats	-0.95 (0.83)	0.253
	Energy intake	426 (132)	0.001
Insomnia (Yes vs. No)	Food weight	116.0 (48.8)	0.018
	Added sugars	1.87 (1.58)	0.235
	% Unsaturated fats	-1.25 (0.50)	0.013
	Energy intake	205 (81)	0.012

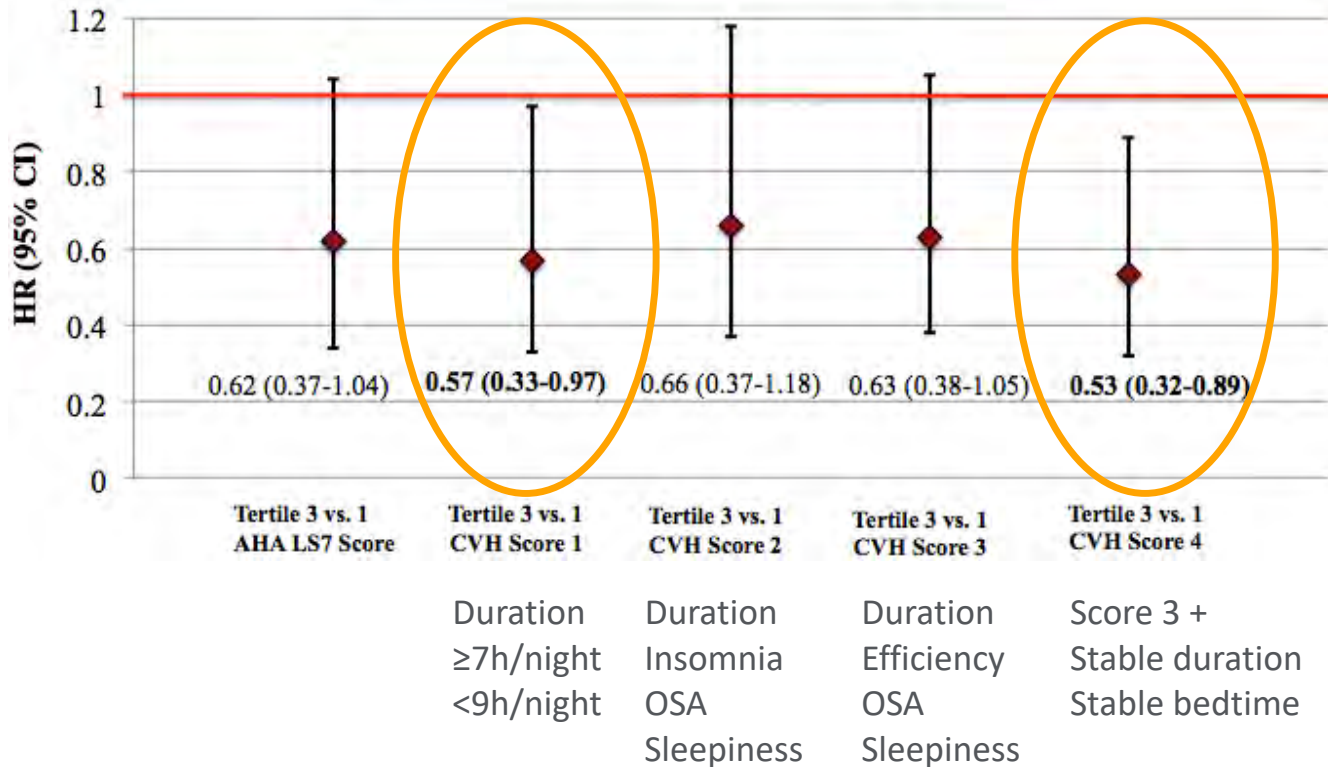
*PSQI: Pittsburg Sleep Quality Index; SOL: Sleep Onset Latency; ISI: Insomnia Severity Index

**Models are adjusted for age, BMI, race/ethnicity, education, and health insurance status

Developing Life's Essential 8



Association of the AHA LS7 Score and Alternate CVH Scores that Include Sleep Metrics with CVD Incidence in Cox Proportional Hazards Models

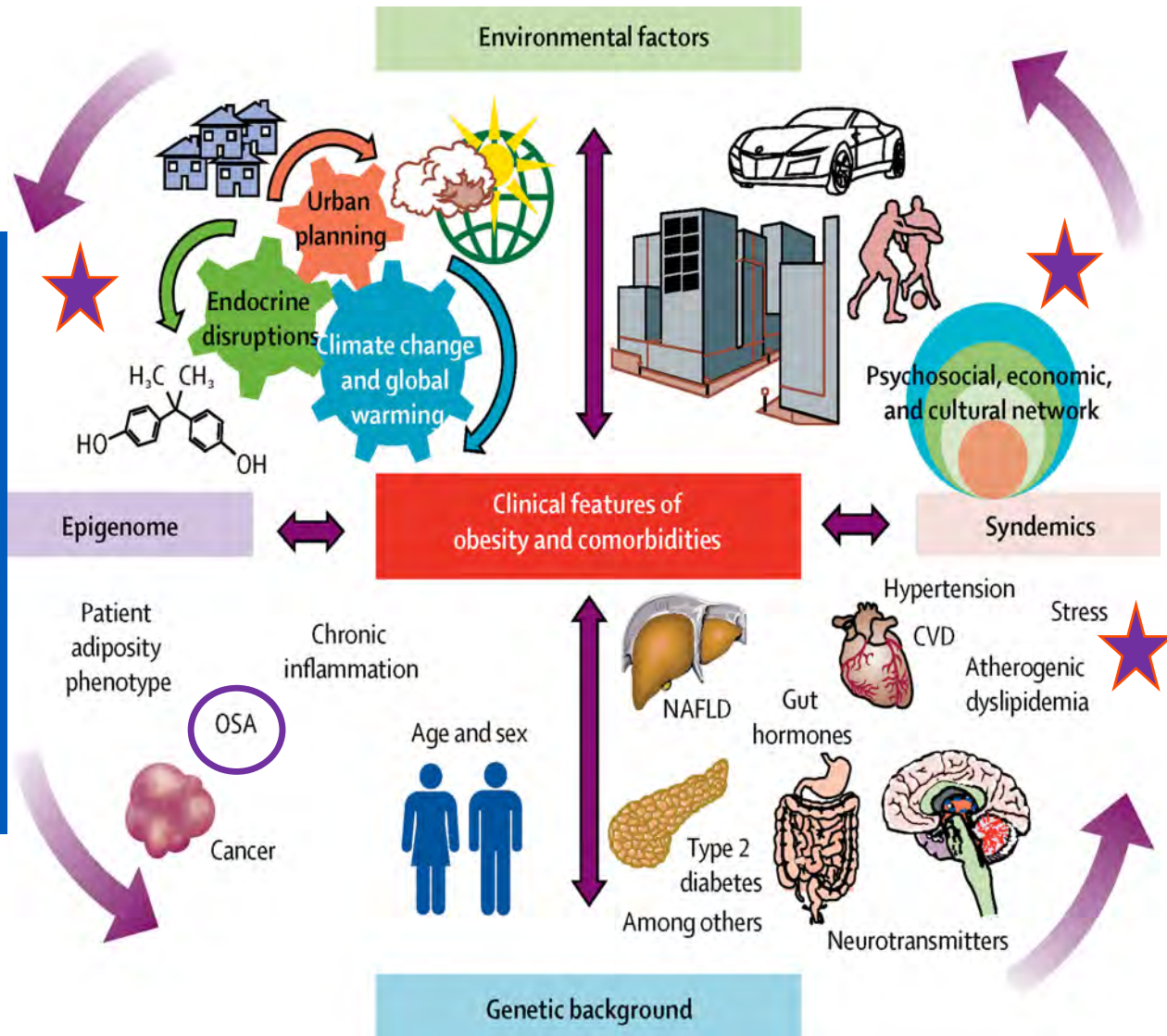


Findings from epidemiological studies

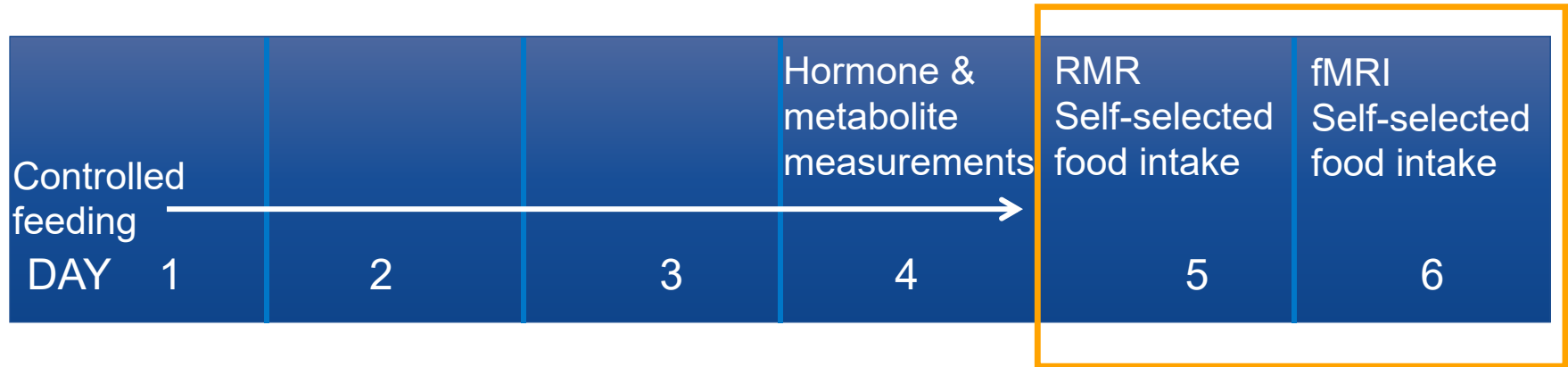
- Short sleepers have risk of obesity than adequate sleepers
- Short & disordered sleep is associated with higher CVD risk
 - Poor sleep is associated with lifestyle behaviors that predict greater CVD risk

But what about causality?

Factors influencing risk of obesity & its comorbidities

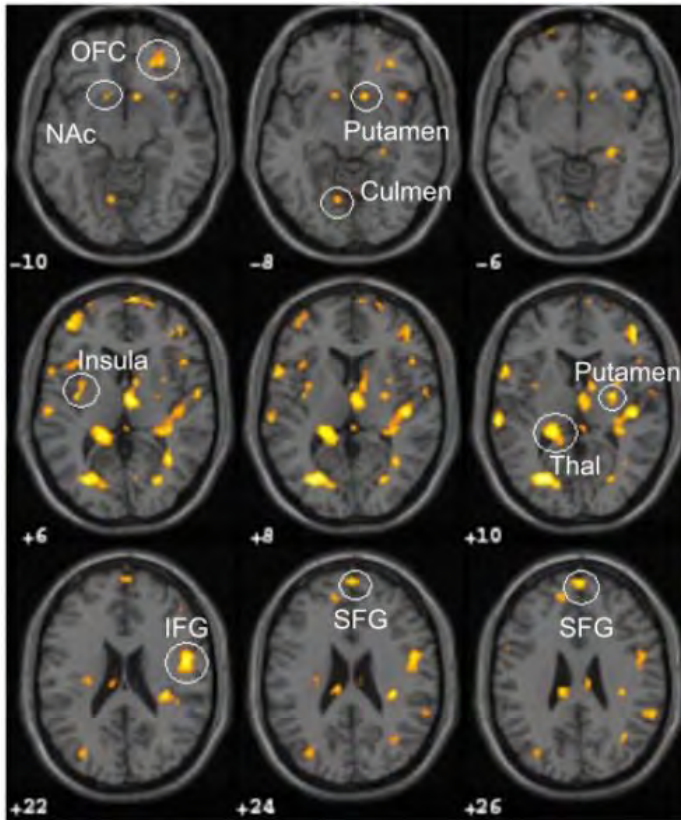


Effects of sleep restriction on energy balance & food intake regulation



Sleep restriction alters neuronal responses to foods

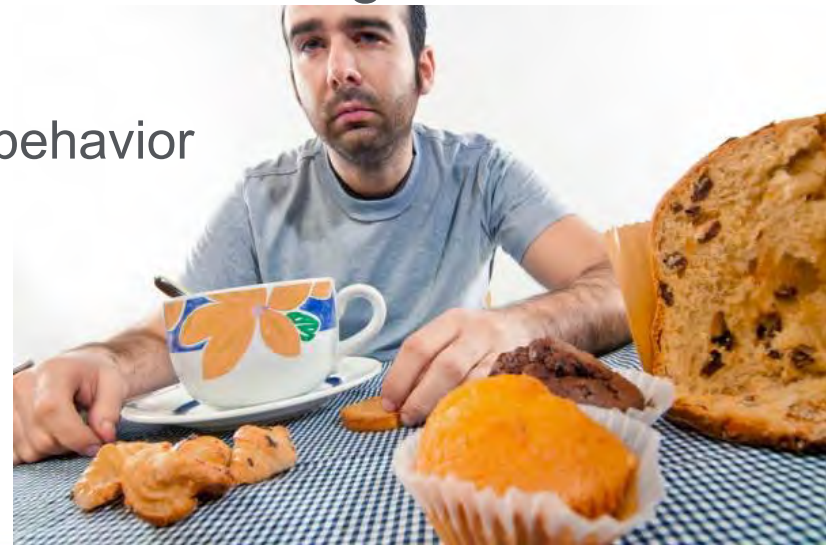
Food>Nonfood Restricted Sleep



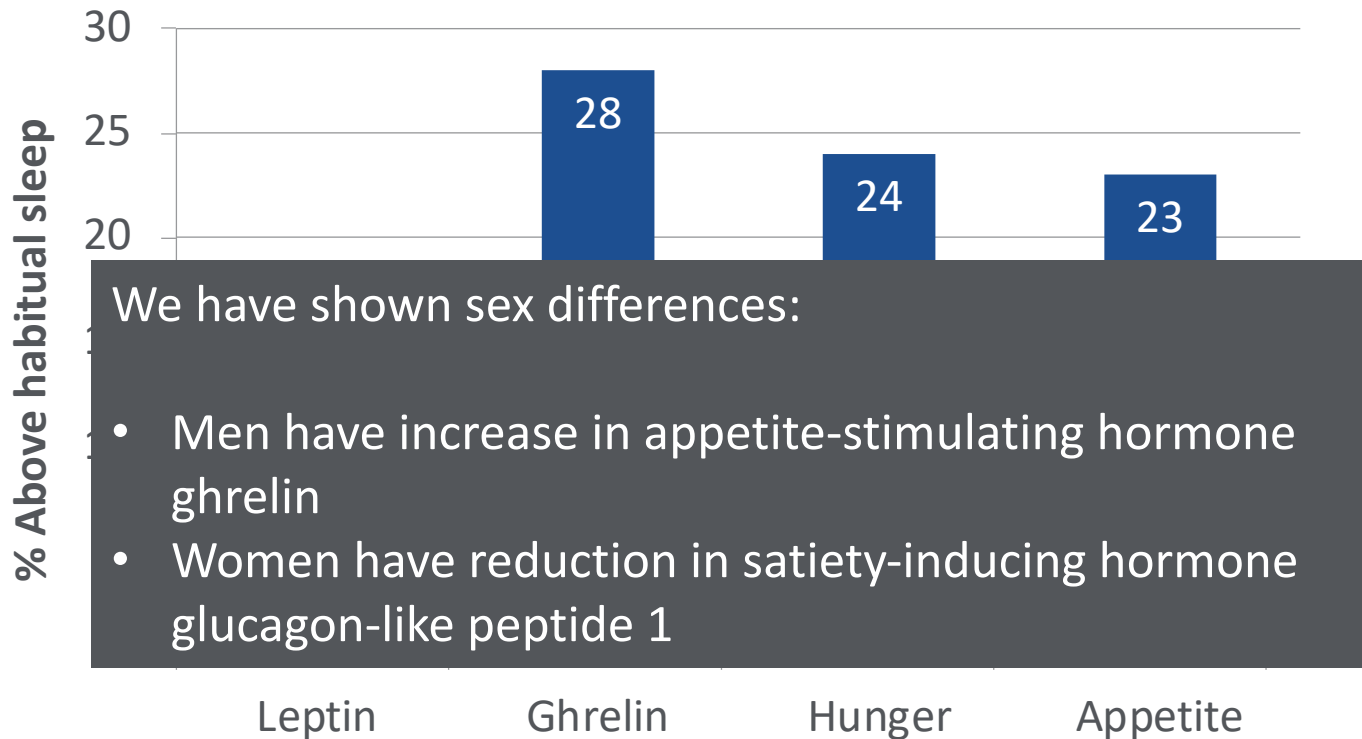
- Food stimuli increased regional brain activity in the OFC, insula, and regions of the basal ganglia and limbic system after restricted sleep
- Restricted sleep induces a state of greater responsiveness to food stimuli and heightened awareness of the rewarding properties of food

How does the brain respond to food stimuli in the sleep restricted state?

- Unhealthy foods activate the areas of the brain associated with reward and hedonic functions
 - Restricting sleep can increase salience of unhealthy food
 - Restricting sleep promotes hedonic hunger
- During habitual sleep, up-regulation of the cognitive control centers
 - Could signify improved food restraint behavior

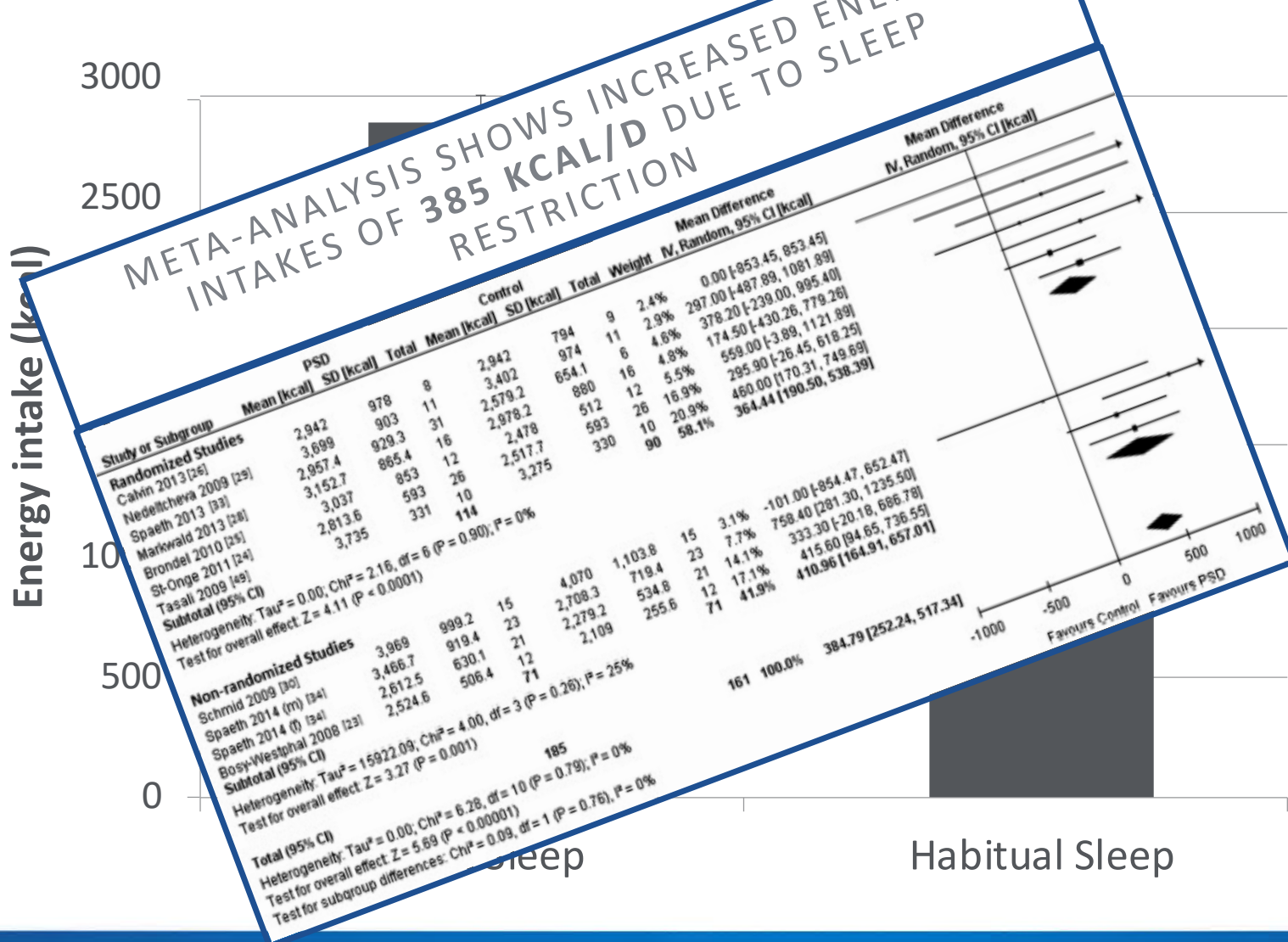


How does sleep influence homeostatic controls of food intake?

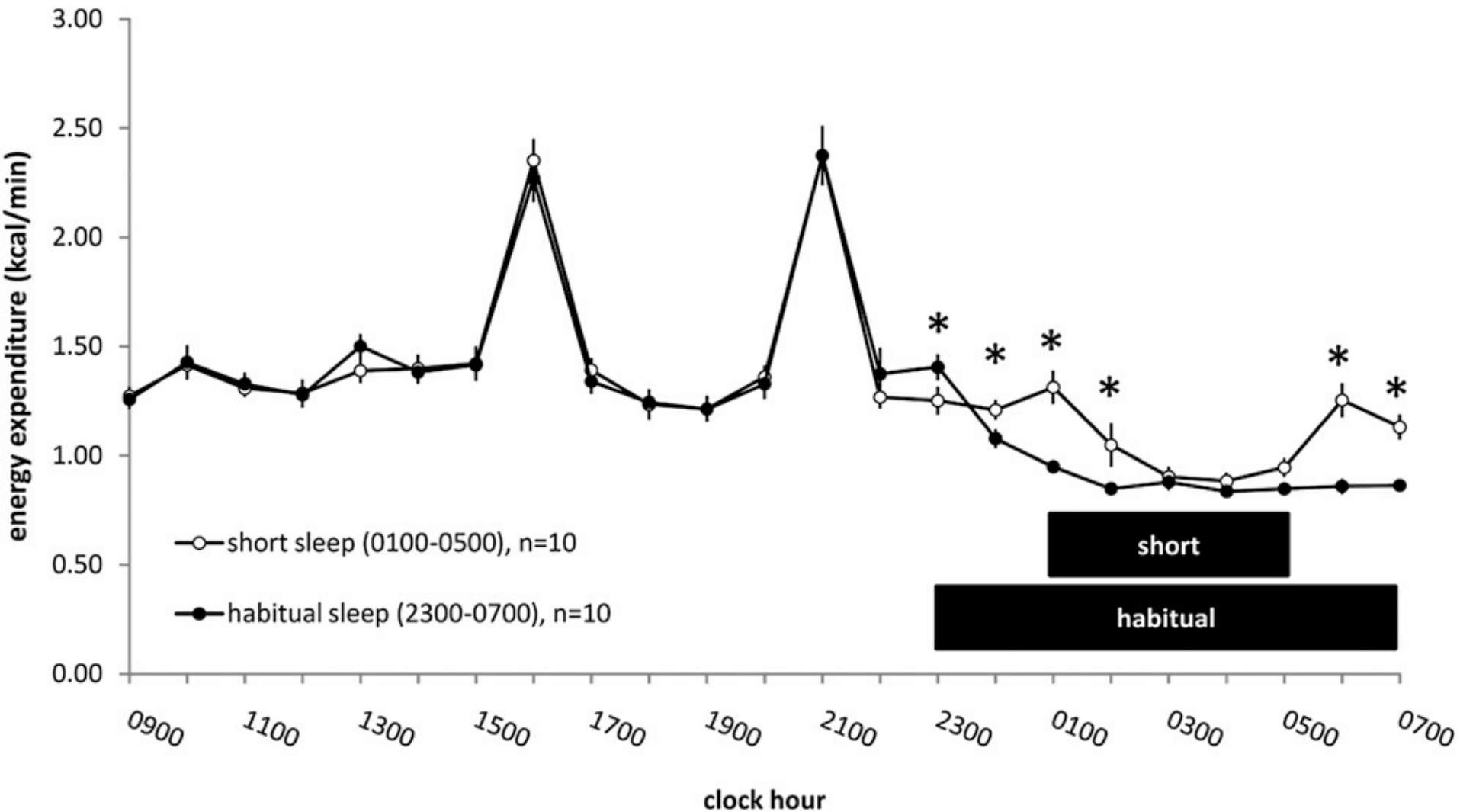


- Increase tended to be greatest for calorie-dense high carbohydrate foods
- Increase in appetite for fruits and vegetables of lesser magnitude

Sleep restriction increases food intake

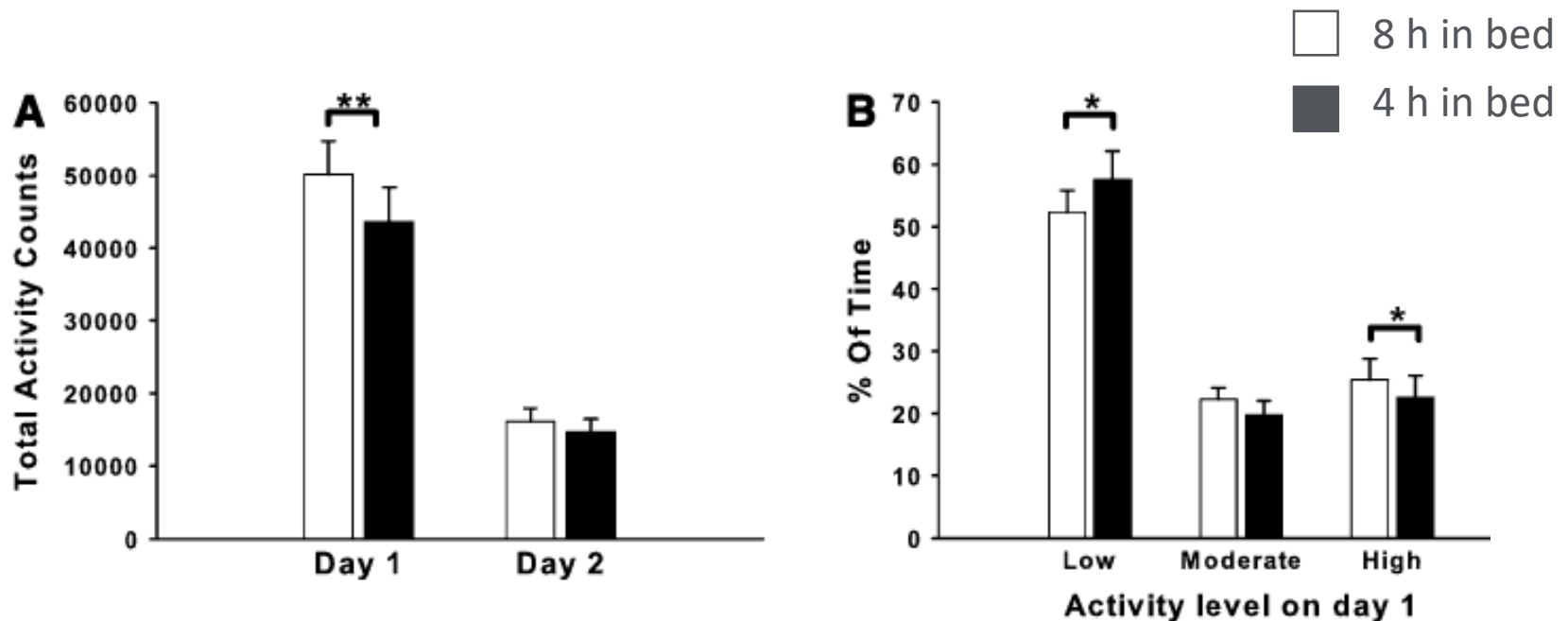


Impact of sleep restriction on 24-hour energy expenditure



Acute sleep restriction reduces physical activity

- Men spent either 4 or 8 h in bed for 2 nights
- Energy expenditure was measured by actigraphy during outpatient (day 1) and inpatient (day 2) days



Evidence of causality: Sleep restriction

2-week inpatient intervention:

- N=12 (9 M)
- Age 26.5 ± 5.8 y
- BMI 24.6 ± 3.7 kg/m²
- Habitual sleep 7.4 ± 1.0 h
 - SR = 4.3 ± 0.4 h/night
 - HS = 8.0 ± 0.5 h/night

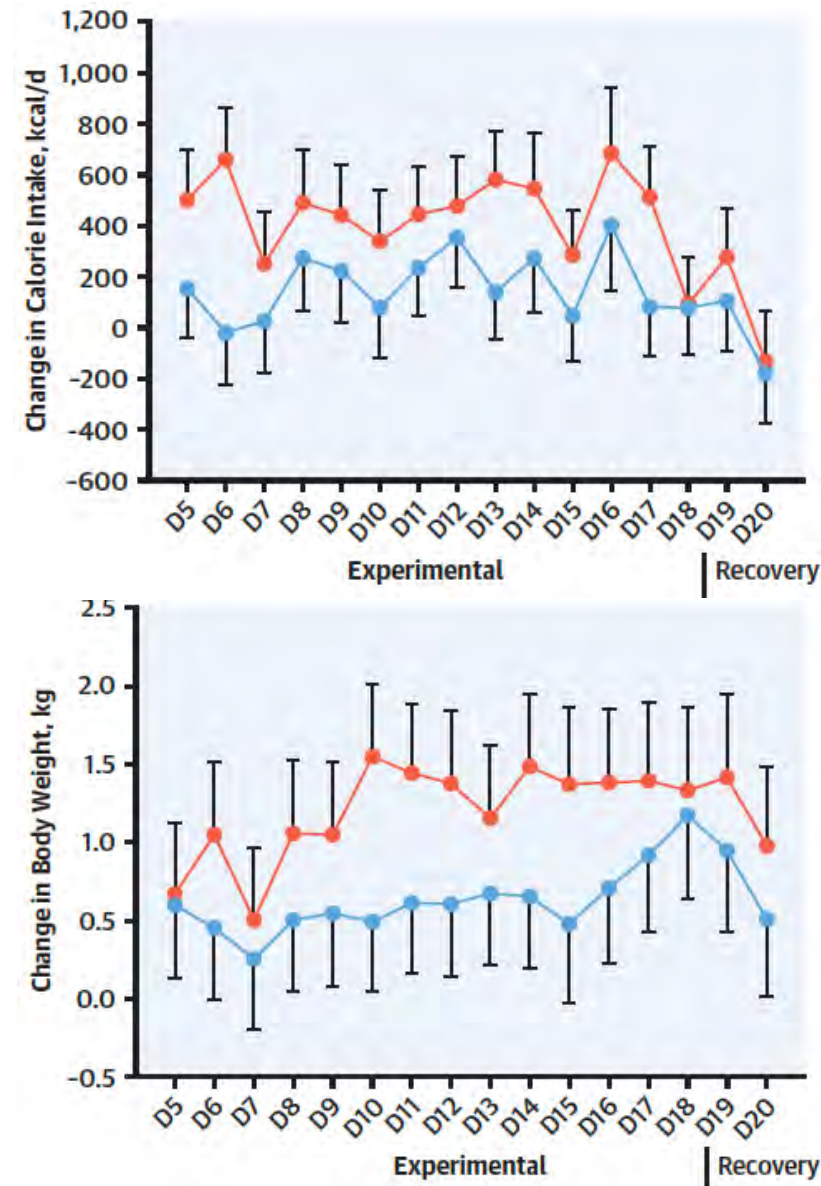
No difference in energy expenditure

Difference in energy intake between conditions:

- 257 kcal/d

Difference in change in body weight:

- 0.5 kg

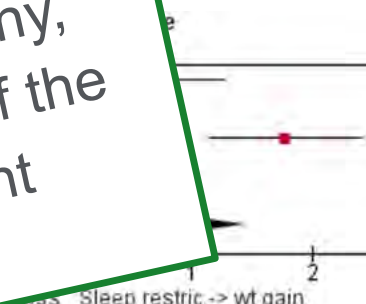


Covassin et al. J Am Coll Cardiol 2022;79:1254-65

Meta-analysis of randomized clinical trials of sleep restriction: Body weight

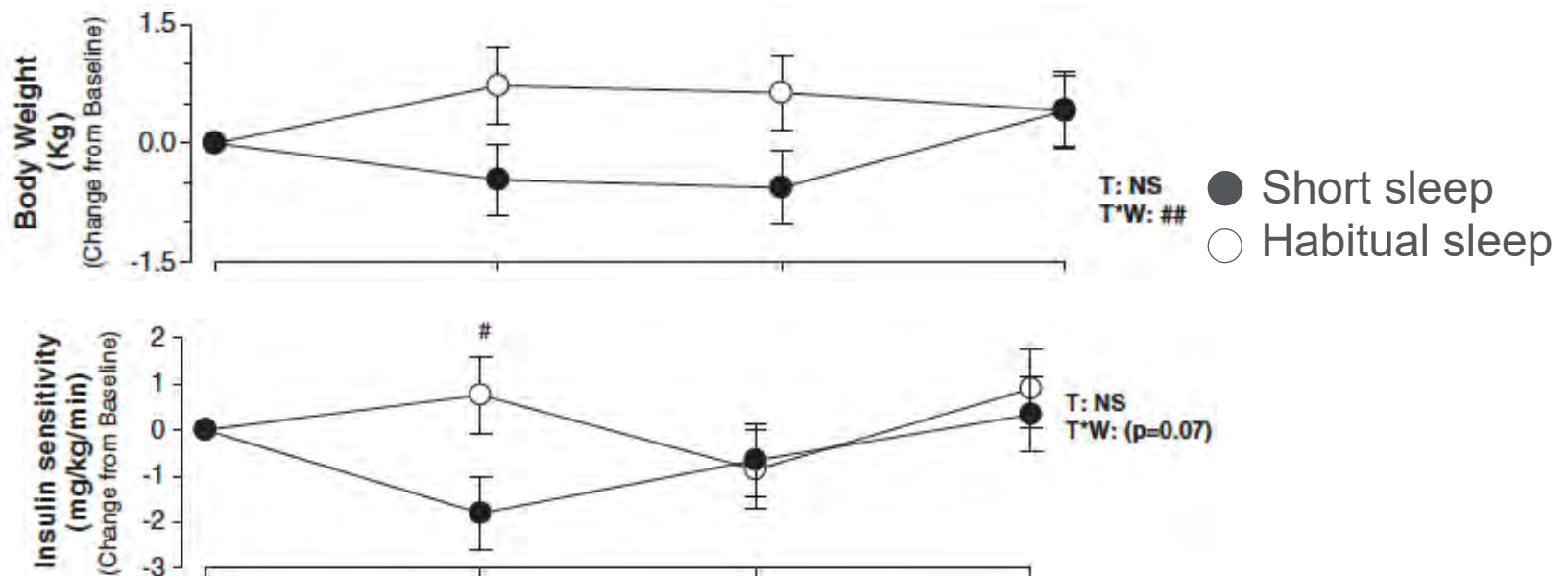
Study or Subgroup
Calvin 2012
Nedelt
O'Keefe
Spaeth
Total (95%
Heterogen
Test for ove

The authors concluded that “findings suggest a potential relationship with weight gain. However, it is unclear whether the degree of net effect, if any, may be observed over longer periods. None of the included studies that reported food intake went beyond 3 weeks in duration.”



Effect of longer, milder sleep restriction on body weight

- Young, healthy males, age 20-30 y, BMI 19-26 kg/m²
- Randomized to maintain regular sleep (7-7.5 h/night) or restrict their sleep by 1.5 h for 3 weeks
 - Actual restriction 1 h:13 min-1 h:30 min



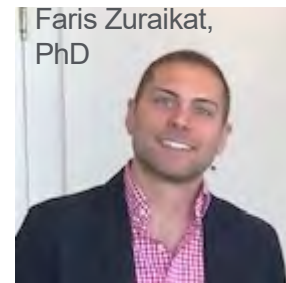
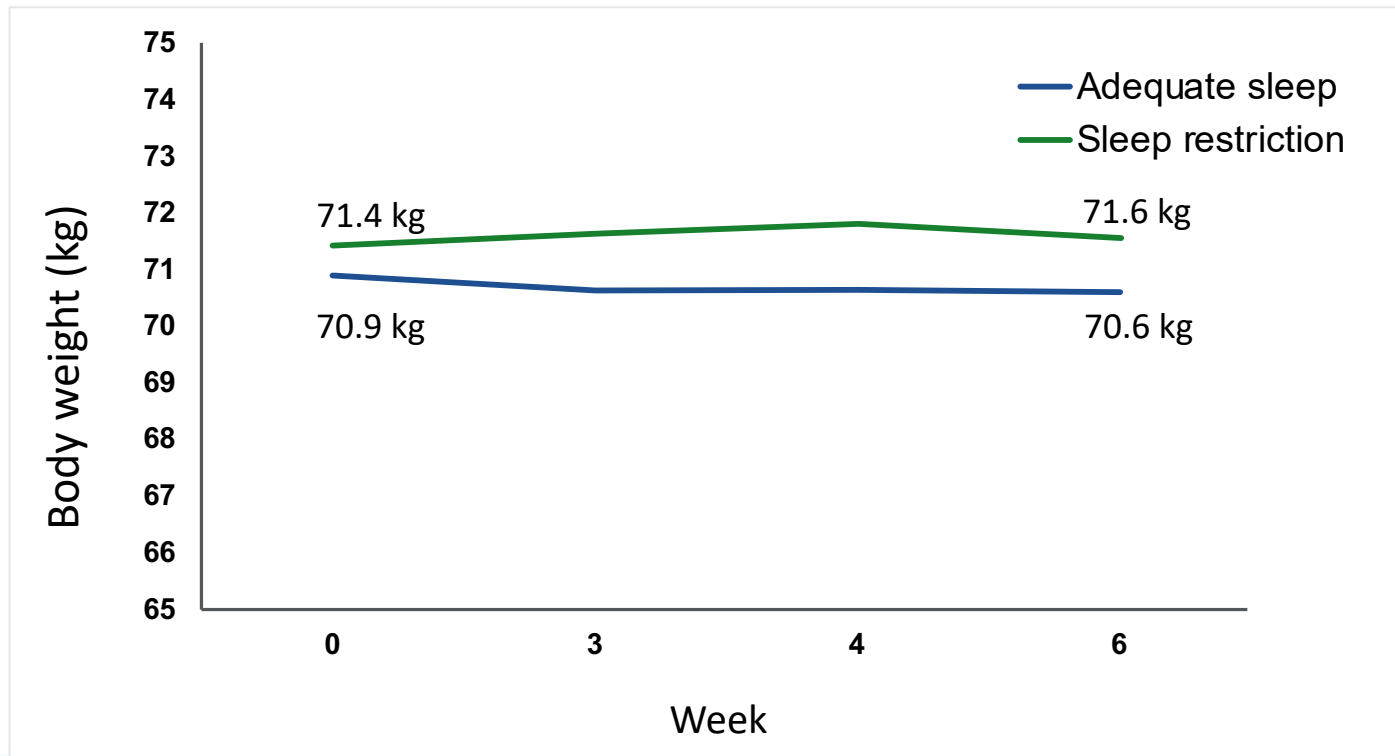
Next step:

What is the impact of 'life-like' sleep restriction conditions?

- To establish if there is a causal relation between sustained, mild sleep restriction (SR) and obesity risk using a randomized crossover clinical intervention
 - 2 phases of 6 weeks with either habitual (adequate) sleep or sleep reduced by 1.5 h (delayed bedtimes)
 - Participants have adequate sleep duration, >7 h/night, at screening
 - Determine effects of SR on body weight, body composition, and other lifestyle behaviors

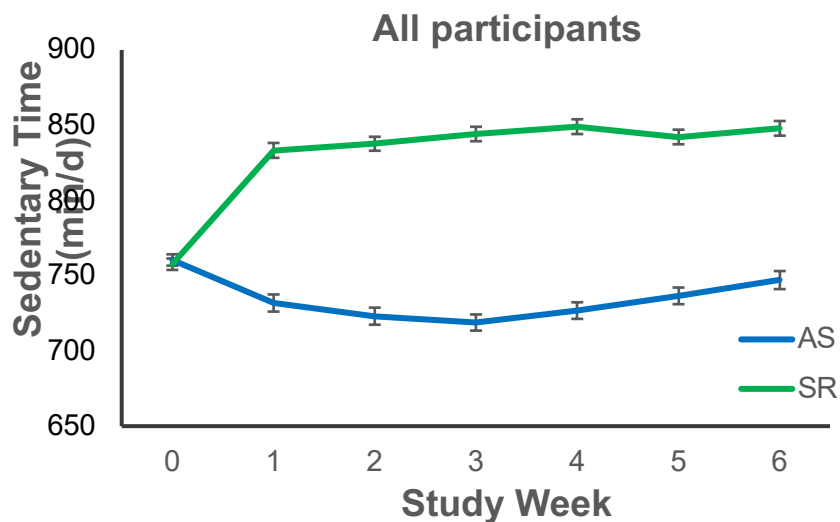
Mild Sleep Restriction Increases Body Weight

- Sleep restriction results more eating occasions and longer eating window
 - This is associated with higher energy intakes and worse diet quality

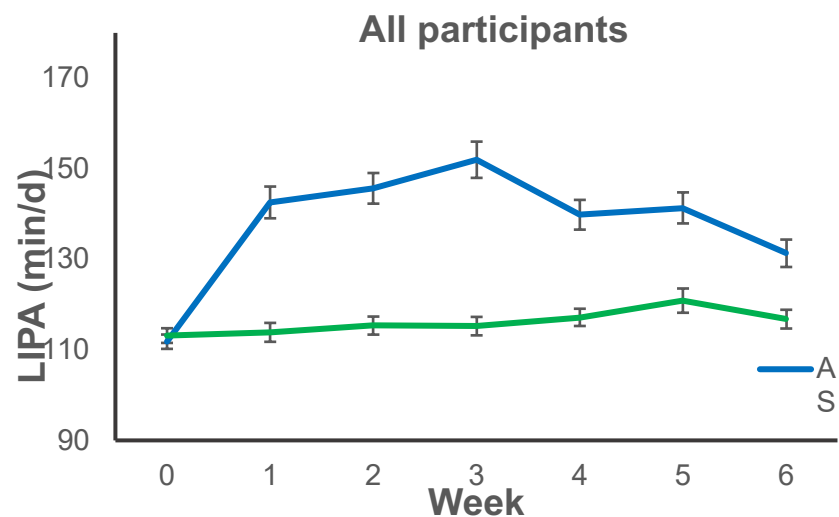


Zuraikat et al., in preparation.

Sleep restriction increases sedentary behavior & light physical activity in men and women

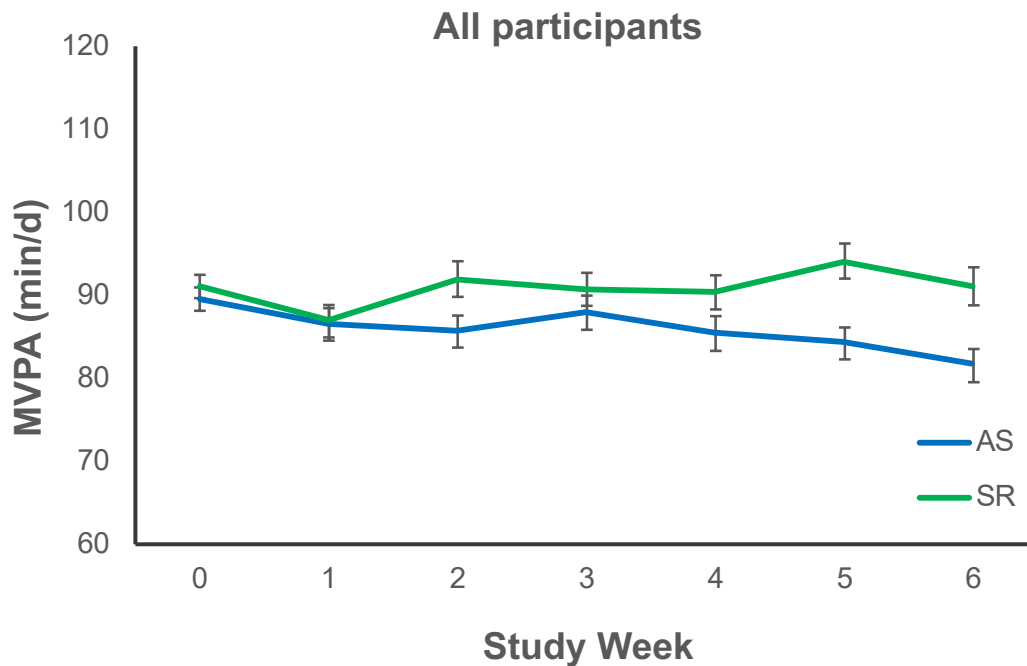


Increased by 12.5 ± 1.1 min/d over 6 wk in SR vs AS ($P < 0.0001$)



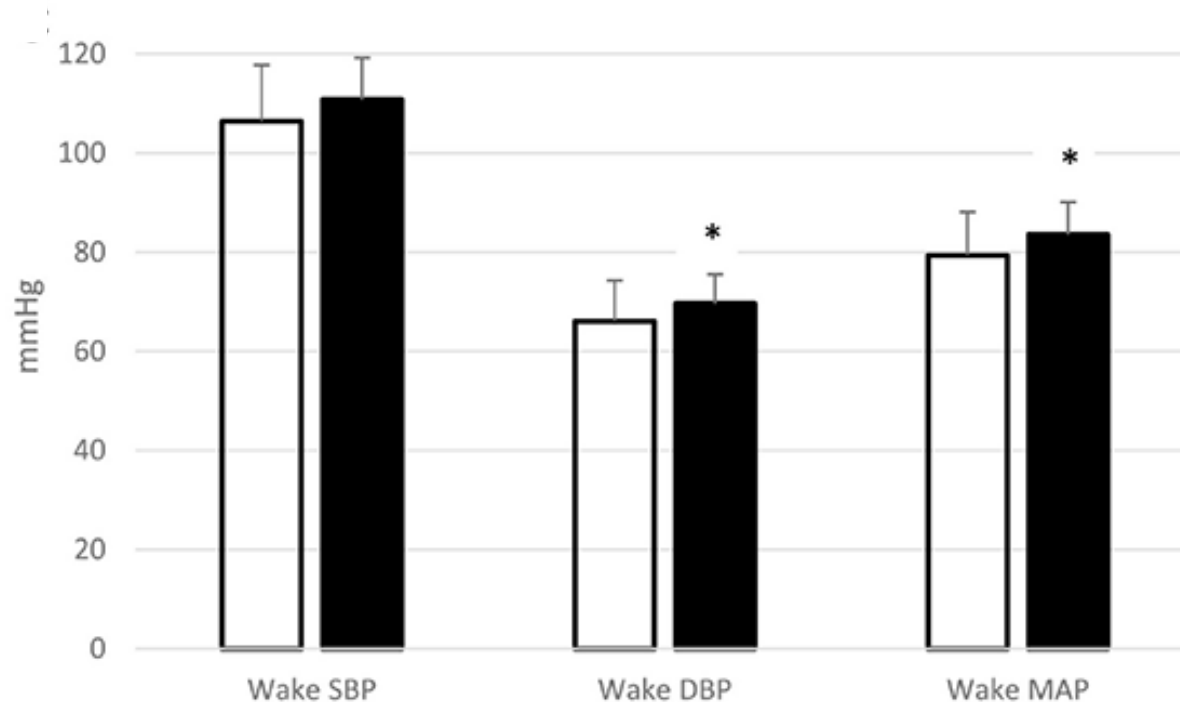
Increased by 1.1 ± 0.4 min/d over 6 wk in SR relative to AS ($P < 0.01$)

Impact of Sleep Restriction on Moderate-to-Vigorous Physical Activity in Men and Women



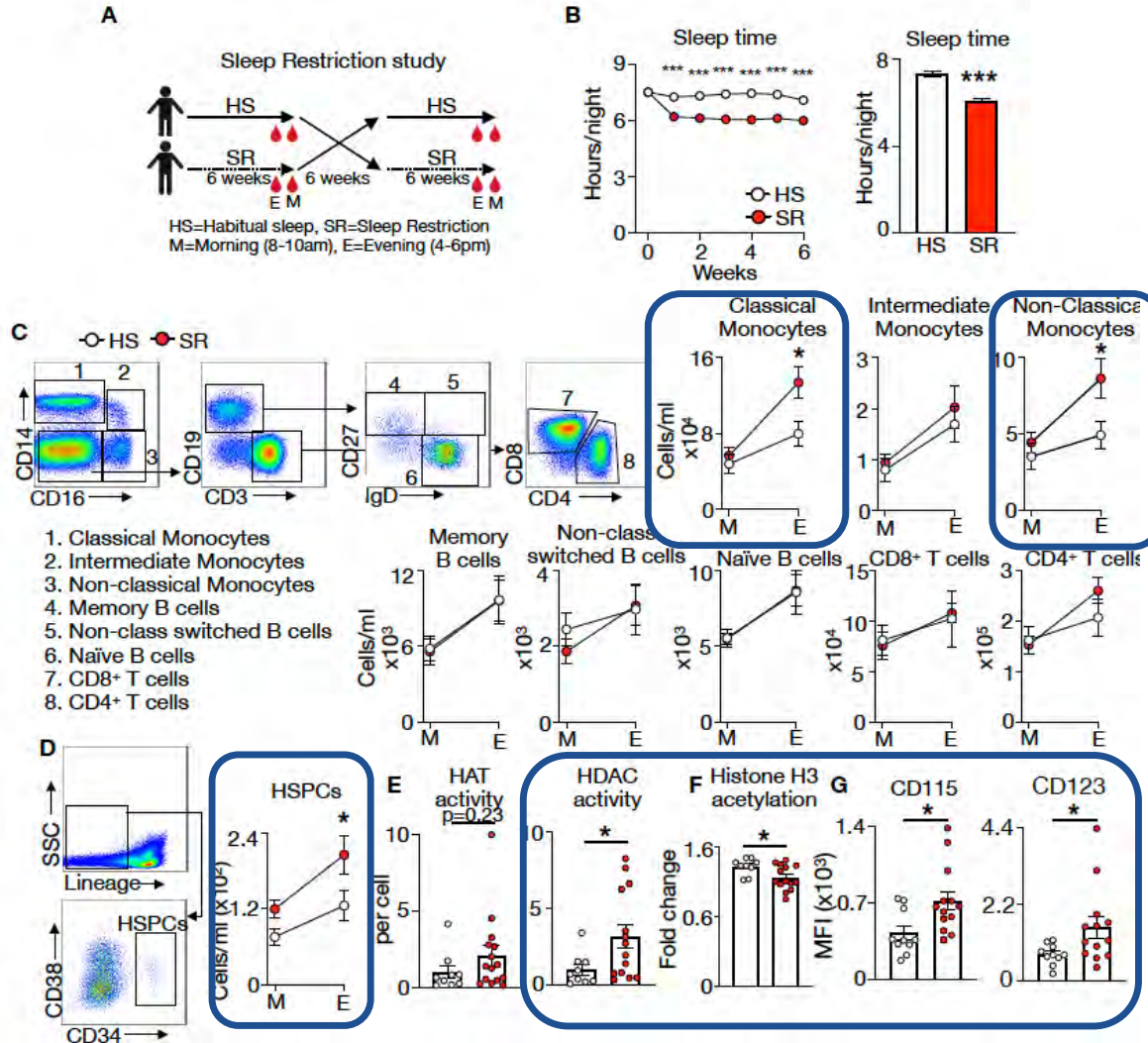
Increased by 1.1 ± 0.4 min/d over 6 wk in SR relative to AS ($P < 0.01$)

Higher blood pressure in women undergoing short sleep duration

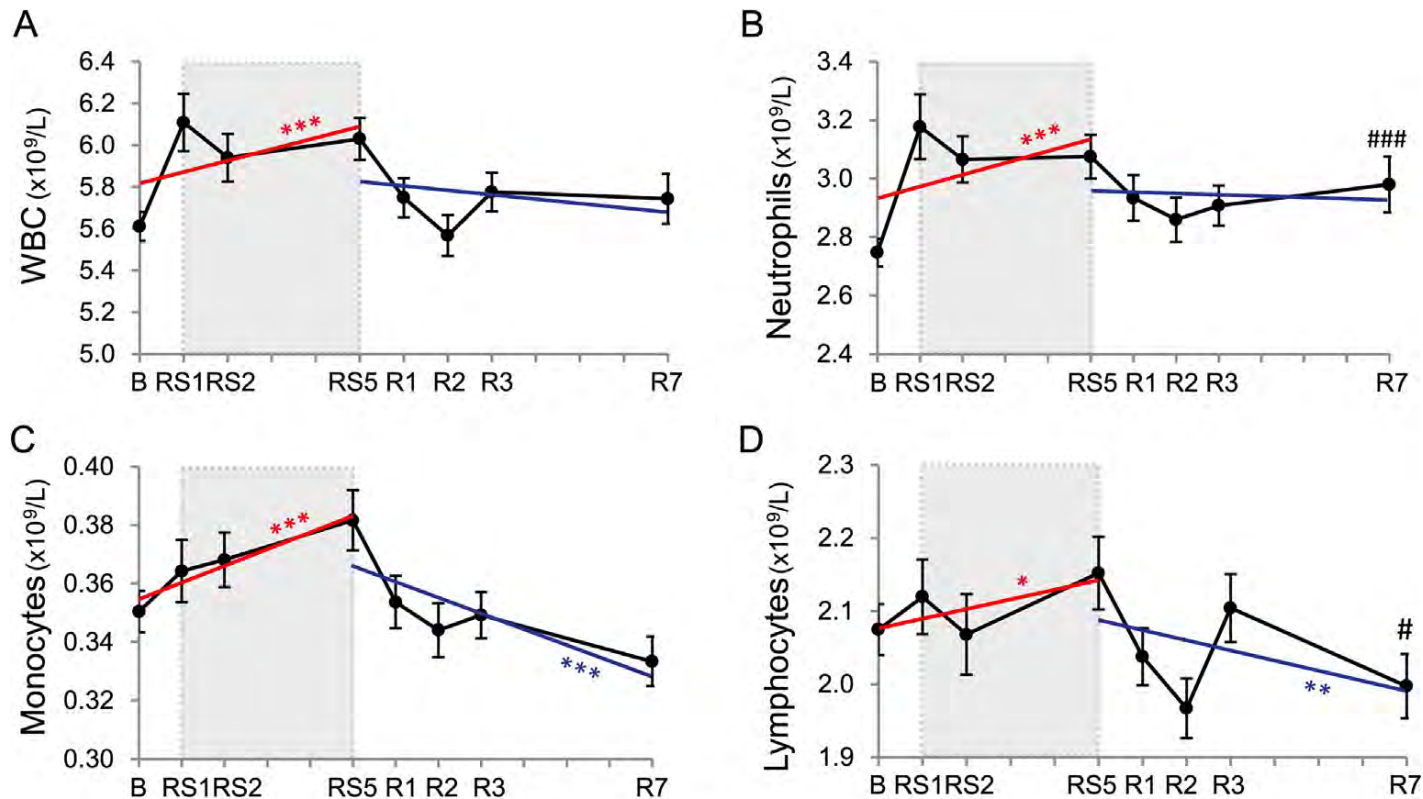


The effects of sleep restriction on 24-h systolic blood pressure were almost twice as high in post-menopausal compared to pre-menopausal women

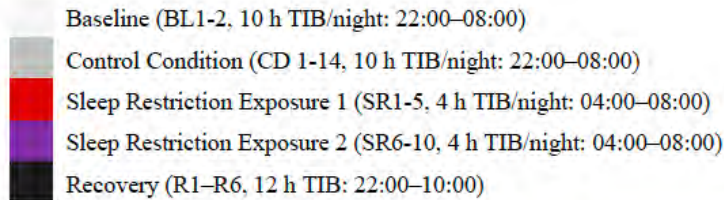
Sleep restriction increases hematopoiesis



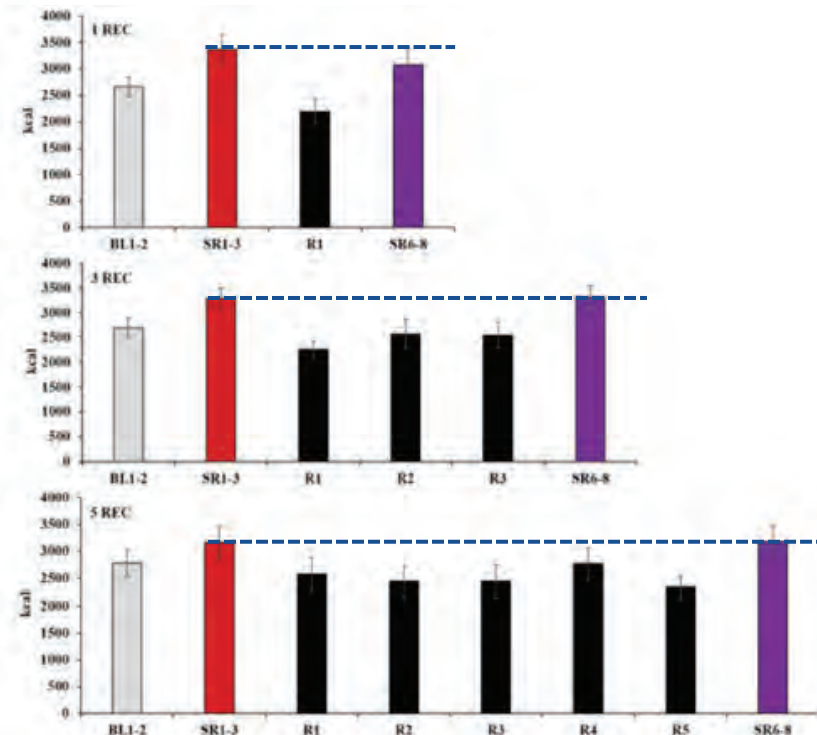
Impact of short & catch-up sleep on cardiometabolic risk factors



Impact of short & catch-up sleep on food intake



No evidence of acclimation from repeated exposures to sleep restriction, regardless of the number of recovery nights



Dietary intakes across different categories of sleep variability in MESA

Diet Outcome	Sleep Duration SD				P-value for trend
	≤ 60 min (n=673)	61-90 min (n=529)	91-120 min (n=392)	> 120 min (n=311)	
aMed Score	4.27 ± 1.82	4.08 ± 1.86	4.11 ± 1.81	4.05 ± 1.80	<0.01
Fruits	1.34 ± 0.98	1.31 ± 1.08	1.31 ± 1.08	1.28 ± 1.09	0.002
Vegetables	1.18 ± 0.85	1.17 ± 0.87	1.19 ± 0.89	1.23 ± 0.95	0.212
Whole grains	0.63 ± 0.50	0.60 ± 0.48	0.59 ± 0.49	0.56 ± 0.52	<0.001
Nuts/Seeds	0.33 ± 0.35	0.27 ± 0.34	0.26 ± 0.34	0.23 ± 0.30	<0.01
Legumes	0.15 ± 0.17	0.17 ± 0.23	0.17 ± 0.21	0.16 ± 0.23	0.802
Red meat	0.24 ± 0.19	0.22 ± 0.20	0.25 ± 0.21	0.24 ± 0.20	0.090
Fish	0.15 ± 0.15	0.16 ± 0.20	0.17 ± 0.16	0.20 ± 0.20	<0.01
Alcohol	3.41 ± 9.32	3.17 ± 7.84	2.69 ± 5.60	2.70 ± 6.18	0.873
MUFA	14.33 ± 4.01	14.14 ± 3.94	14.23 ± 4.05	14.45 ± 4.04	0.358
Saturated fat	10.41 ± 3.21	10.52 ± 3.06	10.16 ± 3.18	10.60 ± 3.22	0.043
Energy intake	1692 ± 770	1688 ± 775	1735 ± 869	1778 ± 914	0.01

Diet variables energy adjusted as: servings/1000 kcal (fruits, vegetables, whole grains, nuts/seeds, legumes, red meat, fish), %kcal (MUFA, SFA), g/1000 kcal (alcohol)

High sleep variability & short sleep duration associated with reduced weight loss

- Participants in PREDIMED-Plus, a Mediterranean diet intervention with vs without caloric restriction
 - Adults, 55-75 y, with BMI 27-40 kg/m²

	Tertiles of sleep variability (h)			Sleep variability (h)		
	1 (lowest)	2	3 (highest)	<i>p</i> Value 2 vs 1	<i>p</i> Value 3 vs 1	<i>p</i> Value
Weight, kg						
<i>n</i>	630	629	629			1888
12-month change	-2.3 (-2.6 to -2.0)	-2.1 (-2.5 to -1.8)	-1.7 (-2.0 to -1.4)			-2.1 (-2.2 to -1.9)
Difference vs first tertile ^a	0 (ref.)	0.1 (-0.3 to 0.5)	0.5 (0.1 to 0.9)	0.553	0.020	0.95 (0.06 to 1.8) 0.037
Difference vs first tertile ^b	0 (ref.)	0.1 (-0.3 to 0.6)	0.5 (0.1 to 0.9)	0.548	0.021	0.88 (-0.01 to 1.8) 0.052
Body mass index, kg/m²						
<i>n</i>	628	628	627			1888
12-month change	-0.8 (-0.9 to -0.7)	-0.8 (-0.9 to -0.6)	-0.6 (-0.7 to -0.5)			-0.7 (-0.8 to -0.7)
Difference vs first tertile ^a	0 (ref.)	0.06 (-0.1 to 0.2)	0.2 (0.04 to 0.4)	0.481	0.016	0.36 (0.03 to 0.7) 0.033
Difference vs first tertile ^b	0 (ref.)	0.05 (-0.1 to 0.2)	0.2 (0.04 to 0.4)	0.507	0.015	0.34 (0.01 to 0.7) 0.043
Waist circumference, cm						
<i>n</i>	600	599	599			1888
12-month change	-2.5 (-3.0 to -2.1)	-2.6 (-3.0 to -2.1)	-1.9 (-2.3 to -1.4)			-2.3 (-2.6 to -2.1)
Difference vs first tertile ^a	0 (ref.)	-0.2 (-0.8 to 0.4)	0.4 (-0.1 to 1.0)	0.496	0.148	0.7 (-0.5 to 1.9) 0.247
Difference vs first tertile ^b	0 (ref.)	-0.1 (-0.7 to 0.4)	0.4 (-0.2 to 1.0)	0.536	0.156	0.6 (-0.6 to 1.7) 0.345

Sleep stability & body composition

N=36 women

Age ≥ 20 y

BMI 20-33 kg/m²

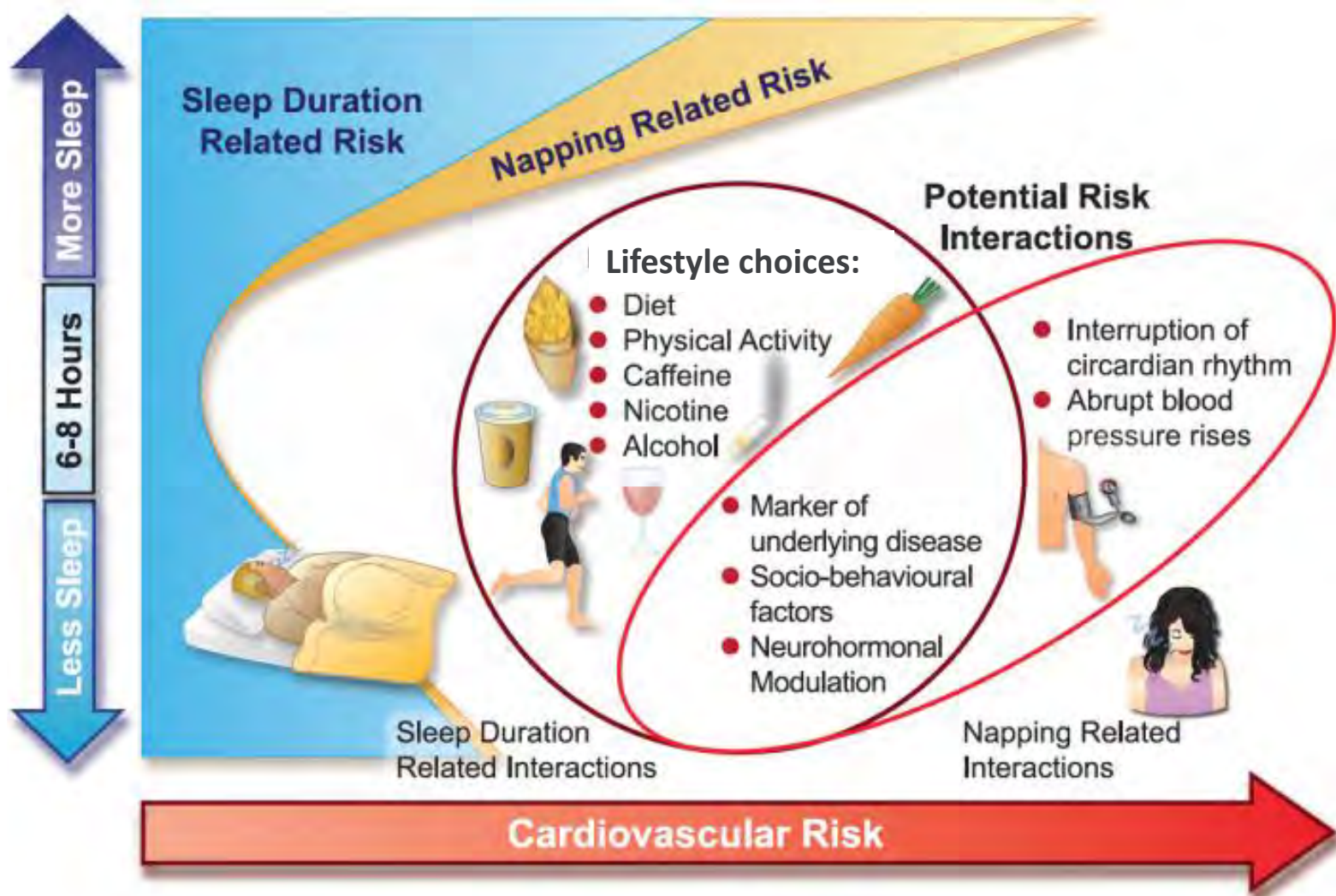
Habitual sleep ≥ 7 h/night

Undergoing 6-wk period of maintained adequate sleep with prescribed bed and wake times based on usual habits

Grouped by change in bedtime variability from screening

Variable	Increased/same bedtime variability (N = 8)	Reduced bedtime variability (N = 29)	P value*
Age, years	36.9 ± 15.0	34.4 ± 11.8	0.621
Race			0.663
White	5 (63)	14 (48)	
Other	3 (37)	15 (52)	
Baseline weight, kg	62.2 ± 5.0	66.5 ± 7.8	0.153
Baseline BMI, kg/m ²	23.5 ± 2.1	25.1 ± 3.0	0.190
Baseline sleep duration, min	453.3 ± 29.3	455.2 ± 30.2	0.875
Baseline bedtime	12:00 a.m.	10:48 p.m.	0.109
Baseline bedtime SD, min	49.7 ± 9.5	57.2 ± 27.4	0.218
Weight change, kg	0.48 ± 1.19	-0.66 ± 1.37	0.059
TAT change, L ^a	0.63 ± 0.41	-0.52 ± 0.98	<0.001
VAT change, L	0.05 ± 0.17	-0.03 ± 0.10	0.297
SAT change, L	0.56 ± 0.31	-0.48 ± 0.86	<0.001
WBV no lungs change, L	0.23 ± 0.91	-0.75 ± 0.90	0.016
IMAT change, L	0.03 ± 0.03	-0.01 ± 0.12	0.134
SM change, L	-0.08 ± 0.49	-0.19 ± 0.47	0.602
Leukocyte platelet aggregates, % ^b	8.42 ± 16.59	-8.42 ± 10.82	0.011

Sleep is Integral to Good Lifestyle Habits Compatible With Cardiovascular Health



Food intake during a controlled diet vs ad lib diet when sleep is sufficient

Nutrient content	Controlled diet	Ad lib diet
Energy, kcal	2055	2518
Protein, %En	17	14
Carbohydrates, %En	53	54.6
Fat, %En	31	32.7
Saturated fat, %En	7.5	10

Sleep after a controlled diet vs ad lib diet

Sleep parameter	Controlled diet	Ad lib diet	P-value
Total sleep time, min	453.5 ± 44.4	455.1 ± 30.2	0.86
Stage 1, min	52.3 ± 21.8	56.2 ± 18.8	0.18
Stage 2, min	240.3 ± 42.9	245.8 ± 35.5	0.45
Slow wave sleep, min	29.3 ± 13.9	24.6 ± 12.8	0.043
Rapid-eye movement sleep, min	91.6 ± 17.8	96.4 ± 18.2	0.19
Sleep onset latency, min	16.9 ± 11.1	29.2 ± 23.1	0.0085
Arousals	143.2 ± 52.1	143.4 ± 51.9	0.98

Relation between diet and sleep after a day of ad lib intakes

Sleep parameter	Fiber, g	Sugar, %En	Non-sugar/non-fiber CHO, %En	Saturated fat, %En
Stage 1, % sleep time	-0.19 ± 0.07	0.08 ± 0.17	0.04 ± 0.03	0.03 ± 0.21
Slow wave sleep, % sleep time	0.26 ± 0.11	-0.18 ± 0.25	-0.04 ± 0.04	-0.71 ± 0.32
Arousals	-0.11 ± 0.81	4.34 ± 1.86	0.66 ± 0.31	2.17 ± 2.40

Data suggest that a high-fiber diet, with low intake of sugars, is associated with better sleep depth and architecture

Could diet improve sleep in those with sleep disorders??

Adherence to Mediterranean Diet associated with sleep quality at 1 y

Predictor	Outcome	β (SE) ^b	<i>p</i> -Value	β (SE) ^c	<i>p</i> -Value
aMed diet score	PSQI total score	-0.30 (0.10)	<0.01	-0.31 (0.08)	<0.0001
	Sleep onset latency	-0.61 (0.65)	0.35	-0.71 (0.59)	0.23
	Sleep efficiency	1.20 (0.35)	<0.001	1.21 (0.33)	<0.001
	Sleep disturbances	-0.30 (0.12)	0.01	-0.35 (0.10)	<0.001
Fruits and vegetables	PSQI total score	-0.16 (0.07)	0.02	-0.19 (0.05)	<0.001
	Sleep onset latency	-0.41 (0.44)	0.36	-0.31 (0.40)	0.44
	Sleep efficiency	0.56 (0.24)	0.02	0.52 (0.22)	0.02
	Sleep disturbances	-0.18 (0.08)	0.03	-0.15 (0.07)	0.02
Legumes	PSQI total score	-0.10 (0.16)	0.55	-0.24 (0.13)	0.06
	Sleep onset latency	-1.13 (1.03)	0.27	-1.21 (0.94)	0.20
	Sleep efficiency	1.36 (0.55)	0.01	1.46 (0.52)	<0.01
	Sleep disturbances	0.17 (0.19)	0.39	-0.08 (0.16)	0.62
Nuts	PSQI total score	0.01 (0.21)	0.96	0.02 (0.17)	0.92
	Sleep onset latency	0.09 (1.35)	0.95	0.25 (1.23)	0.84
	Sleep efficiency	-0.47 (0.72)	0.51	-0.36 (0.68)	0.60
	Sleep disturbances	-0.26 (0.25)	0.31	-0.09 (0.20)	0.65
Dark breads	PSQI total score	-0.68 (0.39)	0.08	-0.55 (0.30)	0.07
	Sleep onset latency	-0.94 (2.48)	0.71	-1.09 (2.26)	0.63
	Sleep efficiency	2.07 (1.33)	0.12	1.96 (1.26)	0.12
	Sleep disturbances	-0.43 (0.47)	0.36	-0.67 (0.38)	0.08

Odds ratio for associations between Mediterranean diet score and sleep: MESA Exam 5

Alternate Mediterranean Diet Score	Sleep duration 6-7 h/night vs <6 h/night	Sleep duration 7-8 h/night vs <6 h/night	Sleep duration >8 h/night vs <6 h/night	Insomnia Symptoms vs None
Moderate-High Score				
Model 1	1.30 (1.03-1.63)	1.05 (0.82-1.34)	0.83 (0.60-1.14)	0.81 (0.67-0.97)
Model 2	1.32 (1.05-1.66)	1.05 (0.82-1.34)	0.84 (0.61-1.16)	0.81 (0.68-0.98)
Model 3	1.38 (1.07-1.78)	1.05 (0.80-1.38)	0.97 (0.68-1.40)	0.82 (0.67-1.00)
Model 4	1.43 (1.08-1.88)	1.05 (0.78-1.40)	0.95 (0.64-1.42)	0.85 (0.68-1.06)

Model 1 is adjusted for age, sex, race/ethnicity

Model 2 is additionally adjusted for education

Model 3 is additionally adjusted for cigarette smoking, intentional exercise, and total energy intake

Model 4 is additionally adjusted for BMI, hypertension, diabetes, depressive symptoms, AHI, anti-depressant and anti-psychotic medications, insomnia symptoms or sleep duration

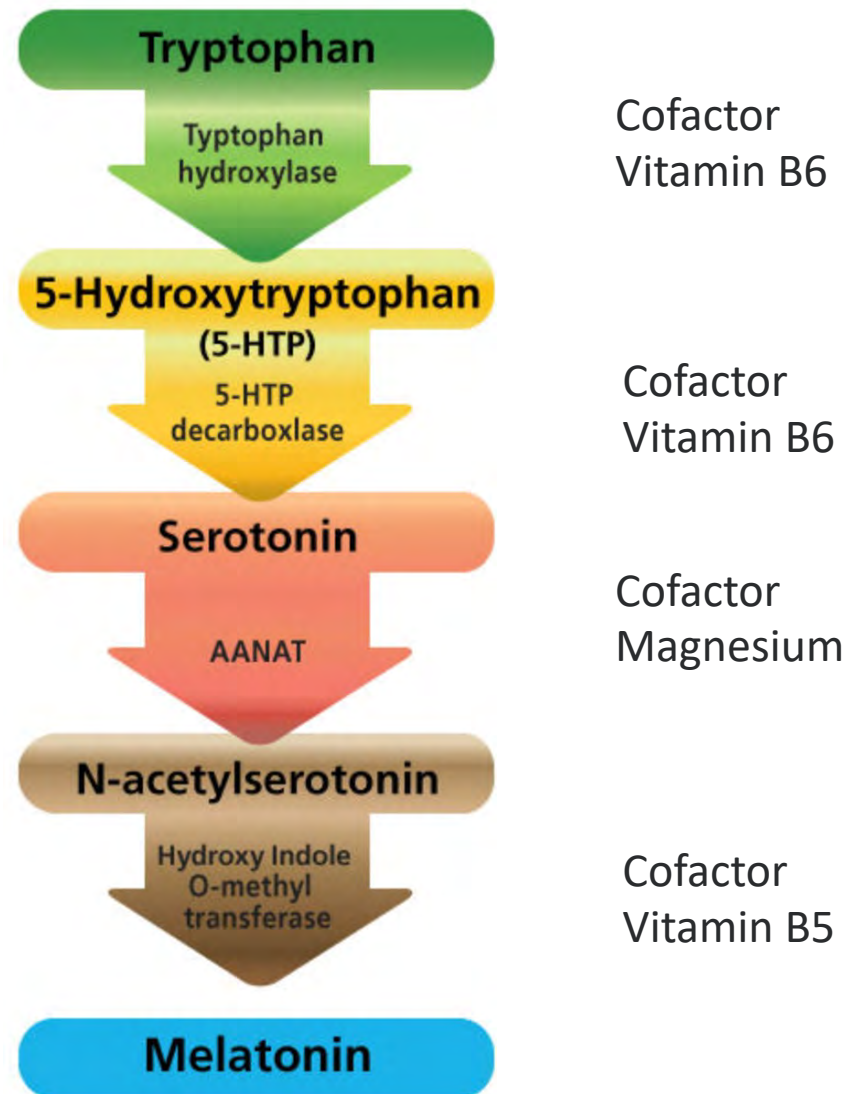
Odds ratio for associations between change in Mediterranean diet score and sleep:

MESA Exams 1 & 5

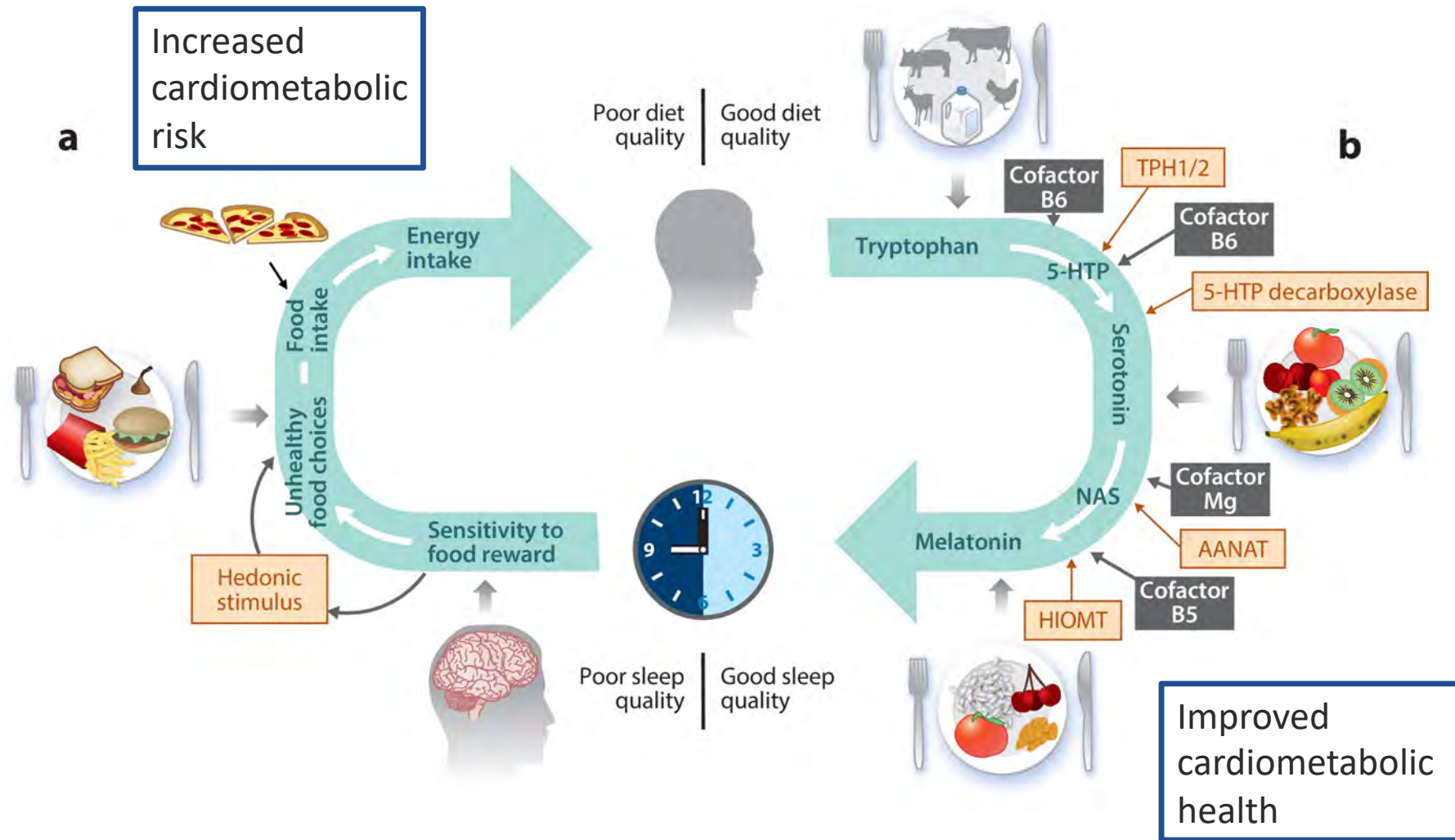
Alternate Mediterranean Diet Score	Sleep duration 6-7 h/night vs <6 h/night	Sleep duration 7-8 h/night vs <6 h/night	Sleep duration >8 h/night vs <6 h/night	Insomnia Symptoms vs None
No change vs decrease				
Model 1	1.05 (0.77-1.45)	1.11 (0.79-1.54)	1.08 (0.71-1.65)	0.64 (0.49-0.83)
Model 2	1.07 (0.78-1.47)	1.10 (0.79-1.54)	1.09 (0.71-1.68)	0.64 (0.49-0.83)
Model 3	1.04 (0.74-1.45)	1.13 (0.80-1.60)	1.13 (0.72-1.80)	0.65 (0.50-0.85)
Model 4	0.98 (0.68-1.40)	1.06 (0.73-1.55)	0.96 (0.57-1.61)	0.61 (0.45-0.82)
Increase vs decrease				
Model 1	1.35 (1.04-1.75)	1.30 (0.98-1.71)	1.04 (0.72-1.50)	0.90 (0.73-1.11)
Model 2	1.36 (1.05-1.76)	1.29 (0.98-1.71)	1.04 (0.72-1.50)	0.90 (0.73-1.11)
Model 3	1.30 (0.99-1.71)	1.26 (0.94-1.69)	1.18 (0.80-1.75)	0.90 (0.72-1.12)
Model 4	1.34 (0.99-1.80)	1.30 (0.95-1.79)	1.15 (0.74-1.77)	0.92 (0.72-1.17)

Biological plausibility for diet impact on sleep

- Tryptophan:
 - Essential amino acid
 - Primary substrate for melatonin synthesis
- Carbohydrates and gut microbiome involved in Trp metabolism
- Various dietary nutrients involved in enzymatic conversions of Trp to melatonin



Cycles of lifestyle behaviors & health



Thank you!

RAs & Fellows:

- Faris Zuraikat, PhD
- Rocio Barragan-Arnal, PhD
- Amy Roberts, Ph.D
- Ayanna Campbell, MS
- Ismel Salazar, MS
- Justin Cochran, MS
- Samantha Scaccia, MS
- Many IHN MS Students

Collaborators:

- Brooke Aggarwal, EdD
- Sanja Jelic, MD
- Blandine Laferrère, MD
- Ari Shechter, Ph.D

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WEBINAR HOST:

Acacia Wright, RD, CD
Sr. Manager of Nutrition
Communications
Orgain, LLC

Acacia.Wright@orgain.com



WEBINAR PRESENTER:

Marie-Pierre St-Onge, Ph.D, CCSH, FAHA
Associate Professor of Nutritional
Medicine, Director of Columbia
University Irving Medical Center Sleep
Center of Excellence

ms2554@cumc.columbia.edu



GENERAL INQUIRIES:

medinfo@orgain.com